

# Vaginal misoprostol for cervical ripening and induction of labour (Review)

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[Intervention Review]

# Vaginal misoprostol for cervical ripening and induction of labour

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## ABSTRACT

### Background

Misoprostol (Cytotec, Searle) is a prostaglandin E1 analogue widely used for off-label indications such as induction of abortion and of labour. This is one of a series of reviews of methods of cervical ripening and labour induction using standardised methodology.

### Objectives

To determine the effects of vaginal misoprostol for third trimester cervical ripening or induction of labour.

### Search strategy

The Cochrane Pregnancy and Childbirth Group's Trials Register (November 2008) and bibliographies of relevant papers. We updated this search on 30 April 2010 and added the results to the awaiting classification section.

### Selection criteria

Clinical trials comparing vaginal misoprostol used for third trimester cervical ripening or labour induction with placebo/no treatment or other methods listed above it on a predefined list of labour induction methods.

### Data collection and analysis

We developed a strategy to deal with the large volume and complexity of trial data relating to labour induction. This involved a two-stage method of data extraction.

We used fixed-effect Mantel-Haenszel meta-analysis for combining dichotomous data.

If we identified substantial heterogeneity ( $I^2$  greater than 50%), we used a random-effects method.

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**Vaginal misoprostol for cervical ripening and induction of labour (Review)**

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## Main results

We included 121 trials. The risk of bias must be kept in mind as only 13 trials were double blind.

Compared to placebo, misoprostol was associated with reduced failure to achieve vaginal delivery within 24 hours (average relative risk (RR) 0.51, 95% confidence interval (CI) 0.37 to 0.71). Uterine hyperstimulation, without fetal heart rate (FHR) changes, was increased (RR 3.52 95% CI 1.78 to 6.99).

Compared with vaginal prostaglandin E2, intracervical prostaglandin E2 and oxytocin, vaginal misoprostol was associated with less epidural analgesia use, fewer failures to achieve vaginal delivery within 24 hours and more uterine hyperstimulation. Compared with vaginal or intracervical prostaglandin E2, oxytocin augmentation was less common with misoprostol and meconium-stained liquor more common.

Lower doses of misoprostol compared to higher doses were associated with more need for oxytocin augmentation and less uterine hyperstimulation, with and without FHR changes.

We found no information on women's views.

## Authors' conclusions

Vaginal misoprostol in doses above 25 mcg four-hourly was more effective than conventional methods of labour induction, but with more uterine hyperstimulation. Lower doses were similar to conventional methods in effectiveness and risks. The authors request information on cases of uterine rupture known to readers. The vaginal route should not be researched further as another Cochrane review has shown that the oral route of administration is preferable to the vaginal route. Professional and governmental bodies should agree guidelines for the use of misoprostol, based on the best available evidence and local circumstances.

## PLAIN LANGUAGE SUMMARY

### Vaginal misoprostol is effective in inducing labour but more research is needed on safety

Sometimes it is necessary to bring on labour artificially because of safety concerns for the mother or baby. Misoprostol is a hormone given by insertion through the vagina or rectum, or by mouth to ripen the cervix and bring on labour. The review of 121 trials found that larger doses of misoprostol are more effective than prostaglandin and that oxytocin is used in addition less often. However, misoprostol also increases hyperstimulation of the uterus. With smaller doses, the results are similar to other methods. The trials reviewed are too small to determine whether the risk of rupture of the uterus is increased. More research is needed into the safety and best dosages of misoprostol. Another Cochrane review has shown that the oral route of administration is preferable to the vaginal route.

## BACKGROUND

Sometimes it is necessary to bring on labour artificially because of safety concerns for the mother or baby. This review is one of a series of reviews of methods of labour induction using a standardised protocol. For more detailed information on the rationale for this methodological approach, please refer to the currently published 'generic' protocol (Hofmeyr 2009). The generic protocol describes how a number of standardised reviews will be combined to compare various methods of preparing the cervix of the uterus and inducing labour.

The main problems experienced during induction of labour are ineffective labour, and excessive uterine activity which may cause

fetal distress. Both problems may lead to an increased risk of caesarean section. Methods of induction of labour include administration of oxytocin, prostaglandins, prostaglandin analogues and smooth muscle stimulants such as herbs or castor oil (Mitri 1987), or mechanical methods such as digital stretching of the cervix and sweeping of the membranes, hygroscopic cervical dilators, extra-amniotic balloon catheters, artificial rupture of the membranes, and nipple stimulation.

Standardised 'scoring' of the cervix prior to labour induction has been recommended (Bishop 1964). Oxytocin has the disadvantage of a high failure rate when the cervix is unfavourable (low cervical

score), and requiring monitored continuous intravenous infusion.

Artificial rupture of membranes is also less effective or may not be possible when the cervix is unfavourable. It may increase the risk of infection if labour does not proceed promptly. Rupture of membranes may also increase the vertical transmission of specific maternal infections such as HIV.

Unsuccessful labour induction is most likely when the cervix is unfavourable and, in this circumstance, prostaglandin preparations have proved to be beneficial (Keirse 1993; MacKenzie 1997). Those prostaglandins that have been registered for cervical ripening and labour induction are expensive and unstable, requiring refrigerated storage. Uterine hyperstimulation has been identified as a particular problem during labour induction with prostaglandins, and has been treated with tocolysis (Egarter 1990).

Misoprostol (Cytotec, Searle) is a methyl ester of prostaglandin E1 additionally methylated at C-16 and is marketed for use in the prevention and treatment of peptic ulcer disease caused by prostaglandin synthetase inhibitors. It is inexpensive, easily stored at room temperature and has few systemic side effects. It is rapidly absorbed orally and vaginally. The reported mean peak serum misoprostol acid following oral administration was 227 pg/ml versus vaginal route 165 pg/ml; the times to peak levels were 34 versus 80 minutes. Vaginally absorbed serum levels are more prolonged (Zieman 1997). Irrespective of serum levels, vaginal misoprostol may have locally mediated effects.

Misoprostol has been shown in several studies to be an effective myometrial stimulant of the pregnant uterus, selectively binding to EP-2/EP-3 prostanoid receptors (Senior 1993).

Misoprostol has been used widely for obstetric and gynaecological indications despite the fact that it has not been registered for such use. It has therefore not undergone the systematic testing for appropriate dosage and safety required for registration.

Misoprostol is an effective abortifacient, both alone and following pretreatment with mifepristone (Norman 1991). Its widespread use in Brazil (Costa 1993) resulted in the identification of teratogenic effects (Fonseca 1991).

Use of misoprostol for second trimester termination of pregnancy has been associated with uterine rupture, particularly when combined with oxytocin infusion. In a report of 803 women admitted with abortion complications in Rio de Janeiro, 458 reported using misoprostol (Costa 1993). There was one maternal death from uterine rupture at 16 weeks' gestation following self-medication with misoprostol.

Third trimester cervical ripening and labour induction with misoprostol have been reported using the oral, vaginal, rectal and buccal/sublingual routes. Clinical experience with misoprostol for labour induction has been reviewed by Wing (Wing 1999b).

Mariani Neto et al (Mariani Neto 1987) first reported using oral misoprostol 400 micrograms (mcg) four hourly for induction of labour following intrauterine death.

In a subsequent paper (Mariani Neto 1988), they described 'uterine tachysystole' with misoprostol use at term, which appeared unrelated to dosage. Since that time, several small studies have confirmed an increased incidence of uterine tachysystole (greater than five contractions per 10 minutes for at least 20 minutes), uterine hypersystole/hypertonus (a contraction of two minutes or more) and/or uterine hyperstimulation syndrome (uterine tachysystole or hypersystole with fetal heart rate (FHR) changes such as persistent decelerations, tachycardia or reduced short term variability). The conclusion from a meta-analysis was that published data confirmed the safety of intravaginal misoprostol for cervical ripening and labour induction. The data showed an increased incidence of uterine tachysystole (odds ratio 2.70, 95% confidence intervals 1.80 to 4.04), but there was no statistically significant increase in adverse fetal outcome (Sanchez-Ramos 1997). Wing et al (088 Wing 1995a; 044 Wing 1995b; 025 Wing 1996; 038 Wing 1997) have suggested that uterine hyperstimulation and meconium passage with vaginal misoprostol may be less frequent using a 25 microgram dose, six hourly.

Merrell and co-workers (Merrell 1995) reported a series of 62 inductions of labour with vaginal misoprostol. There were two stillbirths, one apparently due to a tight nuchal cord, and one unexplained. They commented on rapid onset of contractions and described one woman with induction to delivery interval of only two hours. In a subsequent abstract (Merrell 1996), they described labour inductions with vaginal misoprostol in 345 women with live fetuses and 86 with intrauterine deaths. There was one unexplained maternal death; two uterine ruptures, one of which followed a previous caesarean section; eight caesarean sections for fetal distress and one for uterine hyperstimulation; and 10 perinatal deaths.

There have been several reports of uterine rupture following misoprostol labour induction with and without previous caesarean section (Bennett 1997; Sciscione 1998; Blanchette 1999; Matthews 1999; Khosla 2002). One unpublished case of uterine rupture occurred in a nulliparous woman following misoprostol use (EM Smith, personal communication). At term plus 12 days she received misoprostol 100 mcg vaginally. After six hours her cervix was found to be 7 cm dilated, and she progressed to full dilatation within a further 70 minutes. Fetal distress was suspected. Ventouse application produced no descent, so delivery was effected by caesarean section. The infant showed no signs of life at birth. After resuscitation, life was sustained for a few hours only. A posterior uterine tear arising from the cervix and spiraling up the posterior aspect of the uterus was discovered and repaired. Because such uterine tears are rare in nulliparous women without prolonged labour or syntocinon use, a causal relationship with the use of misoprostol must be considered.

One trial of misoprostol for labour induction in women with prior caesarean section has been terminated prematurely because of disruption of the uterine incision in two of the first 17 misoprostol-treated women (025 Wing 1998a). The dosage of misoprostol used was conservative (25 µg six hourly to a maximum of four doses). Two and three doses were used respectively in the two cases of ruptured uterus.

In a retrospective review, uterine rupture occurred in 5/89 (5.6%) of women with previous caesarean delivery who had labour induced with misoprostol, compared with 1/423 (0.2%) of those who did not (Plaut 1999). In another retrospective review of labour induction in 575 women with previous caesarean section, the rate of uterine rupture was 5/172 (2.9%) for prostaglandin E2 gel; 1/129 (0.76%) for intracervical Foley catheter; and 3/474 (0.74%) for induction not requiring cervical ripening; compared with 7/1544 (0.45%) for spontaneous trial of labour (Ravasia 2000). In a third retrospective review, no uterine ruptures were detected among 48 women with previous caesarean section whose labour was induced with misoprostol 50 mcg vaginally four hourly (Choy-Hee 2001).

Personal discussion with colleagues has revealed several cases of rupture of an unscarred uterus following misoprostol usage, possibly related to higher dosages than have been used in the trials reviewed. These cases are usually not reported. We call on readers to send us details of any such cases known to them, including if possible age, parity, any previous uterine surgery, dosage of misoprostol and details of the uterine rupture. This will enable us to compile a register of such problems.

This review will focus on the effectiveness and safety of misoprostol administered vaginally for cervical ripening and labour induction in the third trimester of pregnancy.

The use of oral (Alfirevic 2006) and buccal/sublingual (Muzonzini 2004) misoprostol for cervical priming and labour induction, compared with other methods including misoprostol administered vaginally, are reviewed separately.

## OBJECTIVES

To determine, from the best available evidence, the effectiveness and safety of misoprostol administered vaginally for third trimester cervical ripening and induction of labour.

## METHODS

### Criteria for considering studies for this review

### Types of studies

Clinical trials comparing misoprostol administered vaginally for cervical ripening or labour induction, with placebo/no treatment or other methods listed above it on a predefined list of methods of labour induction (*see Methods*); the trials included some form of random allocation to either group; they reported one or more of the pre-stated outcomes; reasonable measures were taken to ensure allocation concealment; and violations of allocated management were not sufficient to materially affect outcomes. We have not included quasi-randomised trials.

### Types of participants

Pregnant women due for third trimester induction of labour. We have not excluded multiple pregnancies. Predefined sub-group analyses (*see list below*): previous caesarean section or not; nulliparity or multiparity; membranes intact or ruptured, and cervix unfavourable, favourable or undefined. Only those outcomes with data appear in the analysis tables.

### Types of interventions

Vaginal administration of misoprostol compared with placebo/no treatment or any other method above it on a predefined list of methods of labour induction.

### Primary comparisons

Misoprostol versus placebo/no treatment

Misoprostol versus oxytocin

Misoprostol versus vaginal prostaglandins

Misoprostol versus intracervical prostaglandins

Low dosage misoprostol regimens versus higher dosage regimens

Misoprostol gel versus tablets

In all the studies of misoprostol versus prostaglandins, the prostaglandin used was dinoprostone intravaginally as a gel, tablet or slow-release pessary, or intracervically as a gel. In most of the studies, oxytocin was used with similar protocols for both the misoprostol and the prostaglandin group, except that in one study (175 Kadanali 1996) oxytocin was started if indicated after six hours in the dinoprostone group and only after 24 hours in the misoprostol group. The effective comparison in this trial is therefore misoprostol versus dinoprostone plus early oxytocin. The results are in keeping with those of other studies.

### Types of outcome measures

Two authors of labour induction reviews (Justus Hofmeyr and Zarko Alfirevic) have prespecified clinically relevant outcomes for trials of methods of cervical ripening/labour induction. We have settled differences by discussion.

We chose five primary outcomes as being most representative of the clinically important measures of effectiveness and complications. We limited sub-group analyses to the primary outcomes:

- (1) vaginal delivery not achieved within 24 hours;
- (2) uterine hyperstimulation with FHR changes;
- (3) caesarean section;
- (4) serious neonatal morbidity or perinatal death (e.g. seizures, birth asphyxia defined by trialists, neonatal encephalopathy, disability in childhood);
- (5) serious maternal morbidity or death (e.g. uterine rupture, admission to intensive care unit, septicaemia).

Perinatal and maternal morbidity and mortality are composite outcomes. This is not an ideal solution because some components are clearly less severe than others. It is possible for one intervention to cause more deaths but less severe morbidity. However, in the context of labour induction at term this is unlikely. All these events will be rare, and a modest change in their incidence will be easier to detect if composite outcomes are presented. We have explored the incidence of individual components as secondary outcomes (*see below*).

Secondary outcomes relate to measures of effectiveness, complications and satisfaction.

#### Measures of effectiveness

- (6) Cervix unfavourable/unchanged after 12 to 24 hours;
- (7) oxytocin augmentation.

#### Complications

- (8) Uterine hyperstimulation without FHR changes;
- (9) uterine rupture;
- (10) epidural analgesia;
- (11) instrumental vaginal delivery;
- (12) meconium-stained liquor;
- (13) Apgar score less than seven at five minutes;
- (14) neonatal intensive care unit admission;
- (15) neonatal encephalopathy;
- (16) perinatal death;
- (17) disability in childhood;
- (18) maternal side effects (all);
- (19) maternal nausea;
- (20) maternal vomiting;
- (21) maternal diarrhoea;
- (22) other maternal side effects;
- (23) postpartum haemorrhage (as defined by the trial authors);
- (24) serious maternal complications (e.g. intensive care unit admission, septicaemia but excluding uterine rupture);
- (25) maternal death.

#### Measures of satisfaction

- (26) Woman not satisfied;

- (27) caregiver not satisfied.

'Uterine rupture' will include all clinically significant ruptures of unscarred or scarred uteri. We will exclude trivial scar dehiscence noted incidentally at the time of surgery.

While we sought all the above outcomes, we have included only those with data in the analysis tables.

The terminology of uterine hyperstimulation is problematic (Curtis 1987). In this review we have used the term 'uterine hyperstimulation without FHR changes' to include uterine tachysystole (greater than five contractions per 10 minutes for at least 20 minutes) and uterine hypersystole/hypertonus (a contraction lasting at least two minutes) and 'uterine hyperstimulation with FHR changes' to denote uterine hyperstimulation syndrome (tachysystole or hypersystole with FHR changes such as persistent decelerations, tachycardia or decreased short-term variability). However, due to varied reporting there is the possibility of subjective bias in interpretation of these outcomes. Also, it is not always clear from trials if these outcomes are reported in a mutually exclusive manner.

We included outcomes in the analysis if reasonable measures were taken to minimise observer bias; missing data were insufficient to materially influence conclusions; and data were available for analysis according to original allocation.

#### Search methods for identification of studies

##### Electronic searches

We searched the Cochrane Pregnancy and Childbirth Group's Trials Register by contacting the Trials Search Co-ordinator (November 2008). We updated this search on 30 April 2010 and added the results to Studies awaiting classification for consideration in the next update.

The Cochrane Pregnancy and Childbirth Group's Trials Register is maintained by the Trials Search Co-ordinator and contains trials identified from:

1. quarterly searches of the Cochrane Central Register of Controlled Trials (CENTRAL);
2. weekly searches of MEDLINE;
3. handsearches of 30 journals and the proceedings of major conferences;
4. weekly current awareness alerts for a further 44 journals plus monthly BioMed Central email alerts.

Details of the search strategies for CENTRAL and MEDLINE, the list of handsearched journals and conference proceedings, and the list of journals reviewed via the current awareness service can be found in the 'Specialized Register' section within the editorial information about the [Cochrane Pregnancy and Childbirth Group](#).

Trials identified through the searching activities described above are each assigned to a review topic (or topics). The Trials Search



Co-ordinator searches the register for each review using the topic list rather than keywords.

### Searching other resources

We hand searched the reference lists of trial reports and reviews. We screened and assessed all Chinese papers according to the review protocol by a first-language Chinese speaker (Linan Cheng) and excluded all due to serious methodological limitations. We did not apply any language restrictions.

### Data collection and analysis

To avoid duplication of data, the authors of induction of labour reviews agreed a specific order for labour induction methods, from one to 27. Each primary review included comparisons between one of the methods (from two to 27) with only those methods above it on the list. Thus, this review of intravenous oxytocin (4) included only comparisons with intracervical prostaglandins (3), vaginal prostaglandins (2) or placebo/no treatment (1). The current list is as follows:

- (1) placebo/no treatment;
- (2) vaginal prostaglandins (Kelly 2003);
- (3) intracervical prostaglandins (Boulvain 2008);
- (4) intravenous oxytocin (Kelly 2001a);
- (5) amniotomy (Bricker 2000);
- (6) intravenous oxytocin with amniotomy (Howarth 2001);
- (7) vaginal misoprostol;
- (8) oral misoprostol (Alfirevic 2006);
- (9) mechanical methods including extra-amniotic Foley catheter (Boulvain 2001);
- (10) membrane sweeping (Boulvain 2005);
- (11) extra-amniotic prostaglandins (Hutton 2001);
- (12) intravenous prostaglandins (Luckas 2000);
- (13) oral prostaglandins (French 2001);
- (14) mifepristone (Hapangama 2009);
- (15) oestrogens with or without amniotomy (Thomas 2001);
- (16) corticosteroids (Kavanagh 2006a);
- (17) relaxin (Kelly 2001b);
- (18) hyaluronidase (Kavanagh 2006b);
- (19) castor oil, bath, and/or enema (Kelly 2001c);
- (20) acupuncture (Smith 2004);
- (21) breast stimulation (Kavanagh 2005);
- (22) sexual intercourse (Kavanagh 2001);
- (23) homoeopathic methods (Smith 2003);
- (24) nitric oxide donors (Kelly 2008);
- (25) buccal or sublingual misoprostol (Muzonzini 2004);
- (26) hypnosis;
- (27) other methods for induction of labour.

The reviews were analysed by the following clinical categories of participants:

1. previous caesarean section or not;

2. nulliparity or multiparity;
3. membranes intact or ruptured;
4. cervix favourable, unfavourable or undefined.

For most reviews, the initial data extraction process was conducted centrally. This was co-ordinated from the Clinical Effectiveness Support Unit (CESU) at the Royal College of Obstetricians and Gynaecologists, UK, in co-operation with the Pregnancy and Childbirth Group of The Cochrane Collaboration. This process allowed the data extraction process to be standardised across all the reviews. From 2001, the data extraction was no longer conducted centrally.

The trials were initially reviewed on eligibility criteria, using a standardised form and the basic selection criteria specified above. Following this, a standardised data extraction form was developed and then piloted for consistency and completeness. This pilot process involved the researchers at the CESU and previous review authors in the area of induction of labour. For a description of the methods used to carry out the initial reviews, see Appendix 1.

Due to the large number of trials, double data extraction was not feasible and agreement between the three data extractors was therefore assessed on a random sample of trials to update in 2003. For the same reason, in the 2009 update, the data extraction was checked on around 50% of the trials in a random sample selection. In 2008, the methods and software for carrying out reviews were updated, as a result of which new reviews and updates, where appropriate, use these new methods (Higgins 2008a; RevMan 2008), which are described in the Methods section of all the individual new and updated reviews.

For this update, we used the following methods when assessing the new trials identified by the updated search.

### Selection of studies

One review author (Cynthia Pileggi (CP)) assessed for inclusion all the potential studies we identified as a result of the search strategy. We discussed studies for which there was any uncertainty with a second author (Justus Hofmeyr (GJH)).

### Data extraction and management

We designed a form to extract data. For eligible studies, one review author (CP) extracted the data using the agreed form. GJH independently repeated selection of studies and data extraction on a random sample of studies. We resolved discrepancies through discussion or, if required, we would have consulted the third author. We entered the data into Review Manager software (RevMan 2008) and checked them for accuracy.

When information regarding any of the above was unclear, we attempted to contact authors of the original reports to provide further details.



## Assessment of risk of bias in included studies

One review author (CP) independently assessed risk of bias for each study using the criteria outlined in the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2008a).

### (1) Sequence generation (checking for possible selection bias)

We described for each included study the method used to generate the allocation sequence in sufficient detail to allow an assessment of whether it should produce comparable groups.

We assessed the method as:

- adequate (any truly random process, e.g. random number table; computer random number generator);
- inadequate (any non-random process, e.g. odd or even date of birth; hospital or clinic record number); or
- unclear.

### (2) Allocation concealment (checking for possible selection bias)

We described for each included study the method used to conceal the allocation sequence in sufficient detail and determined whether intervention allocation could have been foreseen in advance of, or during recruitment, or changed after assignment.

We assessed the methods as:

- adequate (e.g. telephone or central randomisation; consecutively numbered sealed opaque envelopes);
- inadequate (open random allocation; unsealed or non-opaque envelopes, alternation; date of birth);
- unclear.

### (3) Blinding (checking for possible performance bias)

We described for each included study the methods used, if any, to blind study participants and personnel from knowledge of which intervention a participant received. We judged studies at low risk of bias if they were blinded, or if we judged that the lack of blinding could not have affected the results. We assessed blinding separately for different outcomes or classes of outcomes.

We assessed the methods as:

- adequate, inadequate or unclear for participants;
- adequate, inadequate or unclear for personnel;
- adequate, inadequate or unclear for outcome assessors.

### (4) Incomplete outcome data (checking for possible attrition bias through withdrawals, dropouts, protocol deviations)

We described for each included study, and for each outcome or class of outcomes, the completeness of data including attrition and exclusions from the analysis. We stated whether attrition and exclusions were reported, the numbers included in the analysis at

each stage (compared with the total randomised participants), reasons for attrition or exclusion where reported, and whether missing data were balanced across groups or were related to outcomes. Where sufficient information was reported, or could be supplied by the trial authors, we would re-include missing data in the analyses. We assessed methods as:

- adequate (less than 5% loss to follow up);
- inadequate;
- unclear.

### (5) Selective reporting bias

We described for each included study how we investigated the possibility of selective outcome reporting bias and what we found. We assessed the methods as:

- adequate (where it is clear that all of the study's pre-specified outcomes and all expected outcomes of interest to the review have been reported);
- inadequate (where not all the study's pre-specified outcomes have been reported; one or more reported primary outcomes were not pre-specified; outcomes of interest are reported incompletely and so cannot be used; study fails to include results of a key outcome that would have been expected to have been reported);
- unclear.

### (6) Other sources of bias

We described for each included study any important concerns we had about other possible sources of bias.

We assessed whether each study was free of other problems that could put it at risk of bias:

- yes;
- no;
- unclear.

### (7) Overall risk of bias

We made explicit judgements about whether studies were at high risk of bias, according to the criteria given in the *Handbook* (Higgins 2008a). With reference to (1) to (6) above, we assessed the likely magnitude and direction of the bias and whether we consider it is likely to impact on the findings. We explored the impact of the level of bias through undertaking sensitivity analyses - see [Sensitivity analysis](#).

## Measures of treatment effect

### Dichotomous data

For dichotomous data, we present results as summary risk ratio with 95% confidence intervals.

### Continuous data

This systematic review did not include continuous data.

### Unit of analysis issues

#### Cluster-randomised trials

We would include cluster-randomised trials in the analyses along with individually randomised trials. We would adjust their sample sizes using the methods described in the *Handbook* (Higgins 2008b) using an estimate of the intra cluster correlation co-efficient (ICC) derived from the trial (if possible), or from another source. If ICCs from other sources are used, we planned to report this and conduct sensitivity analyses to investigate the effect of variation in the ICC. If we identify cluster-randomised trials in addition to the individually-randomised trials, we plan to synthesise the relevant information. We would consider it reasonable to combine the results from both if there is little heterogeneity between the study designs and the interaction between the effect of intervention and the choice of randomisation unit is considered to be unlikely.

We also planned to acknowledge heterogeneity in the randomisation unit and perform a separate meta-analysis.

#### Dealing with missing data

For included studies, we have noted levels of attrition. We explored the impact of including studies with high levels of missing data in the overall assessment of treatment effect by using sensitivity analysis.

For all outcomes, we have carried out analyses, as far as possible, on an intention-to-treat basis, i.e. we attempted to include all participants randomised to each group in the analyses. The denominator for each outcome in each trial is the number randomised minus any participants whose outcomes are known to be missing.

#### Assessment of heterogeneity

We used the  $I^2$  statistic to measure heterogeneity among the trials in each analysis. If we identified substantial heterogeneity ( $I^2$  greater than 50%), we explored it by pre-specified subgroup analysis.

#### Assessment of reporting biases

Where we suspected reporting bias (see 'Selective reporting bias' above), we attempted to contact study authors asking them to provide missing outcome data. Where this was not possible, and the missing data were thought to introduce serious bias, we explored the impact of including such studies in the overall assessment of results by a sensitivity analysis.

### Data synthesis

We carried out statistical analysis using the Review Manager software (RevMan 2008). We used fixed-effect Mantel-Haenszel meta-analysis for combining dichotomous data where trials were examining the same intervention, and we judged the trials' populations and methods sufficiently similar. Where we suspected clinical or methodological heterogeneity between studies sufficient to suggest that treatment effects may differ between trials, we used a random-effects meta-analysis.

If we identified substantial heterogeneity, we noted this and performed the analysis using a random-effects method.

### Subgroup analysis and investigation of heterogeneity

We carried out the following subgroup analyses:

1. previous caesarean section or not;
2. nulliparity or multiparity;
3. membranes intact or ruptured;
4. cervix favourable, unfavourable or undefined.

We used primary outcomes only in subgroup analysis.

For fixed-effect meta-analyses we conducted planned subgroup analyses classifying whole trials by interaction tests as described by Deeks 2001. For random-effects meta-analyses we assessed differences between subgroups by inspection of the subgroups' confidence intervals; non-overlapping confidence intervals indicate a statistically significant difference in treatment effect between the subgroups.

### Sensitivity analysis

We plan to carry out sensitivity analysis by excluding trials with greater risk of bias, particularly with respect to allocation concealment, in a future update of this review.

## RESULTS

### Description of studies

See: [Characteristics of included studies](#); [Characteristics of excluded studies](#); [Characteristics of studies awaiting classification](#); [Characteristics of ongoing studies](#).

#### Included studies

We have included 121 studies in this review. See table of [Characteristics of included studies](#) for details. Because a wide range of misoprostol dosages has been used, we have coded the included studies with a prefix to reflect roughly the dosage of vaginal misoprostol received in the first six hours, calculated as follows: initial

dose + (s x (6 - i)/4), where 's' is a subsequent dose within six hours, and 'i' is the interval in hours. This is based on the approximation that vaginal misoprostol is absorbed uniformly over a four-hour period. Where a subsequent oral dose was used, the oral dose value was halved. Use of a gel preparation is indicated by the letter 'G'. This coding allows approximate ranking of the trials by misoprostol dosage, and enables readers to assess the effect of dosage on results. We detected no discrepancies in the sample of data extraction performed in duplicate.

### Excluded studies

For details of excluded studies, see table of [Characteristics of excluded studies](#).

### Risk of bias in included studies

With the exception of 13 double-blind trials (100G Fletcher 1993; 050 El-Azeem 1997; 043 Farah 1997; 050 Surbek 1997; 050 Gotschall 1998; 025G Srisomboon 1998; 043 Diro 1999; 100 Montealegre 1999; 025 Stitely 2000; 048 Khoury 2001; 058 Ferguson 2002; 038 Meydanli 2003; 050 Ramsey 2003 - blinded low versus high dose misoprostol comparison only), allocation was by means of sealed envelopes or unspecified, and treatment was not blinded. There is therefore a real possibility of bias affecting both the clinical management of the women (e.g. decisions to undertake caesarean section) and the assessment of outcomes. Such biases might operate in either direction (for example, a clinician enthusiastic about the potential of misoprostol might be less likely to perform caesarean section in the misoprostol group, while one anxious about the experimental nature of misoprostol might be more likely to perform caesarean section in this group). We performed limited sensitivity analysis excluding non-blinded studies for primary outcomes with significant heterogeneity and 10 or more trials included. For the comparison misoprostol versus vaginal prostaglandins, all women: the outcome vaginal delivery not achieved in 24 hours was unchanged; the outcome uterine hyperstimulation with FHR changes was no longer statistically significant (small numbers). For the comparison misoprostol versus oxytocin: the outcomes vaginal delivery not achieved in 24 hours and caesarean section were no longer statistically significant (small numbers remaining in the analysis).

The possibility of bias must be kept in mind in the interpretation of the results.

In the study of 050 Le Roux 2002, 93 of 573 enrolled women were excluded for 'protocol violations'. There did not appear to be a selective loss from any group, and the baseline data were similar between groups.

In 050 Pandis 2001, 235/670 were excluded after randomisation, mainly for spontaneous delivery before induction or induction by amniotomy for cervical score seven or more.

In 075 Ghidini 2001, seven of 65 enrolled women were excluded due to emergence of exclusion criteria. The groups were somewhat unbalanced (32 received 50 mcg and 26 received 100 mcg).

In 150 De la Torre 2001, 50 of 410 enrolled were withdrawn for protocol deviation (16), patient withdrawal (7), or missing data (27). The final groups differed in numbers (misoprostol 168, oxytocin 192). This raises the possibility of selective withdrawal from the misoprostol group.

In 088 Garry 2003, 14 women of 200 enrolled were withdrawn for physician request (10), used wrong medication (2), patient request (1) and unknown breech presentation (1). It suggests deviation from the intention to treat analysis.

The 2009 update used the new methodology of the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2008a) regarding the evaluation of sequence generation, allocation concealment, blinding of participants, personnel and outcome assessors, report of incomplete outcome data, selective outcome reporting bias and other sources. We have presented the quality evaluation of each included study in the corresponding risk of bias table. This update includes 54 comparisons with more than 10 study results in the pooled analyses, 19 of them in primary outcomes. Five out of these 19 comparisons present asymmetrical funnel plots suggesting potential publication bias.

### Effects of interventions

We have included 121 studies in this review. We sought all the outcomes listed under 'Types of outcome measures', and sub-groups defined in 'Types of participants'. Only those with data appear in the analysis tables.

#### Vaginal misoprostol versus placebo

##### Primary outcomes

The 10 studies (1141 women) included in this part of the review (100G Fletcher 1993; 100G Srisomboon 1996; 025 Stitely 2000; 050 Thomas 2000; 025 Incerpi 2001; 050 Ortiz 2002; 025 McKenna 2004; 050 Gelisen 2005; 025 Krupa 2005; 025 Oboro 2005) showed a trend towards failure to deliver within 24 hours (five trials, 735 women, average relative risk (RR) 0.56, 95% CI 0.31 to 1.03).

##### Secondary outcomes

We found a clear effect of misoprostol on cervical ripening (two trials, average RR of unchanged cervix at 12 to 24 hours 0.09, 95% confidence interval (CI) 0.03 to 0.24).

Uterine hyperstimulation without FHR changes was increased (six trials, 794 women RR 3.52, 95% CI 1.78 to 6.99). Six trials (100G Fletcher 1993; 025 Stitely 2000; 050 Thomas 2000; 050

[Ortiz 2002](#); [050 Gelisen 2005](#); [025 Krupa 2005](#)) with 814 women showed an unexpected reduction of meconium-stained liquor with the use of misoprostol for labour induction. The numbers studied were too small to assess the impact on obstetric management and maternal and neonatal complications.

### **Vaginal misoprostol versus vaginal prostaglandins**

There were 38 included trials with 7022 participants.

#### **Primary outcomes**

Failure to achieve vaginal delivery within 24 hours (22 trials, average RR 0.77, 95% CI 0.66 to 0.89) was reduced overall, but not in the two trials using less than 50 mcg misoprostol in the first six hours ([038 Wing 1997](#); [048 Khoury 2001](#)).

Uterine hyperstimulation with FHR changes was variable between trials, but overall tended to be more common with misoprostol (31 trials, average RR 1.43, 95% CI 0.97 to 2.09).

Caesarean sections were variable between trials, with a trend to be reduced with vaginal misoprostol (34 trials RR 0.95, 95% CI 0.87 to 1.03).

#### **Secondary outcomes**

Oxytocin augmentation was reduced with misoprostol (36 trials, average RR 0.68, 95% CI 0.60 to 0.76).

Uterine hyperstimulation without FHR changes was more common with misoprostol (26 trials, average RR 1.99, 95% CI 1.41 to 2.79).

Epidural analgesia was used less frequently with misoprostol (eight trials, RR 0.92, 95% CI 0.85 to 0.99).

Meconium-stained liquor was more common with misoprostol (18 trials, RR 1.35, 95% CI 1.13 to 1.61). There were no statistically significant differences in perinatal or maternal outcomes.

Results were similar for the sub-groups of women with unfavourable cervixes and those with intact membranes and unfavourable cervixes.

There were similar trends for women with intact membranes and variable or undefined cervixes, but the numbers were too small for clear outcomes.

For subgroups of primiparous or multiparous women, the numbers were small and no differences in any outcomes were shown, except that for all primiparous women, misoprostol shows reduced caesarean section (RR 0.82, 95% CI 0.68 to 0.99), and a trend to reduced vaginal delivery not achieved in 24h (average RR 0.70, 95% CI 0.46 to 1.05).

The results of [048 Khoury 2001](#) differed somewhat from other studies. This may be because a gel preparation of misoprostol was used, and it is possible that some activity is lost in the preparation or administration (see results of misoprostol gel versus tablets below).

### **Vaginal misoprostol versus intracervical prostaglandins**

There were 27 included trials with 3311 participants.

#### **Primary outcomes**

Failure to achieve vaginal delivery within 24 hours was consistently reduced with misoprostol (13 trials, RR 0.63, 95% CI 0.56 to 0.71).

Uterine hyperstimulation with associated FHR changes was variable between trials, but all were consistent with the pooled result showing an increase with misoprostol (20 trials, RR 2.32, 95% CI 1.64 to 3.28). The latter result was similar for the sub-group of trials studying women with intact membranes and unfavourable cervixes.

Caesarean sections were variable between trials, with no significant differences overall.

#### **Secondary outcomes**

Only one trial reported the outcome 'failure to achieve cervical ripening within 12 hours' ([075 Buser 1997](#)); this was reduced with misoprostol (RR 0.68, 95% CI 0.52 to 0.88).

Oxytocin augmentation was used less often with misoprostol (20 trials, average RR 0.55, 95% CI 0.48 to 0.64).

Uterine hyperstimulation without FHR changes was more common with misoprostol (17 trials, RR 1.95, 95% CI 1.57 to 2.42). The rates of vaginal instrumental delivery were variable between trials. Epidural analgesia was used less frequently with misoprostol (two trials, RR 0.64, 95% CI 0.48 to 0.86). Meconium-stained liquor was increased with misoprostol (14 trials, RR 1.29, 95% CI 1.04 to 1.59). There were no statistically significant differences in perinatal or maternal outcomes.

Most of the trials studied women with unfavourable cervixes, for whom the results were similar to the overall results. Results were also similar for women with intact membranes and unfavourable cervixes. The two trials with intact membranes and variable or undefined cervix also showed a similar pattern of results.

### **Vaginal misoprostol versus oxytocin**

There were 25 trials with 3074 participants.

#### **Primary outcomes**

Misoprostol, in the doses used in these trials, was more effective than oxytocin for labour induction (10 trials, average RR of failure to achieve vaginal delivery within 24 hours 0.65, 95% CI 0.47 to 0.90). Two trials using less than 50 mcg misoprostol showed no reduction ([025 Wing 1998b](#); [025 Haghighi 2006](#)).

Twenty-five studies showed a reduction in caesarean section risk with the use of misoprostol (average RR 0.76, 95% CI 0.60 to 0.96).

## Secondary outcomes

Uterine hyperstimulation without FHR changes was more common with misoprostol (15 trials, RR 2.24 95% CI 1.82 to 2.77 respectively). There was a trend to reduced epidural analgesia with misoprostol (three trials, RR 0.82, 95% CI 0.67 to 1.00). Vaginal instrumental delivery was reduced in the misoprostol group (13 trials, RR 0.74, 95% CI 0.56 to 0.99).

Apgar score less than 7 at five minutes, with 13 studies and 1906 participants in the general group, was substantially reduced with misoprostol use (RR 0.56, 95% CI 0.34 to 0.92). Four studies with 334 participants showed increased risk of maternal side effects

(RR 5.04, 95% CI 1.51 to 16.86). There were no differences in other perinatal or maternal outcomes.

One trial in women with previous caesarean section was stopped when uterine rupture occurred in two of the first 17 women who received misoprostol (025 Wing 1998b) and in another study one uterine rupture occurred in 34 women in the misoprostol group (all women with unfavourable cervix, 050 Abdul 2007).

## Misoprostol lower dosage regimen versus higher dose

There were 21 trials with 2913 participants. The dosages compared are as follows.

Number of studies	Misoprostol low dosage	Misoprostol high dosage	Interval of use
2	12.5 mcg	25 mcg	4 to 6 hours
11	25 mcg	50 mcg	3 to 6 hours
1	35 mcg	50 mcg	4.5 hours
6	50 mcg	100 mcg	4 to 6 hours
Note: 006 Ewert 2006 was not included because of the mode of misoprostol administration.			

## Primary outcomes

There was no significant difference in the risk of failures to achieve delivery within 24 hours. There was less uterine hyperstimulation with FHR changes in the lower dose groups (16 trials, RR 0.51, 95% CI 0.37 to 0.69).

Serious maternal complications were reported in one study (025 Wing 1996): one maternal death occurred in a primiparous woman, nine hours after a single misoprostol dose and shortly after amniotomies and epidural analgesia, from amniotic fluid embolisation. Two caesarean hysterectomies were performed for atonic uterine haemorrhage, 13 and 30 hours after single doses of misoprostol, in one primiparous woman with uncomplicated labour, and in one nulliparous woman who developed chorioamnionitis following prolonged labour induction attempts by oxytocin augmentation. It is not clear whether these three women were allocated to the low (25 mcg) or the higher (50 mcg) dosage regimen misoprostol group.

lower dosage was 50 mcg. There were no differences in mode of delivery, meconium-stained liquor or maternal side effects. There was less uterine hyperstimulation without FHR changes (14 trials, RR 0.57, 95% CI 0.46 to 0.69). There was a trend to fewer babies being admitted to the neonatal intensive care unit (9 trials, RR 0.82, 95% CI 0.64 to 1.05), particularly in the higher dose ranges. Five perinatal deaths were reported (019 Filho 2007; 050 Majoko 2002a). There was one uterine rupture (038 Has 2002) with the use of low dose of misoprostol and two with the use of higher dose of misoprostol (050 Majoko 2002a; 075 Reyna-Villasmil 2005). However, most studies have not specifically reported these outcomes. We have included only those specified in the reports in the data tables.

## Misoprostol gel versus tablets

### Primary outcomes

In one trial with 467 participants reviewed (050G Carlan 1997), uterine hyperstimulation with FHR changes was reduced with the gel preparation (RR 0.49, 95% CI 0.29 to 0.83).

### Secondary outcomes

There was significantly more use of oxytocin (18 trials, average RR 1.30, 95% CI 1.14 to 1.49). This effect was due to the trials with a lower range of doses, and was not seen in the trials in which the



## Secondary outcomes

The use of oxytocin (RR 1.26, 95% CI 1.13 to 1.41) and epidural analgesia (RR 1.19, 95% CI 1.03 to 1.38) were increased. It is possible that in the process of gel preparation some potency is lost or that absorption is reduced.

One study showed no benefit from moistening misoprostol prior to insertion with 11 ml 3% acetic acid, versus dry tablets (Sanchez Ramos 2002).

A cost analysis in a high-income country showed that the reduced cost in the misoprostol group (Sterling mean 2134, SD 574 versus 2202, SD 595 per case) was insignificant in relation to the overall cost of labour induction (050 Rozenberg 2001).

## DISCUSSION

Overall, this systematic review found that vaginal misoprostol is the more effective option for induction of labour and cervical ripening compared with oxytocin, dinoprostone and placebo. It also found that higher doses of vaginal misoprostol have no comparative advantages to the lower doses. There is, in general, considerable consistency between trials, except with respect to caesarean section rates and to the low misoprostol dosage regimens. The trials show that vaginal misoprostol in dosages ranging from 25 mcg two to three hourly, to 50 mcg four hourly (most studies), to 100 mcg six to 12 hourly, appear to be more effective than oxytocin or dinoprostone in the usual recommended doses for induction of labour, but with increased rates of uterine hyperstimulation both without and with associated FHR changes. The rates of caesarean section were inconsistent, tending to be reduced with misoprostol. The indication for caesarean section was not a pre-specified outcome in this review. However, there was a consistent pattern of more operations for fetal distress and fewer for poor labour progress in the misoprostol groups (see 'Characteristics of included studies' table under 'Outcomes').

No differences in perinatal or maternal outcome were shown. However, the trials were not sufficiently large to assess the likelihood of uncommon, serious adverse perinatal and maternal complications. Of particular concern are several reports of uterine rupture following misoprostol use in women with and without previous caesarean section. One maternal death from amniotic fluid embolism following misoprostol induction was reported.

The possibility of inadvertent bias because of the unblinded nature of these studies should be kept in mind.

Lower dosage regimens of misoprostol were not less effective than higher doses in terms of failure to achieve vaginal birth within 24 hours. Adverse effects were reduced, with lower rates of uterine hyperstimulation and a trend to fewer admissions to neonatal intensive care unit.

The finding of a significantly more meconium-stained liquor with misoprostol versus vaginal or intracervical prostaglandins is of interest. Wing et al (088 Wing 1995a) suggested the possibility of meconium passage in response to uterine hyperstimulation or a direct effect of absorbed misoprostol metabolites on the fetal gastrointestinal tract. We have previously observed an increased rate of meconium-stained liquor in women who have ingested castor oil, though causality was not proven, and suggested a possible direct effect of the castor oil metabolites on fetal bowel (Mitri 1987). It is unlikely that the small amount of hydrogenated castor oil found in misoprostol tablets (075 Chuck 1995) would have any pharmacological effect, but the possibility that misoprostol metabolites may directly stimulate fetal bowel is of interest. We have shown an in vitro effect of misoprostol on isolated rat ileum (as well as myometrium) (Matonhodze 2002).

In countries in which misoprostol is being used for non-registered obstetric indications, there is a need for health authorities and professional organisations to clarify the medicolegal implications. Particularly in countries in which conventional prostaglandins are unaffordable, health authorities need to decide whether misoprostol should be used in specific circumstances and, if so, take steps to legalise and regulate such use.

The trials reviewed lacked information on women's views with respect to this method of labour induction.

## AUTHORS' CONCLUSIONS

### Implications for practice

The comparison between oral and vaginal misoprostol is dealt with in a separate Cochrane review (Alfirevic 2006). That review suggests that the optimal route for administration of misoprostol for labour induction is oral, not vaginal. The reasons for this are as follows.

1. Safety. The oral route is associated with considerably less uterine hyperstimulation with FHR changes (9% versus 24.6%; RR 0.37, 95%CI 0.23 to 0.59).
2. Convenience and comfort for the woman.
3. Because of a short half-life, the oral dose can be titrated against the uterine response, commencing with a low dose such as 25 mcg 2-hourly, and increasing if necessary in nulliparous women to a maximum dose of 50 mcg 2-hourly.
4. Accuracy of dosage. In many countries, misoprostol is available only as 200 mcg or 100 mcg tablets. Breaking these tablets into small fragments for vaginal administration carries the risk of inappropriate dosage. Accurate oral dosage can be achieved by dissolving misoprostol in tap water, shaking well and administering as a solution. Left over solution should be discarded 24 hours after preparation.

The relative disadvantages of oral versus vaginal misoprostol are greater need for oxytocin augmentation (RR 1.28, 95% CI 1.11 to 1.48), and increased meconium staining of the amniotic fluid (RR 1.27, 95% CI 1.10 to 1.60).

The results of this review are therefore of limited practical importance: in dosages of 25 mcg three hourly or more, vaginal misoprostol is more effective than conventional methods of cervical ripening and labour induction. However, uterine hyperstimulation with FHR changes are increased. Although no differences in perinatal outcome were shown, the studies were not sufficiently large to exclude the possibility of uncommon serious adverse effects. The increase in meconium-stained liquor is also of concern. Anecdotal reports of uterine rupture following labour induction with misoprostol are cause for concern (Gherman 1999; Daisley 2000; Hill 2000; Majoko 2002b).

The limited information on lower dosage regimens (25 mcg four hourly or less) suggests that they may be as effective as other prostaglandins, without increased uterine hyperstimulation.

Though misoprostol shows promise as a highly effective, inexpensive and convenient agent for labour induction, the lack of registration for this purpose, and thus of well-established regimens, is problematic.

In most countries misoprostol is not registered for use for labour induction. In countries in which its use is considered advantageous, it is important that health authorities provide guidelines for practitioners to ensure the greatest possible level of safety in its use.

## Implications for research

While this review assessed the efficacy and safety of vaginal misoprostol (including its different regimens) based on data from its comparison to oral misoprostol, the vaginal route should not be researched further.

Because of the potential economic and clinical advantages of misoprostol, there is the need for further trials to establish its safety, particularly the relative safety of various dosages of oral administration. On the basis of this review, such trials should have the following features.

- (1) Randomised, double blind.
- (2) Oral or sublingual route of administration.
- (3) Sample size sufficient to detect moderate differences in important uncommon complications such as serious perinatal morbidity/mortality.
- (4) Meconium-stained liquor included as an outcome measure.
- (5) Women's views included as an outcome.

Randomised trials sufficiently large to assess rare events such as uterine rupture are not feasible. Alternative research methods are necessary such as case-control studies and prospective audits of complications in services in which misoprostol is used routinely for labour induction.

We would be grateful to receive reports of rare serious complications such as uterine rupture in order to compile a register of such incidents.

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\* *Indicates the major publication for the study*

## CHARACTERISTICS OF STUDIES

### Characteristics of included studies [ordered by study ID]

#### 006 Ewert 2006

Methods	Double-blinded randomised multicenter study. Randomisation and allocation concealment were computer generated.	
Participants	<p>Inclusion criteria: pregnant women were aged 18 years or older, 37 to 42 weeks of gestation, requiring cervical ripening and labour induction and at least 1 previous delivery with gestation of 37 weeks or more. The current pregnancy had to be a singleton fetus with cephalic presentation and the women had to be classified as a pregnant uncomplicated with Bishop score less than 6.</p> <p>Exclusion criteria: women with plus than 4 terms full deliveries and those with caesarean delivery. Spontaneous labour, tocolytic agents used within 7 days before induction, the use of any other cervical-ripening or labour-inducing agent before enrolment, suspected cephalopelvic disproportion, fetal distress, use of non steroidal anti-inflammatory drugs 4 hours before the drug studied treatment, pyrexia, unexplained genital bleeding after 24 weeks of this pregnancy, pelvic inflammatory disease, placenta previa and known or suspected allergy to misoprostol or other prostaglandins.</p>	
Interventions	A single misoprostol vaginal insert was administered high into the posterior fornix and positioned transversally behind the cervix. Each insert was loaded with 1 of the 4 dose reservoirs of misoprostol being investigated: 25, 50, 100 and 200 mcg. The vaginal insert was designed to release misoprostol at a rate of approximately 1, 2, 4 and 8 mcg per hour respectively.	
Outcomes	The median time to vaginal delivery was 27.5, 19.1, 13.1 and 10.6 hours for the 25-, 50-, 100- and 200-mcg doses, respectively. The percentage of women who delivered within 12 hours was 9%, 14%, 47% and 53% (P < .001 using the 25 mcg group as the comparator) and within 24 hours was 42%, 79%, 81% and 70%(P = .003). Uterine hyperstimulation syndrome occurred in 1 women who received the 25 mcg, 2 women who received the 100 mcg, and 3 women who received 200 mcg dose reservoirs.	
Notes	<p>This trial was conducted at 6 sites in the United Kingdom.</p> <p>At this review the comparison of doses were made grouping lower 2 doses vs higher 2 doses.</p>	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Unblinded only for the outcome assessors

**006 Ewert 2006** (Continued)

Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**013 Papanikolaou 2004**

Methods	Randomised unblinded trial. Allocation not stated.
Participants	Inclusion criteria: women with > 39 completed weeks singleton gestation. Exclusion criteria: previous uterine scar, malpresentation, cervix dilated > 3cm, uterine contraction > 3/10 minutes, any contraindication to vaginal delivery, hypersensitivity to prostaglandins, parity > 5, abnormal antepartum testing, cephalopelvic disproportion, premature rupture of membranes and maternal illnesses.
Interventions	The intervention group received 100 mcg 6 hourly and the comparison group received 50 mcg 4 hourly of vaginal misoprostol until labour.
Outcomes	In two groups the dose used of misoprostol were similar. There was no difference between two groups in mean time to delivery, caesarean rate, Apgar of 5 minutes and meconium passage.
Notes	Department of Obstetrics and Gynaecology, Isparta, Turkey.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	Yes	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate



### 013 Tedesco 2002

Methods	Single blind randomised trial with computer-generated allocation.	
Participants	Inclusion criteria: indication for induction of labour; intact membranes; live fetus; cephalic presentation; cervical score < 6; gestation 37+ weeks. Exclusion criteria: caesarean section or other uterine surgery; antepartum haemorrhage; pyrexia; anaemia; mental incapacity; contraindication to labour; fetal anomaly.	
Interventions	Vaginal misoprostol 12.5 vs 25 mcg in absorbable capsules, repeated once after 6 hours if not in labour.	
Outcomes	Time from first dose to delivery (12.5 mcg: 28.7 +/- 19.6 vs 25 mcg: 23.6 +/-10.3 hours); mode of delivery; cardiotocograph changes; meconium staining; Apgar scores; uterine hyperstimulation; analgesia.	
Notes	June 2000 to July 2001.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

### 019 Filho 2007

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Methods	Double-blinded randomised controlled clinical trial. Allocation by sequentially numbered sealed opaque envelopes.	
Participants	<p>Inclusion criteria: pregnant women with indication to induction of labour, singleton, gestational age from 37 to 42 weeks, reactive non-stress test, vertex presentation and no rupture of membranes or labour.</p> <p>Exclusion criteria: previous uterine scar, premature rupture of membranes, preterm birth, contraindication to vaginal delivery, any vaginal bleeding, doubts in the gestational age and known allergy to prostaglandins.</p>	
Interventions	The patients received 12.5 or 25 mcg vaginal misoprostol 4 hourly until effective labour.	
Outcomes	The two groups did not differ in the mean time from induction to delivery, in the frequency of vaginal delivery, Apgar score and tachysystole frequency.	
Notes	Maternity Sant'Anna of Santa Casa de Misericórdia de Sobral, Fortaleza, Ceara, Brazil.	
<b><i>Risk of bias</i></b>		
<b>Item</b>	<b>Authors' judgement</b>	<b>Description</b>
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate

**019 Filho 2007** (Continued)

Blinding? All outcomes	No	Unblinded only for the outcome assessors
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**025 Elhassan 2005a**

Methods	“An open randomised controlled clinical trial”. The sequence generation and allocation concealment were not stated.
Participants	Inclusion criteria: pregnant women with single babies, unripe cervixes and intact membranes. Exclusion criteria: previous uterine surgery, antepartum haemorrhage, asthma, heart disease and grand multiparity.
Interventions	The intervention group received 25 mcg of misoprostol 6 hourly up to 4 doses. The comparison group received 50 mcg of misoprostol 6 hourly up to 4 doses.
Outcomes	The induction-delivery interval was significantly longer in the 25 mcg group vs. 50 mcg group (21.9 h $\pm$ 4.3 h vs 9.6 h $\pm$ 2.2 h, p = 0.04). More women in the 25 mcg group received oxytocin (61.3% vs. 56.3% p > 0.05). Significantly fewer patients delivered vaginally in the 25 mcg group (61.3% vs. 90.6%, p = 0.05). There were no differences between the two groups at neonatal outcomes.
Notes	This study was conducted at the labour ward of Wad Medani Hospital at Sudan, from January to July 2004.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate

**025 Elhassan 2005a** (Continued)

Free of other bias?	Yes	Adequate
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**025 Haghighi 2006**

Methods	Randomised trial. Allocation concealment made by sequential sealed opaque envelope.
Participants	Inclusion criteria: women with singleton pregnancy, gestational age from 29 to 36 weeks, PPROM, parity of 2 or less, Bishop score $\leq 4$ , Normal FHR reactivity, absence of uterine contractions and vertex presentation. Exclusion criteria: previous uterine scar, fetal growth retardation, pre-eclampsia, evidence of cephalopelvic disproportion, chorioamnionitis and contraindication of prostaglandin treatment.
Interventions	The intervention group received 1 25 mcg dose of misoprostol. If the contractions were not adequate in 3 hours, oxytocin augmentation was started. The comparison group received oxytocin infusion (started with 2 mcU/min and increased every 15 minutes until adequate uterine contraction).
Outcomes	There was a statistically significant difference in the interval between admittance and vaginal delivery (507.68 min $\pm$ 248.01 min in the misoprostol group vs 596.66 min $\pm$ 246.38 min in the control group; $P < .005$ ) and also in the incidence of caesarean section due to failed labour induction (9.2% in the misoprostol group vs 18.5% in the control group).
Notes	Akbarabadi Hospital in Tehran, Iran. The study run from December 2002 to May 2004. The numbers of baseline data and the neonatal outcomes were not stated.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	No	Inadequate
Free of other bias?	No	Inadequate

## 025 Incerpi 2001

Methods	Misoprostol or placebo dispensed by pharmacy according to computer-generated randomisation schedule.
Participants	Inclusion criteria: women with pre-gestational or gestational diabetes; good control of blood glucose levels; singleton gestation; intact membranes; cervical score < 5; uterine contractions < 8/hour; cephalic presentation; gestational age 38.5 weeks or more; normal amniotic fluid index; reactive fetal heart rate pattern; good compliance. Exclusion criteria: multiple pregnancy; estimated fetal weight > 4500 or < 2000 g; ruptured membranes; placenta praevia; vaginal bleeding; active genital herpes; glaucoma; hypersensitivity to prostaglandins; renal, hepatic or cardiovascular disease; severe asthma; parity > 5.
Interventions	Misoprostol 25 mcg or placebo vaginally on day 1; if not in labour after 4 hours, discharged home; repeated on day 4 if enrolment criteria still met; if cervical score > 8, labour induced with amniotomy and oxytocin infusion; if not delivered by day 7, labour induced with amniotomy and oxytocin, or misoprostol 25 mcg 4-hourly.
Outcomes	Primary outcome: delivery within 7 days (misoprostol 31/57 vs placebo 36/63).
Notes	Los Angeles County - University of Southern California Women's and Children's Hospital and Good Samaritan Hospital, August 1996 to November 2000.

### *Risk of bias*

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

## 025 Krupa 2005

Methods	Open randomised controlled trial with allocation concealment using consecutively numbered sealed opaque envelopes.
Participants	Inclusion criteria: pregnant women with PROM confirmed up to 6 hours after occurrence, gestational age $\geq 37$ weeks, cephalic presentation and a live fetus showing no signs of fetal compromise as evaluated by cardiotography. Exclusion criteria: previous caesarean section or uterine surgery, being in labour at admission, presence of fetal malformation of incompatible with life, twin pregnancy or strongly suspected or confirmed chorioamnionitis.
Interventions	The intervention group received 25 mcg of misoprostol 6 hourly up to 4 doses. The comparison group received vital signs monitorisation during 24 hours. After 24 hours, if the women did not initiate labour, oxytocin was started.
Outcomes	The misoprostol group had a significantly shorter latency period (9.4 h vs 15.8 h), a shorter time interval from recruitment to delivery (18.9 h vs 27.5 h), a shorter period of maternal hospitalisation and a slightly higher proportion of alterations of contractility when compared to expectant group. Caesarean section was higher in the observational group. The complications were similar in the groups. Within 24 hours, 44% of the women had delivered in the expectant group against 73% in the misoprostol group.

**025 Krupa 2005** (Continued)

Notes	The study was conducted between January 2000 and May 2003. Faculty of Medical Sciences, Universidade Estadual de Campinas - UNICAMP, São Paulo, Brazil.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**025 Kumar 2001**

Methods	"Patients were randomly assigned..." Sequence generation and allocation concealment were not stated.
Participants	Inclusion criteria: obstetric indications of labour induction including hypertensive disorders of pregnancy, intrauterine growth retardation of fetus and postdatism; medical complications including diabetes and renal disease; no history of previous caesarean section or uterine surgery; absence of active labour or fetal distress and singleton pregnancy with vertex presentation and no contraindication of vaginal delivery.
Interventions	The intervention group received 25 mcg dose of misoprostol six hourly until adequate uterine contractions were achieved. The patients in active phase of labour with arrest of dilatation received oxytocin for augmentation. The comparison group received dinoprostone gel 0.5 mg intra cervically. If there was absence of uterine contraction after 10 hours of dinoprostone, the patient started to receive oxytocin infusion.
Outcomes	The average interval from start to induction of vaginal delivery was shorter in misoprostol group (1315 min $\pm$ 811min) compared to dinoprostone/oxytocin group (1512 min $\pm$ 712 min) ( $p < 0.01$ ). There were no significant difference in route of delivery, incidence of uterine hyperstimulation and perinatal outcome.
Notes	The study was conducted in INHS Asvini from January 1999 to January 2000. Colba, Mubai.

**025 Kumar 2001** (Continued)

<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unblinded for personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**025 Majoko 2001**

Methods	“Non blinded randomized controlled trial”, did not provide information on allocation methods	
Participants	Women admitted for induction of labour, with singleton pregnancy and cephalic presentation	
Interventions	Vaginal misoprostol 50 mcg versus prostaglandin F2 alpha 2 times 8 hourly if needed.	
Outcomes	Need of oxytocin, caesarian section rate and delivery interval.	
Notes	Only abstract available. This trial did not contribute with any data because were provided only percentages.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	Unclear	Unclear

**025 Majoko 2001** (Continued)

Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**025 Majoko 2002c**

Methods	Randomised controlled trial. Randomisation by computer generated table and allocation recorded on a card placed in a sealed opaque envelope .
Participants	Inclusion criteria: women for IOL after 37 weeks with singleton pregnancy in cephalic presentation. Exclusion criteria: previous uterine surgery, abnormal FHR pattern, contra-indication to vaginal delivery and known sensitivity to prostaglandin or misoprostol.
Interventions	Intervention group received vaginal misoprostol 50 mcg at maximum of 2 doses each 8 hours. There were 3 comparison groups: 1. Suspension of oral misoprostol 4 hourly. 2. Prostaglandin F2 $\alpha$ gel 5 mg intra cervically in 2 doses (8 hours interval) if the Foley was well located. 3. Prostaglandin E2 pessary 3 mg, vaginally repeated after 8 hours if necessary.
Outcomes	There was no difference in mode of delivery. There was significantly reduced risk of caesarean section in the intervention group (OR 0.20; 95% CI 0.22 to 0.78). The need of oxytocin augmentation was reduced in the group of prostaglandin E2 pessary, vaginal misoprostol and oral misoprostol. The vaginal misoprostol reduced the interval from induction to delivery, but the 2 misoprostol groups increased the admissions at the neonatal unit.
Notes	Harare Maternity Hospital, Zimbabwe. The vaginal misoprostol was compared with the vaginal prostaglandin. The other groups were evaluated in the specific review.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	Yes	Inadequate

**025 Majoko 2002c** (Continued)

Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**025 McKenna 2004**

Methods	Randomised, double blinded, placebo-controlled trial. Allocation by a centre of randomisation.
Participants	Inclusion criteria: pregnant women older than 18 years old, at list 40 weeks of pregnancy. Bishop < 9. Well dated pregnancy. Exclusion criteria: oligohydramnios, rupture of membranes, malpresentation, vaginal bleeding, multiple gestation, labour and history of prior caesarean section.
Interventions	The intervention group received a 1 time 25 mcg intravaginal. The comparison group received an inert substance as placebo, both in gel capsules.
Outcomes	The mean interval to delivery was significantly lower in the misoprostol group, 4.2 days $\pm$ 4.1 days compared with 6.1 days $\pm$ 3.6 days, $P = .04$ after receiving the insert. The interval to delivery was significantly less in the misoprostol group only for nulliparous women. The survival curves for the interval from intervention to delivery were significantly different ( $P = .04$ ); for misoprostol the median interval was 4.1 days to delivery compared with 9.2 days for placebo.
Notes	Wright-Paterson US Air Force Base Medical Center. 1 patient of the intervention group and 3 of the comparison group were excluded from the trial after randomisation. They did not received the medication of the study.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Yes	Adequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate



**025 Meyer 2005**

Methods	Randomised unblinded controlled trial. Allocation by sequential numbered opaque envelope.
Participants	Inclusion criteria: gestation at term, vertex presentation, Bishop score 6, singleton pregnancies, intact membranes, cephalic presentation and non-stress test previous. Exclusion criteria: Bishop score > 6, labour, any uterine scar and rupture of membranes.
Interventions	The intervention group received 25 mcg of vaginal misoprostol single dose. The comparison group received an intra cervically dose of 0.5 mg of dinoprostone.
Outcomes	Single dose of misoprostol significantly decreased the cumulative dose of oxytocin, the cumulative dose of time of oxytocin administration, and the dose intensity of time oxytocin (dose divided by time).
Notes	The study was conducted from November 1999 to December 2001 at the University of Vermont.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Yes	Inadequate only for personnel
Incomplete outcome data addressed? All outcomes	Unclear	Unclear
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**025 Oboro 2005**

Methods	Randomised controlled trial. Allocation by card drawn from a consecutive series of sealed envelopes.
Participants	Inclusion criteria: women with singleton gestation in cephalic presentation, Bishop score $\leq 8$ , 40 weeks of gestational age. Exclusion criteria: patients with rupture of membranes, vaginal bleeding, a prior uterine incision, non-reactive non-stress test and an estimated fetal weight of > 4500g.
Interventions	The intervention group received one dose of 25 mcg misoprostol vaginally and the comparison group received no treatment.

**025 Oboro 2005** (Continued)

Outcomes	Misoprostol was associated with significant decrease of time to delivery, earlier gestational age at delivery, shorter duration of active labour, without changes at the neonatal outcome.
Notes	The trial was conducted from August 1, 2000 to October 31, 2001 at Zonal General Hospital, Kwale, of Delta state of Nigeria.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	Unclear	Unclear
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**025 Sheela 2007**

Methods	"A total of 150 women were randomised". The sequence generation and the allocation concealment were not stated.
Participants	Inclusion criteria: all women at term, singleton pregnancies, live fetus, cephalic presentation, obstetric or medical indication for induction of labour, intact membranes, Bishop score $\leq 4$ , and reassuring non-stress test. Exclusion criteria: grand multiparas, previous caesarean section, other uterine scar, medical contraindication for prostaglandins and those with maternal or fetal compromise.
Interventions	One group received 0.5 mg intracervical dinoprostone 12 hourly (maximum 3 doses) and the other two groups received 25 or 50 mcg vaginal misoprostol 6 hourly (maximum 5 doses). The patients could receive oxytocin augmentation.
Outcomes	There were no differences between the groups in mode of delivery, neonatal outcomes, caesarean section rate, failed induction, failed induction and hyperstimulation. The women from the misoprostol group required fewer oxytocin augmentation and present shorter interval from induction to delivery.
Notes	Data from oral misoprostol were accessed in a specific review. Department of Obstetrics and Gynaecology, St. John's Medical College Hospital, Karnataka, India.

025 Sheela 2007 (Continued)

<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Unclear	Unclear

025 Stitely 2000

Methods	Double-blind, placebo-controlled in computer generated sequence, permuted block design. Medications prepared independently by pharmacy.
Participants	Inclusion criteria: gestational age 41 weeks to 41 weeks 6 days; singleton pregnancy; intact membranes; cervical score 4 or less; uterine contractions < 8 per hour; amniotic fluid index > 5 cm; reactive cardiotocography; maternal age 18 to 50 years. Exclusion criteria: fetal malpresentation; estimated fetal weight > 4500 g or < 2000 g; placenta praevia; vasa praevia; unexplained vaginal bleeding; active herpes simplex; hypersensitivity to prostaglandins; prior uterine surgery or caesarean section; evidence of intra-amniotic infection; severe asthma or cardiovascular disease; renal or hepatic dysfunction.
Interventions	1. Misoprostol 25 mcg (1/4 tablet). 2. Placebo (dicalcium phosphate 1/5 tablet). Medication placed in posterior vaginal fornix. FHR and uterine activity monitored continuously for 4 hours, then discharged. Repeated after 24 hours if cervical score < 9 and other original criteria unchanged. After a further 24 hours, labour induced with oxytocin if cervical score > 6, pre-induction vaginal misoprostol 25 mcg if < 6.
Outcomes	Primary: number of inpatient inductions needed by study day 3 (3/27 vs 28/33). Secondary: mode of delivery; Apgar score < 7 at 5 minutes; meconium passage; neonatal ICU admission; number of medication doses (1.41 sem 0.1 vs 1.85 sem 0.1); dosing required on study day 2 (15/27 vs 30/33); change in cervical score day 1-2 (2.7 sem 0.8 vs 0.96 sem 0.008); first dose to delivery (36.9 sem 3.8 vs 61.3 sem 3.8 hours).
Notes	Naval Medical Center, Portsmouth VA and National Naval Medical Center, Bethesda, MD; December 7 1997 to December 10 1999.

*Risk of bias*

**025 Stitely 2000** (Continued)

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**025 Wang 1998**

Methods	This study described as random allocation but there were no details in methods.	
Participants	Term pregnant women, singleton pregnancy, vertex presentation and intact membranes. There were no exclusion criteria.	
Interventions	Patients received 50 mcg or 25 mcg vaginal misoprostol.	
Outcomes	The study did not show differences between these 2 dosages of misoprostol.	
Notes	The study was enrolled from November 96 to November 97.	

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Yes	Adequate

**025 Wing 1996**

Methods	Allocation by sequentially numbered sealed opaque envelopes, maintained by the primary investigator. Sequence from a computerised random number generator.	
Participants	Membranes were intact, cervical status unfavourable. Inclusion criteria: women requiring induction of labour for medical or obstetric reasons; singleton pregnancy; cephalic presentation; intact membranes; cervical score < 5; reactive nonstress test; < 8 uterine contractions per hour. Exclusion criteria: estimated fetal weight > 4500 g; evidence of cephalopelvic disproportion; placenta praevia; unexplained vaginal bleeding; vasa praevia; active herpes simplex infection; contraindication to receiving prostaglandins; renal or hepatic dysfunction; suspected chorioamnionitis; previous uterine	

**025 Wing 1996** (Continued)

	<p>surgery; parity &gt; 5.</p> <p>Indications for labour induction: oligohydramnios (6-hourly misoprostol 131, 3-hourly 126); pre-eclampsia (47, 43); post-term pregnancy (32, 35); macrosomia (8, 17); diabetes mellitus (17, 9); abnormal antepartum testing (12, 13); other (18, 12).</p>
Interventions	<p>Misoprostol 25 mcg inserted into the posterior vaginal fornix 6-hourly (n = 259) versus 3-hourly (n = 261), until 3 uterine contractions per 10 minutes, cervical score &gt; 7, cervical dilation &gt; 3 cm or spontaneous rupture of membranes (maximum 24 hours). Artificial rupture of membranes usually performed when the cervix was 80% effaced and 3 cm dilated, or 4 cm dilated. Oxytocin augmentation was used for lack of contractions after maximum dosage or spontaneous rupture of membranes, or for arrested cervical dilation, &gt; 3 hours after the last misoprostol dose.</p> <p>Uterine hyperstimulation was treated in some cases by tocolytic therapy.</p>
Outcomes	<p>Oxytocin augmentation; uterine tachysystole (&gt; 5 contractions per 10 minutes); uterine hypertonus (contraction &gt; 2 minutes) (6-hourly misoprostol 12/259, 3-hourly 11/261); hyperstimulation syndrome (tachysystole or hypertonus with fetal heart rate abnormalities); induction to delivery interval; mode of delivery; vaginal delivery within 24 hours; terbutaline used for hyperstimulation (18/259, 26/261); neonatal resuscitation (83/259, 90/261); days in NICU (mean 9.2 (SD 8.6), 9.8 (12.8)); meconium aspiration syndrome (3/259, 2/261); hyperbilirubinaemia (14/259, 16/261).</p> <p>Tachysystole occurred after the first dose of misoprostol in 43/520 (8.3%) of women.</p> <p>One maternal death occurred in a primiparous woman, 9 hours after a single misoprostol dose and shortly after amnioinfusion and epidural analgesia, from amniotic fluid embolization. Two caesarean hysterectomies were performed for atonic uterine haemorrhage 13 and 30 hours after single doses of misoprostol, in one primiparous woman with uncomplicated labour, and one nulliparous woman following prolonged induction attempt with oxytocin augmentation and chorioamnionitis.</p>
Notes	<p>Los Angeles, California, USA. October 1994 to June 1995.</p> <p>522/535 women agreed to participate. No women withdrew from the protocol. Two women allocated to misoprostol 6-hourly were excluded from the analysis because of deviation from the study protocol.</p>

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**025 Wing 1998a**

Methods	Randomised clinical trial with allocation concealment by consecutive numbered sealed opaque envelopes.
Participants	<p>Inclusion criteria: women with spontaneous ruptured membranes, singleton pregnancies with cephalic presentation, reactive FHR pattern beyond 36 weeks' gestational age, not in labour.</p> <p>Exclusion criteria: women with cervical dilatation in &gt; 3cm, in labour, estimated fetal weight &gt; 4500g or evidence of cephalopelvic disproportion, estimated fetal weight &lt; 1800g, placenta previa, unexplained vaginal bleeding, active herpes simplex, scared uterus, clinical signal of maternal infection, parity &gt; 5, moderate or severe pre-existing</p>

**025 Wing 1998a** (Continued)

	medical disease or any contraindication for use of prostaglandin.
Interventions	1. Misoprostol 25 mcg placed in the posterior vaginal fornix, repeated after 6 hours if necessary, n = 98. 2. Intravenous incremental oxytocin infusion to maximum dose of 22 mU/minute, n = 99.
Outcomes	Induction to delivery interval (misoprostol 811.5 +/- 511.4 vs oxytocin 747.0 +/- 448.0 minutes, p = 0.65); caesarean section; abnormal FHR tracing (29/98 vs 29/99); chorioamnionitis (28/98 vs 26/99); meconium-stained liquor; Apgar score < 7 at 1 and 5 minutes; neonatal resuscitation (24/98 vs 27/99); neonatal ICU admission.
Notes	3 withdrawals not accounted for.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**025 Wing 1998b**

Methods	Women 'assigned by means of a computerised random number generator' using numbered, sealed opaque envelopes.
Participants	38 of 40 women requiring induction of labour for medical or obstetric indications with one immediate prior caesarean delivery agreed to participate. Inclusion criteria: singleton pregnancy; cephalic presentation; intact membranes; cervical score < 6; reactive nonstress test; < 8 uterine contractions per hour. Exclusion criteria: estimated fetal weight > 4500g; evidence of cephalopelvic disproportion; placenta praevia; unexplained vaginal bleeding; vasa praevia; active herpes simplex infection; contraindication to receiving prostaglandins; renal or hepatic dysfunction; suspected chorioamnionitis; previous uterine surgery; parity > 5, prior classical caesarean section.

**025 Wing 1998b** (Continued)

Interventions	1. Misoprostol 25 mcg vaginally every 6 hours until 3 contractions per 10 minutes, cervical score 8 or more, or cervix 3 cm dilated (maximum 4 doses). Amniotomy and oxytocin augmentation used when necessary (n = 17). 2. Intravenous oxytocin 1 mU per minute, increased every 30 minutes of necessary (maximum 22 mU per minute) (n = 21). Management included active amniotomy and continuous fetal heart rate and uterine activity monitoring.
Outcomes	Vaginal delivery within 24 hours; oxytocin augmentation; route of delivery; uterine dehiscence (asymptomatic uterine scar disruption); uterine rupture (separation of uterine scar requiring emergency laparotomy).
Notes	October 28, 1995 to November 18, 1996. Women's Hospital, Los Angeles, USA. Calculated sample size 160. Trial terminated prematurely because of disruption of uterine scar in two women who received misoprostol. Further outcome details received on request from authors. One woman withdrew from protocol requesting caesarean section (misoprostol group) and included in the analysis.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**025G Srisomboon 1998**

Methods	Allocated by 'block randomisation'.
Participants	Pregnant women eligible for labour induction. Inclusion criteria: singleton pregnancy; parity < 4; vertex presentation; obstetric or medical indication for labour induction; intact membranes; cervical score < 5; gestational age > 35 weeks. Exclusion criteria: labour; fetal distress; previous caesarian delivery or other uterine surgery; definite cephalopelvic disproportion; contraindication to the use of prostaglandins.
Interventions	Vaginal misoprostol 25 mcg versus 50 mcg in carboxymethylcellulose gel, 6-hourly until adequate contraction, cervical score > 6, cervical dilation > 3, or spontaneous rupture of membranes (maximum 4 doses). If cervix favourable, amniotomy performed and oxytocin infused if necessary. Continuous cardiotocography was used.
Outcomes	Tachysystole (> 5 contractions per 10 minutes; hypertonus (contraction > 90 seconds); hyperstimulation syndrome.
Notes	November 1995 to May 1996, Chiang Mai University Hospital, Thailand.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**030 Moodley 2003**

Methods	Allocation by sequentially numbered opaque envelopes with computer-generated sequence. Not blinded.
Participants	Inclusion criteria: viable term or near-term pregnancy; indication for induction of labour. Exclusion criteria: previous caesarean section; malpresentation; non-reassuring fetal heart rate pattern; cervical score > 5; parity > 4.
Interventions	Misoprostol 25 mcg into posterior vaginal fornix; if cervix unchanged and not in labour after 4 hours, oral misoprostol solution (200 mcg in 200 ml water), 20 mcg 2-hourly until adequate contractions, up to 3 doses; versus dinoprostone 1 mg into the posterior vaginal fornix 6-hourly for up to 3 doses if contractions inadequate.
Outcomes	Delivery within 24 hours; induction to delivery time; meconium staining of liquor; non-reassuring fetal heart rate pattern; hyperstimulation syndrome (hypersystole and/or tachysystole with abnormal fetal heart rate pattern); mode of delivery; perinatal mortality; average dose requirements.
Notes	King Edward VIII Hospital, Durban, South Africa. A third arm (oral misoprostol) is for consideration in the oral misoprostol for labour induction review.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**038 Aquino 2003**

Methods	This was a randomised, by computer generation, controlled clinical trial with a sequential numbered envelope used for allocation.
Participants	Inclusion criteria: patients with medical indication for induction of labour, singleton gestation, gestational age greater than 36 weeks, vertex presentation, intact membranes, Bishop score < 6, no labour occurring and normal fetal heart rate. Exclusion criteria: pelvic dystocia, estimate fetal weight greater than 4 kg, evidence of cephalopelvic disproportion, placenta previa, any unexplained vaginal bleeding, parity > 5, fetal malformation, previous uterine scar, any situation when vaginal delivery was not indicated or contraindication to use misoprostol.
Interventions	The intervention group used 25 mcg of vaginal misoprostol 4 hourly (maximum 200 mcg) and the comparison group received oxytocin alone.
Outcomes	The caesarean section rate and the time to induction from delivery was significantly shorter in the misoprostol group. Tachysystole was more frequent in the intervention group. There were no differences in the neonatal outcomes between the groups.
Notes	Maternity Hospital Leonor Mendes de Barros, São Paulo, Brazil, from November 1998 to December 2000.

***Risk of bias***



**038 Aquino 2003** (Continued)

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear for personnel, but blinded for participants and outcome assessors
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**038 Cecatti 2000**

Methods	'Randomised controlled trial'.	
Participants	Term pregnant women with indication for labour induction; intact membranes.	
Interventions	Vaginal misoprostol 25 mcg 4-hourly (maximum 8 doses versus oxytocin infusion.	
Outcomes	Latent period; induction to vaginal delivery time; route of delivery; hyperstimulation syndrome.	
Notes	Hospital-Maternity Leonor Mendes de Barros, Sao Paulo, Brazil. November 1998 to August 1999.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**038 Clark 1998**

Methods	Method of women being 'randomised' not described.	
Participants	Inclusion criteria: obstetric or medical indication for labour induction; singleton pregnancy; vertex presentation; reassuring fetal status; cervical score $\geq 5$ . Exclusion criteria: abnormal FHR pattern; placenta praevia; active herpes infection; history of asthma, glaucoma, cardiac or hepatic disease; chorioamnionitis; previous uterine scar; parity $> 5$ .	
Interventions	1. Misoprostol 25 µg to the posterior vaginal fornix every 4 hours (maximum 4 doses) until adequate contraction pattern or active labour, vs 2. dinoprostone gel 0.5 mg intracervically every 4 hours (maximum 4 doses) until adequate contractions	

**038 Clark 1998** (Continued)

	or cervical score > 5. Oxytocin used if labour not established after 4 doses, or labour protracted. Continuous FHR and uterine activity monitoring were used.	
Outcomes	Oxytocin use; induction to delivery interval (misoprostol 1181 +/- 566 vs dinoprostone 1403 +/- 566 minutes, p = 0.02); caesarean section (15% vs 31%); hyperstimulation syndrome.	
Notes	Caesarean section (percentages only given as E1 15% and E2 31%. However previous sentence and relative risk figure given correspond to 31% for E1, 15% for E2).	
<i><b>Risk of bias</b></i>		
<b>Item</b>	<b>Authors' judgement</b>	<b>Description</b>
Allocation concealment?	No	Inadequate

**038 El-Sherbiny 2001**

Methods	Randomised using random number tables.	
Participants	Inclusion criteria: singleton; live fetus; 37 or more weeks' gestation; cephalic presentation; intact membranes. Exclusion criteria: parity > 5; previous uterine surgery.	
Interventions	Vaginal misoprostol gel 25 vs 50 mcg 4-hourly (maximum 6 doses). Prepared by mixing one 200 mcg tablet with 8 or 4 ml hydroxyethyl gel.	
Outcomes	Tachysystole; hypersystole; uterine hyperstimulation; oxytocin used; induction-delivery time; caesarean section; meconium-stained liquor; neonatal outcomes.	
Notes	El-Sherbiny Hospital and El-Salaam General hospital, Egypt, May 1997 to April 1999.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**038 Eroglu 2007**

Methods	Randomised trial with a sequence generation not stated. The allocation concealment was made by sequential envelopes pulled by hospital residents.	
Participants	Inclusion criteria: women with gestational age $\geq$ 37 weeks, singleton pregnancy, Bishop score $\geq$ 6, vertex presentation, no contraindication to vaginal delivery, and obstetric indication for induction of labour. Exclusion criteria: Bishop score > 6, prior caesarean delivery, placenta previa, unexplained vaginal bleeding and contraindication or allergic reaction to the use of prostaglandins.	

**038 Eroglu 2007** (Continued)

Interventions	The study groups used 50 mcg or 25 mcg of vaginal misoprostol 4 hourly (maximum 6 doses). The amniotomy was performed after 2 - 3 hours without labour progress. The patients could receive oxytocin augmentation.
Outcomes	The mean time from induction to delivery was shorter in the 50 mcg group. The use of oxytocin was smaller in the 50 mcg group. The incidence of tachysystole was higher in the 50 mcg group. There were no differences between the two groups in the caesarean section rate and neonatal outcomes.
Notes	The study was conducted from June 2004 and March 2006. Department of Obstetrics and Gynaecology, Baskent University, Ankara, Turkey.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	No	Inadequate
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**038 Gregson 2005**

Methods	Randomised controlled trial. Allocation concealment was made by consecutively numbered, sealed, opaque envelope.
Participants	Inclusion criteria: singleton pregnancy women, cephalic presentation, gestational age at term (37-42 weeks), parity < 4 times and no contraindication to prostaglandin. Exclusion criteria: significant maternal or fetal medical condition, previous uterine surgery and significant uterine activity.
Interventions	The intervention group received 25 mcg dose of misoprostol 4 hourly until 6 doses. The comparison group received dinoprostone vaginal gel 1-2 mg dose 6 hourly repeated at maximum of 3 mg in 24 hours.
Outcomes	Primary: induction to vaginal delivery interval. Secondary: requirements of oxytocin, mode of delivery, number of women delivering < 24 hours, incidence of uterine contrac-

**038 Gregson 2005** (Continued)

	tion abnormalities, incidence of abnormal cardiotocography recordings, 5-minute Apgar scores, umbilical cord pH recordings, analgesia requirements, admission to NICU and blood loss at delivery.	
Notes	The study took place at Queen Mary's Sidcup NHS trust between 18th July and 2nd December 2003. Florence Nightingale School of Nursing and Midwifery, King's College, UK.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear for participants, but blinded for personel and outcome assessors
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**038 Has 2002**

Methods	Tablets in bottles 'A' and 'B' prepared by the pharmacist. Placed in paper packets according to a table, stratified for age and parity. The first allocation in each group by lots, then alternation.
Participants	Inclusion criteria: singleton pregnancy; 37 or more weeks; cervical score 4 or less; vertex presentation; estimated fetal weight 4500 g or more; four or fewer contractions per hour. Exclusion criteria: prior uterine surgery; contraindication to prostaglandins; poor fetal surveillance scores; active genital herpes infection; placenta praevia; abruptio placentae; vasa praevia; unexplained vaginal bleeding.
Interventions	25 mcg versus 50 mcg in tablets reconstituted by the pharmacy, 4-hourly (maximum 6 doses). Oxytocin infusion if not in labour 4 hours after the third dose.
Outcomes	Primary: caesarean section; oxytocin use; tachysystole (6 or more contractions per 10 minutes, for 20 minutes); hypersystole (a contraction lasting 2 minutes); hyperstimulation (tachysystole or hypersystole and fetal heart rate abnormality); fetal distress requiring delivery. Secondary: caesarean section for fetal distress (low dose 6/58 vs high dose 16/56); umbilical artery pH < 7 (0/58 vs 2/56).
Notes	June 1998 to November 2000, Istanbul Medical School, Turkey.
<b><i>Risk of bias</i></b>	

**038 Has 2002** (Continued)

Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

**038 Kidanto 2006**

Methods	Randomised trial. Allocation concealment made by sealed opaque envelope with sequence generation by computer.
Participants	Inclusion criteria: pregnant women with singleton vertex presentation and gestational age > 36 weeks. Exclusion criteria: previous miomectomy, uteroplasty and caesarean section.
Interventions	The intervention group received 25 mcg of misoprostol 4 hourly until four doses. The comparison group received oxytocin infusion and amniotomy.
Outcomes	The interval from induction to delivery was shorter in the misoprostol group (10.86 h) compared to oxytocin group (15.45 h). There were fewer caesarean sections and neonatal ward in the misoprostol group. The induction with misoprostol was effective, safe and cheaper than oxytocin for IOL.
Notes	Muhimbili National Hospital, Dar es Salaam Tanzania. The study was performed between June to December of 2004.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Unclear for participants and for outcome assessors. No blinding for personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Unclear	Unclear

**038 Krithika 2008**

Methods	“Prospective randomized study” with no reference for allocation concealment methods.
Participants	Inclusion criteria: women with singleton pregnancy at more than 34 weeks with Bishop score less than 4 admitted for induction of labour. Exclusion criteria: scared uterus, active medical disorder, antepartum haemorrhage, abnormal fetal heart rate pattern, contracted pelvis, cephalopelvic disproportion, suspected chorioamnionitis and known uterine abnormalities.
Interventions	The intervention group received vaginal misoprostol 25 mcg 4 hourly until 6 doses. The comparison group received 0.5 mg of endocervical prostaglandin E2 and if necessary a second dose 12 hours later.
Outcomes	Induction to delivery interval was significantly shorter in the misoprostol group. The rate of complications were comparable in between groups.
Notes	A total of 100 women were enrolled in this trial. Study developed in the Department of Obstetrics and Gynaecology, Postgraduate Institute of Medical Education and Research, Chandigarh, India.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**038 Meydanli 2003**

Methods	Double-blind study. Random allocation by sealed consecutively numbered opaque envelopes in computer-generated random sequence in varying blocks.
Participants	Inclusion criteria: singleton live gestation; vertex presentation; intact membranes; gestational age > 41 weeks; cervical score < 5; < 4 spontaneous uterine contractions per hour; estimated fetal weight < 4500 g; reactive cardiotocography. Exclusion criteria: hypersensitivity to prostaglandins; previous caesarean section or uterine surgery; body mass index $\geq$ 30 before pregnancy; previous labour induction attempt in the current pregnancy; hypertensive disorders of pregnancy; fetal growth restriction; diabetes.

**038 Meydanli 2003** (Continued)

Interventions	Misoprostol 25 mcg (1/4 100 mcg tablet) compared with 50 mcg (1/4 200 mcg tablet). Labour induction with misoprostol 4-hourly placed in the posterior vaginal fornix, up to 6 doses. Vaginal examination and cardiotocography prior to each dose. No further misoprostol when uterine contractions of 3 per 10 minutes. When in labour or cervical score 8 or more, artificial rupture of membranes. Oxytocin used for poor progress more than 4 hours after last dose of misoprostol. No epidural analgesia or intravenous sedation used.	
Outcomes	Time from first dose to delivery; delivery with 12 hours and 24 hours; mode of delivery; emergency caesarean section for fetal hear rate abnormality; misoprostol doses, oxytocin requirement; tachysystole (> 5 contractions per 10 minutes for 20 minutes; uterine hyperstimulation (tachysystole with fetal tachycardia, late decelerations or reduced variability); birthweight; meconium-stained amniotic fluid; arterial cord pH < 7.16; neonatal intensive care unit admission.	
Notes	July 2001 to June 2002; two tertiary training centres in Turkey.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**038 Murthy 2006**

Methods	“The women enrolled into the study were randomized”. The sequence generation was not stated and the allocation was made by sealed envelopes.	
Participants	Inclusion criteria: singleton gestation, gestational age between 37 - 42 weeks, live uterine fetus, intact membranes, cephalic presentation and Bishop score $\leq 5$ . Exclusion criteria: multiple pregnancy, malpresentation, abnormal fetal heart rate pattern, cephalopelvic disproportion, rupture of membranes, previous caesarean section, parity more than 5 and history of hypersensitivity to prostaglandins.	
Interventions	The intervention group received 25mcg vaginal misoprostol 4 hourly (maximum 200 mcg/patient). The comparison group used 0.5 mg intracervical dinoprostone associated with oxytocin after 6 hours.	
Outcomes	The intervention group has shorter interval from induction to delivery. Fetal distress was more common in the intervention group. Neonatal outcomes were similar. The cost of the therapy was significantly less in the misoprostol group.	
Notes	The study was from December 2003 to May 2004.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description
Adequate sequence generation?	Unclear	Unclear

**038 Murthy 2006** (Continued)

Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**038 Van Gemund 2004**

Methods	Randomised multicentre trial using a random number table. The allocation was by sequentially numbered, opaque and sealed envelope.
Participants	Inclusion criteria: singleton pregnancy, cephalic or breech presentation, gestational age at least 36 completed weeks and Bishop score < 6. Exclusion criteria: contraindications to use prostaglandins, previous caesarean section or uterine incision and no informed consent.
Interventions	The intervention group used 25 mcg of vaginal misoprostol 4 hourly (maximum 3 doses per day until 2 days). The comparison group used 1 mg of vaginal dinoprostone 4 hourly until 3 doses per day.
Outcomes	The median induction-delivery interval was longer in the misoprostol group compared to the dinoprostone group (25 versus 19 hours , P = 0.008). Significantly fewer neonates were admitted at the NICU in the group of misoprostol intervention.
Notes	The study was carried out between 1 November 1999 and 31 December 2002 in the Netherlands.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate



038 Van Gemund 2004 (Continued)

Free of other bias?	Yes	Adequate
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038 Wing 1997

Methods	Allocation by sequentially numbered sealed opaque envelopes, maintained by the primary investigator. Sequence from a computerised random number generator.
Participants	<p>Membranes were intact, cervical status unfavourable. Inclusion criteria: women requiring induction of labour for medical or obstetric reasons; singleton pregnancy; cephalic presentation; intact membranes; cervical score &lt; 5; reactive fetal heart rate pattern; &lt; 8 uterine contractions per hour.</p> <p>Exclusion criteria: abnormal fetal heart rate pattern; malpresentation; estimated fetal weight &gt; 4500g; suspected cephalopelvic disproportion; ruptured membranes; placenta praevia; unexplained vaginal bleeding; vasa praevia; active herpes simplex infection; contraindication to receiving prostaglandins; renal or hepatic dysfunction; suspected chorioamnionitis; previous uterine surgery; parity &gt; 5.</p> <p>Indications for labour induction: oligohydramnios (misoprostol 46, dinoprostone 39); pre-eclampsia (21, 28); post-term pregnancy (16, 10); macrosomia (4, 8); diabetes mellitus (7, 8); impaired fetal growth (4, 3); other (1, 2).</p>
Interventions	<p>Misoprostol 25 mcg inserted into the posterior vaginal fornix 4-hourly, maximum 6 doses or 24 hours (n = 99) versus dinoprostone 10 mg vaginal insert, delivering 0.3 mg per hour (n = 98), until 3 uterine contractions per 10 minutes, cervical score &gt; 7, cervical dilation &gt; 2 cm or spontaneous rupture of membranes. The dinoprostone insert was removed if the above occurred or if there were uterine contraction or fetal heart rate abnormalities, or 24 hours had elapsed. Artificial rupture of membranes usually performed when the cervix was 80% effaced and 3 cm dilated, or 4 cm dilated. Oxytocin augmentation was used for lack of contractions after maximum dosage or spontaneous rupture of membranes, or for arrested cervical dilation, &gt; 2 hours after last misoprostol dose or dinoprostone removal.</p> <p>Continuous fetal heart rate and uterine activity monitoring was used.</p> <p>Uterine hyperstimulation was treated in some cases with terbutaline 250 mcg intravenously or subcutaneously.</p>
Outcomes	<p>Oxytocin augmentation; uterine tachysystole (&gt; 5 contractions per 10 minutes); uterine hypertonus (contraction &gt; 2 minutes); hyperstimulation syndrome (tachysystole or hypertonus with fetal heart rate abnormalities); abnormal fetal heart rate patterns (23/99 vs 35/98); induction to delivery interval (misoprostol mean 1429, SD 793 vs dinoprostone 1484, 866 minutes); mode of delivery; vaginal delivery within 24 hours; terbutaline used for hyperstimulation (3/99 vs 3/98); neonatal resuscitation (29/99 vs 25/98); days in NICU (mean 7.4, SD 5.1 vs 10.0, 13.4); meconium aspiration syndrome (1/99 vs 1/98); hyperbilirubinaemia (13/99 vs 7/98).</p> <p>Frequent uterine contractions occurred in 9 women after a single misoprostol dose, the onset occurring after mean 5.1 (SD 2.4) hours, and 11 after a second dose.</p>
Notes	<p>Los Angeles, California, USA. October 1 1995 to June 1996.</p> <p>200/206 women agreed to participate. No women withdrew from the protocol. 3/101 in the dinoprostone group were excluded from the data analysis because of deviation from the study protocol. In the abstract report all 200 women are included in the analysis, and the induction delivery times are somewhat different.</p>

*Risk of bias*

038 Wing 1997 (Continued)

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

043 Diro 1999

Methods	Double-blind study. Quartered 100 or 200 mcg misoprostol tablets were supplied by the research pharmacy according to random number tables.
Participants	Women undergoing induction of labour for maternal or fetal reasons. Inclusion criteria: cervical score < 5. Exclusion criteria: cardiopulmonary disease; HIV infection; malpresentation; multiple gestation; placenta praevia; previous uterine incision; 2 or more regular contractions in 10 minutes; sickle cell disease.
Interventions	Misoprostol 25 versus 50 mcg to posterior vaginal fornix 3-hourly till the onset of labour (maximum 8 doses). Membranes ruptured as soon as safe and feasible. Women not in labour after 24 hours received PGE2 gel intravaginally or oxytocin infusion.
Outcomes	Induction to onset of active labour; induction to delivery (low dose 1194 SD 785 vs high dose 933 SD 555 minutes); duration of first and second stages of labour; oxytocin augmentation and total dose; tachysystole, hyperstimulation; mode of delivery; perinatal outcome.
Notes	University of Miami/Jackson Memorial Centre, USA, September 1995 to April 1997. Of 284 women randomised, 33 withdrawn: 19 because of increased cervical score between randomisation and start of study; 12 randomised in violation of exclusion criteria; and 2 because of missing data (delivered vaginally).

*Risk of bias*

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

043 Farah 1997

Methods	Allocation by computer generated randomisation table. Pharmacy supplied one-fourth of a 200 µg or 100 µg tablet, which appeared identical.
Participants	Of 430 women enrolled, 29 excluded after randomisation and 2 after administration of misoprostol, all because enrolment criteria were not met. Inclusion criteria: indication for induction of labour; cervical score < 5; singleton third trimester pregnancy; vertex presentation. Exclusion criteria: active labour; fetal distress; previous uterine surgery; contraindication to vaginal delivery; contraindication to use of prostaglandins.

**043 Farah 1997** (Continued)

Interventions	Misoprostol 25 µg or 50 µg into the posterior vaginal fornix 3 hourly until adequate labour achieved (3 contractions per 10 minutes). Maximum 8 applications. Continuous fetal heart rate monitoring. As soon as cervical dilation permitted, amniotomy performed and intrauterine pressure catheter and scalp electrode applied. Oxytocin used if not in labour after 8 doses or for active phase of labour arrest. Epidural analgesia or intravenous sedation offered. Hyperstimulation syndrome was managed with change in position, oxygen by face mask and in some cases subcutaneous terbutaline or vaginal lavage to remove the misoprostol tablet.	
Outcomes	Mode of delivery; delivery intervals (25 µg 970 +/- 684 vs 50 µg 826 +/- 554 minutes); occurrence of active labour (180/192 vs 191/207); vaginal delivery after one dose (48/192 vs 79/207); maternal and perinatal outcomes; cord pH < 7.16 (13/192 vs 27/207); blind assessment of cardiotocograph for tachysystole, hypertonus and hyperstimulation syndrome.	
Notes	University Medical Centre, Jacksonville and Shands Hospital, Gainesville, July 1994 to September 1995. Apparent transposition of results for delivery < 12 hours and delivery < 24 hours in table IV (see text page 366).	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**044 Chen 2005**

Methods	Randomised clinical trial with allocation by sealed sequential envelopes.	
Participants	Singleton pregnant women with gestational age from 36 to 41 weeks.	
Interventions	Vaginal misoprostol 25 mcg 3 hourly versus no intervention.	
Outcomes	Measure of urinary cGMP/creatinine during labour and duration of latent phase.	
Notes	This trial did not contribute any data to this review.	
<i><b>Risk of bias</b></i>		
<b>Item</b>	<b>Authors' judgement</b>	<b>Description</b>
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Inadequate

**044 Chen 2005** (Continued)

Incomplete outcome data addressed? All outcomes	Yes	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**044 Nanda 2007**

Methods	“The patients were divided randomly by drawing lots and were balanced in blocks of 6” and “The investigators were blinded to the allocation”, but the methods were not specified.
Participants	Intervention group: women between 25 - 35 years old, single fetus, cephalic presentation, intact membranes and Bishop score $\leq 6$ . Exclusion criteria: hypersensitivity to prostaglandin, previous caesarean section, asthma, epilepsy, grand multiparae (4 or more gestations), placenta previa, cephalopelvic disproportion and malpresentation.
Interventions	The intervention group received 25 mcg 3 hourly (maximum 200 mcg). The comparison group used 0.5 mg of dinoprostone intra cervically repeated 6 hourly until 3 doses in 24 hours.
Outcomes	The use of misoprostol in pregnant woman for labour induction showed successes compared with dinoprostone. The labour induction with misoprostol showed less need for oxytocin augmentation.
Notes	Department of Obstetrics and Gynecology, Sharma Post Graduate Institute of Medical Sciences, India.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	No	Inadequate
Allocation concealment?	No	Inadequate
Blinding? All outcomes	Unclear	Unclear for outcome assessors, but blinded for participants and personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

# 044 Wing 1995b

Methods	Allocation by sequentially numbered sealed opaque envelopes, maintained by the primary investigator. Sequence from random number table in blocks of 6.	
Participants	<p>Membranes were intact, cervical status unfavourable. Inclusion criteria: women requiring induction of labour for medical or obstetric reasons; singleton pregnancy; cephalic presentation; intact membranes; cervical score &lt; 5; reactive nonstress test; &lt; 4 uterine contractions per hour.</p> <p>Exclusion criteria: estimated fetal weight &gt; 4500 g; evidence of cephalopelvic disproportion; placenta praevia; unexplained vaginal bleeding; vasa praevia; active herpes simplex infection; contraindication to receiving prostaglandins; renal or hepatic dysfunction; suspected chorioamnionitis; previous uterine surgery; parity &gt; 5.</p> <p>Indications for labour induction: oligohydramnios (misoprostol 53, dinoprostone 57); pre-eclampsia (39, 25); post-term pregnancy (19, 24); macrosomia (12, 15); diabetes mellitus (7, 6); abnormal antepartum testing (3, 3); other (5, 7).</p>	
Interventions	<p>Misoprostol 25 mcg inserted into the posterior vaginal fornix 3-hourly, maximum 8 doses or 24 hours (n = 138) versus dinoprostone gel 0.5 mg intracervically 6-hourly, maximum 3 doses (n = 137), until 3 uterine contractions per 10 minutes, cervical score &gt; 7, cervical dilation &gt; 3 or spontaneous rupture of membranes. Artificial rupture of membranes usually performed when the cervix was 80% effaced and 3 cm dilated. Oxytocin augmentation was used for lack of contractions after maximum dosage or spontaneous rupture of membranes, or for arrested cervical dilation, &gt; 3 hours after misoprostol and &gt; 6 hours after dinoprostone.</p> <p>Uterine hyperstimulation was treated in some cases with tocolytic therapy.</p>	
Outcomes	<p>Oxytocin augmentation; uterine tachysystole (&gt; 5 contractions per 10 minutes); uterine hypertonus (contraction &gt; 2 minutes) (misoprostol 0/138, dinoprostone 5/137); hyperstimulation syndrome (tachysystole or hypertonus with fetal heart rate abnormalities); induction to delivery interval; mode of delivery; vaginal delivery within 24 hours; terbutaline used for hyperstimulation (9/138, 6/137); neonatal resuscitation (44/138, 43/137); days in NICU (mean 7.9 (SD 6.1), 9.9 (7.5)); meconium aspiration syndrome (1/138, 3/137); hyperbilirubinaemia (10/138, 13/137).</p> <p>Frequent uterine contractions occurred in 9 women after a single misoprostol dose, the onset occurring after mean 5.1 (SD 2.4) hours, and 11 after a second dose.</p>	
Notes	<p>Los Angeles, California, USA. February to June 1994.</p> <p>276/287 women agreed to participate. No women withdrew from the protocol. One woman allocated to dinoprostone, inadvertently received misoprostol, and was excluded from the analysis.</p> <p>The authors postulate that the increased rate of meconium-stained amniotic fluid may be due to the increased incidence of uterine tachysystole, or to a direct effect of misoprostol on the fetal gastrointestinal tract.</p> <p>The authors do not recommend this dosage for induction of labour.</p>	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

### 048 Khoury 2001

Methods	“Computer-generated” assignment. Examiner blinded to the dose of misoprostol but not to whether misoprostol or dinoprostone was used.
Participants	Inclusion criteria: indication for labour induction; singleton pregnancy; gestational age 37 weeks or more; vertex presentation; intact membranes; cervical score 4 or less; reassuring fetal heart rate pattern. Exclusion criteria: unexplained third trimester vaginal bleeding; placenta praevia; active herpes simplex infection; history of asthma or glaucoma; suspected cephalopelvic disproportion; parity > 5; uterine scar; estimated fetal weight > 4500 g or > 4000 g in a diabetic mother.
Interventions	Misoprostol 35 mcg versus 50 mcg suppositories in fatty base vaginally 4.5 hourly (maximum 6 doses) versus dinoprostone 10 mg vaginal insert, left for 22.5 hours. Insert removed or misoprostol stopped for active labour or cervical score > 7 or cervical dilatation > 2 or spontaneous rupture of membranes or uterine contraction abnormalities or abnormal fetal heart rate pattern. Syntocinon infusion commenced one hour after the last dose of misoprostol or removal of the insert, if not in labour. For misoprostol vs dinoprostone comparison, the two misoprostol regimens have been combined.
Outcomes	Tachysystole (6 contractions per 10 minutes for 20 minutes); hypertonus (a contraction lasting 2 minutes) ; uterine hyperstimulation (tachysystole or hypertonus plus fetal heart rate abnormality).
Notes	Inova Fairfax Hospital, VA, USA, June 1998 to June 1999. Cardiotocograph tracings were evaluated blind to the allocation.

#### *Risk of bias*

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

### 050 Abdul 2007

Methods	Randomised clinical trial. Random sequence generated by computer. Allocation by numbered sealed opaque envelopes.
Participants	Inclusion criteria: consecutive patients with medical indication to induction of labour. Exclusion criteria: patients with scared uterus and multiple gestation.
Interventions	The intervention group used 50 mcg of vaginal misoprostol 6 hourly during 24 hours. The comparison group used oxytocin alone.
Outcomes	There were no differences between the groups about mean time to delivery, Apgar score and perinatal mortality rate.
Notes	The study period was from November 2005 to October 2006. Labour Unit of Federal Medical Center of Azare, Nigeria. One patient of the misoprostol group had a ruptured uterus.

#### *Risk of bias*

**050 Abdul 2007** (Continued)

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 Agarwal 2003**

Methods	Assigned by “computerised random numbers”. Not blinded.	
Participants	Inclusion criteria: singleton pregnancy; cephalic presentation; intact membranes; cervical score < 7; reactive fetal heart rate testing. Exclusion criteria: previous caesarean section; contraindication to prostaglandin; chorioamnionitis.	
Interventions	Misoprostol 50 mcg vaginally versus dinoprostone 0.5 mg intracervically; repeated after 6 hours if cervical score < 7, cervical dilation < 3 cm, and no adequate uterien contractions (3 or more for 40 seconds or more per 10 minutes); maximum 3 doses. Artificial rupture of membranes once cervix 3 or more cm dilated. Intravenous oxytocin if no adequate labour after 3 doses or rupture of membranes.	
Outcomes	Tachysystole (6 or more contractions per 10 minutes for 20 minutes); hypertonus (contraction for 2 minutes or more); hyperstimulation (tachysystole or hypertonus plus abnormal fetal heart rate pattern) ; change in cervical score after 6 hours (2.98 (SD 2.57) vs 2.05 (SD 1.83)); route of delivery; oxytocin augmentation; passage of meconium; fetal heart rate deceleration; single insertion (54/60 vs 52/60); time to onset of contractions (2.03 (SD 1.3) vs 2.04 (SD 1.0) hours); induction to delivery interval (12.89 (SD 6.5) vs 18.01 (SD 8.4) hours); delivery within 12 hours (52.5% vs 35%).	
Notes	June 2001 to February 2002, New Delhi.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description
Allocation concealment?	Unclear	Unclear

**050 Ayad 2002**

Methods	“There were 238 patients randomly assigned”. The sequence generation and the allocation concealment were not stated.
Participants	Inclusion criteria: women with gestational age > 36 weeks, singleton pregnancy, premature rupture of membranes < 24 hours, Bishop score < 8, vertex presentation and fewer contraction than 12 per hour. Exclusion criteria: prior caesarean delivery, parity > 5 and no reassuring fetal monitoring.
Interventions	The intervention group received 50 mcg vaginal misoprostol and the comparison group received 0.5 mg intra cervically dinoprostone. Both groups used the oxytocin if there was inadequate labour.
Outcomes	Mean time from induction to delivery and the need of oxytocin were significantly fewer in the misoprostol group. There were no difference between the groups in spontaneous labour rate, type of delivery and perinatal outcome.
Notes	The study was performed from February 1999 to February 2000. Prince Rashid Ibn Al-Hassan Hospital, Ramtha, Jordan.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 Bounyasong 2000**

Methods	“This prospective randomised study” has no stated sequence generation and allocation concealment.
Participants	Inclusion criteria: singleton pregnancy, gestational age of 38 weeks, vertex presentation, intact membranes and no evidence of fetal distress. Exclusion criteria: abnormal lie, premature rupture of membranes, oligohydramnios, prior uterine scar, uterine contraction, obstetrical complication, contraindication to prostaglandins and severe medical diseases.



**050 Bounyasong 2000** (Continued)

Interventions	The patients were randomised to receiving 50 mcg or 25 mcg of vaginal misoprostol 6 hourly each group. Amniotomy was performed when the cervical dilatation was achieved 3-4cm. Oxytocin augmentation was used if indicated in both groups.	
Outcomes	There were no differences between the groups in caesarean section rate. The neonatal asphyxia and the uterine hyperstimulation were greater in the 50 mcg group (P = 0.15 and 0.0315).	
Notes	Department of Obstetrics and Gynaecology of Srisangwal Hospital, Thailand.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Inadequate
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**050 Calder 2008**

Methods	"This randomized, open" with not stated allocation methods.	
Participants	Inclusion criteria: all women with singleton pregnancy, with 18 years or older, with at least 37 weeks of pregnancy and unfavourable cervix. Exclusion criteria: women with multiple pregnancy, insulin dependency for diabetes control, multiple pregnancy, PROM, ascending infection, systemic infection, placenta praevia, placental abruption, unexplained vaginal bleeding, active cardiac, pulmonary, renal or hepatic disease a contraindication to vaginal delivery or known allergy to prostaglandins.	
Interventions	The intervention group received vaginal misoprostol 25 mcg (or 50 mcg in nulliparous women) followed by further administration of 25 mcg 4 hourly until a total of 3 doses. Comparioson group: vaginal dinoprostone 3 mg followed by a second dose of 3 mg (6 hours later).	
Outcomes	A total of 626 women were enrolled in this trial. The rate of vaginal deliveries achieved within 24 hours of induction did not significantly differ between the misoprostol and dinoprostone groups. Those treatments were comparable for other efficacy measures and for presence of maternal or fetal averse events.	

**050 Calder 2008** (Continued)

Notes	Eighteen NHS study centers were enrolled in this trial. Reprint of BJOG	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	Yes	Inadequate
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**050 Campos Perez 1994**

Methods	Open randomised study. Computer-generated randomisation.	
Participants	Membranes were intact, cervical status variable. Pregnant women with single fetus. Bishop score < 5 in 35/78 misoprostol and 32/75 oxytocin. Pregnancy complications: hypertension, misoprostol 28 vs oxytocin 26; intrahepatic cholestasis 13 vs 16; impaired fetal growth 6 vs 5; prolonged pregnancy 25 vs 27; others 6 vs 4.	
Interventions	Misoprostol 50 mcg tablet intravaginally (n = 78) vs intravenous oxytocin 2-32 mU/minute (n = 75).	
Outcomes	Delivery within 24 hours; induction to delivery interval; uterine hyperstimulation (> 5 contractions in 10 minutes); caesarean section; Apgar scores. Intrauterine pressures recorded in a subset of 10 women in each group: intensity of contractions as mean (standard deviation): misoprostol 48.7 (10.3) vs oxytocin 56.2 (14.5) mmHg; uterine tone 10.5 (3.6) vs 9.6 (4.8). Differences not statistically significant.	
Notes	Valdivia, Chile. One woman excluded from the misoprostol group because of accidental removal of the misoprostol tablet during a vaginal examination 180 minutes after initiation of the induction. Authors conclude misoprostol safe for induction of labour.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

## 050 Charoenkul 2000

Methods	'Randomised' drugs inserted by a resident not involved in the outcome assessment. Randomisation stratified by parity.	
Participants	Pregnant women with indications for labour induction; singleton pregnancy; 37 or more weeks of gestation; cervical score < 7. Exclusion criteria: suspected cephalo-pelvic disproportion; estimated fetal weight > 4000 g; parity > 5; previous caesarean section and other uterine surgery; suspected chorioamnionitis; contraindications to vaginal delivery; contraindications to the use of prostaglandins; moderate to severe medical disease.	
Interventions	Misoprostol 50 mcg or dinoprostone 3 mg vaginally.	
Outcomes	Primary: change of cervical score at 24 hours. Secondary: uterine tachysystole (5 or more contractions per 10 minutes for 20 minutes); hypertonus (a contraction lasting 90 seconds); hyperstimulation syndrome (tachysystole or hypersystole and fetal heart rate changes); vaginal delivery in 24 hours; caesarean section; caesarean section for fetal distress (misoprostol 6/72 vs dinoprostone 3/71).	
Notes	Vajira Hospital, Bangkok, Thailand, November 1998 to December 1999.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

## 050 Denguezli 2007

Methods	Randomised controlled trial with a computer sequence generation and allocation concealment from a predetermined random table.	
Participants	Women with obstetrical indication for cervical ripening and induction of labour. Inclusion criteria: obstetrical indication, medical complication, absence of active labour or fetal distress, no previous caesarean section delivery or other type of uterine surgery, singleton pregnancy with vertex presentation and no contraindication to vaginal delivery. Exclusion criteria: less than 18 years, gestational age less than 36 weeks, PROM, history of dystocia or forceps delivery, history of more than one episode of surgical interruption of pregnancy.	
Interventions	The intervention group received 50 mcg of misoprostol 6 hourly until Bishop favourable (maximum 200 mcg). The comparison group received intracervical prostaglandin 0.5 mg 6 hourly until induction of labour (maximum 2 mg).	
Outcomes	The proportion of vaginal delivery within 24 hours was significantly higher in the misoprostol group (75%) than in the dinoprostone group (53.8%) (RR = 1.40, 95% CI (1.07-1.45), p = 0.02). There was no difference between the mean time interval of delivery in the misoprostol group and the dinoprostone group (14.9 vs. 15.8h) (p = 0.51). Secondary outcomes were evolution of Bishop score during the labour, rate of caesarean delivery, tachysystole, hyperstimulation syndrome, maternal side effects and neonatal outcomes.	

**050 Denguezli 2007** (Continued)

Notes	University Hospital Medical Centre The patients were included from 1 August 2003 to 30 April 2004. No blinding trial.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear for participants and outcome assessors and open for personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 El-Azeem 1997**

1994 to 1995

Methods	Double blind randomised trial.	
Participants	29 women with indication for induction of labour.	
Interventions	Misoprostol 50 µg in an applicator plus placebo gel in a syringe (n = 15), versus 0.5 mg dinoprostone in a syringe plus placebo applicator (n = 14), repeated after 6 hours if necessary. Oxytocin started 6 hours after first or second dose as indicated.	
Outcomes	Mean change in cervical score after 6 hours (misoprostol 4.2 +/- 6.3 versus dinoprostone 2.3+/-3.3, p < 0.05); induction to delivery interval (17.2+/-10.1 versus 32.9+/-18.6, p = 0.03); delivery with prostaglandin alone (46.2% versus 16.7%); oxytocin dosage and duration. No data in format suitable for review.	
Notes	1994 to 1995. The percentages could not be converted to proportions (16.7% of 14 is 2.3). Full report of trial awaited for incorporation of data. Not clear whether vaginal or intracervical dinoprostone.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Elhassan 2004**

Methods	“Prospective randomised controlled clinical trial”. The sequence generation and allocation concealment were not stated.
Participants	Inclusion criteria: women admitted to the labour ward with singleton pregnancy and unripe cervix. Exclusion criteria: previous uterine surgery, antepartum haemorrhage, glaucoma, asthma, heart disease and grand multiparity.
Interventions	The intervention group received 50 mcg of misoprostol 6 hourly until 200 mcg. The comparison group received vaginal prostaglandin 0.5 mg 6 hourly until 2 mg.
Outcomes	The induction-to-delivery interval (mean $\pm$ S.D.) was 17.5 h $\pm$ 7.6h with misoprostol and 19.15 h $\pm$ 6.9 h with dinoprostone(p > 0.05). Secondary outcomes were route of delivery, need of oxytocin augmentation and neonatal outcomes.
Notes	This study was conducted at the labour ward of Wad Medani Hospital at Sudan, from March to December 2001.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unblinded for personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 Elhassan 2005b**

Methods	“An open randomised controlled clinical trial”. The sequence generation and allocation concealment were not stated.
Participants	Included criteria: pregnant women with single babies, favourable cervix and intact membranes. Exclusion criteria: previous uterine surgery, antepartum haemorrhage, asthma, heart disease and grand multiparity.
Interventions	The intervention group received 50 mcg of misoprostol 6 hourly until 200 mcg. The comparison group received oxytocin alone (they started with 2 mcU/min, which was doubled at 30-min interval until labour).

**050 Elhassan 2005b** (Continued)

Outcomes	The induction-delivery interval was significantly shorter in misoprostol group ( 8.2 h ± 1.1 h vs 12.04 h ± 1.5 h) vs oxytocin group. The secondary outcomes were rate of instrumental vaginal delivery, caesarean section and neonatal outcomes.	
Notes	This study was conducted at the labour ward of Wad Medani Hospital at Sudan, from August 2004 to February 2005.	
<i><b>Risk of bias</b></i>		
<b>Item</b>	<b>Authors' judgement</b>	<b>Description</b>
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Unclear	Unclear

**050 Frohn 2002**

Methods	Allocation by computer-generated random number sequence in sealed opaque envelopes, in balanced blocks of 50.
Participants	Inclusion criteria: ruptured membranes; gestation 34 weeks or more; singleton gestation; reassuring fetal heart rate pattern; cephalic presentation; not in labour; cervical dilation < 3 cm; effacement no more than 80%; no contraindication to labour and vaginal delivery. Exclusion criteria: estimated fetal weight > 4500 g; intrauterine infection; contraindication to use of prostaglandins. Women with one previous lower transverse caesarean section were not excluded.
Interventions	Misoprostol 50 mcg vaginally, repeated after 6 hours if not in labour, versus dinoprostone gel 2.5 mg vaginally 6-hourly. If not in labour 12 hours after the first dose, oxytocin infusion started. Continuous cardiotocography was used.
Outcomes	Tachysystole (6 or more contractions per 10 minutes for 20 minutes); hyperstimulation (tachysystole and FHR changes); chorioamnionitis; postpartum endometritis; neonatal sepsis.
Notes	Arnold Palmer Hospital for Women and Children, Florida, USA. January 1995 to December 2000. Not blinded.
<b><i>Risk of bias</i></b>	

**050 Frohn 2002** (Continued)

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Gelisen 2005**

Methods	Randomised controlled trial. The investigators were not involved at the clinical care of the patients and they carried out the randomisation by opening sealed opaque envelope.
Participants	Inclusion criteria: singleton live pregnancy with vertex presentation and intact membranes, gestational age plus than 41 completed weeks, Bishop score < 5, absence of spontaneous uterine contractions, estimated fetal body weight < 4500g, reactive non-stress test and amniotic fluid index $\geq$ 5cm. Exclusion criteria: known hypersensitivity to the use of prostaglandins, previous caesarean delivery or other uterine surgery, non-cephalic presentation, body mass index $\geq$ 30 before conception, any previous attempt at induction of labour during the current pregnancy and low-lying placenta.
Interventions	The intervention group received 50 mcg of misoprostol 6 hourly until 24 hours of labour. The first comparison group received oxytocin alone (the initial doses started with 1 mU/min, increased by 1mU/min every 15 minutes until contractions of 200 - 250 Montevideo units were achieved.) The second comparison group received a Foley catheter balloon inserted to above the cervical OS (inflated with 50 ml of sterile saline). After the expulsion of the catheter the oxytocin was started with the same regime described above. The third comparison group was the spontaneous follow up until 42 weeks of pregnancy. If the women complete 42 weeks, the IOL started with 50 mcg of vaginal misoprostol 6 hourly until 24 h. If the delivery was not achieved, the caesarean section was performed.
Outcomes	The abdominal delivery rate was 19.3% in the intervention, first and second comparison groups, and 22% in the spontaneous follow up. The meconium-stained amniotic fluid and meconium aspiration were significantly higher in the follow-up group. Rates of emergency abdominal delivery, NICU admission and low umbilical artery pH were similar at the groups.
Notes	The trial was performed in a tertiary training centre in Turkey. SSK Ankara Maternity and Women's Health Teaching Hospital, Kocaeli, Turkey. For the analyses of data at this review the mechanical methods to IOL were not included.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Yes	Adequate

**050 Gelisen 2005** (Continued)

Blinding? All outcomes	No	Unclear for participants, inadequate for personnel and blinded for outcome assessors
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 Gotschall 1998**

Methods	Medications prepared by hospital pharmacy in a randomised, double-blind fashion.
Participants	Women presenting for induction of labour.
Interventions	Misoprostol 50 vs 100 mcg intravaginally as a single dose. If not in labour after 6 hours, oxytocin was commenced.
Outcomes	Primary: induction to delivery time. Secondary: need for oxytocin; delivery mode; maternal symptoms; uterine hyperstimulation.
Notes	Three women excluded for failed induction. Not indicated which group(s) they belonged to. Data not included, pending further information.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Kovavisarach 1997**

Methods	"All patients were randomized". The sequence generation and allocation concealment were not stated.
Participants	Inclusion criteria: women with gestational age $\geq 37$ weeks, indication for induction of labour, Bishop score $< 7$ and no contraindication to use prostaglandins. No exclusion criteria.
Interventions	The intervention group used 100 mcg 12/12 hours (maximum 200 mcg). The intervention group used vaginal dinoprostone 3 mg two times.
Outcomes	Bishop score and tachysystole were significantly higher in the misoprostol group. Induction to delivery interval was significantly lower in misoprostol group ( $p < 0.05$ ). Mode of delivery, Apgar score, hyperstimulation, neonatal and maternal complications were



**050 Kovavisarach 1997** (Continued)

	not different in the group.	
Notes	The study enrolled patients from March 15 to September 15, 1995. Department of Obstetrics and Gynaecology, Rajavith Hospital, Ministry of Public Health, Bangkok, Thailand.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unblinded for personnel
Incomplete outcome data addressed? All outcomes	Unclear	Unclear
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**050 Kovavisarach 1998**

Methods	“The woman were randomized in two groups”. The sequence generation and the allocation concealment were not stated.	
Participants	The pregnant women were included if the gestational age was higher than 37 weeks, Bishop score < 7 and no contraindication to use prostaglandin.	
Interventions	The intervention group used 50 mcg vaginal misoprostol 6 hourly (maximum 150 mcg) . The comparison group used 3 mg vaginal dinoprostone 6 hourly (3 times) and both groups could receive oxytocin augmentation.	
Outcomes	Misoprostol group presented significantly higher tachysystole (p < 0.05) and lower induction to delivery interval, induction to oxytocin stimulation interval and oxytocin stimulation to delivery interval. There were no difference of mode of delivery, Apgar score, hyperstimulation syndrome, neonatal and maternal complications.	
Notes	The study was enrolled from December 1, 1996 to August 30,1996. Department of Obstetrics and Gynaecology, Rajavith Hospital, Ministry of Public Health, Bangkok, Thailand.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description

**050 Kovavisarach 1998** (Continued)

Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	Yes	Paucity of information mainly in the neonatal side
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**050 Le Roux 2002**

Methods	“Computer randomisation” to vaginal misoprostol, oral misoprostol or dinoprostone in ration 1:1:2. Stratified by delivery site. Allocations sealed in opaque envelopes opened by labour ward staff. Cervical assessment and cardiotocography before enrolment.
Participants	Women having labour induced with singleton cephalic fetus; 34 weeks or more gestation; intact membranes; no fetal distress on cardiotocography; no painful contractions; age 18 years or more. Exclusion criteria: previous caesarean section; parity > 4; fetal anomaly; fetal death; cervical score > 7 where amniotomy without prostaglandin preparation was possible.
Interventions	Vaginal misoprostol 50 mcg (maximum 4 doses) versus oral misoprostol 50 mcg (maximum 4 doses) versus dinoprostone gel 1 mg vaginally (maximum 2 doses), 6-hourly. Artificial rupture of membranes and incremental oxytocin infusion were used for poor progress in established labour, not for labour induction. Continuous cardiotocography was used. If tachysystole occurred, the next dose was delayed until the tachysystole had resolved. For this review only vaginal misoprostol vs dinoprostone compared.
Outcomes	Primary: vaginal delivery within 24 hours; secondary: mode of delivery; indication for caesarean section; caesarean section for fetal distress (vaginal misoprostol 33/120 vs dinoprostone 33/240); placental abruption (vaginal misoprostol 4/120 vs dinoprostone 6/240); fetal complications (thick meconium-stained liquor; 5 minute Apgar score < 7; admission to neonatal intensive care unit; hypoxic ischaemic encephalopathy); tachysystole (5 or more uterine contractions in 10 minutes, in two 10-minute windows; fetal distress (a fetal heart rate tracing that justified immediate delivery).
Notes	One tertiary and one secondary academic hospital in Cape Town, South Africa. 93/573 enrolled women were excluded for clerical errors (65), exclusion criteria ignored (18), patient withdrew (1), incorrect dosage (5), underage (2). No blinding of clinicians or of the cardiotocographic evaluation.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

### 050 Lokugamage 2003a

Methods	Allocation by opaque numbered sealed envelopes in computer-generated sequence. Not blinded.	
Participants	Inclusion criteria: multiparous women; > 36 weeks' gestation. Exclusion criteria: previous caesarean section; uterine surgery; parity > 5; abnormal FHR pattern; malpresentation; estimated fetal weight > 4500 g; other evidence of cephalopelvic disproportion; placenta praevia; contraindication to prostaglandin therapy.	
Interventions	Vaginal misoprostol 50 mcg single dose versus two doses, the second after 6 hours if no uterine activity and amniotomy not possible. After 24 hours, dinoprostone used if needed. Oxytocin used after rupture of membranes if needed.	
Outcomes	Induction to delivery time (789 (SD 539) vs 576 (SD 331) minutes); delivery within 12 and 24 hours; oxytocin augmentation; tachysystole (6 or more contractions in 10 minutes); hyperstimulation (abnormal FHR pattern related to excessive uterine activity); mode of delivery; failed induction of labour; Apgar score; umbilical venous blood pH (7.33 (7.12-7.49) vs 7.29 (7.11-7.49)); admission to NICU.	
Notes	University College Hospital, London, UK, January 1998 to December 1999.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

### 050 Lokugamage 2003b

Methods	Randomised unblinded trial with allocation concealment in opaque sealed numbered, opened consecutively.	
Participants	Inclusion criteria: nulliparous women at term (37 to 42 weeks of gestation). Exclusion criteria: previous uterine surgery, abnormal FHR, malpresentation, estimated fetal body weight > 4,500 g, or other evidence of cephalopelvic disproportion, placenta previa and any contraindication to prostaglandin therapy.	
Interventions	The intervention group received 50 mcg of misoprostol and then if it was required another dose of 50 mcg after 6 hours. The comparison group received 2 mg of dinoprostone vaginal, amniotomy and additional dose of 1 mg before 24 hours. If there was no success within 24 hours the regimen was repeated for a second day.	
Outcomes	The induction to delivery interval (1047 min vs 1355 min, $p = 0.01$ ), delivery within 12 hours (35.4% vs 18.9%, $p = 0.02$ ) and delivery within 24 hours (83.3% vs 63.3%, $p = 0.82$ ) were all shorter in the misoprostol arm. There was no differences in rates of oxytocin augmentation, tachysystole and hyperstimulation syndrome. There was no difference in neonatal outcomes.	
Notes	The study run from January 1998 to December 1999. Royal Free & University College London Medical School, London, UK.	

**050 Lokugamage 2003b** (Continued)

<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 Majoko 2002a**

Methods	Single-blinded randomised controlled trial. Researcher knew group allocation. Allocated by cards in numbered sealed opaque cards, randomised by random number tables.	
Participants	Inclusion criteria: women with obstetric indication for induction of labour; singleton live fetus; cephalic presentation; 37 weeks' gestation or more. Exclusion criteria: previous uterine surgery; vaginal delivery contraindicated.	
Interventions	Half versus quarter 200 µg misoprostol tablet inserted into posterior vaginal fornix. Repeated after 8 hours if cervical score < 10 (maximum 2 doses).	
Outcomes	Induction to delivery interval; mode of delivery; augmentation with oxytocin; postpartum haemorrhage; uterine trauma; fetal outcomes.	
Notes	Harare Maternity Hospital, Zimbabwe, June to September 1998. Terminated early because of complications in the 100 µg group.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Mosquera 1999**

Methods	Allocation by computer-generated sequence using sealed, opaque, numbered envelopes.
Participants	Inclusion criteria: gestation 42 or more weeks; single fetus, alive, cephalic. Exclusion criteria: uterine contractions; ruptured membranes; placenta praevia, placental abruption, breech presentation, dystocia; previous uterine surgery; fetal anomaly; multiple pregnancy.
Interventions	Misoprostol 50 microgram doses dissolved in 0.5 ml to 1 ml saline intravaginally, versus oxytocin intravenous infusion.
Outcomes	Induction to delivery (misoprostol 15.4 +/- 1.5, oxytocin 18.6 +/- 2.0).
Notes	University Hospital Del Valle, Cali, Colombia, 2 April 1993 to 2 April 1996. Four from the misoprostol group were withdrawn.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Neiger 2001**

Methods	"Randomly assigned".
Participants	Inclusion criteria: indication for labour induction; cervical score < 5; singleton pregnancy; cephalic presentation; term; intact membranes; reactive FHR pattern. Exclusion criteria: contraindication to vaginal delivery; previous uterine surgery; labour; vaginal bleeding; asthma; hypersensitivity to prostaglandins.
Interventions	Misoprostol 50 mcg vaginally versus dinoprostone 0.5 mg intracervically, maximum 2 doses 6 hours apart; continuous cardiotocography; oxytocin infusion when cervical score > 8 or at 2 hours, if not in labour; amniotomy when cervix 3-4 cm dilated.
Outcomes	Number of doses; change in cervical score; induction interval; oxytocin use; caesarean section; uterine hyperstimulation (tachysystole or hypertonus with abnormal FHR pattern).
Notes	University of Tennessee Medical Center, August 1995 to December 1996.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**050 Ortiz 2002**

Methods	Randomised trial with sequence generation and allocation not stated.
Participants	Inclusion criteria: gestation at term, singleton pregnancy, cephalic presentation, rupture of membranes and Bishop score $\leq 4$ . Exclusion criteria: fetal distress, estimated fetal weight > 4 kg, placenta previa, prolapsed cord, fever and diarrhoea.
Interventions	The patients were divided in three groups: I. patients received 50 mcg vaginal misoprostol associated with oxytocin EV until labour; II. patients received intravaginal placebo associated with oxytocin; III. patients received 50 mcg vaginal misoprostol associated with placebo EV until labour.
Outcomes	The frequency of tachysystole, mode of delivery and perinatal outcomes were similar among the three groups. The mean time from induction to delivery was different in the three groups: Group I: 48.75 minutes, Group II: 537.05, Group III: 474.54 ( $p < 0.05$ ).
Notes	The analysis data will include groups I and II considering oxytocin as a co-intervention. Hospital Civil de Culiacán, Universidade Autónoma de Sinaloa.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear for outcome assessors and blinded for participants and personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 Pandis 2001**

Methods	Computer-generated random sequence in balanced blocks of 10. Allocation by contacting the lead researcher.
Participants	Inclusion criteria: age > 16 years; singleton pregnancy; gestational age 37-42 weeks; live fetus; cephalic presentation; intact membranes. Exclusion criteria: history of antepartum haemorrhage; previous uterine surgery; allergy to prostaglandins.
Interventions	Ultrasound and clinical vaginal assessment before induction. Misoprostol 50 mcg vaginally versus dinoprostone (nulliparae with cervical score < 5: 2 mg; score 5-6 and all multipara 1 mg). In both groups:

**050 Pandis 2001** (Continued)

	dose repeated after 6 hours, and process repeated after 24 hours if necessary; score 7 or more had artificial rupture of membranes as well; cardiotocography after the medication and throughout labour; oxytocin augmentation if needed 6 or more hours after last prostaglandin dose.
Outcomes	Primary: vaginal delivery in 24 hours. Secondary: hyperstimulation syndrome (tachysystole, > 5 contractions per 10 minutes for 20 minutes or hypersystole, a contraction > 2 minutes with FHR abnormalities) ; caesarean section; adverse maternal and neonatal outcome.
Notes	September 2000 to September 2001. Universitäts-Frauenklinik, Kantonsspital, Basel, Switzerland; King George, Harold Wood and Southend Hospitals, Essex, UK. 235/67 excluded after randomisation, mainly for spontaneous delivery before induction or induction by amniotomy for cervical score 7 or more.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Ramsey 1998**

Methods	Described as a “prospective randomized blinded phase III clinical trial” .
Participants	No reference to gestational age.
Interventions	Vaginal misoprostol 50 mcg 6 hourly 2 times versus vaginal Prepidil® or Cervidil® 10mg one dose.
Outcomes	Analysis of costs for induction of labour.
Notes	This study did not contribute with any data to this review.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Yes	Adequate
Incomplete outcome data addressed? All outcomes	Unclear	Unclear
Free of selective reporting?	Unclear	Unclear

**050 Ramsey 1998** (Continued)

Free of other bias?	Unclear	Unclear
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**050 Ramsey 2003**

Methods	Prospective randomised blinded phase III clinical trial. Randomisation by pharmacy using dynamic allocation with stratification by parity and initial cervical score (0-2 vs > 2).
Participants	Inclusion criteria: indication for labour induction; cervical score < 6; singleton vertex pregnancy; no contraindication to vaginal delivery; < 4 uterine contractions per hour; reactive cardiotocography. Exclusion criteria: hypersensitivity to prostaglandins; ruptured membranes; suspected chorioamnionitis; parity > 5; previous caesarean delivery or uterine surgery; previous attempted labour induction for this pregnancy.
Interventions	1. Misoprostol 50 µg intravaginally 6-hourly for 2 doses, vs 2. dinoprostone pessary 10 mg intravaginally vs 3. dinoprostone gel 0.5 mg intracervically 6-hourly for 2 doses. Second doses omitted if > 3 uterine contractions in 10 minutes, fetal distress or ruptured membranes. After 12 hours the cervix was reassessed and oxytocin commenced if necessary. Continuous cardiotocography was used. If no cervical change in 24 hours, treatment repeated.
Outcomes	Mean change in cervical score over 12 hours (misoprostol 5.2 +/- 3.1 vs dinoprostone pessary 3.2 +/- 2.3 vs dinoprostone gel 2.2 +/- 1.3); cervical score > 5 after 12 hours (30/38 vs 23/38 vs 14/35); time to vaginal delivery (23.9 +/- 11.1 vs 31.5 +/- 13.5 vs 31.1 +/- 14.2 hours); mean cost (\$2.37 +/- 0.65 vs \$168.23 +/- 0 vs 203.43 +/- 21.84); complete cervical dilation within 24 hours (26/38 vs 19/38 vs 18/35); time to delivery (24.0 +/- 10.8 vs 32.2 +/- 14.7 vs 31.6 +/- 13.4 hours); delivery within 48 hours (37/38 vs 28/38 vs 26/35); additional cervical ripening (4/38 vs 8/38 vs 10/35); caesarean section; obstetric outcomes; neonatal outcomes.
Notes	Mayo Medical Centre, University of Alabama at Birmingham, April 1996 to August 1997. Two earlier abstracts Ramsay 1998 and 2001) assumed to be reports of the same study.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Ramsey 2005**

Methods	"Randomized clinical trial" with method of allocation and sequence generation not stated.
Participants	Inclusion criteria: Bishop score ≤ 5, singleton pregnancy with vertex presentation and no contraindication to vaginal delivery, absence of spontaneous contractions and reactive non-stress test. Exclusion criteria: known hypersensitivity to prostaglandins, ruptures of membranes, suspected chorioamnionitis, parity more than five, previous caesarean delivery or uterine



**050 Ramsey 2005** (Continued)

	surgery and previous attempted induction of labour for this pregnancy.
Interventions	The intervention group received 50 mcg of vaginal misoprostol repeated dosing 6 hours later and there are 2 comparison groups. The first comparison group received 0.5 mg of dinoprostone intra cervically repeated 6 hours late and the second one received 10 mg of dinoprostone intravaginally repeated 12 hours late.
Outcomes	The 55% of the misoprostol treated women demonstrated abnormal tracing cardiotocography event during first 24 hours of induction compared with 21.1% and 31.4% respectively the comparison group.
Notes	Data collected at Mayo Medical Center from April 1996 to August 1997.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Yes	Adequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**050 Rozenberg 2001**

Methods	Computer-generated randomisation in balanced blocks of 4-16, using sealed opaque envelopes. Midwife who administered induction agent asked not to divulge allocation to woman or other staff.
Participants	Inclusion criteria: singleton; cephalic; cervical score > 6; < 4 uterine contractions per hour. Exclusion criteria: age < 18 years; fetal distress; feto-pelvic disproportion; placental praevia; previous caesarean section; allergy to prostaglandins.
Interventions	Misoprostol 50 mcg versus dinoprostone gel 2 mg, vaginally and repeated after 6 hours if not in labour; if not in labour after 24 hours, misoprostol or dinoprostone (2 mg, 1 mg, 1 mg) repeated 4-hourly for 3 doses; amniotomy after 2-3 cm cervical dilation; oxytocin augmentation for poor progress of labour. If not in labour after 48 hours, amniotomy and syntocinon if cervical score > 6, or caesarean section.
Outcomes	Primary: vaginal delivery within 24 hours. Secondary: vaginal delivery within 12 hours; time from randomisation to delivery; cervical score after 12 hours; caesarean section; caesarean section for fetal distress (misoprostol 24/184 vs dinoprostone 13/185); forceps delivery; thick meconium-stained liquor; fetal dis-

**050 Rozenberg 2001** (Continued)

	tress; hypertonus (a contraction of 2 minutes); hyperstimulation (tachysystole or hypertonus with FHR abnormality); uterine tachysystole (> 5 contractions/10 minutes for 20 minutes); Apgar scores; arterial cord pH; meconium aspiration (3/184 vs 3/185); NICU admission; poor neonatal outcome (6/184 vs 9/185); vaginal pain (3/184 vs 34/185); maternal complications; cost (Sterling 2134, SD 574 vs 2202, SD 595).	
Notes	Poissy Hospital, France, July 1997 to April 1999. One withdrawal after enrolment in the misoprostol group.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Not used

**050 Rozenberg 2004**

Methods	Randomised clinical trial. Sequence generation by computer with list permuted blocks of 4 blinded to the investigators and allocation by sealed opaque envelopes.	
Participants	Inclusion criteria: singleton pregnancy, cephalic presentation, Bishop score < 5 and < 4 spontaneous uterine contraction per hour. Exclusion criteria: age < 18 years old, fetal distress before induction, fetal pelvic disproportion, placenta previa, premature rupture of membranes, breech or transverse lie, previous caesarean delivery and known allergy to prostaglandin.	
Interventions	The intervention group received 50 mcg of vaginally misoprostol 6 hourly until 250 mcg in 24 hours. The comparison group received 10 mg of vaginal dinoprostone repeated 12 hours later or until 48 hours after the primary dose.	
Outcomes	Neonatal tolerance was similar in the 2 groups, with no difference in the caesarean delivery rate for fetal distress or in the incidence of meconium-stained amniotic fluid. Time to vaginal delivery was shortened by misoprostol.	
Notes	Data collected from February 2000 and February 2001. Poissy Hospital, University Versailles-St Quentin, France.	
<i><b>Risk of bias</b></i>		
<b>Item</b>	<b>Authors' judgement</b>	<b>Description</b>
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Blinded for participants, unblinded for personnel and unclear for the outcome assessors

**050 Rozenberg 2004** (Continued)

Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	No	Inadequate
Free of other bias?	Yes	Adequate

**050 Sahu 2004**

936 Santa 2004

Methods	“The women were randomly allocated”. The sequence generation and the allocation concealment were not stated.	
Participants	Inclusion criteria: singleton pregnancy, vertex presentation, Bishop score < 5 and intact membranes. Exclusion criteria: multiple pregnancy, parity > 4, breech presentation, previous uterine scar, hypersensitivity to prostaglandins, probable cephalopelvic disproportion, vaginal bleeding in second pattern and vaginal or cervical infection.	
Interventions	The intervention group received 50 mcg vaginal misoprostol repeated if no cervical ripening after 6 hours. The comparison group received 0.5 mg intracervical dinoprostone repeated after 12 hours if necessary. Both groups could use oxytocin augmentation.	
Outcomes	There were no differences between the groups in time interval from induction to delivery, in the incidence of tachysystole, hypersystole and caesarean section rate. The incidence of delivery before 12 hours was higher in the misoprostol group (p < 0.02).	
Notes	The study was carried from January to April 2002. Rajah Muthaih Medical College and Hospital, Annamalai University, Tamil Nadu.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Blinded only for personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Unclear	Unclear

## 050 Sifakis 2007

Methods	“A sequence from a computerized random number generator was used for the allocation of patients to each group.”
Participants	Inclusion criteria: singleton pregnancy with vertex presentation, intact membranes, Bishop score $\leq 4$ , reactive non-stress test and absence of labour. Exclusion criteria: cephalopelvic disproportion, suspected fetal distress, renal or hepatic dysfunction, contraindication to prostaglandin administration, previous caesarean delivery or uterine surgery and parity greater than 5.
Interventions	The intervention group received 50 mcg of vaginal misoprostol 6 hourly at the maximum of 3 doses. The comparison group received 3 mg of dinoprostone intravaginally at the maximum of 3 doses.
Outcomes	The mean interval from labour induction to delivery and the average use of oxytocin was smaller in the misoprostol group.
Notes	Department of Obstetrics and Gynecology, University of Crete, Greece.

### *Risk of bias*

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	No	Inadequate

## 050 Surbek 1997

Methods	Random number-generated table used by hospital pharmacy to allocate identical-looking vaginal suppositories with quick-release, white gelatin capsules filled with the commercially available drugs. The code was not broken till study closure.
Participants	Of 103 enrolled, 3 excluded for protocol violations. Inclusion criteria: viable term pregnancy; induction of labor required; willing to participate; cervical score $< 6$ ; reactive cardiotocography; singleton vertex presentation; labour absent. Misoprostol group had significantly lower cervical scores. Exclusion criteria: malpresentation; previous uterine surgery; contraindication to receive prostaglandins.

**050 Surbek 1997** (Continued)

Interventions	Misoprostol 50 µg vs prostaglandin E2 3 mg into the posterior vaginal fornix, repeated if not in labour after 6, 24 and 30 hours; if not in labour after 48 hours, intravenous oxytocin was given; the cervix was assessed before each administration; oxytocin augmentation (1.25 to 20 mU per minute) was used at the discretion of the attending midwives and residents; preferably late artificial rupture of membranes was performed. External cardiotocographyc monitoring was used. Analgesia included spasmolytic and epidural analgesia.	
Outcomes	Primary: delivery within 24 hours. Secondary: intrapartum complications, maternal side effects (vomiting 2/50 vs 3/50, diarrhoea 1/50 vs 1/50, fever 2/50 vs 4/50), fetal outcome. Tachysystole (6 or more contractions per 10 minutes); hyperstimulation syndrome (tachysystole plus FHR abnormalities), both assessed blind; uterine contraction pain in the latent or early active phase of the first stage of labour, assessed 12 hours after delivery using a pain analogue scale (7.4 (3-10) vs 7.7 (1-10); FHR anomalies (16/50 vs 17/50); epidural analgesia (22/50 vs 25/50); spasmolytic therapy (27/50 vs 30/50).	
Notes	Basel University Hospital, January to November 1995.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**050 Thomas 2000**

Methods	“Randomized”, method not stated.	
Participants	Women with term pre-labour spontaneous rupture of membranes; reassuring fetal status; vertex presentation.	
Interventions	Intravaginal misoprostol 50 mcg vaginally, repeated if necessary after 6 hours, versus placebo. Oxytocin used if not in labour after 12 hours.	
Outcomes	Induction to delivery interval (misoprostol 15.5 SD 7.3 vs placebo 19.0 SD 6.7 hours); oxytocin; oxytocin dose (7.6 SD 7.9 vs 11.6 SD 9.5 mU/minute); caesarean section; chorioamnionitis (2/27 vs 5/25); fetal meconium passage.	
Notes	Abstract only.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description
Allocation concealment?	Unclear	Unclear

### 050G Carlan 1997

Methods	Opaque sealed envelopes in computer-generated random sequence.
Participants	Inclusion criteria: singleton pregnancy; cervical score < 7; estimated fetal weight < 4500 g; live fetus; gestational age > 26 weeks. Exclusion criteria: vaginal bleeding; non-reassuring FHR pattern; breech presentation; uterine contractions 4 or more per 20 minutes; contraindication to vaginal birth. Women with previous caesarean section or ruptured membranes were not excluded.
Interventions	Misoprostol gel (50 mcg mixed with 1 ml saline and 4 ml hydroxy ethylcellulose gel) vs tablets (moistened with 4-8 drops of saline); inserted vaginally 8-hourly in dosages of 50 mcg x 2 then 100 mcg, until labour, cervical score 7 or more, intervention required or 6 doses given.
Outcomes	Hyperstimulation; time from administration to labour or start of induction (gel 18.2 SD 16.6 vs tablet 13.8 SD 11.4); and delivery (29.0 SD 19 vs 22.4 SD 15); caesarean section and oxytocin use.
Notes	Arnold Palmer Hospital for women and children, Tampa, Florida, USA. August 1 1995 to February 1 1997. The physicians were not blinded to the treatment allocation.

#### *Risk of bias*

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

### 058 Ferguson 2002

Methods	Double-masked, allocation by random number tables.
Participants	Inclusion criteria: membranes intact, modified cervical score < 5, singleton vertex presentation. Exclusion criteria: > 5 contractions per hour; active cardiovascular disease; vaginal bleeding; glaucoma; asthma; hypersensitivity to prostaglandins or beta-adrenergic agents; abnormal FHR pattern; any other contraindication to vaginal delivery; (after 1998): previous caesarean section.
Interventions	Vaginal misoprostol 50 mcg stat then 25 mcg 4-hourly, versus syntocinon infusion 1 mU/minute, increased hourly -2 -4 or reduced if > 3 contractions/10 minutes; continuous electronic FHR monitoring; induction of labour after 16 hours, or sooner if modified cervical score > 6 or spontaneous rupture of membranes; early amniotomy and intrauterine pressure catheter monitoring.
Outcomes	Uterine tachysystole (> 5 contractions in 10 minutes for 20 minutes); hypertonia (contraction lasting > 2 minutes); hyperstimulation (above plus abnormal FHR pattern, managed with lateral position, oxygen and terbutaline.
Notes	Recruitment 1996 to 2000.

#### *Risk of bias*

Item	Authors' judgement	Description
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**058 Ferguson 2002** (Continued)

Allocation concealment?	Unclear	Unclear
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**063 Varaklis 1995**

Methods	Open, randomised trial. Allocation by numbered, sealed opaque envelopes, in sequence from random number tables.
Participants	Membranes were intact, cervical status unfavourable. Women at term with a medical indication for induction of labour. Exclusion criteria: severe oligohydramnios (< 2 cm pocket); non-reactive non-stress test; prior uterine surgery; malpresentation; multiple pregnancy; 3 or more contractions in 10 minutes; cervical score > 5.
Interventions	Misoprostol 25 mcg 2-hourly administered vaginally, maximum 6 doses (n = 36), versus prostaglandin E2 0.5 mg intracervically 6-hourly, maximum 2 doses (n = 32); administration stopped when uterine contractions reached 3 in 10 minutes; membranes ruptured when cervix 3 cm dilated, and scalp electrode and intrauterine pressure catheter placed; oxytocin infusion commenced 12 hours after the first dose if not in progressive labour.
Outcomes	Frequent uterine contractions (> 5 per 10 minutes over 20 minutes); hyperstimulation (frequent contractions or prolonged contraction > 2 minutes, with fetal tachycardia, late decelerations or loss of short-term variability); times to labour onset, rupture of membranes, full cervical dilation and delivery; use of oxytocin; mode of delivery; neonatal outcome.
Notes	Maine, USA. After enrolment, 11/80 (14%) excluded because of protocol violations (5), receiving prostaglandin E2 vaginal gel rather than the trial preparation (2), prematurity (1), HELLP syndrome (1), spontaneous labour (2). The 2 cases of uterine hyperstimulation occurred after the 2nd and 3rd misoprostol doses, and the one of frequent contractions after the 3rd dose. The authors recommend further study of the dosing interval.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**075 Buser 1997**

Methods	Allocation by numbered opaque sealed envelopes. Sequence taken from random number table in blocks of 6.
Participants	Membranes were intact, cervical status unfavourable. Inclusion criteria: women requiring induction of labour; singleton pregnancy at term, cephalic presentation; reassuring FHR tracing; cervical score < 6. Indications for labour induction: post-dates pregnancy (misoprostol 31, dinoprostone 23); pre-eclampsia (21, 23); decreased amniotic fluid (9, 6); macrosomia (3, 12); gestational diabetes (2, 2); fetal growth restriction (3, 2); other (6, 11).

**075 Buser 1997** (Continued)

	Exclusion criteria: ruptured membranes; low-lying placenta; previous caesarean section; parity > 5; suspected feto pelvic disproportion; history of asthma, glaucoma or cardiac disease.
Interventions	<p>Misoprostol 50 mcg placed in the posterior vaginal fornix 4-hourly (n = 76), versus dinoprostone gel 0.5 mg intracervically 6-hourly (n = 79), unless adequate uterine contractions (3 per 10 minutes lasting &gt; 40 seconds), cervix 3 cm dilated and 100% effaced, or spontaneous rupture of membranes, maximum 3 doses. Oxytocin was used when necessary for augmentation of labour, commencing at least 4 hours after misoprostol or 6 hours after dinoprostone.</p> <p>If the cervix was unchanged and no adequate uterine contractions occurred after 3 doses, the women were offered the option of returning another day for attempted labour induction. For the purposes of analysis, they remained in the group originally allocated.</p> <p>Artificial rupture of membranes was generally performed when the cervix was 3-4 cm dilated and at least 80% effaced. A fetal scalp electrode and intrauterine pressure transducer were placed at the discretion of the attending physician.</p>
Outcomes	Cervical ripening (score improved by > 3) (misoprostol 38/76, dinoprostone 21/79); active labour; caesarean delivery; induction-delivery interval; oxytocin augmentation; instrumental vaginal delivery; dystocia; non-reassuring FHR patterns; uterine tachysystole; uterine hypersystole; uterine hyperstimulation; Apgar score < 6 at 5 minutes; maternal and neonatal complications.
Notes	<p>St Louis, Missouri, USA. July 1994 to December 1995.</p> <p>Private community hospital with more than 35 attending obstetricians. The authors suggest that the high rate of caesarean section in the misoprostol group may be due to the tendency for the obstetricians to react to unfamiliar situations such as uterine hyperstimulation with an experimental drug, by performing caesarean section, rather than pharmacological management as has been described in other studies (eg Wing 1995a, Wing 1995b, Wing 1996).</p>

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**075 Chang 1997**

Methods	"Randomly divided into two groups".
Participants	<p>Membranes were intact, cervical status variable, nulliparous women.</p> <p>Inclusion criteria: term singleton pregnancy; scheduled for induction of labour.</p> <p>Exclusion criteria: known contraindications to vaginal prostaglandins; previous uterine surgery; antepartum haemorrhage; fetal distress; premature rupture of membranes; abnormal lie; cephalopelvic disproportion; maternal illness for which induction of labour was inappropriate.</p>
Interventions	<p>Misoprostol 50 mcg into posterior vaginal fornix 4-hourly, maximum 600 mcg (n = 30) versus dinoprostone 3 mg vaginal tablets 6 hourly, maximum 9 mg (n = 30). Cervical scores were evaluated every 4 hours. If cervical score was 9 or more and uterine contractions inadequate, oxytocin infusion was started.</p>



**075 Chang 1997** (Continued)

Outcomes	Maternal temperature, pulse, blood pressure and side effects; cervical scores after 12 hours (misoprostol mean 9.7, standard deviation 3.1 vs dinoprostone 7.3, 2.5, p < 0.05); induction to delivery interval (16.5, 2.7 vs 25.7, 3.8 hours, p < 0.001); occurrence of spontaneous labour (86% vs 77%); uterine hyperstimulation (13.4% vs 8.9%); meconium staining (10% vs 13%); mode of delivery; maternal complications (none); umbilical artery blood flow velocity waveforms (all normal); cord arterial blood gases (7.29, 0.73 vs 7.32, 0.91); Apgar scores at 1 and 5 minutes ('the same'); neonatal complications.	
Notes	Tainan, Taiwan. July 1994 to June 1995. Some data reported as percentages only. Not clear what denominator was used.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**075 Chuck 1995**

Methods	Open, randomised trial. Computer-generated random allocation using opaque envelopes.
Participants	<p>Membranes were intact or ruptured, cervical status unfavourable. Pregnant women at 35 to 42 weeks' gestation, admitted for induction of labour.</p> <p>Exclusion criteria: non-vertex presentation; uterine scar other than prior low transverse caesarean section; ominous FHR tracing; multiple gestation; complete cervical effacement.</p> <p>Reasons for induction (misoprostol, dinoprostone): post-dates pregnancy (6, 12); oligohydramnios (7, 3); hypertensive disorders (4, 6); growth impairment (1, 4); premature rupture of membranes (16, 11); diabetes mellitus (10, 7); non-reassuring antepartum surveillance (4, 4); other (1, 3).</p> <p>Other characteristics: initial cervical score 3 or less (26, 26); nulliparous (23, 21); prior caesarean section (5, 10).</p>
Interventions	Application 4-hourly if uterine contractions not adequate for up to 5 doses of misoprostol 50 mcg into the posterior vaginal fornix (n = 49), vs dinoprostone 0.5 mg intracervically (n = 50); oxytocin started after 5th dose if progress of labour unsatisfactory; membranes ruptured when cervix 3 cm dilated and fully effaced.
Outcomes	Induction to delivery interval; number of doses (misoprostol mean 1.8 (SD 1.1) vs dinoprostone 2.5 (1.4) ); number of women delivered after a single dose (20/49 vs 11/50); oxytocin augmentation; mode of delivery; fever > 100 degrees Fahrenheit (2/49 vs 4/50); gastrointestinal symptoms; uterine tachysystole (> 5 contractions per 10 minutes without FHR changes); uterine hyperstimulation (frequent or prolonged contractions with FHR decelerations or tachycardia); postpartum haemorrhage; Apgar scores < 7; NICU admission.
Notes	<p>Los Angeles, USA. September 1993 to January 1994. Of 103 women enrolled, 4 (3.9%) were excluded because of protocol violations.</p> <p>Authors conclude misoprostol is "apparently safe ... and may become the drug of choice for induction of labor".</p>
Risk of bias	

**075 Chuck 1995** (Continued)

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**075 Danielian 1999**

Methods	Random numbers allocated by pharmacy.
Participants	Inclusion criteria: 37 to 42 weeks; singleton cephalic presentation; intact membranes. Exclusion criteria: > 1 caesarean section; uterine surgery; significant fetal or maternal medical condition; known prostaglandin hypersensitivity; cervical score 8 or more.
Interventions	Misoprostol 50 mcg vaginally 4-hourly for maximum 4 doses (n = 105), compared with dinoprostone 1 mg vaginally 6-hourly, maximum 3 doses) (n = 106); given until cervical score > 7 or labour ensued or membranes ruptured. Amniotomy when cervical score 8 or more. Oxytocin augmentation if indicated.
Outcomes	Median induction - delivery interval misoprostol 14.4 vs dinoprostone 22.9 hours. Delivered after single dose (81/105 vs 52/106). No vaginal delivery in 12 hours (67/105 vs 91/106). Median visual analogue pain score (n = 18 vs 24) at induction (20 vs 16), before analgesia given (84 vs 66), during second stage (90 vs 77) and at delivery (79 vs 75).
Notes	Clinicians not blinded. No post-randomisation exclusions or withdrawals.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**075 Escudero 1997**

Methods	Open trial. Assigned by means of a randomised table. No further details given.
Participants	Membranes were intact or ruptured, cervical status variable. Pregnant women who required labour induction for obstetric or medical reasons. Inclusion criteria: singleton pregnancy; vertex presentation. Exclusion criteria: labour; fetal distress; previous uterine surgery; contraindication to vaginal delivery. Indications for labour induction: post-dates pregnancy (misoprostol 15, oxytocin 15); pre-eclampsia (22, 30); ruptured membranes (15, 15); fetal demise (4, 0); anencephaly (0, 1); other (1, 2). Unfavourable cervix was not specified as a criterion, though the mean cervical score was low (2.6, SD 1.5).
Interventions	Misoprostol 50 mcg placed in the posterior vaginal fornix 4-hourly until 3 uterine contractions of at least 40 seconds per 10 minutes, maximum 600 mcg; artificial rupture of membranes as soon as possible; arrest in cervical dilation at 5 or more cm cervical dilation managed with oxytocin infusion (n = 57); versus labour induction with oxytocin according to standard protocol of the centre (n = 63). Continuous electronic FHR monitoring in all women. Cervical assessments were repeated every 4 hours. Uterine

**075 Escudero 1997** (Continued)

	hyperstimulation was managed by left lateral positioning, nasal oxygen administration, nifedipine 10 mg sublingually and flushing the misoprostol from the vagina with saline or stopping the oxytocin infusion. Of 22 women with unsuccessful oxytocin labour induction after 24 hours, 9 had labour induced with misoprostol and all had vaginal deliveries without complications.
Outcomes	Cervical score 4 hours after application of the drug (misoprostol mean 5.3, SD 3.6 vs oxytocin 5.5, 3.9); time from induction to delivery (11.3, 6.9 h vs 8.4, 4.1 h); oxytocin use; route of delivery; Apgar scores (9.1, 0.9 vs 9.0, 1.3 at 5 minutes); complications during labour induction and after delivery; uterine hyperstimulation with and without FHR changes.
Notes	Lima, Peru. September 1994 to March 1995. Of 63 women enrolled in the misoprostol group, 3 (4.8%) were excluded because of protocol violations.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

**075 Fuchs 2006**

Methods	"Prospective double-blind, controlled trial" with no reference to allocation concealment methods.
Participants	Inclusion criteria: singleton pregnancies undergoing induction of labour. Exclusion criteria: women with Bishop score < 5, PROM, in labour women, non-reassuring fetal testing and scared uterus.
Interventions	The intervention group received vaginal prostaglandin E1 gel 50 mcg, 4 hourly and continuous infusion of placebo solution intra venally. Control group: intravaginal placebo gel, 4 hourly and oxytocin titrated to a maximum infusion of 30 mU per minute.
Outcomes	Time to delivery was not significantly different in between groups.
Notes	Only abstract available. No data included in this systematic review.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear for data assessment

**075 Fuchs 2006** (Continued)

Incomplete outcome data addressed? All outcomes	Yes	Inadequate
Free of selective reporting?	Unclear	Unclear
Free of other bias?	Unclear	Unclear

**075 Ghidini 2001**

Methods	Allocated by random number generated at the hospital pharmacy.
Participants	Women admitted for induction of labour; singleton pregnancy; 37 weeks' gestation. Exclusion criteria: non-vertex presentation; uterine scar other than from previous lower transverse caesarean section; non-reassuring cardiotocography; cervix dilated > 3 cm; uterine contractions > 3/10 minutes; contraindication for vaginal delivery; hypersensitivity to prostaglandins; parity < 5.
Interventions	Vaginal misoprostol 50 mcg 4-hourly versus 100 mcg 6-hourly until adequate contraction pattern or dilatation > 3 cm or forewater amniotomy or signs of uterine hyperstimulation. Intravenous oxytocin infusion after 24 hours.
Outcomes	Primary: caesarean section. Secondary: vaginal delivery < 24 hours; induction to delivery interval; oxytocin augmentation; fetal and neonatal morbidity; hyperstimulation (tachysystole of 6 contractions/10 minutes for 20 minutes, or hypertonus of a contraction lasting for 2 minutes, with changes in the FHR; caesarean section for fetal distress (2/32 vs 3/26) umbilical artery pH < 7.1 (50 mcg 0/32 vs 100 mcg 1/26).
Notes	Inova Alexandria Hospital, Virginia, USA. Seven of 65 enrolled women excluded due to emergence of exclusion criteria. The groups were somewhat unbalanced (32 50 mcg and 26 100 mcg).

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	
Allocation concealment?	Unclear	Women were assigned by means of a random number generated at the Hospital Pharmacy to receive either misoprostol 100 µg every 6 hours or 50 µg every 4 hours.
Blinding? All outcomes	No	
Incomplete outcome data addressed? All outcomes	No	7 of 65 women withdrawn because misoprostol was not administered for clinical reasons.

**075 Kolderup 1999**

Methods	Assignment in envelopes according to random number tables.
Participants	Inclusion criteria: singleton pregnancy 31 or more weeks; cervical score < 6; vertex presentation; < 12 contractions per hour. Exclusion criteria: prior uterine surgery; ruptured membranes; contraindication for vaginal delivery; parity > 5; non-reassuring fetal surveillance.
Interventions	1. Misoprostol 50 mcg vaginally, repeated 4-hourly if necessary to maximum of 6 doses (81 women); 2. dinoprostone gel 0.5 mg intracervically, repeated 6-hourly if necessary, to a maximum of 4 doses (78 women). Fetal and uterine monitoring for at least 1 hour after treatment. Amniotomy attempted when cervix 3-4 cm dilated. If not in adequate labour 4 hours after last dose or arrest of cervical dilation for more than 2 hours after 4 cm dilation, oxytocin infusion commenced or the woman was crossed over to the other group.
Outcomes	Primary: caesarean delivery, induction to delivery time (misoprostol 19.8 hours SD 11.5 vs dinoprostone 28.9 SD 14.8, $p = 0.005$ ), oxytocin use, hyperstimulation syndrome (tachysystole with fetal bradycardia, or FHR pattern requiring treatment), fetal distress requiring delivery. Secondary outcomes: late decelerations and fetal bradycardia; meconium stained liquor; tachysystole; number of doses used (1.4 SD 1.0 vs 2.2 SD 1.3, $p = 0.0005$ ).
Notes	3 hospitals in California. January 1994 to December 1996.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**075 Lemancewicz 1999**

Methods	Computer-generated randomisation table.
Participants	Inclusion criteria: indication for induction of labour; singleton; cephalic; reactive cardiotocograph; 41 weeks of pregnancy. Exclusion criteria: estimated fetal weight > 4500 g; evidence of cephalopelvic disproportion; placenta praevia; unexplained vaginal bleeding; vasa praevia; renal or hepatic dysfunction; contraindication to prostaglandins; suspected chorioamnionitis; previous uterine surgery.
Interventions	Misoprostol 50 mcg in posterior vaginal fornix 4-hourly till progressive labour, contractions 3 per 10 minutes, ruptured membranes or delivery (maximum 300 mcg, $n = 44$ ); versus intravenous oxytocin at 1 mU per minute, increase every 30 minutes by 2 mU per minute till uterine activity adequate (maximum 17 mU per minute $n = 47$ ).
Outcomes	Doppler velocimetry of umbilical, uterine and arcuate arteries before and 2-3 hours after induction (no significant differences between groups). Induction to delivery (misoprostol 615, SD 65 vs oxytocin 711 SD 70 minutes); caesarean section; FHR changes; meconium.

**075 Lemancewicz 1999** (Continued)

Notes	Bialystok University Hospital, Poland. 3-way study. Misoprostol vs intracervical dinoprostone published as Urban 2003.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

**075 Magtibay 1998**

Methods	Method of randomisation not specified.	
Participants	Pregnant women with an indication for labour induction. Inclusion criteria: singleton; cephalic; live fetus; cervical score < 5. Exclusion criteria: previous caesarean section.	
Interventions	Intravaginal misoprostol 50 mcg 4-hourly (n = 17) compared with intracervical dinoprostone 0.5 mg and oxytocin infusion (n = 19).	
Outcomes	Change in cervical score after 12 hours (misoprostol median 4 versus dinoprostone 1); delivery within 36 hours (15/17 vs 9/19); time to complete dilatation (n = 30; 17 h vs 24 h); caesarean section.	
Notes	Two women allocated to the dinoprostone group refused treatment. Outcome measures assessed blind to allocation.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

**075 Megalo 2004**

Methods	Randomised non-blinded trial with allocation concealment through sealed opaque envelopes.	
Participants	Inclusion criteria: pregnant women between 36 - 42 weeks of gestation that presents indication for cervical ripening, with singleton pregnant, cephalic presentation, normal FHR tracing, unscarred uterus, absence of fetal anomalies, no contraindication for vaginal delivery, absence of chorioamnionitis and no hypersensitivity to prostaglandin.	
Interventions	The intervention group received 50 mcg of vaginal misoprostol 4 hourly (maximum 5 times) and the comparison group received 0.5 mcg of dinoprostone intracervical 6 hourly up to 3 times. If necessary they received 3 mg of dinoprostone intravaginal (maximum 2 times).	

**075 Megalo 2004** (Continued)

Outcomes	Time induction-to-delivery and the need of oxytocin were reduced with misoprostol (p < 0.05). Pathological CTG tracing was more frequent in the misoprostol treated group (p > 0.001).	
Notes	The study was conducted during 18 months in the Department of Obstetrics and Gynaecology, Lausanne University Hospital (CHUV).	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Only blinded for outcome assessors
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**075 Mundle 1996**

Methods	Open, randomised trial. Allocation by sequentially numbered opaque sealed envelopes, prepared from randomisation tables in blocks of four. Caregivers were unaware of the blocking.
Participants	Membranes were intact, cervical status variable. Inclusion criteria: indication for labour induction; single live fetus; gestation greater than 37 weeks; cephalic presentation; intact membranes. Exclusion criteria: non-reassuring FHR tracing; prior uterine surgery; known hypersensitivity to prostaglandins; contraindications to vaginal birth.
Interventions	Misoprostol 50 mcg in the upper vagina 4 hourly until progressive labour, contraction frequency of 3 per 10 minutes, membranes ruptured, non-reassuring FHR tracing, delivery or maximum of 16 doses ( $n = 111$ ); versus physician-chosen combinations of dinoprostone 0.5 mg intracervically for cervical ripening or 1-2 mg intravaginal gel for induction, and oxytocin infusion ( $n = 111$ ). Artificial rupture of membranes in both groups was at the discretion of the attending physician. After membrane rupture, augmentation of labour was by oxytocin infusion. Oxytocin was not allowed within 4 hours of the last misoprostol or 6 hours of the last dinoprostone dose. Continuous electronic FHR and uterine contraction monitoring were used.
Outcomes	Labour induction to vaginal delivery time (misoprostol mean 753, standard deviation 588 vs dinoprostone/oxytocin 941, 506 minutes); oxytocin use (22/111 vs 46/111); method of delivery; epidural use (34/111 vs 55/111); no analgesia (13/111 vs 9/111); meconium; scalp pH done (12/111 vs 9/111); intact

**075 Mundle 1996** (Continued)

	<p>perineum (17/111 vs 18/111).</p> <p>Neonatal assessments blind to group allocation: Apgar scores; cord pH (mean 7.28, standard deviation 0.09 vs 7.28, 0.10); neurological and general physical assessment; birth asphyxia (profound metabolic or mixed acidaemia, 5 minutes Apgar score 3 or less, neonatal neurologic abnormality and dysfunction of one other major body system).</p> <p>Oxytocin use has not been included in the review outcomes because oxytocin use was an option for induction of labour in only the dinoprostone group, and would therefore be expected to be used more frequently in that group.</p>
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Notes	<p>St John's, Newfoundland. Canada. March to September 1994.</p> <p>There were no losses to follow up.</p>
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***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**075 Reyna-Villasmil 2005**

Methods	Randomised trial with allocation method and sequence generation not stated.
Participants	<p>Inclusion criteria: singleton pregnancy, gestational age &gt; 37 weeks, Bishop score &lt; 4, cephalic presentation, vertex and estimated fetal weight &lt; 4 kg.</p> <p>Exclusion criteria: uterine surgery, placenta previa, contraindication to use prostaglandins, fetal distress, herpes simplex active genital infection and abruptio placentae.</p>
Interventions	The intervention group received 50 mcg or 100 mcg vaginal misoprostol 4 hourly (maximum 6 doses each group). Oxytocin augmentation could be used if there were no labour after third dose of misoprostol (4 hours after third dose).
Outcomes	The interval from induction to delivery was shorter in the 100 mcg group ( $p < 0.05$ ). The need of oxytocin augmentation was higher in the 50 mcg group. The caesarean section was double in the 100 mcg group ( $p < 0.05$ ).
Notes	The study was enrolled from June 2002 to November 2003 at the Service of Obstetrics from Central Hospital "Dr. Urquinaona", Maracaibo, Venezuela.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear



**075 Reyna-Villasmil 2005** (Continued)

Blinding? All outcomes	Unclear	Unblinded for personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**075 S-Ramos 1997**

Methods	Open randomised trial. Allocations in numbered, sealed envelopes, in computer-generated random sequence.	
Participants	Membranes were ruptured, cervical status variable. Inclusion criteria: women admitted for induction of labour for term premature rupture of membranes; pregnancy 36 weeks or more; singleton vertex presentation; no evidence of labour; cervical dilation < 2 cm and effacement no more than 80%; reassuring FHR tracing; spontaneous rupture of membranes. Exclusion criteria: intrauterine infection; contraindication to labour or vaginal delivery; previous uterine surgery.	
Interventions	Misoprostol 50 mcg into the posterior vaginal fornix 4-hourly until labour established (at least 3 contractions in 10 minutes), maximum 12 doses, augmented with oxytocin for active labour arrest (n = 70); versus oxytocin infusion commenced at 1-2 mU per minute (n = 71). Continuous FHR monitoring in all women. Intrauterine pressure catheter and scalp electrode applied as soon as possible. Episodes of hyperstimulation were treated with position change, oxygen therapy and tocolysis with terbutaline 250 mcg subcutaneously. Fetal scalp blood sampling was performed when indicated.	
Outcomes	Delivery within 24 hours; uterine tachysystole, hypertonus and hyperstimulation; chorioamnionitis (misoprostol 4/70, oxytocin 5/71); postpartum endometritis; retained placenta (1/70, 2/71).	
Notes	Jacksonville, Florida, USA. November 1992 to October 1993.	
Risk of bias		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**075 Sahin 2002**

Methods	Computer-generated random number table, in numbered, sealed envelopes.	
Participants	Inclusion criteria: women with pre-eclampsia/eclampsia in whom induction of labour was anticipated; cervical score < 5. Exclusion criteria: abnormal lie; non-cephalic presentation; abnormal FHR pattern; abnormal umbilical diastolic velocities; multiple pregnancy; previous caesarean section; antepartum haemorrhage; premature rupture of membranes; expected cephalopelvic disproportion; maternal illness which contraindicated induction with prostaglandins.	
Interventions	Misoprostol 50 mcg vaginally 4-hourly (maximum 4 doses), versus oxytocin infusion at 1-30 mU per minute.	
Outcomes	Cervical score at 12 hours; induction to delivery time, Apgar scores; in labour within 12 hours; caesarean sections; admission to NICU.	
Notes	Women not in labour after 12 hours were excluded from further analysis in the report. As this would introduce bias, only the caesarean section rates, which could be re-calculated for all women, have been included for this review.	
<i><b>Risk of bias</b></i>		
<b>Item</b>	<b>Authors' judgement</b>	<b>Description</b>
Allocation concealment?	Yes	Adequate

**075 Saleen 2006**

Methods	“The study designed as a randomised trial” with sequence generation and allocation concealment methods not stated.	
Participants	Women with singleton alive pregnancies and Bishop score 5 requiring induction of labour between 37 to 42 weeks gestational.	
Interventions	There were 3 groups: A. 50 mcg vaginal misoprostol 4 hourly (maximum 200 mcg); B. 3 mg vaginal dinoprostone pessary 6 hourly (2 times); C. supra-cervical Foley catheter. Each patient could use oxytocin augmentation.	
Outcomes	There were no significant difference in outcomes between those groups.	
Notes	The mechanical method are accessed in a specific review. The trial was conducted at Hamdard University Hospital and Patel Hospital from July 2004 to June 2005.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description

**075 Saleen 2006** (Continued)

Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**075 Tabor 1995**

Methods	“Prospective randomised trial”.	
Participants	Membranes were intact or ruptured, cervical status unfavourable. Women with cervical score < 5.	
Interventions	Misoprostol 50 mcg administered vaginally 4-hourly, maximum 12 doses (n = 68) vs prostaglandin E2 gel 0.5 mg intracervically 6-hourly, maximum 3 doses (n = 59). Oxytocin commenced when cervix favourable or having > 3 uterine contractions per 10 minutes.	
Outcomes	Induction to delivery times; oxytocin use; caesarean sections; maternal and neonatal morbidity.	
Notes	Fort Worth, Texas, USA. Authors conclude that misoprostol shortens induction time without increased morbidity.	
<i>Risk of bias</i>		
Item	Authors’ judgement	Description
Allocation concealment?	No	Inadequate

**075 Urban 2003**

Methods	Allocation by computer-generated randomisation table, consecutively numbered sealed opaque envelopes.
Participants	Inclusion criteria: singleton pregnancy; reassuring FHR tracing; 41 weeks’ gestation; cervical score < 6. Exclusion criteria: multiple pregnancy; malpresentation; estimated fetal weight > 4500 g; evidence of cephalopelvic disproportion; placenta praevia; unexplained vaginal bleeding; vasa praevia; renal or hepatic dysfunction; allergy to prostaglandins; suspected chorioamnionitis; previous caesarean delivery; uterine surgery.

**075 Urban 2003** (Continued)

Interventions	Misoprostol 50 mcg vaginally 4-hourly (maximum 300 mcg) versus dinoprostone 0.5 mg intracervically 6-hourly (maximum dose 1 mg), until progressive labour or contraction frequency of 3 in 10 minutes, or ruptured membranes or delivery. Oxytocin infusion was used for failure to progress in labour after adequate cervical ripening. Continuous FHR monitoring was used.
Outcomes	Doppler velocimetry of umbilical, uterine and arcuate arteries; mean time to delivery (misoprostol 615 (SEM 65) versus dinoprostone 772 (82) minutes, $p = 0.14$ ); oxytocin augmentation; caesarean section; umbilical artery pH; abnormal FHR patterns; uterine hyperstimulation.
Notes	3-way trial, data on misoprostol versus oxytocin published previously as Lemancewicz 1999. No mention in 1999 paper that it was a three-way trial, but communication with first author of 2003 report confirmed that was the case.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**075 Webb 1997**

Methods	"Randomised", no details given (abstract report). 100 enrolled women reviewed.
Participants	Inclusion criteria: indication for labour induction in third trimester; cervical score < 6. Exclusion criteria: contraction frequency > 6 per hour; multiple pregnancy; breech; uterine scar other than single lower transverse; parity > 4.
Interventions	Prostaglandin E2 4 mg vaginal suppositories 4 mg to posterior fornix vs dinoprostone 0.5 mg intracervical gel vs misoprostol 50 µg to posterior fornix; 4-hourly until cervix 2 cm dilated, contraction frequency 3 per 10 minutes or membrane rupture associated with contractions. Oxytocin used when indicated, only after discontinuation of the study drug.
Outcomes	Time to 4 cm cervical dilation: prostaglandin E2 19.4 hours vs dinoprostone 17.7 vs misoprostol 10.8; time to delivery 27 hours vs 30.5 vs 15.3; oxytocin requirement 86% vs 82% vs 56%; caesarean section 35% vs 41% vs 23%. Dinoprostone use was associated with significantly fewer episodes of terbutaline administration and stated hyperstimulation. Misoprostol use was associated with significantly less fever.
Notes	No comment on exclusions. Data in abstract inadequate for inclusion in tables. Further data awaited for inclusion.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

### 075 Zeteroglu 2004

Methods	Randomised trial. Allocation method and sequence generation not stated.
Participants	Inclusion criteria: grand multiparas, Bishop score < 6 and gestational age $\geq$ 35 weeks. Exclusion criteria: Abnormal lie, non-cephalic presentation, abnormal heart rate patterns, abnormal umbilical diastolic velocities, gestational age < 35 weeks, history of previous caesarean section, antepartum haemorrhage, expectation of cephalopelvic disproportion and any maternal illness or contraindication for prostaglandins.
Interventions	The intervention group received 50 mcg of vaginal misoprostol 4 hourly (maximum 4 doses) and the comparison group received oxytocin alone.
Outcomes	The mean time from induction to delivery was significantly shorter in the misoprostol group ( $p = 0.02$ ).
Notes	University of Yüzüncü Yıl, Medical Faculty, Van, Turkey.

#### *Risk of bias*

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

### 075 Zeteroglu 2006a

Methods	Randomised trial with allocation not stated. Computer generation of random assignment.
Participants	Inclusion criteria: singleton pregnancies, gestational age > 37 weeks, no contraindication for induction of labour, premature rupture of membranes > 24 hours. Exclusion criteria: previous labour induction failure, non-cephalic presentation, chorioamnionitis, prior uterine surgery, contraindication to prostaglandins.
Interventions	The intervention group received 50 mcg of vaginal misoprostol 4 hourly (maximum 200 mcg) and the comparison group received oxytocin alone.

075 Zeteroglu 2006a (Continued)

Outcomes	The mean time from induction to delivery was shorter in the misoprostol group (p = 0.063).	
Notes	The trial was conducted during 2 years. University of Yünzücü Yil, Medical Faculty, Van, Turkey.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

075 Zeteroglu 2006b

Methods	Randomised trial with allocation by sealed opaque numbered envelopes. Sequence generation made by computer.	
Participants	Inclusion criteria: grand multiparas pregnancies ( $\geq 10$ pregnancies), admission for whom after 35 weeks to induction of labour, well dated with US ( $< 22$ weeks) and Bishop $\leq 6$ . Exclusion criteria: abnormal position, non cephalic presentation, abnormal heart rate pattern, $< 35$ gestational weeks, multiple pregnancies, history of caesarean section, antepartum haemorrhage, expectation of cephalopelvic disproportion or any maternal illness or which induction was contraindicated with prostaglandins.	
Interventions	The intervention group received 50 mcg vaginal misoprostol with oxytocin augmentation if necessary. The comparison group received oxytocin associated with amniotomy.	
Outcomes	There was no difference between the groups in rates of vaginal delivery and mean time from induction to delivery.	
Notes	University of Yünzücü Yil, Medical Faculty, Van, Turkey.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description

**075 Zeteroglu 2006b** (Continued)

Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**075 Zeteroglu 2006c**

Methods	Randomised trial. Computer generated random number table. Allocation method not described.
Participants	Inclusion criteria: advanced maternal age (> 35 years), Bishop score < 6, indication for induction of labour at 35 weeks or more, singleton and vertex presentation. Exclusion criteria: abnormal lie, non-cephalic presentation, abnormal umbilical diastolic velocities, age below 35 weeks, multiple pregnancies, previous caesarean section, antepartum haemorrhage, expectation of cephalopelvic disproportion, or any maternal contraindication to use prostaglandins.
Interventions	The intervention group received 50 mcg vaginal misoprostol 4 hourly (maximum: 200 mcg) and the comparison group received oxytocin alone.
Outcomes	The induction to delivery interval was significantly shorter in the misoprostol group. The vaginal delivery rate was higher in the misoprostol group.
Notes	University of Yüzüncü Yıl, Medical Faculty, Van, Turkey.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Unclear only for outcome assessors
Incomplete outcome data addressed? All outcomes	Yes	Inadequate

**075 Zeteroglu 2006c** (Continued)

Free of selective reporting?	No	Inadequate
Free of other bias?	No	Inadequate

**088 Garry 2003**

Methods	Randomised trial with allocation concealment made according to a computer-generated schedule prepared by a hospital pharmacy.
Participants	Inclusion criteria: pregnant women with obstetric indication for IOL, Bishop $\leq 4$ , cephalic presentation and reassuring fetus status. Exclusion criteria: known fetal structural or chromosomal anomalies, fetal death, refusal to participate, multiple gestation and any contraindication to vaginal delivery.
Interventions	The intervention group received 50 mcg of misoprostol 3 hourly until 400 mcg. The comparison group received dinoprostone 10 mg vaginal repeated if was necessary after 12 hours.
Outcomes	The interval from start of induction to vaginal delivery was significantly shorter in the misoprostol group ( $794.5 \pm 408$ minutes vs $1005.3 \pm 523$ minutes; $p < 0.02$ ). A non reassuring FHR tracing was the indication for 71.4% of caesarean deliveries in the misoprostol group, compared to 40% in the dinoprostone group ( $p = 0.03$ ). There were no significant differences in neonatal outcomes.
Notes	Winthrop University Hospital, Mineola, NY, USA. The study ran from 1 April 1998 to 6 June 1999.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Yes	Adequate
Blinding? All outcomes	No	Only blinded for personnel
Incomplete outcome data addressed? All outcomes	Yes	Inadequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate



**088 Pi 1999**

Methods	Randomised by computer sequence generation. Allocation methods not stated.
Participants	Inclusion criteria: term pregnancy, low platelets, premature rupture of membranes and women needed to IOL. Exclusion criteria: scarred uterus, antepartum haemorrhage, abnormal cardiotocography, transverse lie, fetus not engaged, patients with asthma and with cardiac problem.
Interventions	Interventions groups used 50 mcg rectal or vaginal misoprostol 3 hourly until effective contraction. Comparison group used oxytocin alone.
Outcomes	There was no difference on use of vaginal or rectal misoprostol for labour induction. The results indicate that misoprostol had a higher rate of successful induction for labour than oxytocin.
Notes	October 1997 to May 1998. Department of Obstetrics and Gynaecology.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unclear
Incomplete outcome data addressed? All outcomes	No	Inadequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	No	Inadequate

**088 Saggaf 2001**

Methods	Sealed envelopes with computer-generated randomisation.
Participants	Inclusion criteria: women admitted for induction of labour; cephalic presentation; singleton; intact membranes; cervical score < 6; reassuring cardiotocography. Exclusion criteria: previous uterine surgery; antepartum haemorrhage; ruptured membranes; malpresentation; estimated fetal weight > 4500 g; chorioamnionitis; known hypersensitivity to prostaglandins.
Interventions	Misoprostol 50 mcg vaginally every 3 hours, maximum 6 doses, versus dinoprostone 3 mg vaginally 6-hourly, maximum 3 doses. If no labour, oxytocin started 3 hours after last misoprostol or 6 hours after last dinoprostone dose. There was continuous cardiotocography during labour.

**088 Saggaf 2001** (Continued)

Outcomes	Mode of delivery; time from insertion to labour (misoprostol 6.3 SD 3.4 vs dinoprostone 10.3 SD 7.1 hours; time from labour to delivery (4.9 SD 2.6 vs 5.6 SD 4.9 hours); tachysystole (6 or more contractions per 10 minutes for 20 minutes); hypertonus (a contraction lasting 2 or more minutes; hyperstimulation (tachysystole or hypersystole with non-reassuring cardiotocograph); Apgar scores (5 min 9.4 SD 0.6 vs 8.3 SD 1.9).	
Notes	King Fahad Military Hospital, Saudi Arabia, April 1999 to August 2000.	
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**088 Sanchez Ramos 1998**

Methods	Computer-generated randomisation table, maintained in the research section of the hospital pharmacy. Physicians were not blinded to allocation. Review of labour tracings was not stated to have been performed blindly. Analysis was according to intention to treat.
Participants	Inclusion criteria: obstetric or medical indications for labour induction; no active labour; no fetal distress; no previous uterine surgery; singleton pregnancy; vertex presentation; no contraindication to vaginal delivery; informed consent (n = 223).
Interventions	<p>1. Misoprostol 50 µg to the posterior vaginal fornix 3-hourly until adequate uterine contractions (maximum 8 doses); oxytocin for arrest in the active phase of labour, versus</p> <p>2. dinoprostone 10 mg vaginal insert in the posterior fornix, which was removed when adequate uterine contractions or hyperstimulation occurred, or after 12 hours.</p> <p>Continuous electronic FHR and uterine activity monitoring. As soon as cervical dilation permitted, artificial rupture of the membranes was performed and an intrauterine pressure catheter and scalp electrode applied. If not in labour after 24 hours, oxytocin was used. Uterine hyperstimulation was managed with left lateral positioning, removal of the tablet or insert, oxygen administration and subcutaneous terbutaline 250 µg.</p>
Outcomes	Induction to vaginal delivery interval (misoprostol median 698, interquartile range 395-1053 vs dinoprostone 1041, 792-1531, p < .001); induction to delivery (699, 395-1053 vs 1053, 780-1590, p < .001; successful induction (98/108 vs 77/115, p < .001; vaginal delivery within 12 hours (44/108 vs 22/115, p < .001); vaginal delivery within 24 hours; need for oxytocin; maximum oxytocin rate (8, 4-13.2 vs 13, 6-20 mU/minute, p < .001); uterine tachysystole; hyperstimulation; need for scalp pH sampling (4/108 vs 5/115); assisted delivery; caesarean section; cord pH (mean 7.29 +/- 0.09 vs 7.30 +/- 0.08); cord pH < 7.16 (9/97 vs 4/99); Apgar score < 7 at 1 minute (11/108 vs 8/115) and 5 minutes; admission to NICU; maternal side effects (none noted in either group); average costs of induction (\$85 vs \$606).
Notes	University of Florida Health Sciences Centre, Jacksonville. February to October 1996.
Risk of bias	

**088 Sanchez Ramos 1998** (Continued)

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**088 Wing 1995a**

Methods	Allocation by sequentially numbered sealed opaque envelopes, maintained by the primary investigator. Sequence from random number table in blocks of 6.
Participants	<p>Membranes were intact, cervical status unfavourable. Inclusion criteria: women requiring induction of labour for medical or obstetric reasons; singleton pregnancy; cephalic presentation; intact membranes; cervical score &lt; 5; reactive nonstress test; &lt; 4 uterine contractions per hour.</p> <p>Exclusion criteria: estimated fetal weight &gt; 4500 g; evidence of cephalopelvic disproportion; placenta praevia; unexplained vaginal bleeding; vasa praevia; active herpes simplex infection; contraindication to receiving prostaglandins; renal or hepatic dysfunction; suspected chorioamnionitis; previous uterine surgery; parity &gt; 5.</p> <p>Indications for labour induction: oligohydramnios (misoprostol 37, dinoprostone 41); pre-eclampsia (14, 5); post-term pregnancy (5, 9); macrosomia (4, 5); abnormal antepartum testing (3, 1); other (5, 6).</p>
Interventions	<p>Misoprostol 50 mcg inserted into the posterior vaginal fornix 3-hourly, maximum 6 doses (n = 68) versus dinoprostone gel 0.5 mg intracervically 6-hourly, maximum 3 doses (n = 67), until 3 uterine contractions per 10 minutes, cervical score &gt; 7, cervical dilation &gt; 3 or spontaneous rupture of membranes. Artificial rupture of membranes usually performed when the cervix was 80% effaced and 3 cm dilated. Oxytocin augmentation was used for lack of contractions after maximum dosage or spontaneous rupture of membranes, or for arrested cervical dilation, &gt; 3 hours after misoprostol and &gt; 6 hours after dinoprostone. Uterine hyperstimulation was treated in some cases by tocolytic therapy.</p>
Outcomes	<p>Oxytocin augmentation; uterine tachysystole (&gt; 5 contractions per 10 minutes); uterine hypertonus (contraction &gt; 2 minutes) (misoprostol 1/68, dinoprostone 2/67); hyperstimulation syndrome (tachysystole or hypertonus with FHR abnormalities); induction to delivery interval; mode of delivery; vaginal delivery within 24 hours; terbutaline used for hyperstimulation (6/68, 3/67); diarrhoea (2/68, 2/67); fever (2/68, 0/67); neonatal resuscitation (15/68, 5/67); days in NICU (mean 11.2 (SD 8.1), 5.8 (2.4)); meconium aspiration syndrome (3/68, 1/67); hyperbilirubinaemia (8/68, 2/67).</p>
Notes	<p>Los Angeles, California, USA. October to November 1993.</p> <p>135/140 women agreed to participate. There were no withdrawals from the protocol.</p> <p>The authors postulate that the increased rate of meconium-stained amniotic fluid may be due to the increased incidence of uterine tachysystole, or to a direct effect of misoprostol on the fetal gastrointestinal tract.</p> <p>The authors do not recommend this dosage for induction of labour.</p>

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

## 100 Fletcher 1994

Methods	Randomised by pulling sealed odd and even numbers from a box.
Participants	Membranes were intact, cervical status variable. Women scheduled for induction of labour. Exclusion criteria: contraindications to vaginal prostaglandins (including uterine scar); antepartum haemorrhage; fetal distress; premature rupture of the membranes; abnormal lie; cephalopelvic disproportion; maternal illness for which induction of labour was contraindicated. Indications for induction of labour: hypertension (misoprostol 11, dinoprostone 13); postdates (10, 11); diabetes (4, 3); other (7, 4).
Interventions	Insertion into the posterior vaginal fornix of misoprostol 100 mcg (n = 32) vs dinoprostone 3 mg (n = 31). If not in labour after 12 hours, oxytocin was commenced (sometimes delayed because of staff shortage).
Outcomes	Cervical score 12 hours after drug insertion (misoprostol mean 9.1 (SD 1.1) vs dinoprostone 7.7 (2.4)); insertion to delivery interval; spontaneous labour; uterine hyperstimulation (> 5 contractions per 10 minutes with fetal bradycardia); mode of delivery; maternal complications; Apgar scores (5 minutes mean 8.8 vs 9.1); perinatal death (excluding one induction of labour for intrauterine death).
Notes	Kingston, Jamaica. September to October 1992. One woman (1/32, 3%) excluded from dinoprostone group because of inadvertently receiving misoprostol. The authors conclude that larger studies are needed to confirm safety.

### *Risk of bias*

Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

## 100 Herabutya 1997

Methods	"Blocked randomisation".
Participants	Membranes were intact or ruptured, cervical status unfavourable. Women with indications for labour induction and unfavourable cervixes.
Interventions	Misoprostol 100 mcg placed in the posterior vaginal fornix (n = 60), versus prostaglandin E2 1.5 mg in gel placed in the endocervix (n = 50). If not in labour after 24 hours, labour induced with amniotomy and oxytocin.
Outcomes	Induction to delivery interval (misoprostol mean 19.1, SD 10.6 vs prostaglandin E2 21.4, 13.1 hours); induction of labour after 24 hours (5/60 vs 13/50); oxytocin augmentation; caesarean section; uterine hyperstimulation; Apgar scores; admission to NICU.
Notes	Bangkok, Thailand. Data from abstract; full report awaited.

### *Risk of bias*

Item	Authors' judgement	Description
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**100 Herabutya 1997** (Continued)

Allocation concealment?	No	Inadequate
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**100 Howarth 1996**

Methods	Consecutively numbered opaque sealed envelopes in computer-generated random sequence. The trial medication was administered by an investigator not involved in ongoing care of the women. Labour ward staff were blind to the allocation.
Participants	Membranes were intact, cervical status unfavourable. Women due for induction of labour; singleton pregnancy; cephalic presentation; no fetal distress; estimated fetal weight > 2000 g; intact membranes; unfavourable cervix. Exclusion criteria: contraindication to vaginal delivery; previous caesarean section; parity > 4; contraindication to prostaglandin administration including asthma or glaucoma. Indications for induction of labour: hypertension (misoprostol 16, dinoprostone 18); post-dates pregnancy (15, 9); other (5, 9).
Interventions	Administration into posterior vaginal fornix (and repeated after 6 hours if the cervix remained unfavourable) of misoprostol 100 mcg vs dinoprostone 1 mg.
Outcomes	Frequent uterine contractions (> 5 per 10 minutes); uterine hyperstimulation (frequent contractions with suspicious or ominous changes in the FHR pattern); analgesia use (misoprostol 24/36 vs dinoprostone 26/36); oxytocin augmentation; delivery within 6 (12/36 vs 3/36) and 12 hours (30/36 vs 13/36); not in labour within 12 hours; induction to delivery interval; caesarean section; Apgar score at 5 minutes. FHR tracings were examined blind to the group allocation.
Notes	Pretoria, South Africa. April to June 1995. The authors conclude that larger trials are needed to address dosage and safety.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**100 Montealegre 1999**

Methods	Double blind, double placebo randomised trial.
Participants	Inclusion criteria: indication for labour induction (pre labour rupture of membranes, hypertension, prolonged pregnancy); cervical score 6 or less; consent. Exclusion criteria: previous caesarean section or other uterine surgery; chorioamnionitis; dystocia; fetal distress; placenta praevia; medical and surgical contraindications to labour; multiple pregnancy, parity > 5.
Interventions	Misoprostol 100 mcg vaginally plus intravenous lactated Ringer solution at 2 mU per minute, increasing 2 mU every 20 minutes (maximum 30 mU or 8 hours), versus oxytocin solution plus placebo tablet.

**100 Montealegre 1999** (Continued)

Outcomes	Time to delivery; method of delivery; uterine hypersystole, fetal distress.	
Notes		
<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**100 Ozsoy 2004**

Methods	Randomised unblinded trial. Allocation not stated.	
Participants	Inclusion criteria: women with > 39 completed weeks singleton gestation. Exclusion criteria: previous uterine scar, malpresentation, cervix dilated > 3cm, uterine contraction > 3/10 minutes, any contraindication to vaginal delivery, hypersensitivity to prostaglandins, parity > 5, abnormal antepartum testing, cephalopelvic disproportion, premature rupture of membranes and maternal illnesses.	
Interventions	The intervention group received 100 mcg 6 hourly and the comparison group received 50 mcg 4 hourly of vaginal misoprostol until labour.	
Outcomes	In two groups the dose used of misoprostol were similar. There was no difference between two groups in mean time to delivery, caesarean rate, Apgar of 5 minutes and meconium passage.	
Notes	Department of Obstetrics and Gynaecology, Isparta, Turkey.	
<i><b>Risk of bias</b></i>		
<b>Item</b>	<b>Authors' judgement</b>	<b>Description</b>
Adequate sequence generation?	Yes	Adequate
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	No	Inadequate
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Unclear	Unclear

**100G Fletcher 1993**

Methods	Randomised, double blind trial. Coded sachets of powdered medication used. Code broken at end of study.
Participants	Membranes were intact, cervical status unfavourable. Women due for induction of labour in the third trimester with unfavourable cervix; no contraindication to prostaglandin use. Indications for induction of labour: post dates pregnancy (misoprostol 12, placebo 11); pre-eclampsia (7, 5); pre-eclampsia with intrauterine death (1, 1); diabetes mellitus (1, 2); other (3, 2).
Interventions	Powdered misoprostol 100 mcg (n = 24) vs ethinyl estradiol 0.05 mg ('placebo') (n = 21), each mixed with hydroxyethyl gel 2.7 mg, administered with a syringe into the posterior vaginal fornix; if not in labour after 12 hours, oxytocin induction was commenced or planned.
Outcomes	Results were as follows (misoprostol vs placebo, mean values (standard deviation) or proportions): insertion to delivery in hours (15.6 (12.5) vs 43.2 (20.5)); no improvement in cervical score (3/24 vs 13/21); oxytocin used (7/24 vs 13/21), all significant $p < 0.05$ . Complications: instrumental vaginal delivery (1/24 vs 1/21); caesarean section (2/24 vs 3/21); meconium-stained liquor (2/24 vs 0/21); fetal tachycardia (0/24 vs 2/21); uterine hyperstimulation (1/24 vs 0/21); postpartum haemorrhage (1/24 vs 0/21).
Notes	Kingston, Jamaica. Three women allocated to the placebo group excluded because of damage to the sachets. The authors conclude that misoprostol is effective and safe for cervical ripening.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

**100G Gottschall 1997**

Methods	Allocated by sequentially numbered, sealed opaque envelopes. Used random number tables in permuted blocks of 6.
Participants	Inclusion criteria: intended pre-induction cervical ripening or labour induction; singleton fetus, cephalic presentation; intact membranes; reactive FHR tracing; no contraindications to vaginal delivery. Out of 262 women for induction of labour, 75 enrolled. Exclusion criteria: uterine scar; known allergy to prostaglandins.
Interventions	Misoprostol 100 µg or prostaglandin E2 (dinoprostone) gel prepared by the hospital pharmacy 5 mg, into the posterior vaginal fornix. After 6 hours if in labour, amniotomy; if not, oxytocin 0.6 to 20 mU per minute.
Outcomes	Primary outcome: time to delivery (misoprostol 14.7 +/- 6.4 hours vs prostaglandin E2 20.4 +/- 10.2). Secondary outcomes: need for oxytocin; change in cervical score at 6 hours; uterine tachysystole (6 per 10 minutes); uterine hypertonus (> 2 minutes); hyperstimulation syndrome.
Notes	New Britain General Hospital, Connecticut, November 1995 to August 1996.

**100G Gottschall 1997** (Continued)

<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**100G Srisomboon 1996**

Methods	Allocation by "blocked randomisation".
Participants	Inclusion criteria: obstetric or medical indication for labour induction; singleton pregnancy; parity 3 or less; vertex presentation; cervical score 4 or less; gestational age > 35 weeks. Exclusion criteria: labour; fetal distress; previous uterine surgery; definite cephalopelvic disproportion; history of 2nd or 3rd trimester n = haemorrhage; contraindication to use of prostaglandins.
Interventions	1. Misoprostol 100 mcg crushed and mixed with 5 ml hydroxyethyl cellulose gel into the posterior vaginal fornix (32). 2. Gel alone (30). All women received continuous cardiotocography. Oxytocin was started after 12 hours if not in labour.
Outcomes	Change in cervical length measured by transvaginal ultrasound (24 (SD 9.1) vs 2.2 (4.3), $p < .001$ ); cervical score (Bishop 1964) 12 hours after gel insertion (8.1 (2.7) vs 0.9 (1.2), $p < .001$ ); time to vaginal delivery (h) (12.0 (8.3) vs 25.5 (6.7), $p < 0.001$ ); analgesia (23/32 vs 22/30, NS); fetal distress (2/32 vs 2/30, NS); 5-minute Apgar score (9.9 (0.4) vs 9.9 (0.4), NS); side effects; uterine hyperstimulation.
Notes	

<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**100G Srisomboon 1997**

Methods	Allocation by "blocked randomisation".
Participants	Membranes were intact, cervical status unfavourable. Inclusion criteria: singleton pregnancy; parity > 4; vertex presentation; obstetric or medical indication for labour induction; intact membranes with no prior stripping; cervical score > 5; gestation > 35 weeks. Exclusion criteria: labour; fetal distress; previous uterine surgery; evidence of cephalopelvic disproportion; placenta praevia; cord presentation; vasa praevia; contraindication to the use of prostaglandins. Indications for labour induction: gestational hypertension/pre-eclampsia (intravaginal 19, intracervical 12); impaired fetal growth (13, 13); post-term pregnancy (14, 20); other (4, 5).



**100G Srisomboon 1997** (Continued)

Interventions	<p>Misoprostol gel (100 mcg crushed and mixed with 3 ml hydroxyethyl cellulose gel in a 10 ml syringe fitted with a nylon feeding tube, 8" 5 FR), instilled under vision into the posterior vaginal fornix (n = 50) versus within the endocervical canal during slow withdrawal of the feeding tube (n = 50).</p> <p>Continuous external cardiotocography in all women. No oxytocin, pelvic examination or amniotomy within 12 hours of instillation. After 12 hours, if the cervix remained unfavourable, misoprostol was repeated; if the cervical score was &gt; 6, an amniotomy was performed and oxytocin infusion instituted if necessary, starting at 1-2 mU per minute. Oxytocin was also used if there were no cervical changes or regular uterine contractions after the second dose of misoprostol.</p> <p>The definition of hypertonus was a uterine contraction &gt; 90 rather than 120 seconds. Uterine hyperstimulation with FHR changes was treated with left lateral positioning, oxygen by nasal catheter, and terbutaline 250 mcg intravenously or subcutaneously.</p>
Outcomes	Cervical score 12 hours after misoprostol administration (intravaginal mean 10.1, SD 2.7, intracervical 9.9, 2.9) insertion to vaginal delivery (16.4, 8.6 vs 17.0, 8.6); vaginal delivery in 24 hours; uterine hyperstimulation without FHR changes; oxytocin use; analgesia (40/5 vs 38/50).
Notes	Chiang Mai, Thailand. August 1994 to September 1995.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

**138 Nunes 1999**

Methods	Randomisation using sealed, opaque envelopes in computer-generated sequence.
Participants	Inclusion criteria: term singleton pregnancy, cervical score < 5, reassuring FHR pattern; exclusion criteria: previous caesarean section or other uterine scar, vaginal bleeding, parity > 4, polyhydramnios, cephalopelvic disproportion, previous fetal death, fetal growth restriction.
Interventions	Vaginal misoprostol 100 mcg, if necessary 50 or 100 mcg after 3 hours, 50 mcg after a further 6 hours (n = 95); versus vaginal dinoprostone 2 mg, if necessary 2 or 1 or 0.5 mg after 6 hours, 1 mg after a further 6 hours (n = 94).
Outcomes	Change in cervical score (no significant difference), caesarean deliveries; interval from initial dose to the active phase of labor (9.8 +/- 5.8 and 14.2 +/- 10.2 hours, p < .01), interval from initial dose to delivery (15.3 +/- 9.8 and 19.1 +/- 13.2 hours, p = .027) for the misoprostol and dinoprostone groups, respectively.
Notes	

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

## 150 De la Torre 2001

Methods	Allocation by hospital pharmacy using computer-generated randomisation schedule. Attendants not blinded after randomisation. FHR patterns were analysed blind to allocation.
Participants	<p>Inclusion criteria: medical indication for labour induction; single live fetus; cephalic presentation.</p> <p>Exclusion criteria: abnormal FHR tracing; uterine contractions every 5 minutes or less; previous uterine surgery; uterine overdistension; intra amniotic infection; hypersensitivity to prostaglandins; medical contraindications; contraindication to vaginal delivery.</p> <p>Use of fetal scalp sampling, and tocolytic therapy in response to non-reassuring FHR tracings was left to the discretion of the managing physician.</p> <p>Indications for induction of labour: pre-eclampsia (misoprostol 23, oxytocin 29); post-term pregnancy (21, 15); diabetes mellitus (1, 1); oligohydramnios (8, 6); other (7, 15).</p>
Interventions	<p>Misoprostol 100 mcg in the posterior vaginal fornix every 4 hours until adequate uterine contractions achieved (maximum 5 doses), versus intravenous oxytocin commencing at 1 mU per minute, increase by 1 mU every 30 minutes to achieve adequate uterine activity (&gt; 200 Montevideo units), maximum 36 mU per minute.</p> <p>Women in the misoprostol group received oxytocin augmentation if required, more than 4 hours after the last dose of misoprostol. Analysis was by intention to treat.</p> <p>All women were monitored continuously with electronic tocodynamometry, intrauterine pressure monitoring in 52%. Amniotomy was generally performed when cervical dilation was about 3-4 cm.</p>
Outcomes	<p>Primary: caesarean section. Secondary: vaginal delivery in 24 hours (data from Kramer 1997 used); epidural analgesia (data from Kramer 1997 used); induction to delivery time; maternal and neonatal outcomes; uterine tachysystole (&gt; 5 contractions/10 minutes for 20 minutes); hypertonus (a contraction lasting 2 minutes); hyperstimulation syndrome (tachysystole or hypertonus with FHR abnormality); caesarean section for fetal distress (misoprostol 23/168 vs oxytocin 15/192); use of terbutaline (Kramer 1997: misoprostol 12/60, oxytocin 2/66); abnormal FHR patterns; meconium stained amniotic fluid; epidural analgesia; Apgar scores; admission to NICU.</p>
Notes	<p>University of New Mexico Health Sciences Centre, June 1995 to July 1998. Of 410 enrolled, 50 withdrawn for protocol deviation (16), patient withdrawal (7), or missing data (27). The final groups differed in numbers (misoprostol 168, oxytocin 192). This raises the possibility of selective withdrawal from the misoprostol group.</p> <p>In the abstract report, 58 women in the misoprostol group are reported on rather than 60.</p>

### *Risk of bias*

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

## 150 Kulshreshtha 2007

Methods	"A total of 40 women were randomly selected". The sequence generation and the allocation concealment were not stated.
Participants	<p>Inclusion criteria: primigravida and multigravida, gestational age <math>\geq 34</math> weeks, singleton gestation, cephalic presentation with indication for induction of labour.</p> <p>Exclusion criteria: abnormal FHR, multigravida (more than 3), cephalopelvic dispro-</p>

**150 Kulshreshtha 2007** (Continued)

	portion, multiple pregnancy, unexplained vaginal bleeding, previous uterine surgery, malpresentation and contraindication to use prostaglandin.
Interventions	The intervention group received 100 mcg vaginal misoprostol 6 hourly (maximum 6 doses). The comparison group received 0.5 mg intra-cervically dinoprostone 4 hourly (maximum 6 doses).
Outcomes	The induction delivery interval was shorter in the misoprostol group ( $p < 0.05$ ). There were very few maternal side effects in this study and no differences between the groups in neonatal outcomes.
Notes	Department of Obstetrics & Gynaecology and Pharmacology of S. N. Medical College and Hospital, Agra.

***Risk of bias***

Item	Authors' judgement	Description
Adequate sequence generation?	Unclear	Unclear
Allocation concealment?	Unclear	Unclear
Blinding? All outcomes	Unclear	Unblinded only for personnel
Incomplete outcome data addressed? All outcomes	No	Adequate
Free of selective reporting?	Yes	Adequate
Free of other bias?	Yes	Adequate

**150 Ngai 2000**

Methods	Allocation using sealed envelopes.
Participants	Inclusion criteria: singleton pregnancy; > 7 weeks; cephalic presentation; ruptured membranes. Exclusion criteria: previous caesarean section; meconium-stained liquor; allergy to prostaglandins.
Interventions	Misoprostol 100 mcg 4-hourly (maximum 3 doses; if no contractions after 12 hours or poor progress, syntocinon used; versus oxytocin 1 mU per minute, increasing every 15 minutes (maximum 32 mU per minute).
Outcomes	See analyses.
Notes	86 women enrolled. One excluded (undiagnosed breech presentation). Cervical scores missing on 5 women.

## 150 Ngai 2000 (Continued)

<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	Yes	Adequate

## 175 Kadanali 1996

Methods	"Random" allocation by sealed envelopes.
Participants	Membranes were intact or ruptured, cervical status unfavourable. Women with medical or obstetric indications for labour induction. Inclusion criteria: singleton vertex presentation; 37 to 42 weeks' gestation. Exclusion criteria: active labour; fetal distress; cervical score > 5. Indications for labour induction: post-dates pregnancy (misoprostol 42, oxytocin 50); pre-eclampsia (28, 22); premature rupture of membranes (10, 14); diabetes (6, 4); impaired fetal growth (8, 6); other (18, 16).
Interventions	Misoprostol 100 mcg into the posterior vaginal fornix, repeated orally every 2 hours until labour established (3 contractions in 10 minutes); oxytocin if not in active labour after 24 hours (n = 112); versus dinoprostone (Cerviprost) instilled into the cervix; oxytocin commenced if after 6 hours if indicated according to a uniform protocol. All women had continuous electronic FHR monitoring. Uterine hyperstimulation was managed by changing the mother's position to left lateral, oxygen by nasal catheter and intravenous ritodrine at 0.3 mg/minute.
Outcomes	Labour induction to delivery interval (misoprostol mean 9.2, SD 2.4 vs dinoprostone/oxytocin 15.2, 3.2 hours); cervical score after 6 hours (6.5, 3.2 vs 6.0, 3.6); 5 cm cervical dilation to delivery interval (1.6, 1.2 vs 7.8, 2.4 hours); fetal distress (4/112 vs 4/112); delivered within 12 hours (72/112 vs 28/112); uterine hyperstimulation with and without FHR changes; vacuum delivery; caesarean section; Apgar score < 5 at 5 minutes (2/112 vs 2/112); cord pH < 7.16 (8/112 vs 10/112).
Notes	Erzurum, Turkey. March to August 1995.

<i>Risk of bias</i>		
Item	Authors' judgement	Description
Allocation concealment?	No	Inadequate

## 200 Lee 1997

Methods	Allocated by coded numbers in sealed envelopes. An independent clinician administered the allocated vaginal medication according to a master list. The rest of the staff were kept blind to the treatment given.
Participants	Inclusion criteria: para 3 or less; singleton pregnancy; cephalic presentation; no previous caesarean section; no contraindication to prostaglandin therapy; uncomplicated pregnancy; cervical score 6 or less.

**200 Lee 1997** (Continued)

Interventions	Misoprostol 200 mcg tablet versus dinoprostone 3 mg, inserted vaginally 6-hourly, maximum 2 doses. The cervix was assessed every 6 hours. When 'ready for labour', transferred to the labour ward. If no labour ensued, oxytocin given. If the cervix remained unfavourable after 24 hours, caesarean section was performed.
Outcomes	Established labour (misoprostol 23/25 vs dinoprostone 16/25); induction to delivery (676 SD 411 vs 875 SD 406 minutes); cervical score increase over 6 hours (3.3 SD 2 vs 2.0 SD 2); delivered within 6 hours (5/25 vs 3/25); delivered within 12 hours (18/25 vs 7/25); second dose of prostaglandin used (10/25 vs 17/25); estimated blood loss (180 SD 48 vs 246 SD 336 ml).
Notes	Kuantan General Hospital, Malaysia.

***Risk of bias***

Item	Authors' judgement	Description
Allocation concealment?	Unclear	Unclear

**200 Rowlands 2001**

Methods	Random allocation using consecutively numbered sealed opaque envelopes.
Participants	Inclusion criteria: nulliparous; singleton; cephalic; gestation > 37 weeks; no known contraindication to vaginal delivery; cervical score < 6. Exclusion criteria: known medical complications in pregnancy; fetal compromise; previous uterine surgery; active labour; rupture of membranes; previous attempt at induction of labour; contraindication to prostaglandins.
Interventions	Misoprostol 200 mcg vs dinoprostone 2 mg pessary vaginally; repeated if necessary after 24 and 30 hours. When cervical score 6 or more, membranes artificially ruptured and oxytocin administered. Continuous cardiotocography.
Outcomes	Primary: induction to vaginal delivery interval (misoprostol 926 SD 569 vs dinoprostone 1578 SD 791); vaginal delivery < 12 hours (45/49 vs 36/47); duration of active labour to vaginal delivery (354 SD 221 vs 497 SD 266). Secondary outcomes: repeated doses (1/49 vs 16/47); artificial rupture of membranes; oxytocin augmentation; mode of delivery; non-reassuring cardiotocograph; side effects; Apgar score; admission to NICU; epidural analgesia (21/49 vs 29/47); blood loss at delivery (263 SD 168 vs 268 SD 138).
Notes	January 1996 to November 1998. Three Australian obstetric units. Women not blinded. Obstetrician responsible for labour care not informed of group allocation. One woman excluded from the misoprostol group after enrolment because cervical score 7.

***Risk of bias***

Item	Authors' judgement	Description
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**200 Rowlands 2001** (Continued)

Allocation concealment?	Yes	Adequate
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CTG: cardiotocograph

FHR: fetal heart rate

HELLP: haemolysis, elevated liver enzymes and low platelets

ICU: intensive care unit

IOL: induction of labour

mcg: micrograms

mU: milliunit

NICU: neonatal intensive care unit

NS: non-stress

para: paragraph

sem: standard error of the mean

SD: standard deviation

vs: versus

**Characteristics of excluded studies** [ordered by study ID]

Study	Reason for exclusion
Adair 1998	Comparison of oral versus vaginal misoprostol. For consideration for inclusion in the review of oral misoprostol for labour induction.
Aggarwal 2006	Excluded because of methodological inconsistencies.
Arrieta 2000	Randomised comparison of misoprostol in escalating dosages with oxytocin for labour induction. Excluded because the first dose of misoprostol was intracervical, not vaginal.
Azeem 2006	Excluded because of methodological inconsistencies.
Balintona 2001	Provisionally excluded pending full report, because data not available in usable format (abstract only). Misoprostol 50 mcg 6-hourly x 2 doses vs Prepidil 0.5 mg 6-hourly x 2 doses vs Cervidil 10 mg insert for 12 hours, intravaginally. Adverse cardiotocographic event in 50% vs 14 % vs 11.1 %).
Belfrage 2000	Excluded because of exclusions from analysis. 4/110 women were excluded from the misoprostol group and 10/100 from the dinoprostone group due to lack of compliance with the protocol, and a further 8 in the dinoprostone group were excluded from further analysis because of failure of cervical ripening after 24 hours.
Bi 2000	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Bolnick 2002a	Excluded because misoprostol 25 mcg 4-hourly was compared with oxytocin, with concurrent dinoprostone (Cervidil) in both groups. There were no differences between the groups.

(Continued)

Bugalho 1995	Allocation not random. A comparison of vaginal misoprostol and oxytocin for induction of labour.
Butler 2004	For consideration in the oral misoprostol for induction of labour.
Cecatti 2001	Abstract only. Brazilian 25 mcg tablet (Prostokos) compared with 1/8 200 mcg Cytotec tablet, 6-hourly during the day for 48 hours. No differences in effectiveness.
Cecatti 2006	The study compares 2 presentations of misoprostol (the vaginal specific 25 mcg presentation and the oral tablet divided in small parts:1/8 of oral presentation 200 mcg). Both groups used the same regimen and doses.
Cetin 1997	Excluded because data not in prespecified format. Random allocation to misoprostol 100 mcg (n = 34) versus dinoprostone 0.5 mg intracervically (n = 34). Induction was 'successful' in 31/34 vs 28/34. Induction delivery interval was shorter with misoprostol (8.57 SD 4.03 vs 11.12 SD 2.07 hours). Uterine hyperstimulation occurred in 2/34 vs 1/34 respectively.
Chang 2003	Excluded because misoprostol was administered intracervically, not vaginally.
Chen 2000	Excluded because data analysis was not based on the intention to treat.
Chen 2001	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Chen 2003	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Chen 2004	For consideration in the review of augmentation methods.
Cui 2001	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Dai 2005	Excluded because data analysis was not based on the intention to treat.
Delaney 2001	Excluded provisionally, pending full report (insufficient information in abstract).
Ding 2001	Excluded because data analysis was not based on the intention to treat.
Ding 2005	Excluded because it is a brief communication with a lack of information on methods. No information concerning randomisation.
Ding 2006	Excluded because data analysis was not based on the intention to treat.
Du 2000	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Dundas 2000	Excluded because no prespecified outcomes given in this abstract publication. To await fuller report. Misoprostol 50 mcg, followed after 12 hours by 50 mcg 6-hourly (maximum 4 doses), versus dinoprostone 2

(Continued)

	mg repeated if necessary after 12 hours and 1 mg after a further 6 hours. Allocation 'at random'. 11/257 withdrawn because of incomplete data.
Duru 1997	Excluded because pregnancy termination in the second trimester was studied.
Echeverria 1995	'Randomised comparative study' - excluded because method of randomisation not specified - ?alternation. Inclusion criteria: postdates pregnancy (41 to 42 weeks, singleton, cephalic presentation, no cephalopelvic disproportion. Exclusion criteria: previous uterine surgery, placental insufficiency diagnosed by fetal monitoring and amniotic fluid volume). Interventions: vaginal misoprostol 100 mcg vs syntocinon infusion 2 to 32 mU per minute. Outcomes: delivery within 24 hours, time from induction to delivery, polysystole. Setting: Alto Riesgo Obstetric Clinic, 1 December 1994 to 30 April 1995.
Eftekhavi 2002	Comparison of vaginal misoprostol with oxytocin for labour induction. Excluded because only percentages given in published abstract. Awaiting full report.
El-Din 2000	Excluded provisionally, pending full report (insufficient information in abstract).
Escalante 1993	Excluded because does not fit the pre-stated comparisons of this review. Labour was induced 'randomly' by either vaginal (n = 68) or intracervical (n = 32) misoprostol 100 mcg, repeated if necessary (in 4 women) after 24 hours. No statistically significant differences regarding cervical ripening and pregnancy outcome were found, though most of the data presented are for the whole group of 100 women. Maternal side effects occurred in 4 women. Uterine hyperstimulation occurred in 11 women, of whom 1 developed fetal distress which resolved with tocolytic therapy. There were 12 caesarean sections.
Fonseca 2007	This study was excluded because of the inclusion of preterm pregnancies which were induced.
Girija 2006	Only abstract available. There is no information on inclusion criteria, randomisation and allocation concealment.
Gorzelac 1999	Excluded because there are no methodological details.
Gorzelac 2001	Excluded because there is no information on randomisation.
Harms 2001	Excluded provisionally, pending full report (insufficient information in abstract).
Hoesli 2003	Abstract report only. Random allocation to misoprostol or dinoprostone for labour induction. Ultrasound measurement of cervical length. Vaginal delivery within 24 hours was compared for those with cervical length 0-28 mm vs 29-50 mm. Unable to use data as percentages only given. Awaiting full report.
How 2001	Abstract only. Excluded because the comparison group was oral misoprostol. For consideration in the oral misoprostol review
Hu 2005	Excluded because data analysis was not based on the intention to treat.
Jackson 1999	Excluded provisionally because results not available in the abstract.
Jazayeri 2003	Excluded because the inclusion/exclusion criteria are unclear and the results are presented using only percentages. Information on the size of each group is not stated.



(Continued)

Jouatte 2000	Excluded because a retrospective study.
Kwon 1999	For consideration for inclusion in the review of oral misoprostol.
Li 2003	Excluded because data analysis was not based on the intention to treat.
Li XQ 2003	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Liu 1998	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Liu 2004	Excluded because data analysis was not based on the intention to treat.
Lulu 1999	Randomisation and concealment are not reported in the text, only in the abstract.
Luo 2000	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Majoko 2002b	For consideration in the review of mechanical methods of labour induction.
Megalo 1998	Abstract only. Excluded because the comparison group included both intracervical and intravaginal prostaglandins. There was an unexplained imbalance between the group size (89 vs 73). Full publication awaited.
Megalo 1999	Excluded because no clinical outcomes reported. Comparison of cardiotocogram findings following labour induction with misoprostol or PGE2.
Molina 2000	Excluded provisionally, pending full report (insufficient information in abstract).
Ngai 1996	Excluded because misoprostol administered orally, not vaginally. To be considered for inclusion in the review 'Oral misoprostol for induction of labour'.
Nuthalapaty 2005	Women between 14 and 24 weeks of pregnancy were included in this trial.
Ozgur 1997	Excluded because 'randomisation' was according to odd and even hospital numbers. Large risk of allocation bias.
Patel 2000	Excluded because misoprostol used for augmentation, not induction of labour.
Perry 1998	Excluded because comparison was between vaginal misoprostol and a combination of an intracervical balloon catheter and dinoprostone.
Perry 1999	Excluded because categorical data not given in abstract. Awaiting full report.
Porojanova 2005	Excluded because there no reference to methods and to primary outcomes.

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Roy 2003	Study on the medical termination of pregnancies between 15 and 23 weeks of pregnancy.
Rust 1999	For consideration for inclusion in the review of mechanical methods of labour induction.
Rust 2000	For consideration for inclusion in the review of mechanical methods of labour induction.
Sabra 2000	Excluded provisionally, pending full report (insufficient information in abstract).
Sanchez Ramos 1993	<p>Excluded because the intervention oxytocin preceded by prostaglandins when necessary was not a pre-stated comparison.</p> <p>Random allocation by consecutively numbered, sealed opaque envelopes. Sequence generated by coin toss. Participants: membranes were intact or ruptured, cervical status variable. Women with obstetric or medical indications for induction of labour; singleton pregnancy; vertex presentation.</p> <p>Indications for labour induction: post-dates pregnancy (misoprostol 14, oxytocin 10); pre-eclampsia (22, 22); diabetes (3, 6); abnormal fetal testing (3, 9); ruptured membranes (9, 8); other (13, 10).</p> <p>Exclusion criteria: active labour; fetal distress; previous uterine surgery; contraindication to vaginal delivery.</p> <p>Interventions: misoprostol 50 mcg introduced into the posterior vaginal fornix 4-hourly until adequate labour was achieved (maximum 12 doses, maximum actually used 4 doses); arrest of labour progress at 5 or more cm cervical dilation was managed with oxytocin augmentation; compared with oxytocin infusion commencing at 1-2 mU per minute (preceded by cervical ripening with prostaglandin E2 gel if cervical score &lt; 5, in 29/65 women).</p> <p>In both groups, artificial rupture of membranes was performed and intrauterine pressure monitoring and scalp electrode monitoring applied as soon as cervical dilation permitted.</p> <p>Fetal scalp sampling was performed for persistent FHR changes.</p> <p>Outcomes: undelivered 24 hours after initiating misoprostol or oxytocin; frequent uterine contractions (&gt; 5 per 10 minutes); hyperstimulation (frequent contractions or prolonged contraction of 2 or more minutes, with fetal tachycardia, late decelerations or reduced short-term variability).</p> <p>Jacksonville, Florida, USA. January to August 1992.</p> <p>Of 130 women enrolled, 1 (0.8%) was excluded after randomisation because of breech presentation.</p> <p>The authors conclude that the trial corroborates the effectiveness and apparent safety of misoprostol for labour induction.</p>
Sanchez Ramos 2002	<p>Excluded because not a pre-specified comparison. Misoprostol moistened with 1 ml 3% acetic acid compared with dry misoprostol, 50 mcg 4-hourly vaginally, maximum 6 doses. Data on 162/177 randomised women given. No significant differences in time from start to vaginal delivery (moistened mean 1004, SD 636 versus dry 1130, 636 minutes); oxytocin (45/82 vs 48/80); no vaginal delivery within 24 hours 23/82 vs 32/80); hyperstimulation (6/82 vs 8/80); Caesarean section (12/82 vs 12/80); Apgar &lt;7 at 5 minutes (1/82 vs 1/80); NICU admission (13/82 vs 14/80).</p>
Sharma 2005	2 women undergoing caesarean section before the second dose of misoprostol excluded from the analysis.
Sheela 2006	Excluded because no information was provided on the randomisation status, eligibility criteria or allocation strategy.
Shen 2003	Excluded because data analysis was not based on the intention to treat.
Shi 2003a	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.

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Shi 2003b	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Su 1998	Excluded because data analysis was not based on the intention to treat. The randomisation and concealment were not reported in the text, only in the abstract.
Su 2003	Excluded because data analysis was not based on the intention to treat.
Thach 2000	Excluded provisionally, pending full report (insufficient information in abstract).
Tian 2003	Excluded because data analysis was not based on the intention to treat.
Toppozada 1997	For consideration for inclusion in review 'Oral misoprostol for induction of labour'. Women with cervical score < 5 were randomly allocated to induction of labour with vaginal misoprostol 100 mcg 3-hourly, increasing to 200 mcg after 2 doses (maximum 1 mg), versus oral misoprostol 100 mcg 3-hourly, increased to 200 mcg after the first dose. The induction to delivery interval was: vaginal mean 7.15 (standard deviation 4.39) hours vs oral 9.93 (3.68) ; side effects (nausea and vomiting) 2/20 vs 4/20; uterine hyperstimulation 8/20 vs 0/20; FHR changes 10/20 vs 1/20; caesarean section: 2/20 vs 4/20; instrumental vaginal delivery: 4/20 vs 2/20.
Varaklis 1994	Excluded provisionally because no results available in the abstract.
Wang 2000	Excluded because data analysis was not based on the intention to treat.
Wang 1997a	Excluded because the comparison group management does not fit into any of the pre-defined comparison categories. Women with cervical score 4 or below, intact membranes and requiring labour induction were 'randomly divided' into 2 groups of 30. The study group received misoprostol 50 mcg vaginally 3-hourly until labour was established (maximum 3 doses). The control group received 40 ml ricinus oil and 5 fried egg yolks; if not in labour after 12 hours, oxytocin was used; if the induction failed (not in labour after 24 hours) , misoprostol was used. The number of women in labour with cervix 2 cm or more dilated after 24 hours was 28/30 for the study and 23/30 for the control group. The time from induction to vaginal delivery was 12.2 +/-3.5 vs 18.1 +/- 3.2 hours respectively. Oxytocin during labour was used in 4/30 vs 12/30. Caesarean section was performed in 6/30 vs 6/30. Uterine hyperstimulation occurred in 5/30 vs 1/30. Diarrhoea or vomiting occurred in 2/30 vs 4/30, and low-grade fever in 2/30 vs 0/30.
Wang 1997b	No predefined outcomes for this review reported. Vaginal misoprostol 100 mcg (n = 43) compared with placebo (n = 42). Oxytocin used after 12 hours if not in labour. The changes in cervical score at 12 hours were 4.4 +/- 2.2 vs 1.0 +/- 0.9; labour within 12 hours 29/43 vs 6/42; and medication to labour time 17.2 +/- 21.1 vs 40.6 +/- 26.0 hours respectively. No changes were noted in fetal blood flow indices. There were no differences in histology of 13 and 5 placentae studied. Uterine hyperstimulation occurred in 4/43 women in the misoprostol group.
Wang 1997c	Excluded because data analysis was not based on the intention to treat.
Wang 2005	Excluded because data analysis was not based on the intention to treat.

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Wicker 1995	Study reported as an abstract only. Data are not available in a format suitable for analysis. The first author has been written to for further information. Intravaginal misoprostol gel 25 mcg 6 hourly was compared with 0.5 mg intracervical dinoprostone gel 6 hourly (maximum 3 doses), in 117 women with cervical scores of 5 or less and reassuring antenatal testing. Women receiving misoprostol had higher cervical scores after the first dose, shorter time from induction to oxytocin use and lower number of doses needed. No differences in complications were noted.
Wilk 2001	Excluded because the analysis was not based on intention to treat.
Windrim 1997	Excluded because misoprostol administered orally, not vaginally. To be considered for inclusion in the review 'Oral misoprostol for induction of labour'.
Wing 1999	For consideration for inclusion in the 'oral misoprostol' review.
Yang 2000	Excluded because data analysis was not based on the intention to treat.
Young 2001	Excluded provisionally, pending full report (insufficient information in abstract).
Zang 1997	Excluded because data analysis was not based on the intention to treat.
Zang 2003	Excluded because data analysis was not based on the intention to treat.
Zhao 2003	Excluded because data analysis was not based on the intention to treat.
Zhu 1998	Excluded because data analysis was not based on the intention to treat.
Zhuang 2000	Excluded because allocation not randomised.

vs: versus

## Characteristics of studies awaiting assessment *[ordered by study ID]*

### Abedi-Asl 2007

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Bebbington 2003a**

Methods	Randomised controlled trial.
Participants	No description, only abstract available.
Interventions	Vaginal misoprostol 50 mcg 4 hourly until 3 doses vs dinoprostone 10 mg 4 hourly until 3 doses.
Outcomes	Presence of tachysystole, no references to primary outcomes of the review.
Notes	Awaiting full report.

**Bebbington 2003b**

Methods	Randomised controlled trial.
Participants	No description, only abstract available.
Interventions	Vaginal misoprostol 50 mcg 4 hourly until 3 doses vs dinoprostone 10 mg 4 hourly until 3 doses.
Outcomes	Interval to active labour, need of oxytocin augmentation, mean of infused dose, route of delivery, Apgar score and NICU admissions.
Notes	Awaiting full report.

**Bolnick 2002b**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Bricker 2007**

Methods	"Randomized" study.
Participants	Women with > 34 weeks of pregnancy with PROM.
Interventions	"Misoprostol regimen" versus "vaginal dinoprostone and/ or intravenous oxytocin".
Outcomes	Rates of caesarian section and vaginal delivery not archived in 24h.
Notes	Awaiting full report for more precise methodological and intervention descriptions.

**Bricker 2008**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Deng 1999**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**ElSedeek 2009**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Gupta 2006**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Li 2000**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Lughmani 2009**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Milchev 2003**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Moodley 2003**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Nigam 2010**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Ozkan 2009**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Pevzner 2008**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Pevzner 2009a**

Methods	
Participants	
Interventions	
Outcomes	
Notes	



**Pevzner 2009b**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Prager 2008**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Tabasi 2007**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Wang 2000**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Wing 2008**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Yin 2006**

Methods	
Participants	
Interventions	
Outcomes	
Notes	

**Characteristics of ongoing studies** *[ordered by study ID]***Botero 1998**

Trial name or title	Oxytocin versus misoprostol for labour induction: a double blind randomised controlled trial.
Methods	
Participants	Pregnant women, aged between 15 and 45 years old. Indications for induction of labour: premature rupture of membranes at term; pregnancy-induced hypertension at term; prolonged pregnancy; normal FHR tracing.
Interventions	Intravenous oxytocin, continuous infusion, 2 mU per minute (maximum 16 mU per minute) versus intravaginal misoprostol 100 mcg single dose.
Outcomes	Cesarean section, time to delivery, Apgar, fetal distress, tachysystole.
Starting date	July 1995.
Contact information	Luis Botero - lfbotero@jkavercol.javeriana.edu.co.
Notes	

**Gregson 2003**

Trial name or title	To compare the safety and efficacy of low dose vaginal misoprostol and dinoprostone vaginal gel for induction of labour at term
Methods	Randomised trial.
Participants	Women at term (37 - 42 completed weeks of gestation), single fetus, cephalic presentation, membrane may be intact or ruptured and reactive fetal heart tracing.
Interventions	IOL with vaginal misoprostol.
Outcomes	Uterine tachysystole, hyperstimulation, meconium stained liquor, Apgar score at 5 minutes, umbilical arterial pH and base deficit, neonatal unit admission, induction-delivery interval, method of delivery, Bishop score at onset of labour, oxytocin requirements, mode of delivery and analgesia requirements in labour.
Starting date	01 of July 2000 to 31 of December of 2003.
Contact information	Ms Sarah Gregson - Queen Mary's Sidcoup NHS Trust.
Notes	Awaiting full publication.

**Jackson 2000**

Trial name or title	
Methods	
Participants	
Interventions	
Outcomes	
Starting date	
Contact information	
Notes	

FHR: fetal heart rate  
IOL: induction of labour  
mU:

## DATA AND ANALYSES

### Comparison 1. Misoprostol versus placebo/no treatment: all women

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved in 24 hours	5	769	Risk Ratio (M-H, Random, 95% CI)	0.56 [0.31, 1.03]
2 Uterine hyperstimulation with FHR changes	5	777	Risk Ratio (M-H, Fixed, 95% CI)	2.38 [0.95, 5.99]
3 Caesarean section	10	1141	Risk Ratio (M-H, Fixed, 95% CI)	0.81 [0.63, 1.05]
4 Neonatal encephalopathy	1	150	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Cervix unfavourable/unchanged after 12-24 hours	2	107	Risk Ratio (M-H, Random, 95% CI)	0.10 [0.01, 0.64]
6 Oxytocin augmentation	5	429	Risk Ratio (M-H, Random, 95% CI)	0.62 [0.38, 1.02]
7 Uterine hyperstimulation without FHR changes	6	794	Risk Ratio (M-H, Fixed, 95% CI)	3.52 [1.78, 6.99]
8 Uterine rupture	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
9 Instrumental vaginal delivery	3	184	Risk Ratio (M-H, Fixed, 95% CI)	1.07 [0.65, 1.77]
10 Meconium-stained liquor	6	814	Risk Ratio (M-H, Fixed, 95% CI)	0.56 [0.35, 0.87]
11 Apgar score < 7 at 5 minutes	4	717	Risk Ratio (M-H, Fixed, 95% CI)	2.0 [0.34, 11.80]
12 Neonatal intensive care unit admission	6	852	Risk Ratio (M-H, Fixed, 95% CI)	0.94 [0.60, 1.48]
13 Perinatal death	2	122	Risk Ratio (M-H, Fixed, 95% CI)	0.34 [0.01, 8.14]
14 Maternal side effects	1	62	Risk Ratio (M-H, Fixed, 95% CI)	2.82 [0.12, 66.62]
15 Postpartum haemorrhage	3	184	Risk Ratio (M-H, Fixed, 95% CI)	0.95 [0.19, 4.62]
16 Serious maternal complication	3	272	Risk Ratio (M-H, Fixed, 95% CI)	0.68 [0.12, 3.87]
17 Maternal death	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable

### Comparison 2. Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved in 24 hours	4	619	Risk Ratio (M-H, Random, 95% CI)	0.70 [0.23, 2.15]
2 Uterine hyperstimulation with FHR changes	4	627	Risk Ratio (M-H, Fixed, 95% CI)	2.05 [0.73, 5.71]
3 Caesarean section	7	862	Risk Ratio (M-H, Fixed, 95% CI)	0.95 [0.69, 1.30]
4 Cervix unfavourable/unchanged after 12-24 hours	2	107	Risk Ratio (M-H, Random, 95% CI)	0.10 [0.01, 0.64]
5 Oxytocin augmentation	2	107	Risk Ratio (M-H, Fixed, 95% CI)	0.39 [0.26, 0.58]
6 Uterine hyperstimulation without FHR changes	5	714	Risk Ratio (M-H, Fixed, 95% CI)	3.47 [1.63, 7.38]
7 Uterine rupture	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
8 Instrumental vaginal delivery	2	107	Risk Ratio (M-H, Fixed, 95% CI)	1.02 [0.50, 2.12]

9 Meconium-stained liquor	4	612	Risk Ratio (M-H, Fixed, 95% CI)	0.53 [0.31, 0.89]
10 Apgar score < 7 at 5 minutes	3	567	Risk Ratio (M-H, Fixed, 95% CI)	2.0 [0.34, 11.80]
11 Neonatal intensive care unit admission	3	505	Risk Ratio (M-H, Fixed, 95% CI)	0.84 [0.35, 2.05]
12 Perinatal death	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
13 Maternal side effects	1	62	Risk Ratio (M-H, Fixed, 95% CI)	2.82 [0.12, 66.62]
14 Postpartum haemorrhage	2	107	Risk Ratio (M-H, Fixed, 95% CI)	0.91 [0.13, 6.37]
15 Serious maternal complication	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
16 Maternal death	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable

### Comparison 3. Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved in 24 hours	2	112	Risk Ratio (M-H, Fixed, 95% CI)	0.40 [0.22, 0.70]
2 Uterine hyperstimulation with FHR changes	3	227	Risk Ratio (M-H, Fixed, 95% CI)	2.31 [0.52, 10.16]
3 Caesarean section	5	355	Risk Ratio (M-H, Fixed, 95% CI)	1.16 [0.75, 1.79]
4 Epidural analgesia	1	50	Risk Ratio (M-H, Fixed, 95% CI)	0.98 [0.77, 1.26]
5 Cervix unfavourable/unchanged after 12-24 hours	2	107	Risk Ratio (M-H, Random, 95% CI)	0.10 [0.01, 0.64]
6 Oxytocin augmentation	2	107	Risk Ratio (M-H, Fixed, 95% CI)	0.39 [0.26, 0.58]
7 Uterine hyperstimulation without FHR changes	3	167	Risk Ratio (M-H, Fixed, 95% CI)	10.11 [1.91, 53.60]
8 Uterine rupture	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
9 Instrumental vaginal delivery	2	107	Risk Ratio (M-H, Fixed, 95% CI)	1.02 [0.50, 2.12]
10 Meconium-stained liquor	2	105	Risk Ratio (M-H, Fixed, 95% CI)	0.71 [0.28, 1.77]
11 Apgar score < 7 at 5 minutes	1	60	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
12 Neonatal intensive care unit admission	2	105	Risk Ratio (M-H, Fixed, 95% CI)	0.41 [0.04, 3.70]
13 Perinatal death	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
14 Maternal side effects	1	62	Risk Ratio (M-H, Fixed, 95% CI)	2.82 [0.12, 66.62]
15 Postpartum haemorrhage	2	107	Risk Ratio (M-H, Fixed, 95% CI)	0.91 [0.13, 6.37]
16 Serious maternal complication	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
17 Maternal death	1	45	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable

**Comparison 4. Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	1	150	Risk Ratio (M-H, Fixed, 95% CI)	4.0 [0.46, 34.96]
2 Vaginal delivery not achieved in 24 hours	2	257	Risk Ratio (M-H, Fixed, 95% CI)	0.48 [0.31, 0.73]
3 Caesarean section	4	386	Risk Ratio (M-H, Fixed, 95% CI)	0.58 [0.36, 0.91]
4 Apgar score < 7 at 5 minutes	2	257	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Neonatal intensive care unit admission	2	227	Risk Ratio (M-H, Fixed, 95% CI)	1.03 [0.07, 15.82]
6 Neonatal encephalopathy	1	150	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
7 Oxytocin augmentation	2	202	Risk Ratio (M-H, Random, 95% CI)	0.62 [0.29, 1.32]
8 Serious maternal complications	2	227	Risk Ratio (M-H, Fixed, 95% CI)	0.68 [0.12, 3.87]
9 Perinatal death	1	77	Risk Ratio (M-H, Fixed, 95% CI)	0.34 [0.01, 8.14]
10 Instrumental delivery	1	77	Risk Ratio (M-H, Fixed, 95% CI)	1.12 [0.56, 2.22]
11 Postpartum haemorrhage	1	77	Risk Ratio (M-H, Fixed, 95% CI)	1.03 [0.07, 15.82]
12 Meconium-stained liquor	3	309	Risk Ratio (M-H, Fixed, 95% CI)	0.83 [0.38, 1.81]
13 Uterine hyperstimulation without FHR changes	1	107	Odds Ratio (M-H, Fixed, 95% CI)	0.76 [0.17, 3.38]

**Comparison 5. Misoprostol versus placebo/no treatment: all primiparae**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	1	39	Risk Ratio (M-H, Fixed, 95% CI)	1.09 [0.49, 2.41]

**Comparison 6. Misoprostol versus placebo/no treatment: all primiparae and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	1	39	Risk Ratio (M-H, Fixed, 95% CI)	1.09 [0.49, 2.41]

**Comparison 7. Misoprostol versus placebo/no treatment: all primiparae, intact membranes and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	1	39	Risk Ratio (M-H, Fixed, 95% CI)	1.09 [0.49, 2.41]

**Comparison 9. Misoprostol versus placebo/no treatment: all multiparous and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	1	29	Risk Ratio (M-H, Fixed, 95% CI)	0.62 [0.06, 6.05]

**Comparison 10. Misoprostol versus placebo/no treatment: all multiparous, intact membranes and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	1	29	Risk Ratio (M-H, Fixed, 95% CI)	0.62 [0.06, 6.05]

**Comparison 11. Misoprostol versus vaginal prostaglandin: all women**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	22	5229	Risk Ratio (M-H, Random, 95% CI)	0.77 [0.66, 0.89]
2 Uterine hyperstimulation with FHR changes	31	5830	Risk Ratio (M-H, Random, 95% CI)	1.43 [0.97, 2.09]
3 Caesarean section	34	6855	Risk Ratio (M-H, Fixed, 95% CI)	0.95 [0.87, 1.03]
4 Serious neonatal morbidity or perinatal death	3	500	Risk Ratio (M-H, Fixed, 95% CI)	5.98 [0.25, 145.59]
5 Serious maternal morbidity or death	2	223	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
6 Cervix unfavourable/unchanged after 12-24 hours	2	303	Risk Ratio (M-H, Random, 95% CI)	0.75 [0.39, 1.44]
7 Oxytocin augmentation	38	7022	Risk Ratio (M-H, Random, 95% CI)	0.68 [0.61, 0.76]
8 Uterine hyperstimulation without FHR changes	26	4804	Risk Ratio (M-H, Random, 95% CI)	1.99 [1.41, 2.79]
9 Uterine rupture	5	1464	Risk Ratio (M-H, Fixed, 95% CI)	1.00 [0.06, 15.88]
10 Epidural analgesia	8	2141	Risk Ratio (M-H, Fixed, 95% CI)	0.92 [0.85, 0.99]

11 Instrumental vaginal delivery	19	3593	Risk Ratio (M-H, Fixed, 95% CI)	1.03 [0.89, 1.18]
12 Meconium-stained liquor	18	3991	Risk Ratio (M-H, Fixed, 95% CI)	1.35 [1.13, 1.61]
13 Apgar score < 7 at 5 minutes	17	3969	Risk Ratio (M-H, Fixed, 95% CI)	1.01 [0.72, 1.41]
14 Neonatal intensive care unit admission	20	4530	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.82, 1.15]
15 Neonatal encephalopathy	1	360	Risk Ratio (M-H, Fixed, 95% CI)	5.98 [0.25, 145.59]
16 Perinatal death	6	1315	Risk Ratio (M-H, Fixed, 95% CI)	0.85 [0.24, 3.00]
17 Serious maternal complications	2	303	Risk Ratio (M-H, Fixed, 95% CI)	0.5 [0.05, 5.39]
18 Maternal side effects (eg nausea, vomiting, diarrhoea)	10	2698	Risk Ratio (M-H, Fixed, 95% CI)	0.94 [0.87, 1.02]
19 Postpartum haemorrhage	8	1385	Risk Ratio (M-H, Fixed, 95% CI)	0.89 [0.65, 1.21]

## Comparison 12. Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	18	4491	Risk Ratio (M-H, Random, 95% CI)	0.77 [0.67, 0.89]
2 Uterine hyperstimulation with FHR changes	26	5010	Risk Ratio (M-H, Fixed, 95% CI)	1.48 [1.17, 1.87]
3 Caesarean section	28	5832	Risk Ratio (M-H, Fixed, 95% CI)	0.95 [0.86, 1.05]
4 Serious neonatal morbidity or perinatal death	5	1344	Risk Ratio (M-H, Fixed, 95% CI)	0.92 [0.23, 3.70]
5 Uterine rupture	2	844	Risk Ratio (M-H, Fixed, 95% CI)	1.00 [0.06, 15.88]
6 Cervix unfavourable/unchanged after 12-24 hours	1	197	Risk Ratio (M-H, Fixed, 95% CI)	0.52 [0.27, 0.98]
7 Oxytocin augmentation	21	4476	Risk Ratio (M-H, Random, 95% CI)	0.67 [0.57, 0.79]
8 Uterine hyperstimulation without FHR changes	16	2683	Risk Ratio (M-H, Random, 95% CI)	2.02 [1.21, 3.36]
9 Serious maternal morbidity or death	2	223	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
10 Epidural analgesia	3	992	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.71, 1.11]
11 Instrumental vaginal delivery	12	2053	Risk Ratio (M-H, Fixed, 95% CI)	1.10 [0.91, 1.34]
12 Meconium-stained liquor	11	2346	Risk Ratio (M-H, Fixed, 95% CI)	1.32 [1.05, 1.65]
13 Apgar score < 7 at 5 minutes	10	2399	Risk Ratio (M-H, Fixed, 95% CI)	1.00 [0.69, 1.45]
14 Neonatal intensive care unit admission	10	2348	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.73, 1.07]
15 Neonatal encephalopathy	1	360	Risk Ratio (M-H, Fixed, 95% CI)	5.98 [0.25, 145.59]
16 Perinatal death	3	887	Risk Ratio (M-H, Fixed, 95% CI)	0.98 [0.17, 5.66]
17 Serious maternal complications	2	303	Risk Ratio (M-H, Fixed, 95% CI)	0.5 [0.05, 5.39]
18 Maternal side effects (eg nausea, vomiting, diarrhoea)	9	2344	Risk Ratio (M-H, Fixed, 95% CI)	0.94 [0.87, 1.02]
19 Postpartum haemorrhage	6	581	Risk Ratio (M-H, Fixed, 95% CI)	0.84 [0.37, 1.91]



**Comparison 13. Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	8	1995	Risk Ratio (M-H, Random, 95% CI)	0.78 [0.67, 0.91]
2 Uterine hyperstimulation with FHR changes	13	2309	Risk Ratio (M-H, Fixed, 95% CI)	1.88 [1.29, 2.72]
3 Caesarean section	14	3011	Risk Ratio (M-H, Fixed, 95% CI)	1.01 [0.88, 1.15]
4 Serious neonatal morbidity or perinatal death	1	360	Risk Ratio (M-H, Fixed, 95% CI)	5.98 [0.25, 145.59]
5 Cervix unfavourable/unchanged after 12-24 hours	1	197	Risk Ratio (M-H, Fixed, 95% CI)	0.52 [0.27, 0.98]
6 Oxytocin augmentation	10	2276	Risk Ratio (M-H, Random, 95% CI)	0.69 [0.56, 0.86]
7 Uterine hyperstimulation without FHR changes	8	1590	Risk Ratio (M-H, Random, 95% CI)	1.97 [1.02, 3.82]
8 Uterine rupture	1	360	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
9 Epidural analgesia	2	311	Risk Ratio (M-H, Fixed, 95% CI)	0.81 [0.57, 1.15]
10 Instrumental vaginal delivery	5	1156	Risk Ratio (M-H, Fixed, 95% CI)	1.05 [0.81, 1.36]
11 Meconium-stained liquor	6	949	Risk Ratio (M-H, Fixed, 95% CI)	1.41 [0.98, 2.04]
12 Apgar score < 7 at 5 minutes	4	868	Risk Ratio (M-H, Fixed, 95% CI)	1.00 [0.61, 1.63]
13 Neonatal intensive care unit admission	6	1100	Risk Ratio (M-H, Fixed, 95% CI)	1.27 [0.89, 1.81]
14 Neonatal encephalopathy	1	360	Risk Ratio (M-H, Fixed, 95% CI)	5.98 [0.25, 145.59]
15 Perinatal death	1	156	Risk Ratio (M-H, Fixed, 95% CI)	2.85 [0.12, 68.95]
16 Maternal side effects (eg nausea, vomiting, diarrhoea)	3	883	Risk Ratio (M-H, Fixed, 95% CI)	1.03 [0.53, 1.99]
17 Postpartum haemorrhage	2	148	Risk Ratio (M-H, Fixed, 95% CI)	1.0 [0.14, 6.91]

**Comparison 14. Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	2	138	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.28, 3.37]
2 Caesarean section	2	138	Risk Ratio (M-H, Fixed, 95% CI)	0.60 [0.27, 1.32]
3 Oxytocin augmentation	2	138	Risk Ratio (M-H, Fixed, 95% CI)	0.70 [0.51, 0.97]
4 Uterine hyperstimulation without FHR changes	1	75	Risk Ratio (M-H, Fixed, 95% CI)	5.84 [0.74, 46.21]
5 Epidural analgesia	2	311	Risk Ratio (M-H, Fixed, 95% CI)	0.81 [0.57, 1.15]
6 Instrumental vaginal delivery	1	63	Risk Ratio (M-H, Fixed, 95% CI)	8.73 [0.49, 155.62]
7 Perinatal death	1	62	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable

**Comparison 15. Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	1	238	Risk Ratio (M-H, Fixed, 95% CI)	1.60 [0.98, 2.59]
2 Uterine hyperstimulation with FHR changes	1	238	Risk Ratio (M-H, Fixed, 95% CI)	5.08 [0.60, 42.87]
3 Uterine hyperstimulation without FHR changes	1	238	Risk Ratio (M-H, Fixed, 95% CI)	3.84 [1.93, 7.65]
4 Instrumental vaginal delivery	1	238	Risk Ratio (M-H, Fixed, 95% CI)	1.61 [0.82, 3.17]
5 Apgar score < 7 at 5 minutes	1	238	Risk Ratio (M-H, Fixed, 95% CI)	1.02 [0.44, 2.35]
6 Neonatal intensive care unit admission	1	238	Risk Ratio (M-H, Fixed, 95% CI)	0.81 [0.51, 1.31]
7 Oxytocin augmentation	1	238	Odds Ratio (M-H, Fixed, 95% CI)	0.37 [0.22, 0.64]

**Comparison 16. Misoprostol versus vaginal prostaglandin: all primiparae**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	5	736	Risk Ratio (M-H, Random, 95% CI)	0.70 [0.46, 1.05]
2 Uterine hyperstimulation with FHR changes	7	879	Risk Ratio (M-H, Fixed, 95% CI)	1.70 [1.00, 2.91]
3 Caesarean section	8	1279	Risk Ratio (M-H, Fixed, 95% CI)	0.82 [0.68, 0.99]
4 Uterine hyperstimulation without FHR changes	4	646	Risk Ratio (M-H, Random, 95% CI)	1.51 [0.80, 2.82]
5 Apgar score < 7 at 5 minutes	4	894	Risk Ratio (M-H, Fixed, 95% CI)	1.41 [0.62, 3.20]
6 Neonatal intensive care unit admission	3	731	Risk Ratio (M-H, Random, 95% CI)	1.03 [0.47, 2.25]
7 Oxytocin augmentation	7	1258	Risk Ratio (M-H, Random, 95% CI)	0.76 [0.58, 0.99]
8 Serious neonatal morbidity	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
9 Perinatal death	1	163	Risk Ratio (M-H, Fixed, 95% CI)	0.35 [0.01, 8.36]
10 Serious maternal morbidity or death	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
11 Instrumental vaginal delivery	4	527	Risk Ratio (M-H, Fixed, 95% CI)	1.30 [0.96, 1.76]
12 Meconium-stained liquor	4	554	Risk Ratio (M-H, Fixed, 95% CI)	1.54 [1.02, 2.33]
13 Uterine rupture	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
14 Postpartum haemorrhage	2	303	Risk Ratio (M-H, Fixed, 95% CI)	1.19 [0.39, 3.63]
15 Serious maternal complications	2	303	Risk Ratio (M-H, Fixed, 95% CI)	0.5 [0.05, 5.39]
16 Epidural analgesia	1	140	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.88, 1.07]
17 Maternal side effects (eg nausea, vomiting, diarrhoea)	3	363	Risk Ratio (M-H, Fixed, 95% CI)	0.58 [0.31, 1.09]

**Comparison 17. Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	3	393	Risk Ratio (M-H, Random, 95% CI)	0.64 [0.29, 1.40]
2 Uterine hyperstimulation with FHR changes	7	494	Risk Ratio (M-H, Fixed, 95% CI)	4.13 [1.66, 10.28]
3 Caesarean section	5	888	Risk Ratio (M-H, Fixed, 95% CI)	0.77 [0.59, 1.00]
4 Apgar score < 7 at 5 minutes	3	703	Risk Ratio (M-H, Fixed, 95% CI)	1.40 [0.55, 3.51]
5 Serious neonatal morbidity	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
6 Neonatal intensive care admission	2	540	Risk Ratio (M-H, Fixed, 95% CI)	0.87 [0.62, 1.22]
7 Oxytocin augmentation	2	212	Risk Ratio (M-H, Fixed, 95% CI)	0.51 [0.38, 0.69]
8 Perinatal death	1	163	Risk Ratio (M-H, Fixed, 95% CI)	0.35 [0.01, 8.36]
9 Serious maternal morbidity or death	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
10 Uterine hyperstimulation without FHR changes	2	303	Risk Ratio (M-H, Fixed, 95% CI)	3.23 [1.33, 7.85]
11 Instrumental vaginal delivery	3	375	Risk Ratio (M-H, Fixed, 95% CI)	1.43 [0.97, 2.11]
12 Meconium-stained liquor	3	363	Risk Ratio (M-H, Fixed, 95% CI)	1.38 [0.83, 2.31]
13 Uterine rupture	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
14 Postpartum haemorrhage	2	303	Risk Ratio (M-H, Fixed, 95% CI)	1.19 [0.39, 3.63]
15 Serious maternal complications	2	303	Risk Ratio (M-H, Fixed, 95% CI)	0.5 [0.05, 5.39]
16 Epidural analgesia	1	140	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.88, 1.07]
17 Maternal side effects (eg nausea, vomiting, diarrhoea)	3	363	Risk Ratio (M-H, Fixed, 95% CI)	0.58 [0.31, 1.09]

**Comparison 18. Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	3	393	Risk Ratio (M-H, Random, 95% CI)	0.64 [0.29, 1.40]
2 Uterine hyperstimulation with FHR changes	4	488	Risk Ratio (M-H, Fixed, 95% CI)	4.13 [1.66, 10.28]
3 Caesarean section	4	488	Risk Ratio (M-H, Fixed, 95% CI)	0.79 [0.53, 1.18]
4 Uterine hyperstimulation without FHR changes	2	303	Risk Ratio (M-H, Fixed, 95% CI)	3.23 [1.33, 7.85]
5 Apgar score < 7 at 5 minutes	2	303	Risk Ratio (M-H, Fixed, 95% CI)	2.37 [0.36, 15.76]
6 Neonatal intensive care unit admission	1	140	Risk Ratio (M-H, Fixed, 95% CI)	0.44 [0.14, 1.38]
7 Oxytocin augmentation	3	363	Risk Ratio (M-H, Random, 95% CI)	0.64 [0.39, 1.05]
8 Postpartum haemorrhage	2	303	Risk Ratio (M-H, Fixed, 95% CI)	1.19 [0.39, 3.63]
9 Serious maternal complication	2	303	Risk Ratio (M-H, Fixed, 95% CI)	0.5 [0.05, 5.39]
10 Epidural analgesia	1	140	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.70 [0.21, 2.26]

11 Meconium-stained liquor	3	363	Risk Ratio (M-H, Fixed, 95% CI)	1.38 [0.83, 2.31]
12 Serious maternal morbidity or death	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
13 Serious neonatal morbidity	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
14 Perinatal death	1	163	Odds Ratio (M-H, Fixed, 95% CI)	0.34 [0.01, 8.51]
15 Uterine rupture	1	163	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
16 Maternal side effects (eg nausea, vomiting, diarrhoea)	3	360	Risk Ratio (M-H, Fixed, 95% CI)	0.58 [0.31, 1.09]

#### Comparison 19. Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, variable or undefined cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	1	48	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
2 Caesarean section	1	48	Risk Ratio (M-H, Fixed, 95% CI)	0.76 [0.35, 1.66]

#### Comparison 20. Misoprostol versus vaginal prostaglandin: all primiparae with ruptured membranes and unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Oxytocin augmentation	1	152	Odds Ratio (M-H, Fixed, 95% CI)	0.17 [0.08, 0.35]
2 Instrumental vaginal delivery	1	152	Risk Ratio (M-H, Fixed, 95% CI)	1.38 [0.69, 2.78]

#### Comparison 21. Misoprostol versus vaginal prostaglandin: all multiparae

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	2	223	Risk Ratio (M-H, Fixed, 95% CI)	1.17 [0.69, 1.96]
2 Uterine hyperstimulation with FHR changes	2	143	Risk Ratio (M-H, Fixed, 95% CI)	0.46 [0.08, 2.57]
3 Caesarean section	3	424	Risk Ratio (M-H, Fixed, 95% CI)	0.74 [0.38, 1.43]
4 Oxytocin augmentation	2	397	Risk Ratio (M-H, Fixed, 95% CI)	1.58 [1.19, 2.09]
5 Uterine hyperstimulation without FHR changes	1	116	Risk Ratio (M-H, Fixed, 95% CI)	0.77 [0.37, 1.61]
6 Instrumental delivery	1	116	Risk Ratio (M-H, Fixed, 95% CI)	0.67 [0.20, 2.24]
7 Apgar score < 7 at 5 minutes	1	281	Risk Ratio (M-H, Fixed, 95% CI)	0.50 [0.05, 5.49]

8 Neonatal intensive care unit admission	1	281	Risk Ratio (M-H, Fixed, 95% CI)	0.52 [0.32, 0.83]
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#### Comparison 22. Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	1	107	Risk Ratio (M-H, Fixed, 95% CI)	1.25 [0.68, 2.31]
2 Caesarean section	1	281	Risk Ratio (M-H, Fixed, 95% CI)	0.76 [0.37, 1.54]
3 Apgar score < 7 at 5 minutes	1	281	Risk Ratio (M-H, Fixed, 95% CI)	0.50 [0.05, 5.49]
4 Neonatal intensive care unit admission	1	281	Risk Ratio (M-H, Fixed, 95% CI)	0.52 [0.32, 0.83]
5 Oxytocin augmentation	5	915	Risk Ratio (M-H, Random, 95% CI)	0.70 [0.48, 1.03]

#### Comparison 23. Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved in 24 hours	1	107	Risk Ratio (M-H, Fixed, 95% CI)	1.25 [0.68, 2.31]
2 Instrumental vaginal delivery	2	223	Risk Ratio (M-H, Fixed, 95% CI)	1.46 [0.91, 2.33]

#### Comparison 24. Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, variable or undefined cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	1	27	Risk Ratio (M-H, Fixed, 95% CI)	2.44 [0.11, 54.97]
2 Caesarean section	1	27	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
3 Caesarean section	1	29	Risk Ratio (M-H, Fixed, 95% CI)	0.62 [0.06, 6.05]

**Comparison 25. Misoprostol versus intracervical prostaglandin: all women**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	13	1627	Risk Ratio (M-H, Fixed, 95% CI)	0.63 [0.56, 0.71]
2 Uterine hyperstimulation with FHR changes	20	2224	Risk Ratio (M-H, Fixed, 95% CI)	2.32 [1.64, 3.28]
3 Caesarean section	27	3311	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.84, 1.11]
4 Serious neonatal morbidity/perinatal death	2	272	Risk Ratio (M-H, Fixed, 95% CI)	0.4 [0.09, 1.87]
5 Cervix unfavourable/unchanged after 12-24 hours	1	155	Risk Ratio (M-H, Fixed, 95% CI)	0.68 [0.52, 0.88]
6 Oxytocin augmentation	20	2316	Risk Ratio (M-H, Random, 95% CI)	0.55 [0.48, 0.64]
7 Uterine hyperstimulation without FHR changes	17	2178	Risk Ratio (M-H, Fixed, 95% CI)	1.95 [1.57, 2.42]
8 Uterine rupture	1	99	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
9 Epidural analgesia	2	321	Risk Ratio (M-H, Fixed, 95% CI)	0.64 [0.48, 0.86]
10 Instrumental vaginal delivery	13	1900	Risk Ratio (M-H, Fixed, 95% CI)	1.05 [0.84, 1.32]
11 Meconium-stained liquor	14	2018	Risk Ratio (M-H, Fixed, 95% CI)	1.29 [1.04, 1.59]
12 Apgar score < 7 at 5 minutes	15	2114	Risk Ratio (M-H, Fixed, 95% CI)	0.93 [0.51, 1.70]
13 Neonatal intensive care unit admission	13	1773	Risk Ratio (M-H, Fixed, 95% CI)	1.11 [0.81, 1.52]
14 Perinatal death	2	378	Risk Ratio (M-H, Fixed, 95% CI)	2.85 [0.12, 68.95]
15 Maternal side effects (eg nausea, vomiting, diarrhoea)	7	731	Risk Ratio (M-H, Fixed, 95% CI)	1.15 [0.68, 1.95]
16 Postpartum haemorrhage	3	320	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.22, 4.24]

**Comparison 26. Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	10	1287	Risk Ratio (M-H, Fixed, 95% CI)	0.61 [0.54, 0.69]
2 Uterine hyperstimulation with FHR changes	19	2124	Risk Ratio (M-H, Fixed, 95% CI)	2.33 [1.64, 3.30]
3 Caesarean section	21	2499	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.83, 1.13]
4 Serious neonatal morbidity	1	50	Risk Ratio (M-H, Fixed, 95% CI)	0.4 [0.09, 1.87]
5 Cervix unfavourable/unchanged after 12-24 hours	1	155	Risk Ratio (M-H, Fixed, 95% CI)	0.68 [0.52, 0.88]
6 Oxytocin augmentation	16	1733	Risk Ratio (M-H, Random, 95% CI)	0.58 [0.50, 0.67]
7 Uterine hyperstimulation without FHR changes	16	1978	Risk Ratio (M-H, Fixed, 95% CI)	1.99 [1.60, 2.47]
8 Uterine rupture	1	99	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
9 Epidural analgesia	1	99	Risk Ratio (M-H, Fixed, 95% CI)	0.71 [0.41, 1.25]
10 Instrumental vaginal delivery	11	1478	Risk Ratio (M-H, Fixed, 95% CI)	0.87 [0.67, 1.13]

11 Meconium-stained liquor	11	1496	Risk Ratio (M-H, Fixed, 95% CI)	1.25 [0.98, 1.61]
12 Apgar score < 7 at 5 minutes	10	1331	Risk Ratio (M-H, Fixed, 95% CI)	0.77 [0.36, 1.64]
13 Neonatal intensive care unit admission	10	1273	Risk Ratio (M-H, Fixed, 95% CI)	0.99 [0.70, 1.41]
14 Perinatal death	1	156	Risk Ratio (M-H, Fixed, 95% CI)	2.85 [0.12, 68.95]
15 Maternal side effects (eg nausea, vomiting, diarrhoea)	6	561	Risk Ratio (M-H, Fixed, 95% CI)	0.69 [0.41, 1.15]
16 Postpartum haemorrhage	3	320	Risk Ratio (M-H, Fixed, 95% CI)	1.79 [0.39, 8.33]

**Comparison 27. Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	8	1088	Risk Ratio (M-H, Fixed, 95% CI)	0.64 [0.56, 0.73]
2 Uterine hyperstimulation with FHR changes	13	1433	Risk Ratio (M-H, Fixed, 95% CI)	3.62 [2.22, 5.90]
3 Caesarean section	12	1433	Risk Ratio (M-H, Fixed, 95% CI)	0.99 [0.81, 1.20]
4 Maternal side effects	3	252	Risk Ratio (M-H, Fixed, 95% CI)	0.65 [0.34, 1.21]
5 Serious neonatal morbidity	1	50	Risk Ratio (M-H, Fixed, 95% CI)	0.4 [0.09, 1.87]
6 Cervix unfavourable/unchanged after 12-24 hours	1	155	Risk Ratio (M-H, Fixed, 95% CI)	0.68 [0.52, 0.88]
7 Oxytocin augmentation	10	1075	Risk Ratio (M-H, Random, 95% CI)	0.53 [0.43, 0.67]
8 Uterine hyperstimulation without FHR changes	12	1422	Risk Ratio (M-H, Fixed, 95% CI)	2.47 [1.88, 3.23]
9 Instrumental vaginal delivery	8	1045	Risk Ratio (M-H, Fixed, 95% CI)	0.85 [0.64, 1.14]
10 Meconium-stained liquor	7	989	Risk Ratio (M-H, Fixed, 95% CI)	1.41 [1.05, 1.89]
11 Apgar score < 7 at 5 minutes	6	814	Risk Ratio (M-H, Fixed, 95% CI)	0.66 [0.24, 1.84]
12 Neonatal intensive care unit admission	8	1074	Risk Ratio (M-H, Fixed, 95% CI)	0.99 [0.69, 1.42]
13 Perinatal death	1	156	Risk Ratio (M-H, Fixed, 95% CI)	2.85 [0.12, 68.95]
14 Postpartum haemorrhage	1	73	Risk Ratio (M-H, Fixed, 95% CI)	2.77 [0.12, 65.82]

**Comparison 28. Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved in 24 hours	1	100	Risk Ratio (M-H, Fixed, 95% CI)	0.61 [0.38, 0.96]
2 Uterine hyperstimulation with FHR changes	1	100	Risk Ratio (M-H, Fixed, 95% CI)	2.0 [0.19, 21.36]
3 Caesarean section	2	322	Risk Ratio (M-H, Fixed, 95% CI)	1.16 [0.65, 2.05]

4 Serious neonatal morbidity/perinatal death	1	222	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Neonatal intensive care unit admission	1	100	Risk Ratio (M-H, Fixed, 95% CI)	3.0 [0.13, 71.92]
6 Oxytocin augmentation	2	322	Risk Ratio (M-H, Fixed, 95% CI)	0.49 [0.34, 0.71]
7 Epidural analgesia	1	222	Risk Ratio (M-H, Fixed, 95% CI)	0.62 [0.44, 0.87]
8 Instrumental vaginal delivery	1	222	Risk Ratio (M-H, Fixed, 95% CI)	2.23 [1.23, 4.06]
9 Meconium-stained liquor	2	322	Risk Ratio (M-H, Fixed, 95% CI)	1.28 [0.84, 1.95]
10 Apgar score < 7 at 5 minutes	2	322	Risk Ratio (M-H, Fixed, 95% CI)	2.33 [0.35, 15.60]
11 Perinatal death	1	222	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable

#### Comparison 29. Misoprostol versus oxytocin: all women

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	10	1397	Risk Ratio (M-H, Random, 95% CI)	0.65 [0.47, 0.90]
2 Uterine hyperstimulation with FHR changes	9	1419	Risk Ratio (M-H, Fixed, 95% CI)	1.87 [1.20, 2.91]
3 Caesarean section	25	3074	Risk Ratio (M-H, Random, 95% CI)	0.76 [0.60, 0.96]
4 Serious neonatal morbidity	2	274	Risk Ratio (M-H, Fixed, 95% CI)	0.96 [0.57, 1.60]
5 Serious maternal morbidity or death	2	190	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
6 Maternal side effects (eg nausea, vomiting, diarrhoea)	4	334	Risk Ratio (M-H, Fixed, 95% CI)	5.04 [1.51, 16.86]
7 Oxytocin augmentation	3	386	Risk Ratio (M-H, Random, 95% CI)	0.51 [0.28, 0.95]
8 Uterine hyperstimulation without FHR changes	15	2050	Risk Ratio (M-H, Fixed, 95% CI)	2.24 [1.82, 2.77]
9 Uterine rupture	2	100	Risk Ratio (M-H, Fixed, 95% CI)	4.12 [0.50, 34.25]
10 Epidural analgesia	3	315	Risk Ratio (M-H, Fixed, 95% CI)	0.82 [0.67, 1.00]
11 Instrumental vaginal delivery	13	1639	Risk Ratio (M-H, Fixed, 95% CI)	0.74 [0.56, 0.99]
12 Meconium-stained liquor	12	1694	Risk Ratio (M-H, Fixed, 95% CI)	1.05 [0.82, 1.35]
13 Apgar score < 7 at 5 minutes	13	1906	Risk Ratio (M-H, Fixed, 95% CI)	0.56 [0.34, 0.92]
14 Neonatal intensive care unit admission	11	1491	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.69, 1.13]
15 Serious maternal complications	1	100	Risk Ratio (M-H, Fixed, 95% CI)	0.6 [0.15, 2.38]
16 Perinatal death	3	334	Risk Ratio (M-H, Fixed, 95% CI)	0.78 [0.14, 4.39]
17 Postpartum haemorrhage	6	568	Risk Ratio (M-H, Fixed, 95% CI)	0.53 [0.23, 1.23]



**Comparison 30. Misoprostol versus oxytocin: all women with unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	5	660	Risk Ratio (M-H, Random, 95% CI)	0.87 [0.56, 1.35]
2 Uterine hyperstimulation with FHR changes	3	507	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.32, 2.39]
3 Caesarean section	14	1598	Risk Ratio (M-H, Random, 95% CI)	0.81 [0.61, 1.08]
4 Perinatal death	2	162	Risk Ratio (M-H, Fixed, 95% CI)	1.24 [0.22, 6.88]
5 Serious maternal morbidity or death	1	38	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
6 Maternal side effects	4	334	Risk Ratio (M-H, Fixed, 95% CI)	5.04 [1.51, 16.86]
7 Postpartum haemorrhage	2	162	Risk Ratio (M-H, Fixed, 95% CI)	0.18 [0.02, 1.52]
8 Uterine hyperstimulation without FHR changes	10	1192	Risk Ratio (M-H, Fixed, 95% CI)	2.43 [1.66, 3.58]
9 Uterine rupture	2	100	Risk Ratio (M-H, Fixed, 95% CI)	4.12 [0.50, 34.25]
10 Serious maternal complications	1	100	Risk Ratio (M-H, Fixed, 95% CI)	0.6 [0.15, 2.38]
11 Instrumental vaginal delivery	8	853	Risk Ratio (M-H, Fixed, 95% CI)	0.42 [0.25, 0.70]
12 Meconium-stained liquor	8	904	Risk Ratio (M-H, Fixed, 95% CI)	0.92 [0.66, 1.29]
13 Apgar score < 7 at 5 minutes	6	738	Risk Ratio (M-H, Fixed, 95% CI)	1.71 [0.69, 4.27]
14 Neonatal intensive care unit admission	7	707	Risk Ratio (M-H, Fixed, 95% CI)	1.25 [0.77, 2.04]
15 Serious neonatal morbidity	1	64	Risk Ratio (M-H, Fixed, 95% CI)	1.0 [0.15, 6.67]

**Comparison 31. Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Instrumental delivery	1	140	Risk Ratio (M-H, Fixed, 95% CI)	0.5 [0.20, 1.26]
2 Vaginal delivery not achieved in 24 hours	1	210	Risk Ratio (M-H, Fixed, 95% CI)	0.53 [0.33, 0.84]
3 Caesarean section	5	736	Risk Ratio (M-H, Random, 95% CI)	0.58 [0.37, 0.90]
4 Uterine hyperstimulation with FHR changes	1	210	Risk Ratio (M-H, Fixed, 95% CI)	0.75 [0.17, 3.27]
5 Uterine hyperstimulation without FHR changes	1	210	Risk Ratio (M-H, Fixed, 95% CI)	3.0 [1.48, 6.07]
6 Meconium-stained liquor	1	210	Risk Ratio (M-H, Fixed, 95% CI)	1.4 [0.65, 3.01]
7 Serious neonatal morbidity	1	210	Risk Ratio (M-H, Fixed, 95% CI)	0.95 [0.56, 1.63]
8 Apgar score < 7 at 5 minutes	1	210	Risk Ratio (M-H, Fixed, 95% CI)	2.0 [0.18, 21.72]

**Comparison 32. Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	2	278	Risk Ratio (M-H, Fixed, 95% CI)	0.46 [0.30, 0.68]
2 Uterine hyperstimulation with FHR changes	1	360	Risk Ratio (M-H, Fixed, 95% CI)	3.43 [1.66, 7.08]
3 Caesarean section	3	652	Risk Ratio (M-H, Random, 95% CI)	0.43 [0.13, 1.45]
4 Serious maternal morbidity or death	1	152	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Uterine hyperstimulation without FHR changes	2	512	Risk Ratio (M-H, Fixed, 95% CI)	2.33 [1.74, 3.11]
6 Epidural analgesia	1	126	Risk Ratio (M-H, Fixed, 95% CI)	0.69 [0.51, 0.92]
7 Instrumental vaginal delivery	2	500	Risk Ratio (M-H, Random, 95% CI)	0.86 [0.36, 2.06]
8 Meconium-stained liquor	1	360	Risk Ratio (M-H, Fixed, 95% CI)	1.19 [0.71, 2.00]
9 Apgar score < 7 at 5 minutes	2	512	Risk Ratio (M-H, Random, 95% CI)	0.97 [0.06, 15.26]
10 Neonatal intensive care unit admission	1	360	Risk Ratio (M-H, Fixed, 95% CI)	1.32 [0.65, 2.69]
11 Perinatal death	1	152	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable

**Comparison 33. Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved in 24 hours	1	108	Risk Ratio (M-H, Fixed, 95% CI)	2.0 [0.73, 5.46]
2 Uterine hyperstimulation without FHR changes	1	108	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
3 Caesarean section	2	290	Risk Ratio (M-H, Random, 95% CI)	0.72 [0.20, 2.60]
4 Maternal side effects	1	108	Risk Ratio (M-H, Fixed, 95% CI)	11.0 [0.62, 194.17]

**Comparison 34. Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	3	323	Risk Ratio (M-H, Fixed, 95% CI)	1.19 [0.41, 3.41]
2 Caesarean section	4	520	Risk Ratio (M-H, Fixed, 95% CI)	0.95 [0.60, 1.51]
3 Oxytocin augmentation	2	282	Risk Ratio (M-H, Fixed, 95% CI)	0.38 [0.30, 0.49]
4 Uterine hyperstimulation without FHR changes	3	323	Risk Ratio (M-H, Fixed, 95% CI)	1.67 [1.04, 2.67]

5 Epidural analgesia	1	85	Risk Ratio (M-H, Fixed, 95% CI)	0.68 [0.12, 3.88]
6 Instrumental vaginal delivery	3	323	Risk Ratio (M-H, Fixed, 95% CI)	0.76 [0.45, 1.27]
7 Meconium-stained liquor	1	197	Risk Ratio (M-H, Fixed, 95% CI)	0.90 [0.36, 2.23]
8 Apgar score < 7 at 5 minutes	4	520	Risk Ratio (M-H, Fixed, 95% CI)	0.65 [0.17, 2.43]
9 Neonatal intensive care unit admission	3	379	Risk Ratio (M-H, Fixed, 95% CI)	0.78 [0.52, 1.18]
10 Postpartum haemorrhage	2	238	Risk Ratio (M-H, Fixed, 95% CI)	1.83 [0.40, 8.43]
11 Vaginal delivery not achieved within 24 hours	1	197	Risk Ratio (M-H, Fixed, 95% CI)	0.89 [0.55, 1.45]

### Comparison 35. Misoprostol versus oxytocin: all women with previous caesarean section

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	1	38	Risk Ratio (M-H, Fixed, 95% CI)	1.35 [0.81, 2.24]
2 Caesarean section	1	38	Risk Ratio (M-H, Fixed, 95% CI)	1.65 [0.71, 3.83]
3 Serious maternal morbidity or death	1	38	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
4 Uterine hyperstimulation without FHR changes	1	38	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
5 Uterine rupture	1	38	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
6 Meconium-stained liquor	1	38	Risk Ratio (M-H, Fixed, 95% CI)	0.41 [0.09, 1.79]
7 Apgar score < 7 at 5 minutes	1	38	Risk Ratio (M-H, Fixed, 95% CI)	3.67 [0.16, 84.66]
8 Neonatal intensive care unit admission	1	38	Risk Ratio (M-H, Fixed, 95% CI)	1.85 [0.62, 5.52]

### Comparison 36. Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	1	38	Risk Ratio (M-H, Fixed, 95% CI)	1.35 [0.81, 2.24]
2 Caesarean section	1	38	Risk Ratio (M-H, Fixed, 95% CI)	1.65 [0.71, 3.83]
3 Serious maternal morbidity or death	1	38	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
4 Uterine hyperstimulation without FHR changes	1	38	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
5 Uterine rupture	1	38	Risk Ratio (M-H, Fixed, 95% CI)	6.11 [0.31, 119.33]
6 Meconium-stained liquor	1	38	Risk Ratio (M-H, Fixed, 95% CI)	0.41 [0.09, 1.79]
7 Apgar score < 7 at 5 minutes	1	38	Risk Ratio (M-H, Fixed, 95% CI)	3.67 [0.16, 84.66]
8 Neonatal intensive care unit admission	1	38	Risk Ratio (M-H, Fixed, 95% CI)	1.85 [0.62, 5.52]

**Comparison 37. Misoprostol versus oxytocin: all multiparae**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	3	268	Risk Ratio (M-H, Fixed, 95% CI)	0.76 [0.44, 1.31]
2 Serious neonatal morbidity	1	64	Risk Ratio (M-H, Fixed, 95% CI)	1.0 [0.15, 6.67]
3 Uterine hyperstimulation without FHR changes	2	164	Risk Ratio (M-H, Fixed, 95% CI)	1.67 [0.41, 6.74]
4 Instrumental delivery	3	268	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.07, 1.62]
5 Meconium-stained liquor	3	268	Risk Ratio (M-H, Fixed, 95% CI)	1.05 [0.59, 1.88]
6 Neonatal intensive care unit admission	3	268	Risk Ratio (M-H, Fixed, 95% CI)	1.17 [0.40, 3.37]
7 Perinatal death	1	100	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
8 Maternal side effects (eg nausea, vomiting, diarrhoea)	2	164	Risk Ratio (M-H, Fixed, 95% CI)	4.33 [0.75, 24.89]
9 Postpartum haemorrhage	2	164	Risk Ratio (M-H, Fixed, 95% CI)	0.25 [0.03, 2.19]
10 Serious maternal complications	1	100	Risk Ratio (M-H, Fixed, 95% CI)	0.6 [0.15, 2.38]

**Comparison 38. Misoprostol versus oxytocin: all multiparae with unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Caesarean section	3	268	Risk Ratio (M-H, Fixed, 95% CI)	0.76 [0.44, 1.31]
2 Serious neonatal morbidity	1	64	Risk Ratio (M-H, Fixed, 95% CI)	1.0 [0.15, 6.67]
3 Uterine hyperstimulation without FHR changes	2	164	Risk Ratio (M-H, Fixed, 95% CI)	1.67 [0.41, 6.74]
4 Instrumental delivery	3	268	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.07, 1.62]
5 Neonatal intensive care unit admission	3	268	Risk Ratio (M-H, Fixed, 95% CI)	1.17 [0.40, 3.37]
6 Maternal side effects	2	164	Risk Ratio (M-H, Fixed, 95% CI)	4.33 [0.75, 24.89]
7 Postpartum haemorrhage	2	164	Risk Ratio (M-H, Fixed, 95% CI)	0.25 [0.03, 2.19]
8 Meconium-stained liquor	3	268	Risk Ratio (M-H, Fixed, 95% CI)	1.05 [0.59, 1.88]
9 Perinatal death	1	100	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
10 Serious maternal complications	1	100	Risk Ratio (M-H, Fixed, 95% CI)	0.6 [0.15, 2.38]

**Comparison 39. Misoprostol lower versus higher dose: all women**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	12	1887	Risk Ratio (M-H, Fixed, 95% CI)	1.08 [0.96, 1.21]
2 Uterine hyperstimulation with FHR changes	16	2540	Risk Ratio (M-H, Fixed, 95% CI)	0.51 [0.37, 0.69]
3 Caesarean section	21	2913	Risk Ratio (M-H, Fixed, 95% CI)	0.96 [0.83, 1.12]
4 Serious maternal morbidity or death	4	8	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Cervix unfavourable/unchanged after 12-24 hours	1	69	Risk Ratio (M-H, Fixed, 95% CI)	0.93 [0.62, 1.39]
6 Oxytocin augmentation	18	2753	Risk Ratio (M-H, Random, 95% CI)	1.30 [1.14, 1.49]
7 Uterine hyperstimulation without FHR changes	14	2085	Risk Ratio (M-H, Fixed, 95% CI)	0.57 [0.46, 0.69]
8 Uterine rupture	3	336	Risk Ratio (M-H, Fixed, 95% CI)	0.65 [0.11, 3.87]
9 Epidural	1	104	Risk Ratio (M-H, Fixed, 95% CI)	1.04 [0.63, 1.72]
10 Instrumental vaginal delivery	14	2116	Risk Ratio (M-H, Fixed, 95% CI)	1.09 [0.82, 1.44]
11 Meconium-stained liquor	13	1673	Risk Ratio (M-H, Fixed, 95% CI)	0.90 [0.69, 1.19]
12 Apgar score < 7 at 5 minutes	13	2045	Risk Ratio (M-H, Fixed, 95% CI)	0.77 [0.44, 1.37]
13 Neonatal intensive care unit admission	9	1795	Risk Ratio (M-H, Fixed, 95% CI)	0.82 [0.64, 1.05]
14 Perinatal death	2	187	Risk Ratio (M-H, Fixed, 95% CI)	0.17 [0.02, 1.34]
15 Maternal side effects (eg nausea, vomiting, diarrhoea)	9	1653	Risk Ratio (M-H, Fixed, 95% CI)	0.85 [0.58, 1.24]
16 Postpartum haemorrhage	5	893	Risk Ratio (M-H, Fixed, 95% CI)	0.76 [0.43, 1.35]

**Comparison 40. Misoprostol lower versus higher dose: all women with unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	8	1563	Risk Ratio (M-H, Fixed, 95% CI)	1.08 [0.96, 1.22]
2 Uterine hyperstimulation with FHR changes	13	2174	Risk Ratio (M-H, Fixed, 95% CI)	0.53 [0.38, 0.73]
3 Caesarean section	15	2214	Risk Ratio (M-H, Fixed, 95% CI)	0.95 [0.81, 1.13]
4 Uterine rupture	1	95	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Serious maternal morbidity or death	4	8	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
6 Perinatal death	1	60	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.01, 7.87]
7 Oxytocin augmentation	11	2049	Risk Ratio (M-H, Random, 95% CI)	1.39 [1.15, 1.67]
8 Uterine hyperstimulation without FHR changes	8	1569	Risk Ratio (M-H, Fixed, 95% CI)	0.60 [0.47, 0.76]
9 Epidural	1	104	Risk Ratio (M-H, Fixed, 95% CI)	1.04 [0.63, 1.72]
10 Instrumental vaginal delivery	9	1703	Risk Ratio (M-H, Fixed, 95% CI)	1.01 [0.73, 1.39]

11 Meconium-stained liquor	7	1115	Risk Ratio (M-H, Fixed, 95% CI)	0.94 [0.66, 1.33]
12 Apgar score < 7 at 5 minutes	9	1609	Risk Ratio (M-H, Fixed, 95% CI)	0.79 [0.40, 1.58]
13 Neonatal intensive care unit admission	5	1290	Risk Ratio (M-H, Fixed, 95% CI)	0.83 [0.62, 1.10]
14 Maternal side effects (eg nausea, vomiting, diarrhoea)	4	1143	Risk Ratio (M-H, Fixed, 95% CI)	0.77 [0.45, 1.30]
15 Postpartum haemorrhage	1	399	Risk Ratio (M-H, Fixed, 95% CI)	1.08 [0.27, 4.25]

#### Comparison 41. Misoprostol lower versus higher dose: all women with intact membranes

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved in 24 hours	1	48	Risk Ratio (M-H, Fixed, 95% CI)	0.78 [0.05, 11.72]
2 Caesarean section	1	48	Risk Ratio (M-H, Fixed, 95% CI)	0.65 [0.23, 1.84]
3 Uterine hyperstimulation without FHR changes	1	48	Risk Ratio (M-H, Fixed, 95% CI)	0.16 [0.02, 1.23]
4 Meconium-stained liquor	1	48	Risk Ratio (M-H, Fixed, 95% CI)	0.78 [0.12, 5.07]

#### Comparison 42. Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	4	779	Risk Ratio (M-H, Fixed, 95% CI)	1.12 [0.97, 1.30]
2 Uterine hyperstimulation with FHR changes	7	1035	Risk Ratio (M-H, Fixed, 95% CI)	0.54 [0.31, 0.95]
3 Caesarean section	8	1098	Risk Ratio (M-H, Fixed, 95% CI)	1.09 [0.86, 1.39]
4 Serious maternal morbidity or death	1	50	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
5 Oxytocin augmentation	5	929	Risk Ratio (M-H, Fixed, 95% CI)	1.13 [0.96, 1.33]
6 Uterine hyperstimulation without FHR changes	3	700	Risk Ratio (M-H, Fixed, 95% CI)	0.78 [0.51, 1.18]
7 Instrumental vaginal delivery	5	929	Risk Ratio (M-H, Fixed, 95% CI)	0.97 [0.60, 1.57]
8 Meconium-stained liquor	4	740	Risk Ratio (M-H, Fixed, 95% CI)	1.01 [0.67, 1.51]
9 Apgar score < 7 at 5 minutes	4	740	Risk Ratio (M-H, Fixed, 95% CI)	1.15 [0.42, 3.12]
10 Neonatal intensive care unit admission	2	640	Risk Ratio (M-H, Fixed, 95% CI)	0.90 [0.65, 1.23]
11 Maternal side effects (eg nausea, vomiting, diarrhoea)	3	700	Risk Ratio (M-H, Fixed, 95% CI)	0.73 [0.36, 1.46]

**Comparison 43. Misoprostol lower versus higher dose: all multiparae**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	1	104	Risk Ratio (M-H, Fixed, 95% CI)	5.19 [0.26, 105.59]
2 Caesarean section	1	104	Risk Ratio (M-H, Fixed, 95% CI)	3.12 [0.66, 14.74]
3 Oxytocin augmentation	1	104	Risk Ratio (M-H, Random, 95% CI)	7.27 [0.93, 57.06]
4 Uterine hyperstimulation without FHR changes	1	104	Risk Ratio (M-H, Fixed, 95% CI)	1.43 [0.63, 3.26]
5 Epidural	1	104	Risk Ratio (M-H, Fixed, 95% CI)	1.04 [0.63, 1.72]
6 Instrumental vaginal delivery	1	104	Risk Ratio (M-H, Fixed, 95% CI)	0.42 [0.08, 2.05]
7 Meconium-stained liquor	1	104	Risk Ratio (M-H, Fixed, 95% CI)	0.74 [0.25, 2.19]
8 Apgar score < 7 at 5 minutes	1	104	Risk Ratio (M-H, Fixed, 95% CI)	3.12 [0.13, 74.76]
9 Neonatal intensive care unit admission	1	104	Risk Ratio (M-H, Fixed, 95% CI)	1.73 [0.44, 6.88]
10 Maternal side effects (eg nausea, vomiting, diarrhoea)	1	104	Risk Ratio (M-H, Fixed, 95% CI)	0.94 [0.44, 2.03]

**Comparison 44. Misoprostol lower versus higher dose: all multiparae with unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Vaginal delivery not achieved within 24 hours	1	124	Risk Ratio (M-H, Fixed, 95% CI)	1.67 [0.98, 2.84]
2 Uterine hyperstimulation with FHR changes	2	228	Risk Ratio (M-H, Random, 95% CI)	0.85 [0.04, 20.36]
3 Caesarean section	2	228	Risk Ratio (M-H, Fixed, 95% CI)	1.84 [0.64, 5.31]
4 Oxytocin augmentation	2	228	Risk Ratio (M-H, Random, 95% CI)	2.61 [0.84, 8.04]
5 Uterine hyperstimulation without FHR changes	2	228	Risk Ratio (M-H, Random, 95% CI)	0.60 [0.07, 5.08]
6 Epidural	1	104	Risk Ratio (M-H, Fixed, 95% CI)	1.04 [0.63, 1.72]
7 Instrumental vaginal delivery	2	228	Risk Ratio (M-H, Fixed, 95% CI)	0.51 [0.13, 2.00]
8 Meconium-stained liquor	2	228	Risk Ratio (M-H, Fixed, 95% CI)	0.85 [0.38, 1.89]
9 Apgar score < 7 at 5 minutes	2	228	Risk Ratio (M-H, Fixed, 95% CI)	3.06 [0.32, 28.96]
10 Neonatal intensive care unit admission	1	104	Risk Ratio (M-H, Fixed, 95% CI)	1.73 [0.44, 6.88]
11 Postpartum haemorrhage	1	124	Risk Ratio (M-H, Fixed, 95% CI)	0.86 [0.31, 2.41]
12 Maternal side effects (eg nausea, vomiting, diarrhoea)	2	228	Risk Ratio (M-H, Fixed, 95% CI)	1.13 [0.64, 1.98]

**Comparison 45. Misoprostol gel versus tablet: all women**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	1	467	Risk Ratio (M-H, Fixed, 95% CI)	0.49 [0.29, 0.83]
2 Caesarean section	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.07 [0.79, 1.45]
3 Oxytocin augmentation	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.26 [1.13, 1.41]
4 Epidural analgesia	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.19 [1.03, 1.38]
5 Instrumental vaginal delivery	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.12 [0.74, 1.70]
6 Apgar score < 7 at 5 minutes	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.16 [0.56, 2.38]
7 Neonatal intensive care unit admission	1	467	Risk Ratio (M-H, Fixed, 95% CI)	0.74 [0.44, 1.24]

**Comparison 46. Misoprostol gel versus tablet: all women with unfavourable cervix**

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Uterine hyperstimulation with FHR changes	1	467	Risk Ratio (M-H, Fixed, 95% CI)	0.49 [0.29, 0.83]
2 Caesarean section	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.07 [0.79, 1.45]
3 Oxytocin augmentation	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.26 [1.13, 1.41]
4 Epidural analgesia	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.19 [1.03, 1.38]
5 Instrumental vaginal delivery	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.12 [0.74, 1.70]
6 Apgar score < 7 at 5 minutes	1	467	Risk Ratio (M-H, Fixed, 95% CI)	1.16 [0.56, 2.38]
7 Neonatal intensive care unit admission	1	467	Risk Ratio (M-H, Fixed, 95% CI)	0.74 [0.44, 1.24]

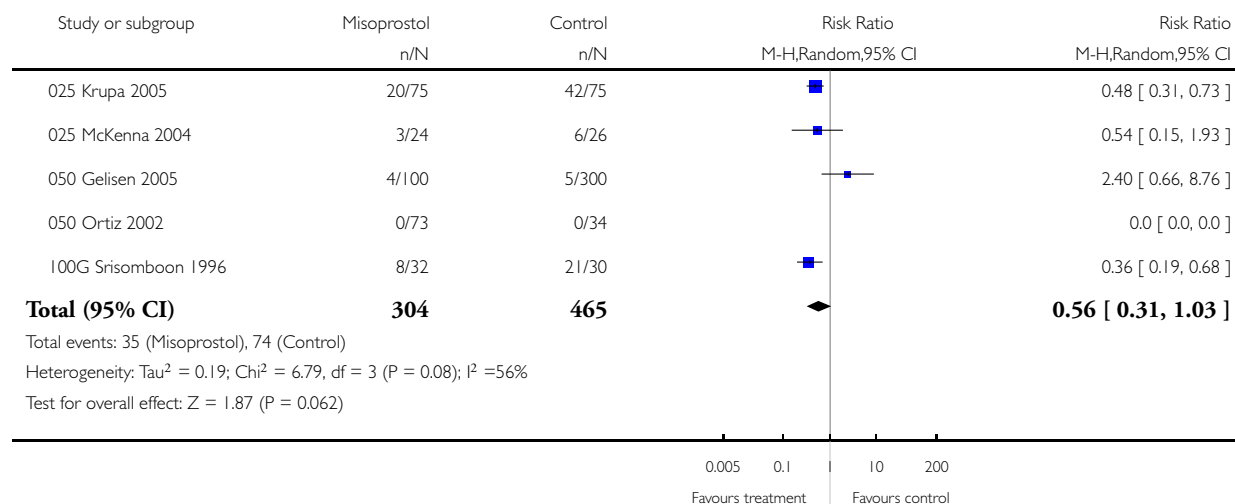


# **Analysis 1.1. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 1 Vaginal delivery not achieved in 24 hours.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 1 Vaginal delivery not achieved in 24 hours

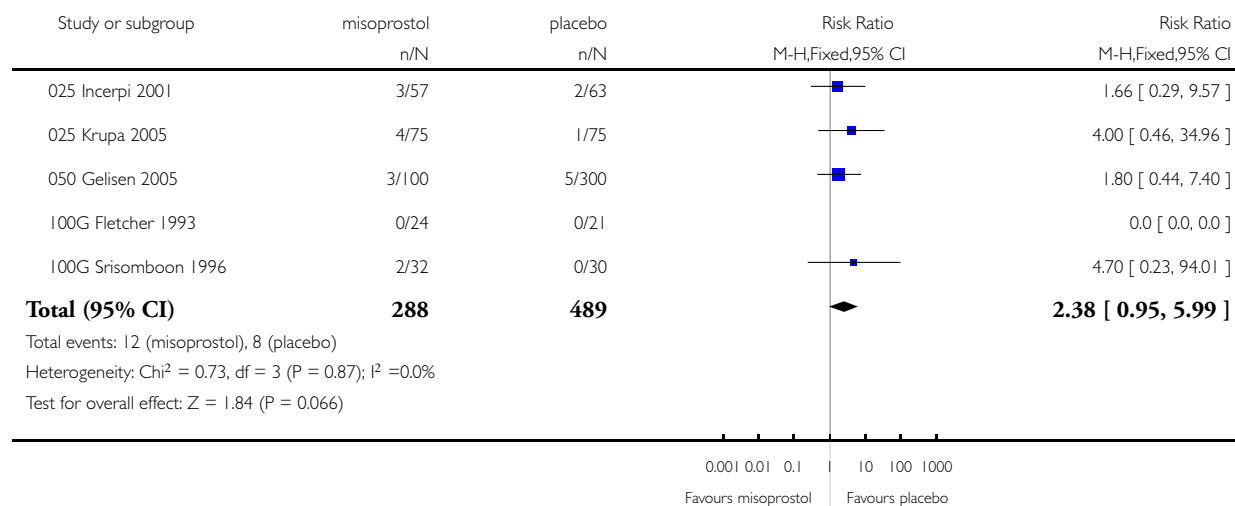


## Analysis 1.2. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 2 Uterine hyperstimulation with FHR changes

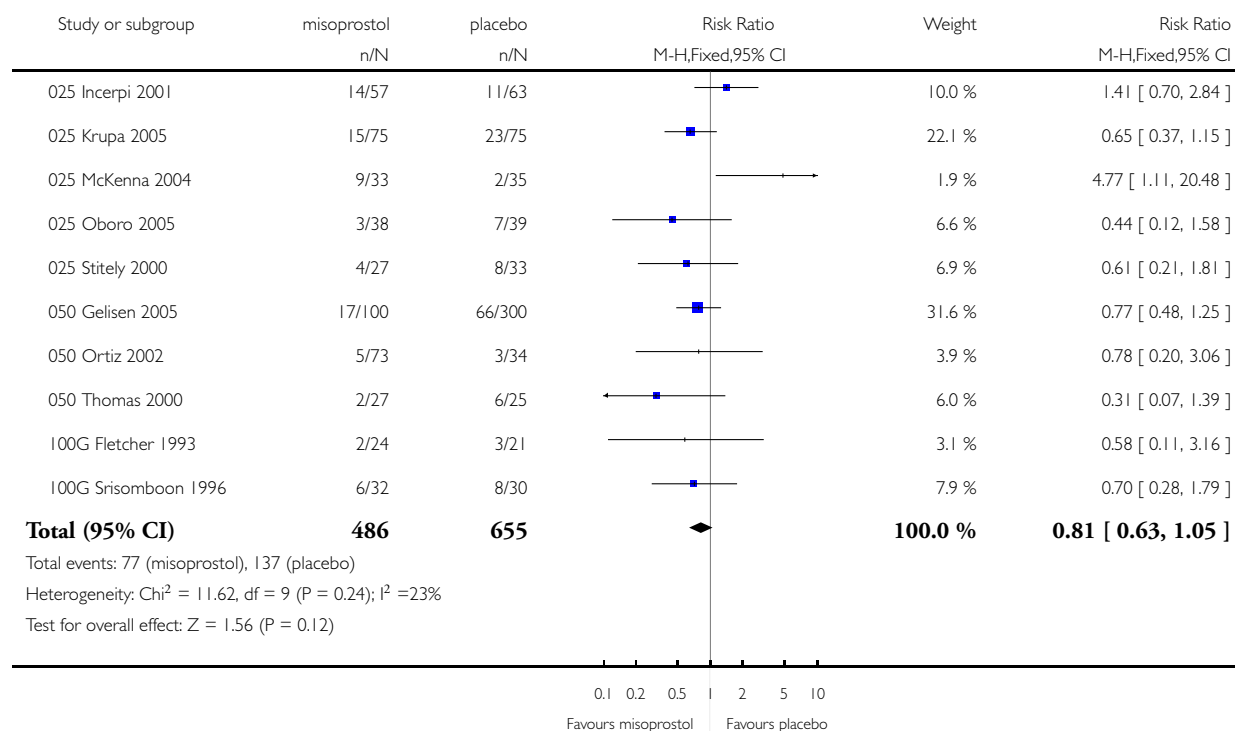


### Analysis 1.3. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 3 Caesarean section

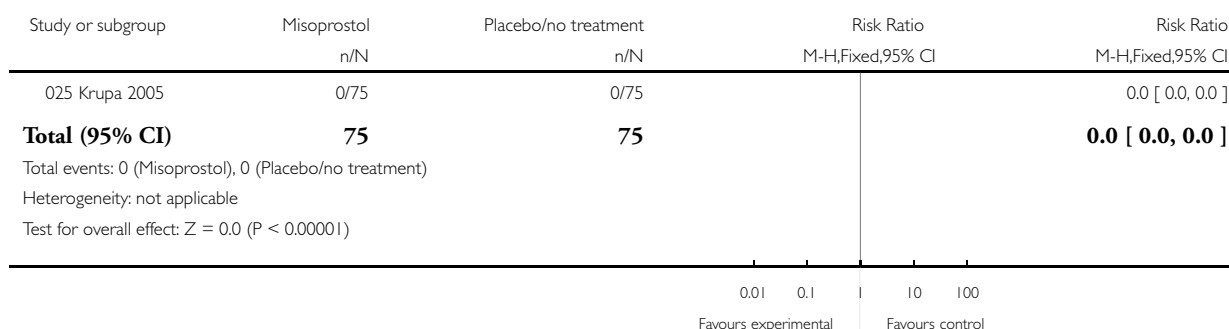


#### Analysis 1.4. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 4 Neonatal encephalopathy.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 4 Neonatal encephalopathy

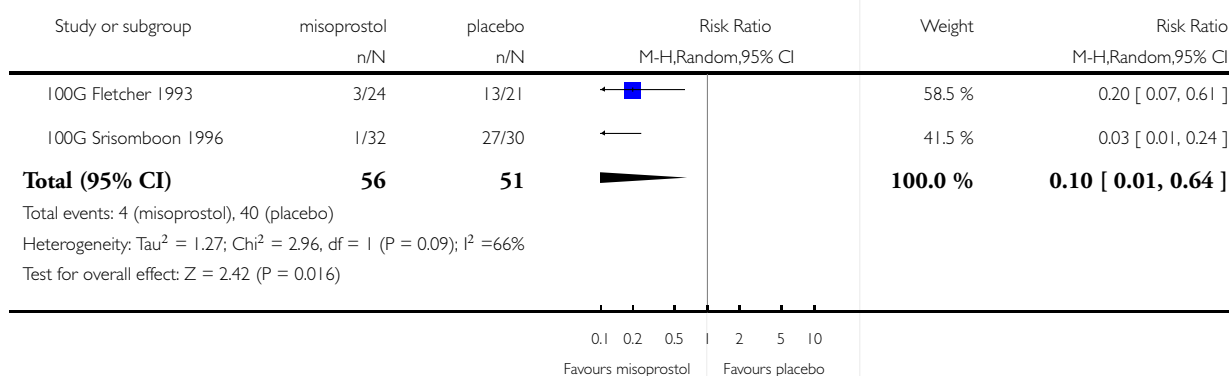


#### Analysis 1.5. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 5 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 5 Cervix unfavourable/unchanged after 12-24 hours

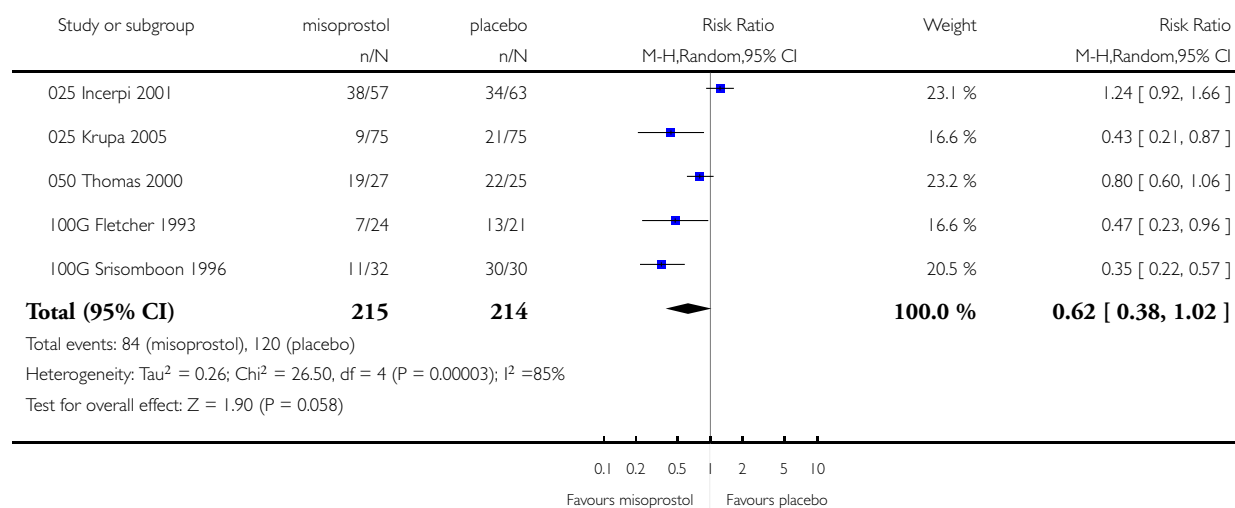


## Analysis 1.6. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 6 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 6 Oxytocin augmentation

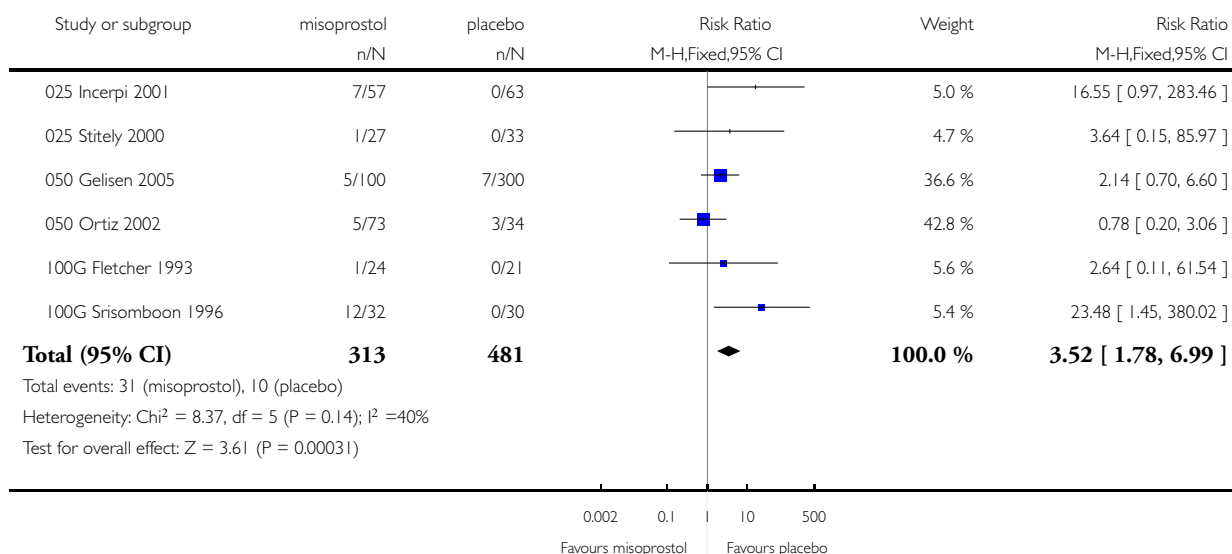


### Analysis 1.7. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 7 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 7 Uterine hyperstimulation without FHR changes

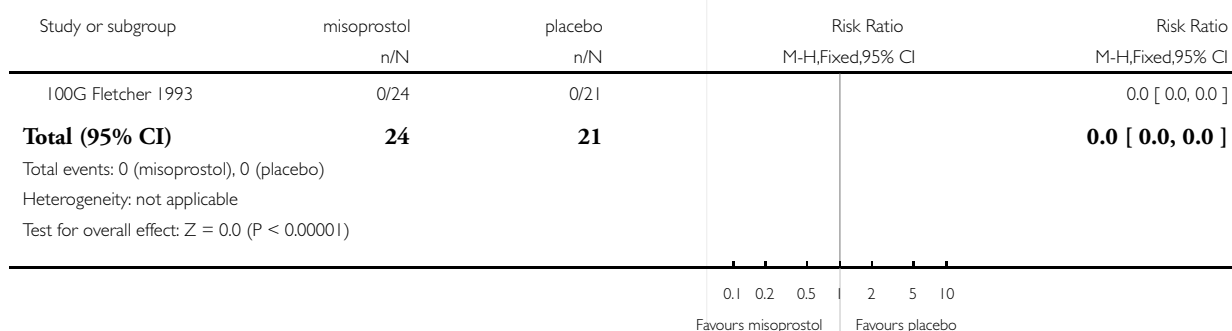


### Analysis 1.8. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 8 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 8 Uterine rupture

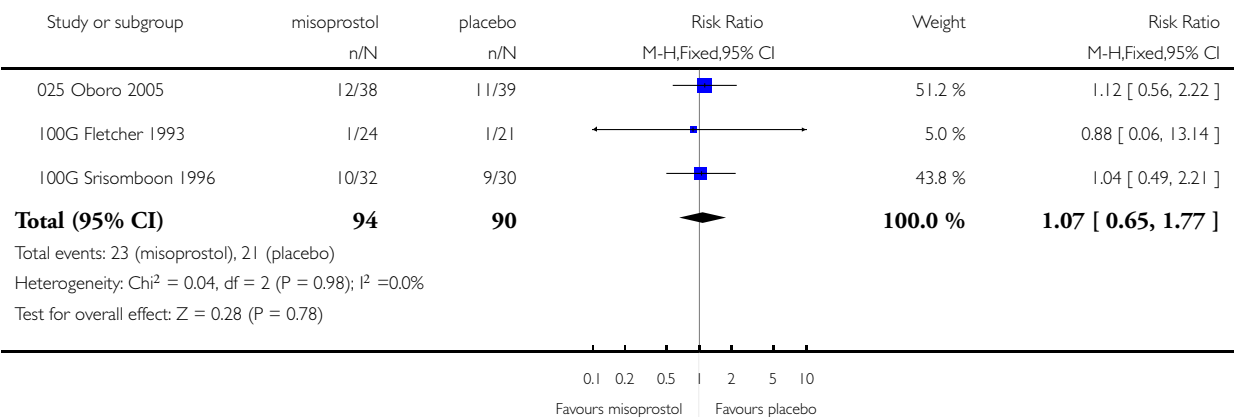


# **Analysis 1.9. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 9 Instrumental vaginal delivery.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 9 Instrumental vaginal delivery

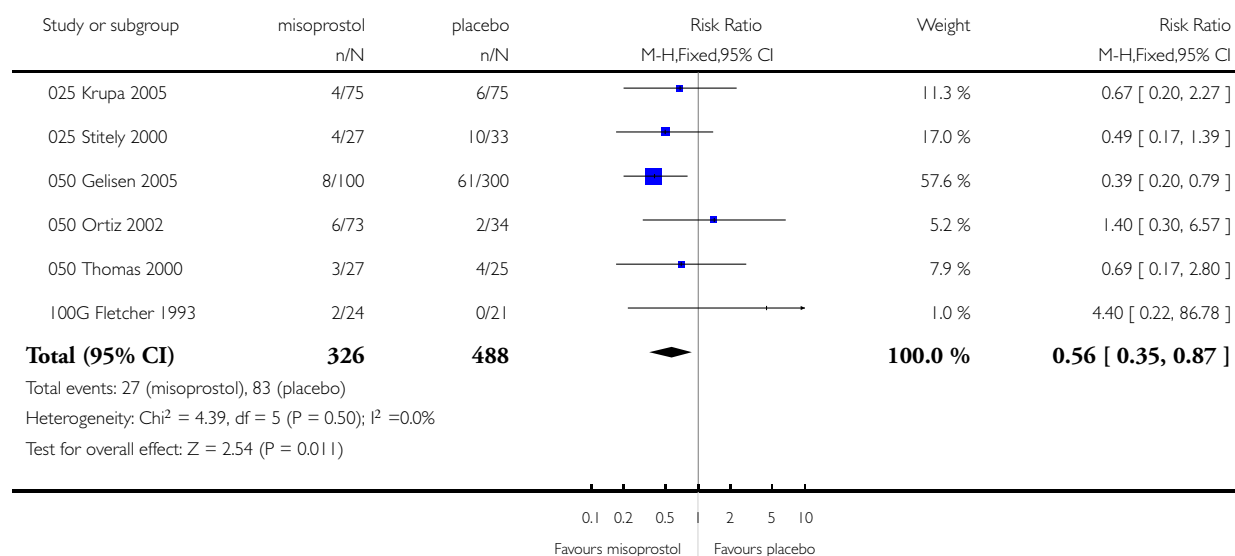


# **Analysis 1.10. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 10 Meconium-stained liquor.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 10 Meconium-stained liquor



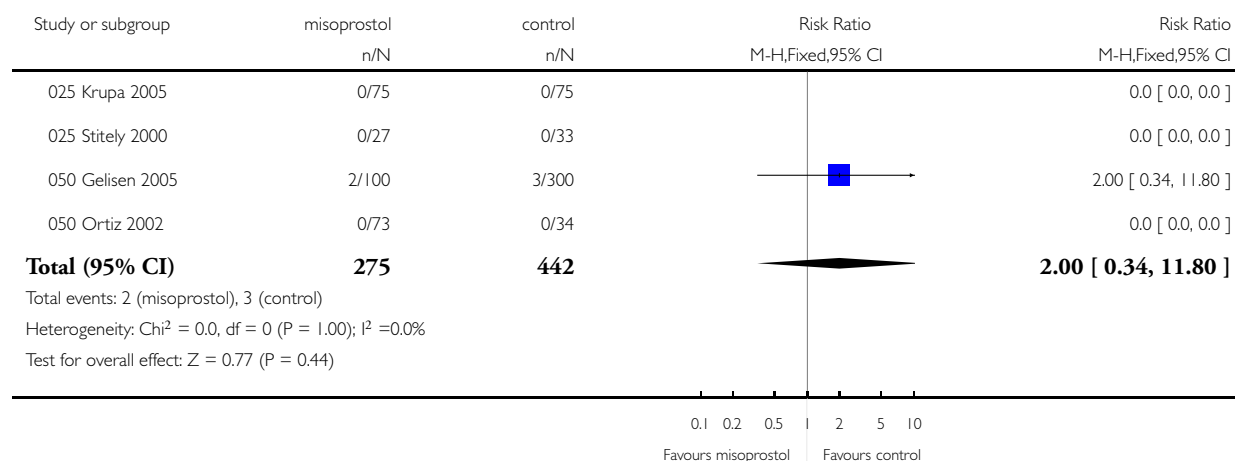


### Analysis 1.11. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 11 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 11 Apgar score < 7 at 5 minutes

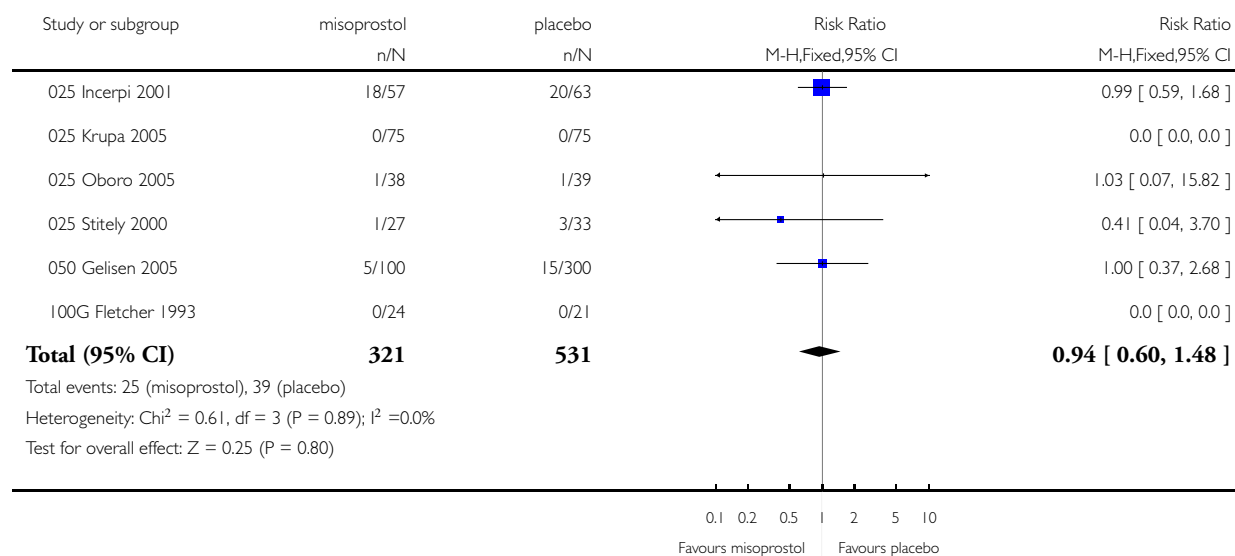


### Analysis 1.12. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 12 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 12 Neonatal intensive care unit admission

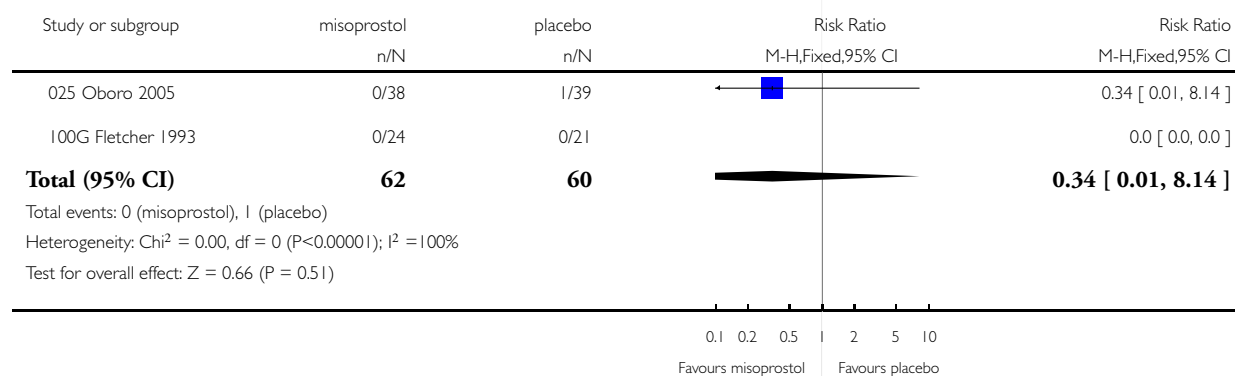


### Analysis 1.13. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 13 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 13 Perinatal death

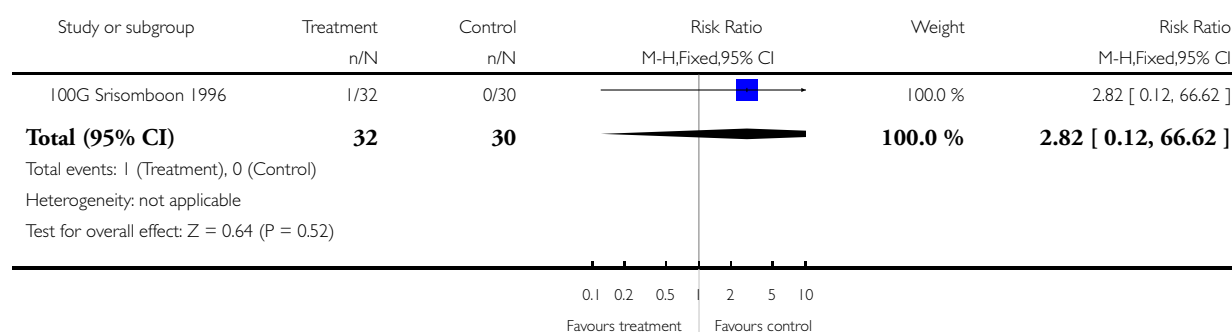


### Analysis 1.14. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 14 Maternal side effects.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 14 Maternal side effects

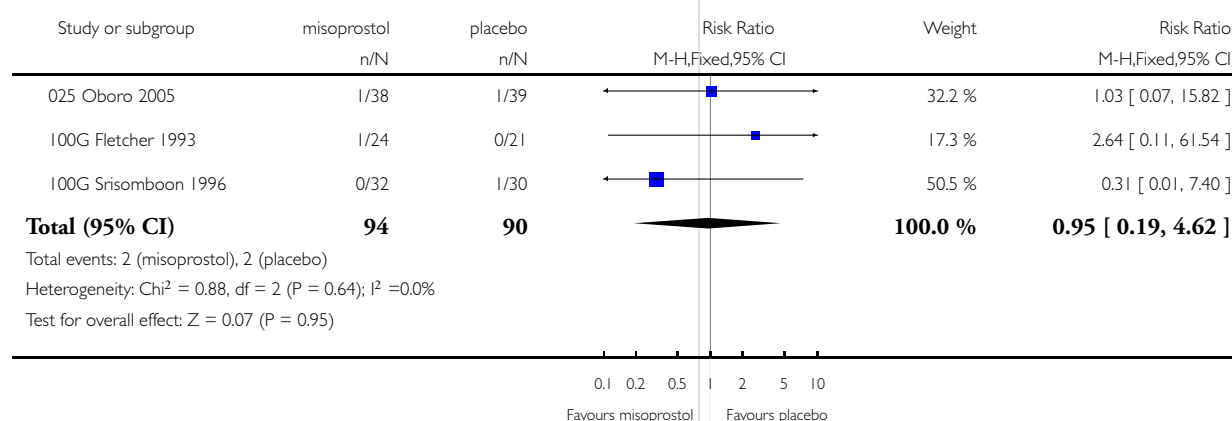


### Analysis 1.15. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 15 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 15 Postpartum haemorrhage

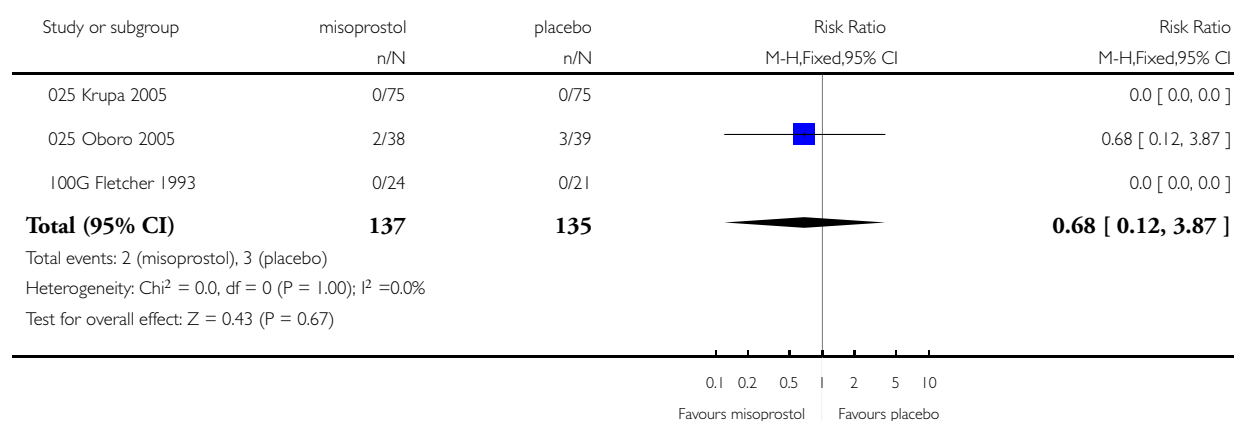


### Analysis 1.16. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 16 Serious maternal complication.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 16 Serious maternal complication

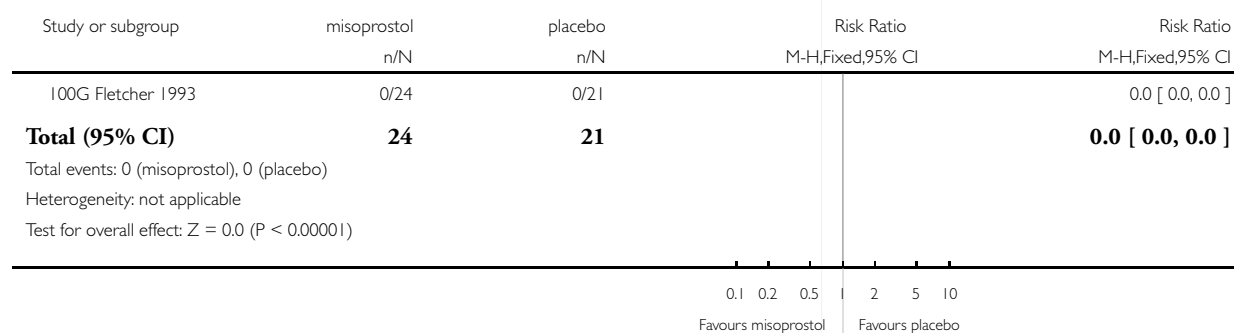


### Analysis 1.17. Comparison 1 Misoprostol versus placebo/no treatment: all women, Outcome 17 Maternal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 1 Misoprostol versus placebo/no treatment: all women

Outcome: 17 Maternal death

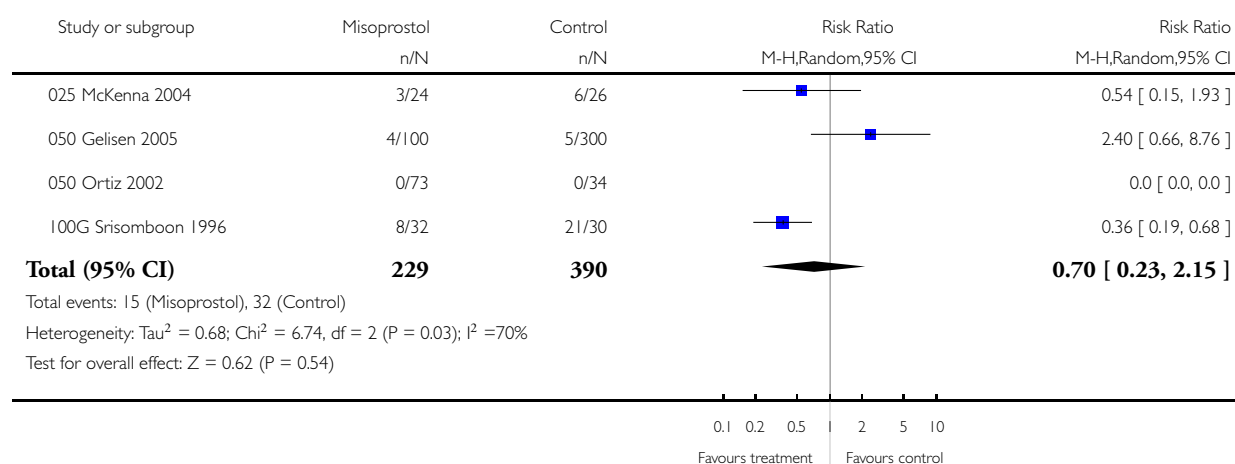


## Analysis 2.1. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 1 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved in 24 hours

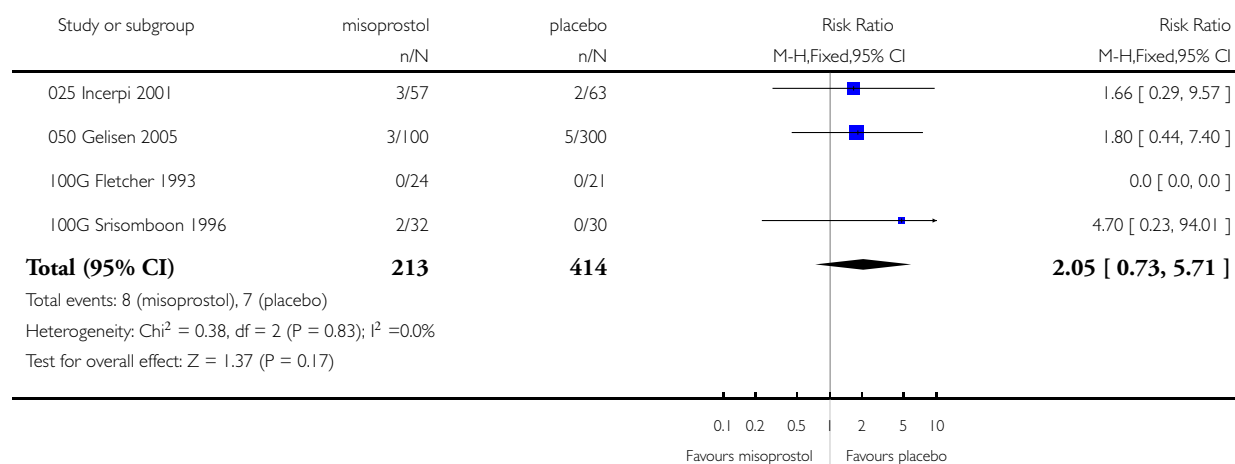


## Analysis 2.2. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

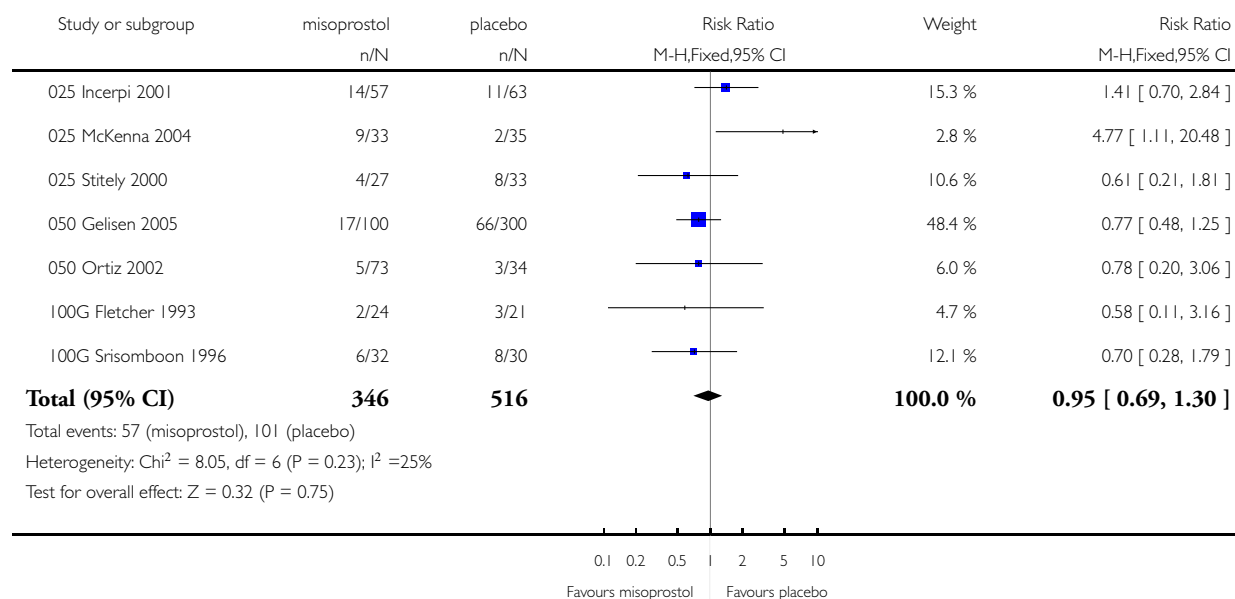


### Analysis 2.3. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 3 Caesarean section

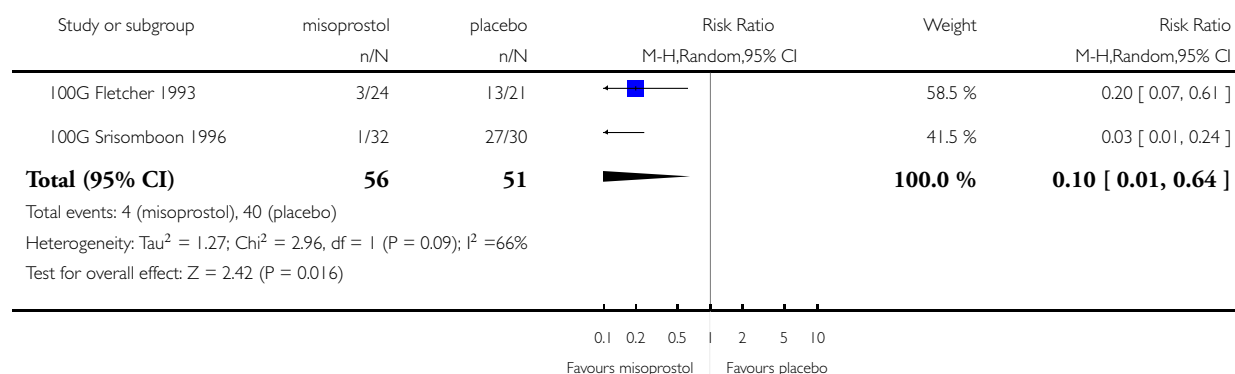


#### Analysis 2.4. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 4 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 4 Cervix unfavourable/unchanged after 12-24 hours

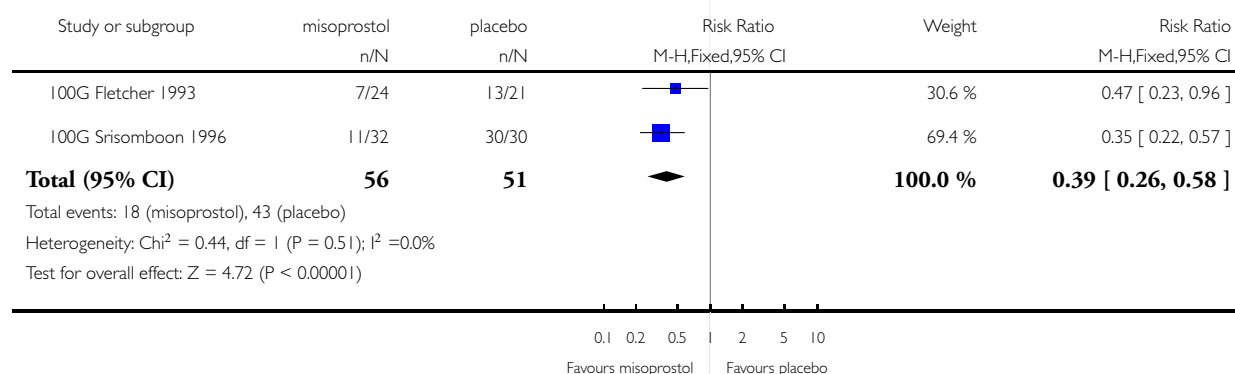


#### Analysis 2.5. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 5 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 5 Oxytocin augmentation



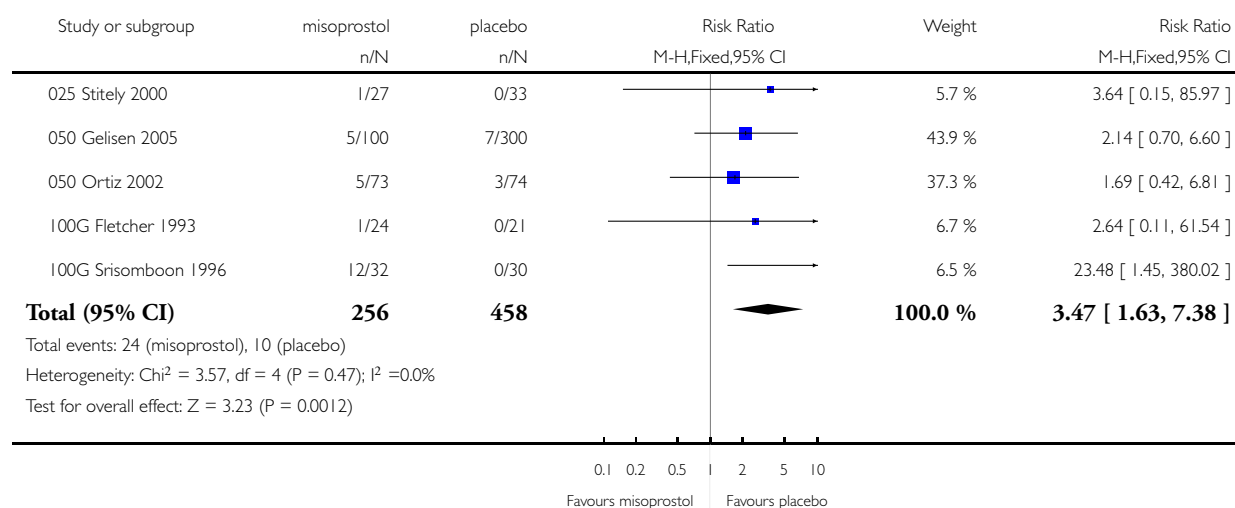


## Analysis 2.6. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 6 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 6 Uterine hyperstimulation without FHR changes

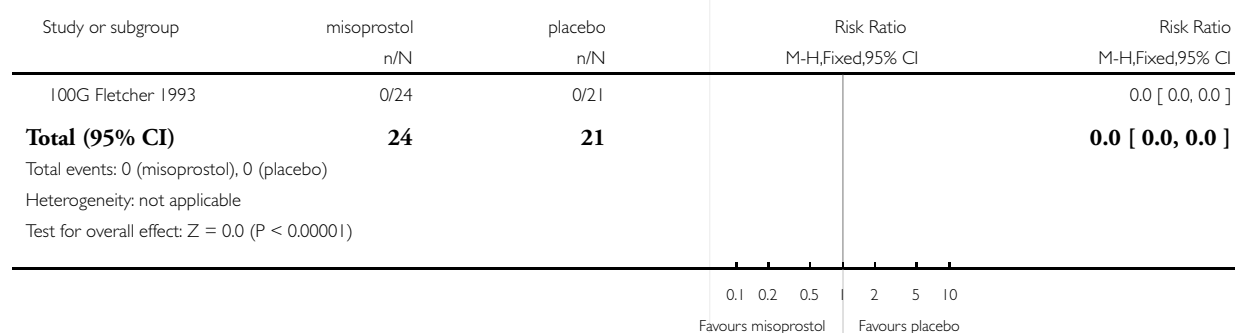


## Analysis 2.7. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 7 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 7 Uterine rupture

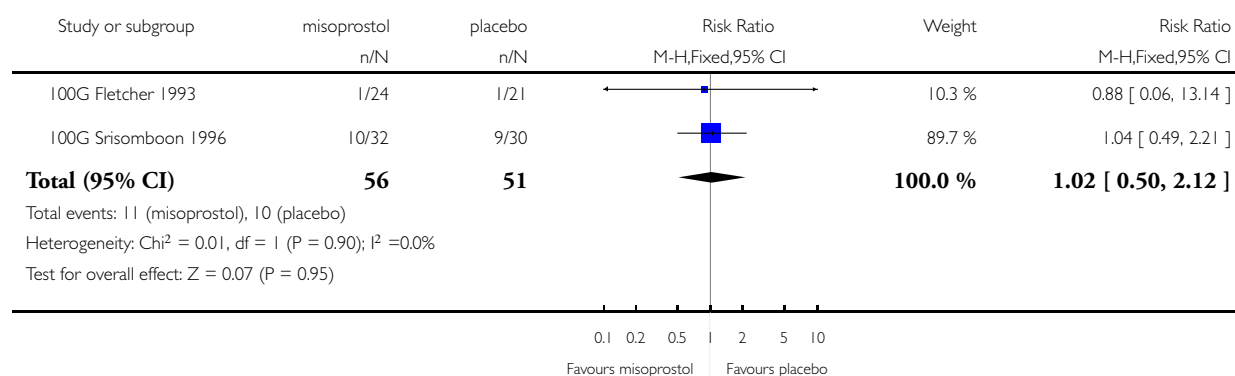


## Analysis 2.8. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 8 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 8 Instrumental vaginal delivery

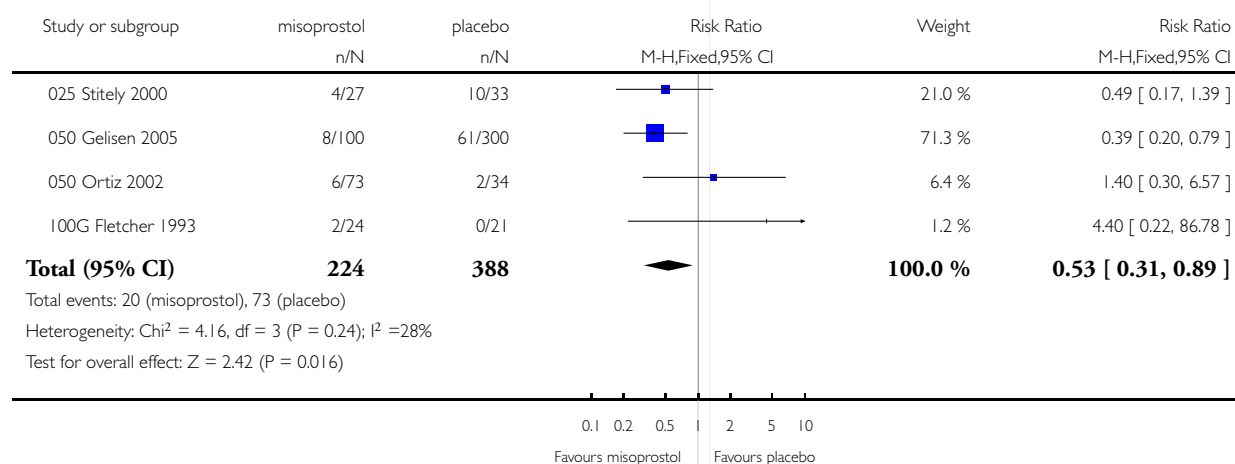


## Analysis 2.9. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 9 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 9 Meconium-stained liquor

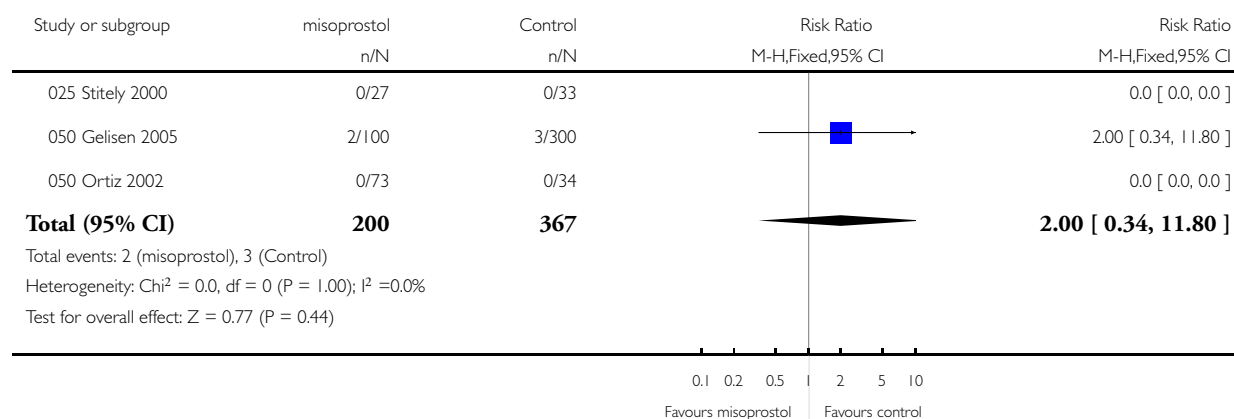


### Analysis 2.10. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 10 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 10 Apgar score < 7 at 5 minutes

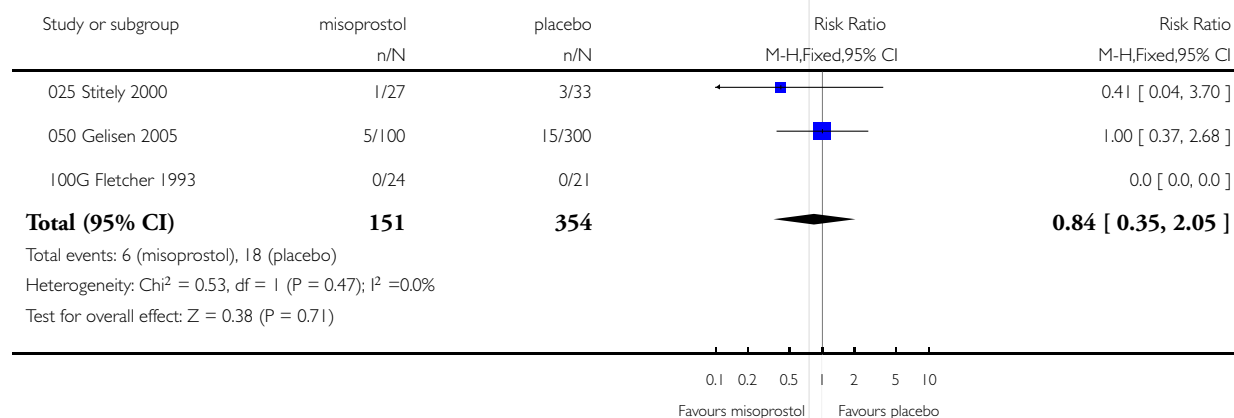


### Analysis 2.11. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 11 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 11 Neonatal intensive care unit admission

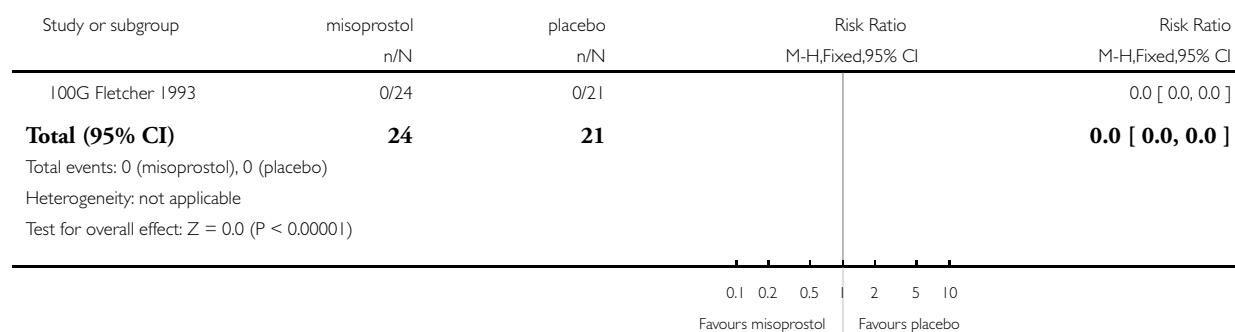


## Analysis 2.12. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 12 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 12 Perinatal death

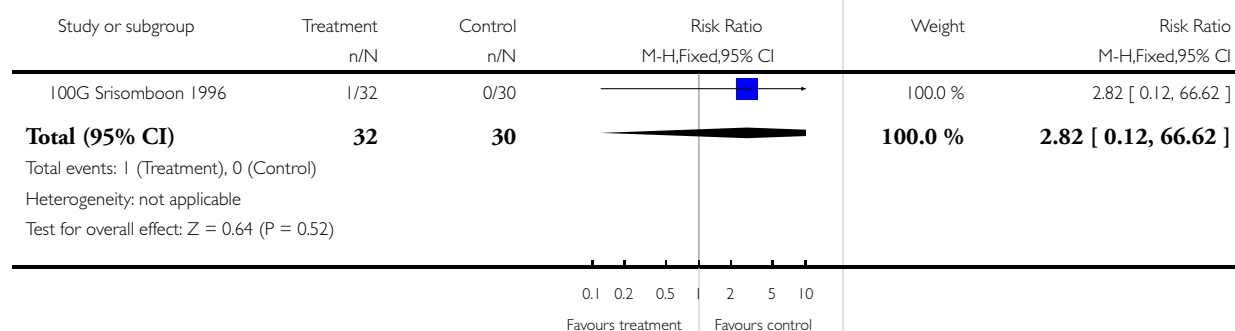


## Analysis 2.13. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 13 Maternal side effects.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 13 Maternal side effects

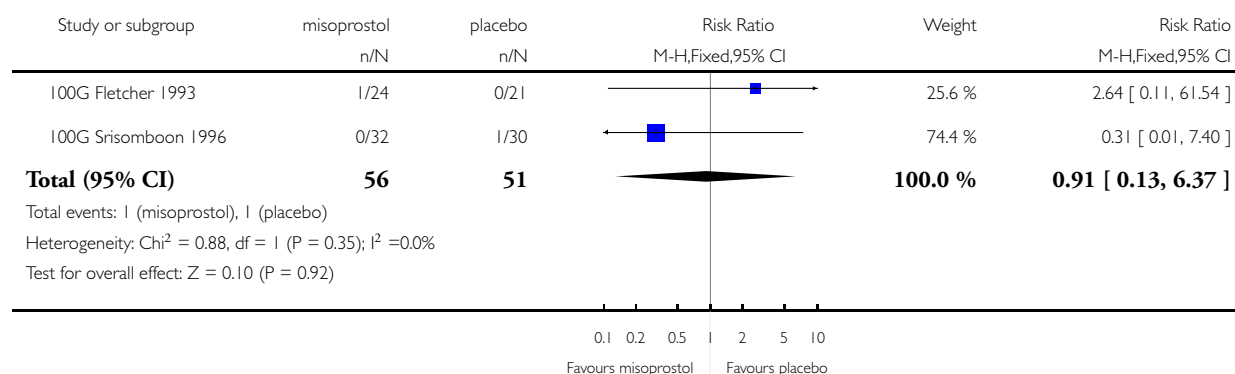


### Analysis 2.14. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 14 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 14 Postpartum haemorrhage

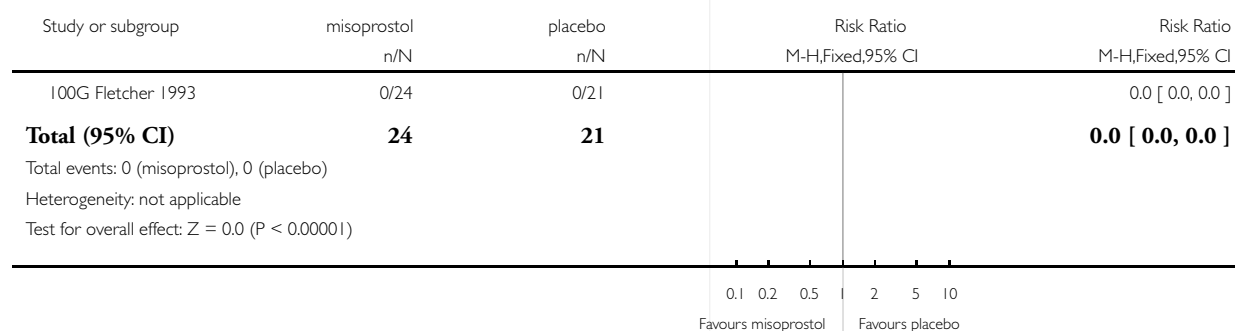


### Analysis 2.15. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 15 Serious maternal complication.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 15 Serious maternal complication

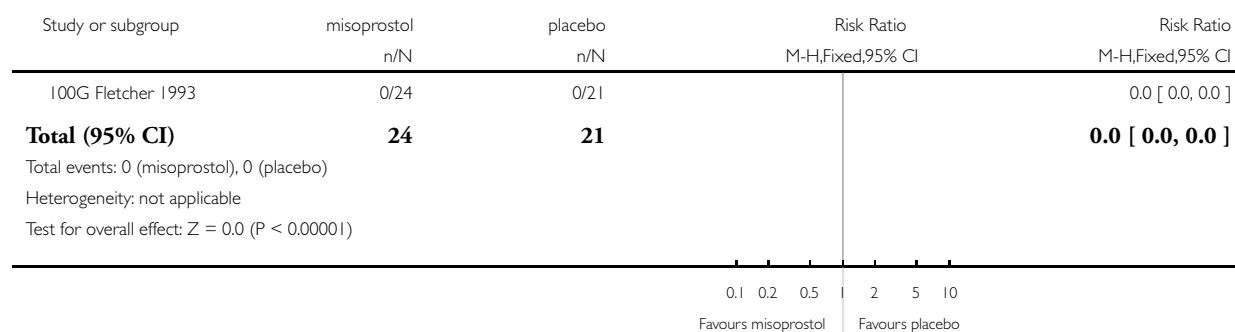


## Analysis 2.16. Comparison 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix, Outcome 16 Maternal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 2 Misoprostol versus placebo/no treatment: all women, unfavourable cervix

Outcome: 16 Maternal death

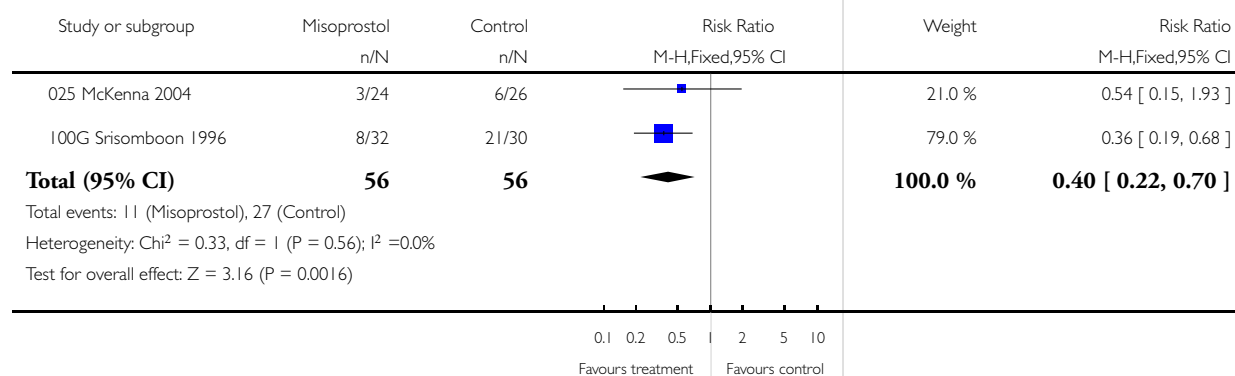


## Analysis 3.1. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 1 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved in 24 hours

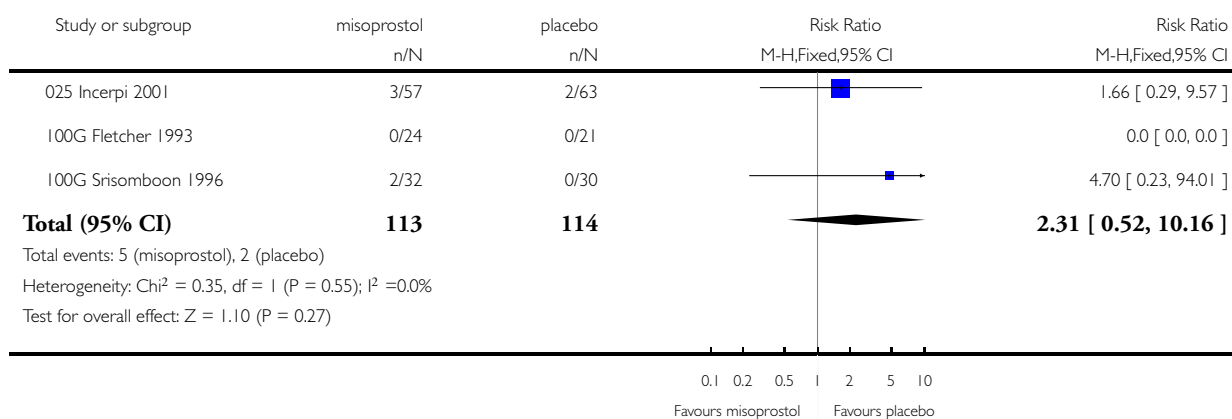


### Analysis 3.2. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

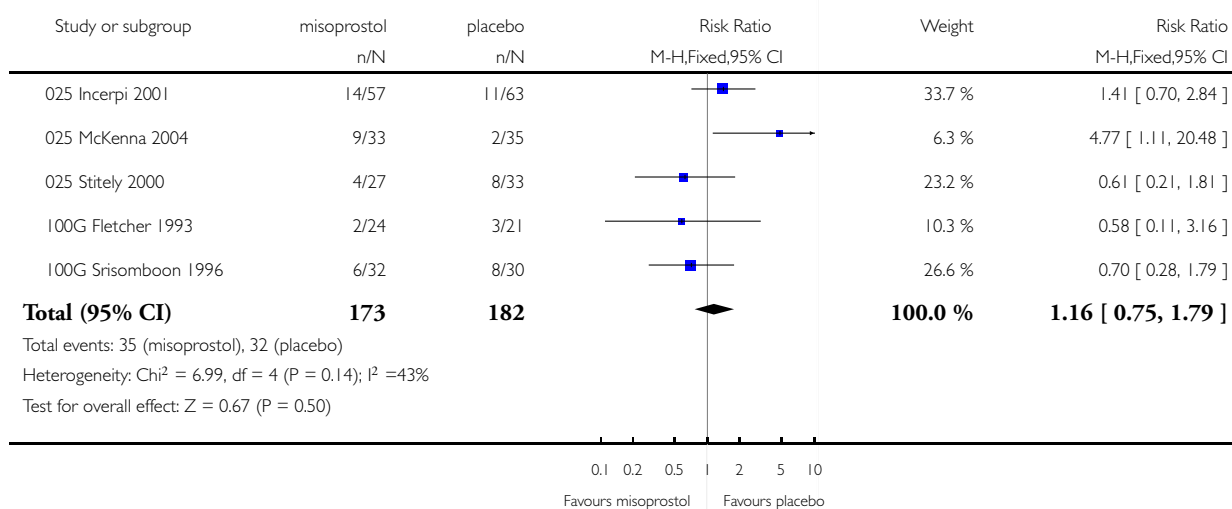


### Analysis 3.3. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 3 Caesarean section

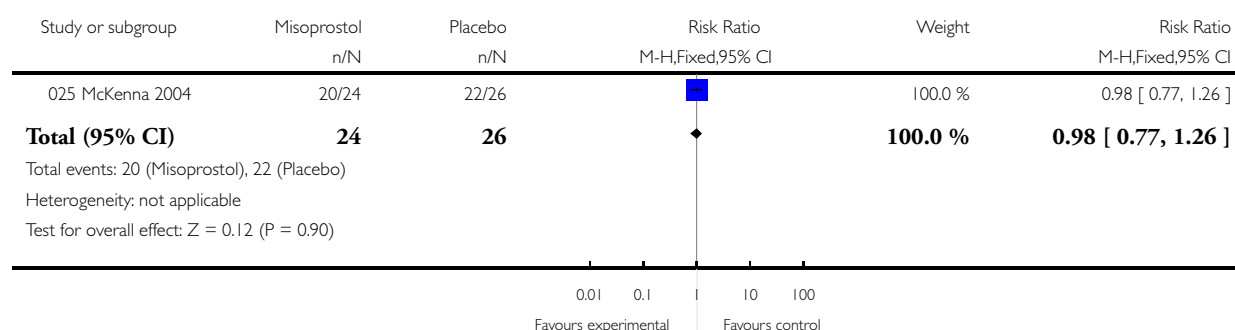


### Analysis 3.4. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 4 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 4 Epidural analgesia

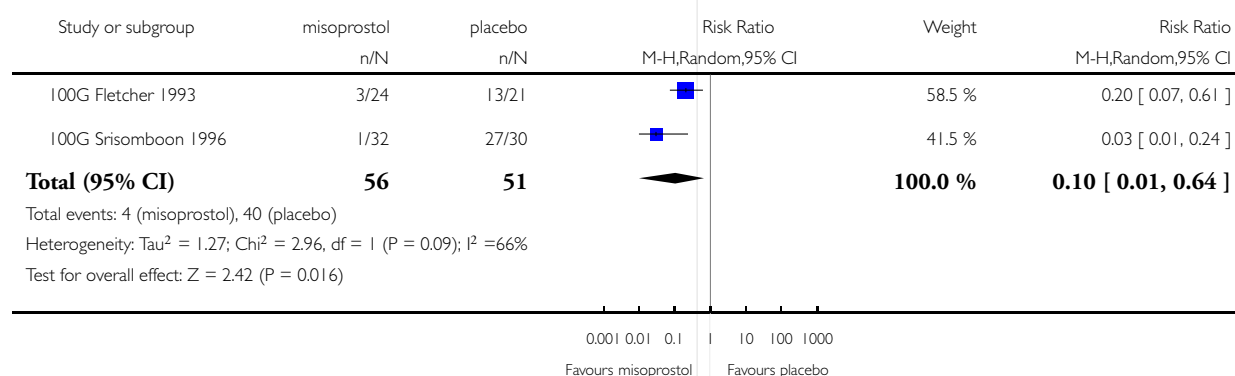


### Analysis 3.5. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 5 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 5 Cervix unfavourable/unchanged after 12-24 hours



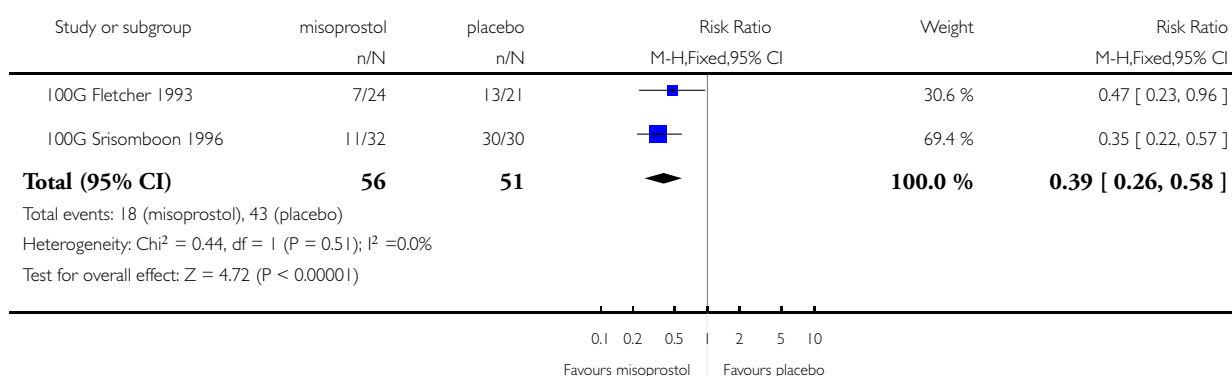


### Analysis 3.6. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 6 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 6 Oxytocin augmentation

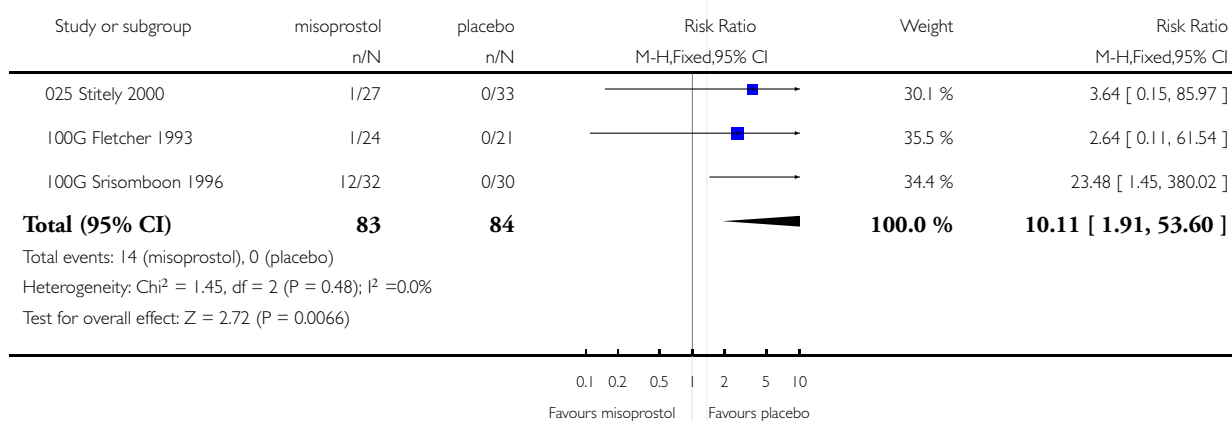


### Analysis 3.7. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 7 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 7 Uterine hyperstimulation without FHR changes

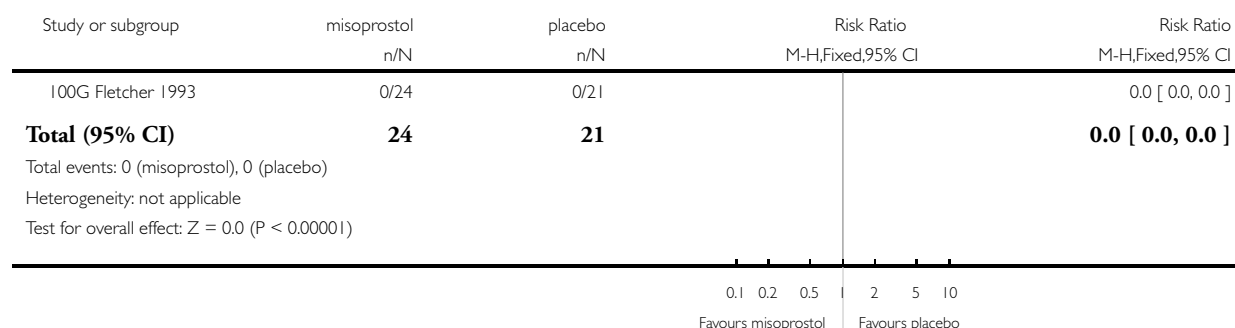


### Analysis 3.8. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 8 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 8 Uterine rupture

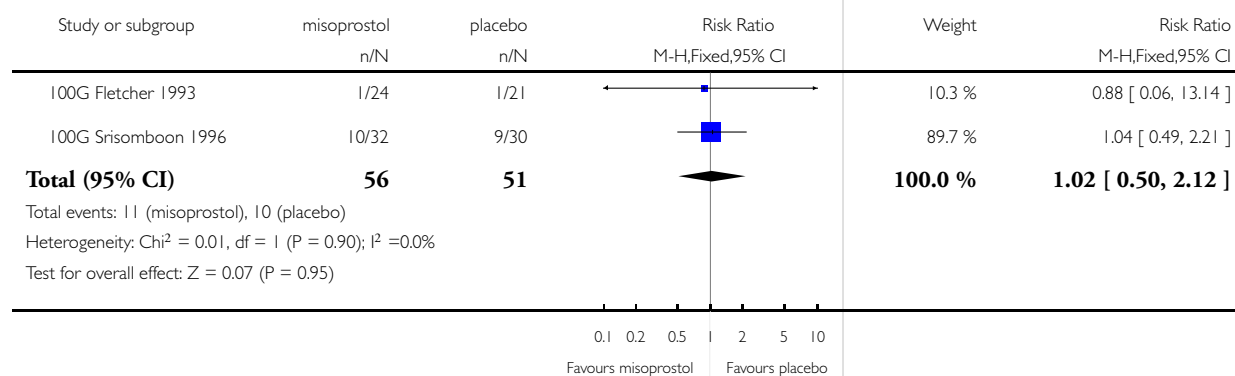


### Analysis 3.9. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 9 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 9 Instrumental vaginal delivery

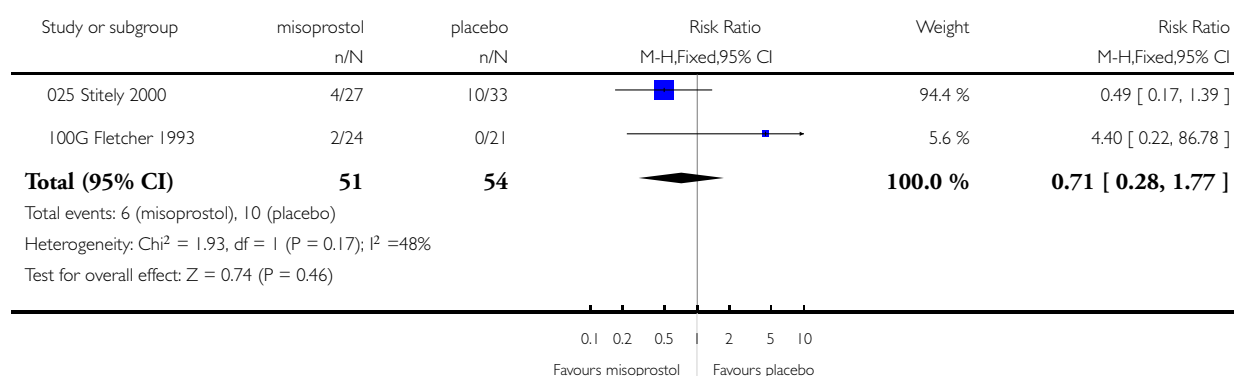


### Analysis 3.10. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 10 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 10 Meconium-stained liquor

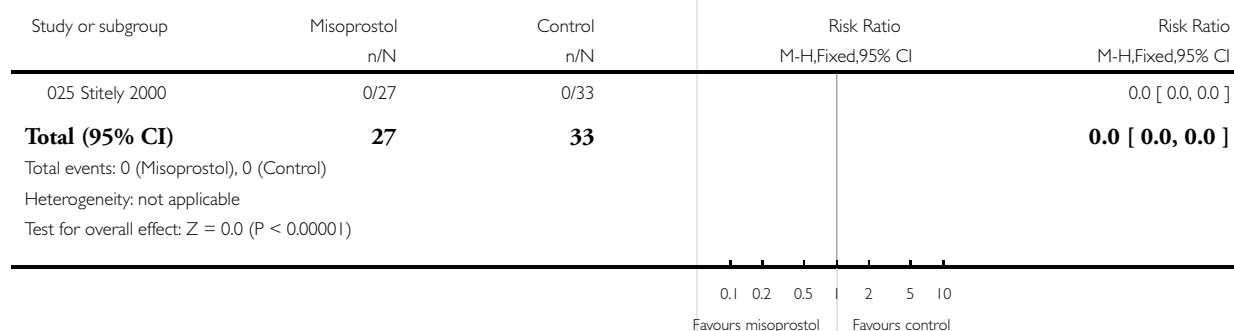


### Analysis 3.11. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 11 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 11 Apgar score < 7 at 5 minutes

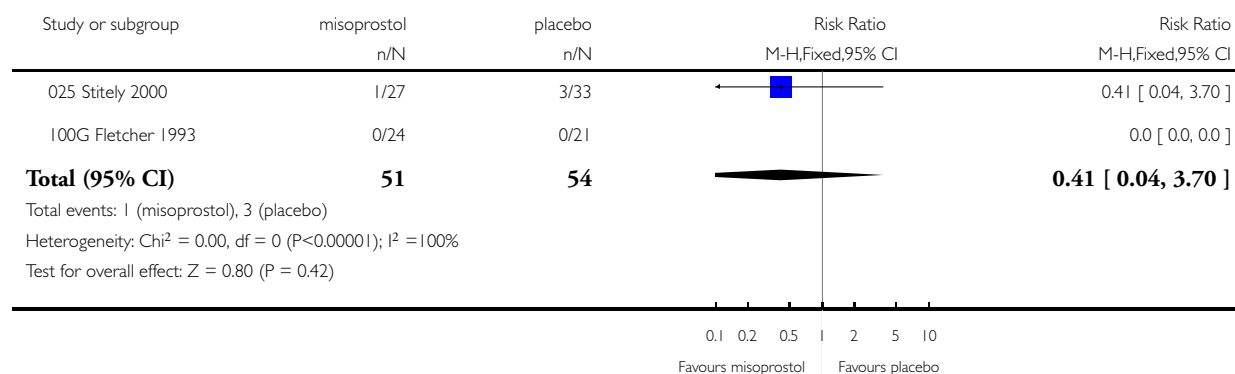


### Analysis 3.12. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 12 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 12 Neonatal intensive care unit admission

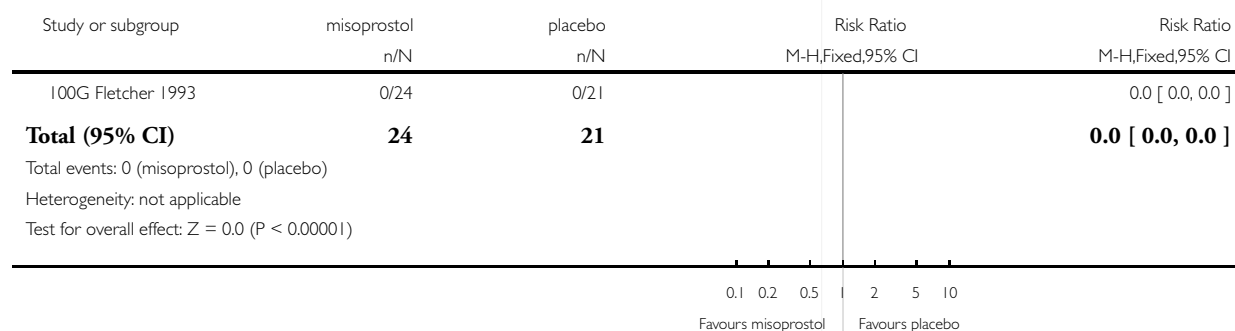


### Analysis 3.13. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 13 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 13 Perinatal death

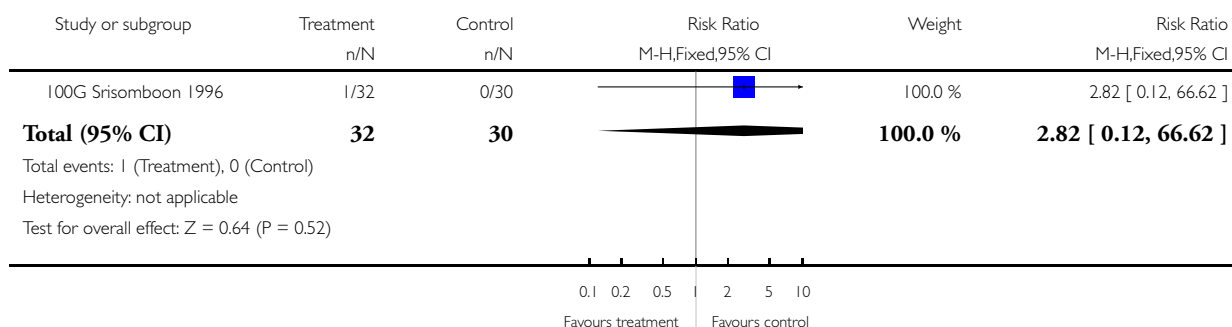


### Analysis 3.14. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 14 Maternal side effects.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 14 Maternal side effects

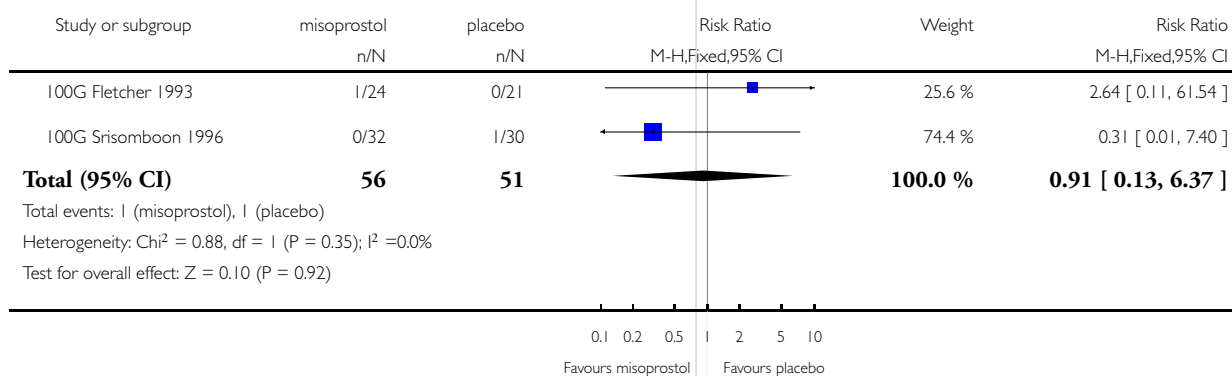


### Analysis 3.15. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 15 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 15 Postpartum haemorrhage

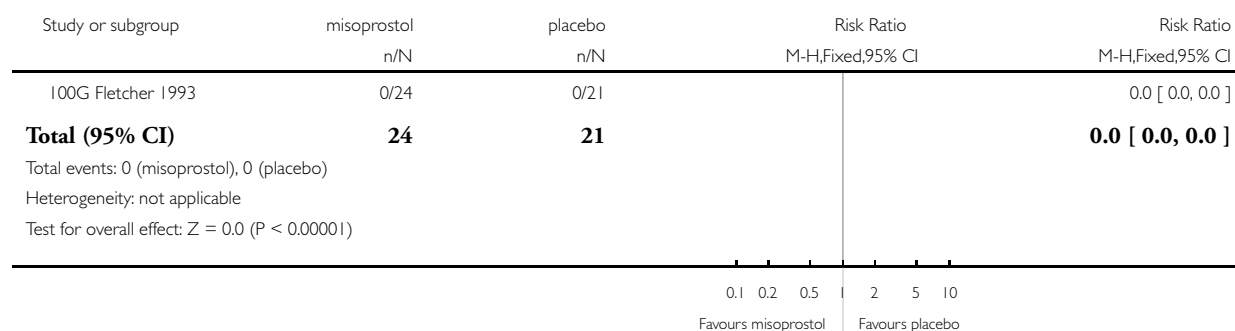


### Analysis 3.16. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 16 Serious maternal complication.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 16 Serious maternal complication

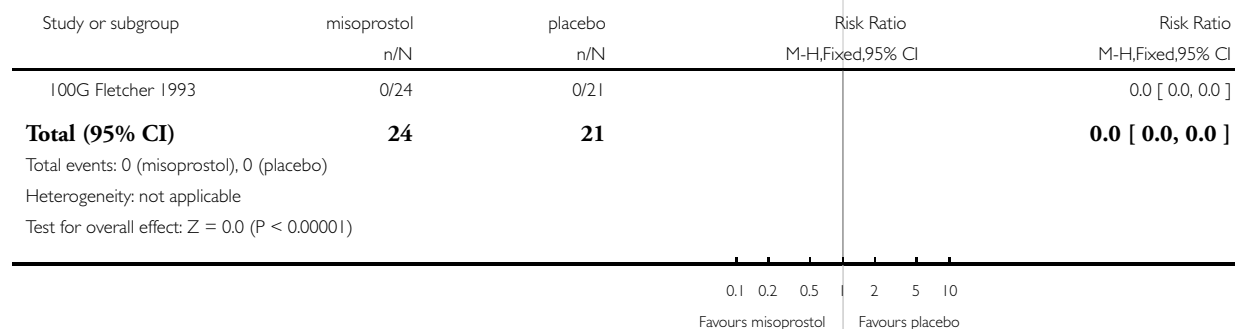


### Analysis 3.17. Comparison 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix, Outcome 17 Maternal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 3 Misoprostol versus placebo/no treatment: all women, intact membranes, unfavourable cervix

Outcome: 17 Maternal death

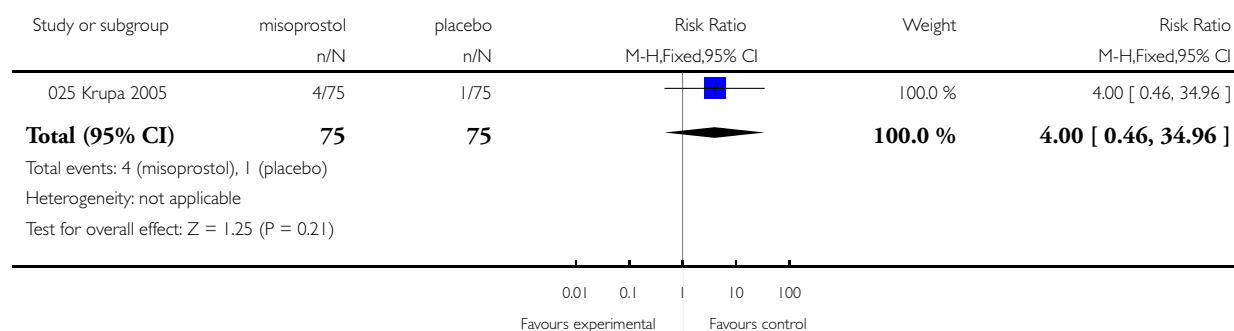


#### Analysis 4.1. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 1 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 1 Uterine hyperstimulation with FHR changes

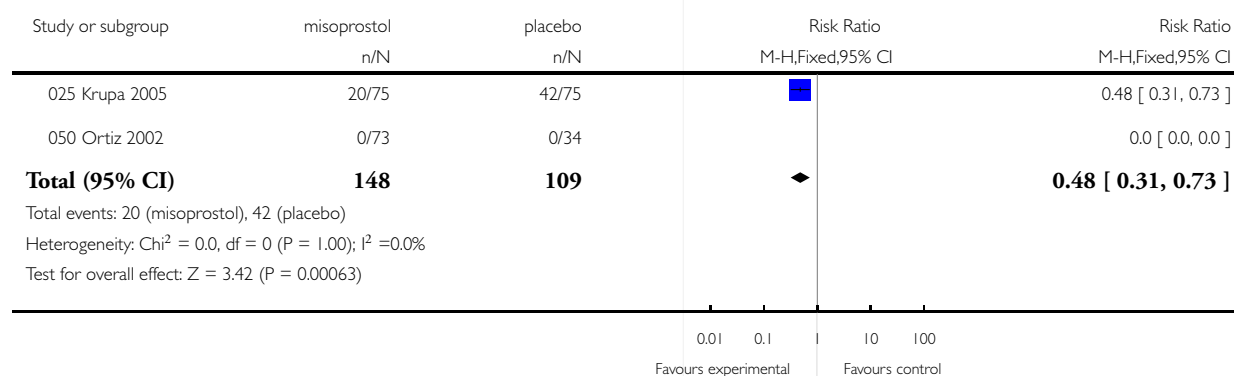


#### Analysis 4.2. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 2 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 2 Vaginal delivery not achieved in 24 hours

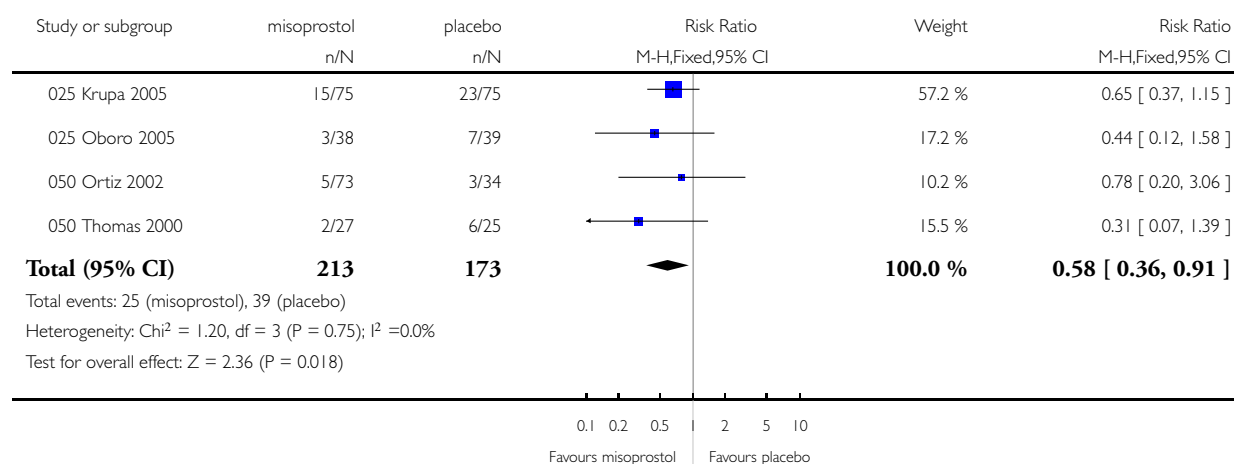


### Analysis 4.3. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 3 Caesarean section

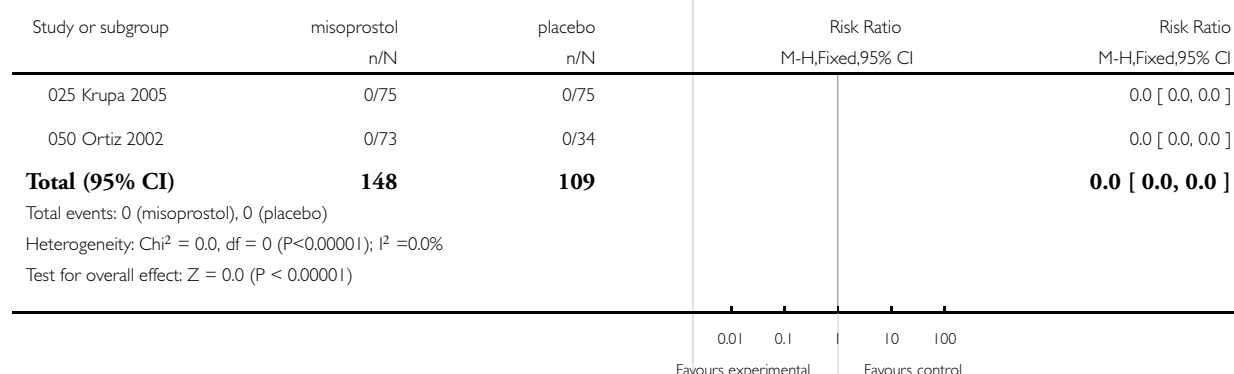


### Analysis 4.4. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 4 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 4 Apgar score < 7 at 5 minutes



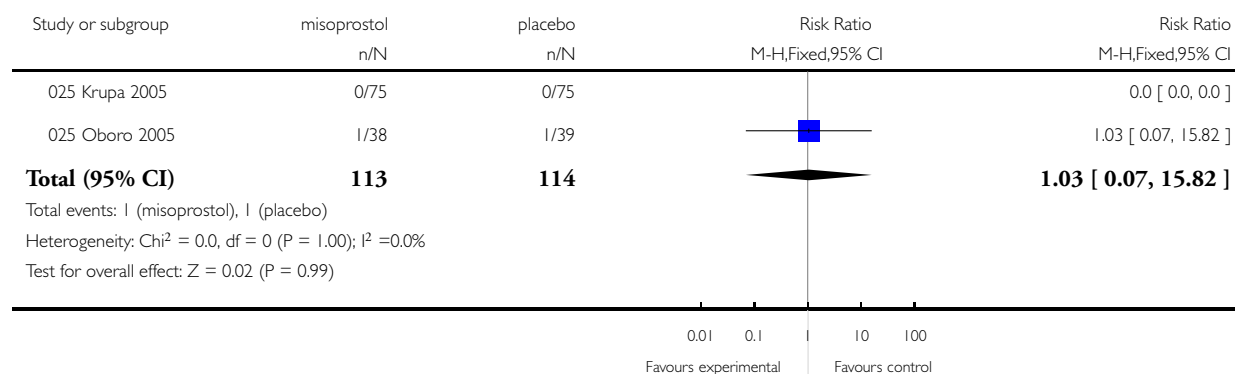


#### Analysis 4.5. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 5 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 5 Neonatal intensive care unit admission

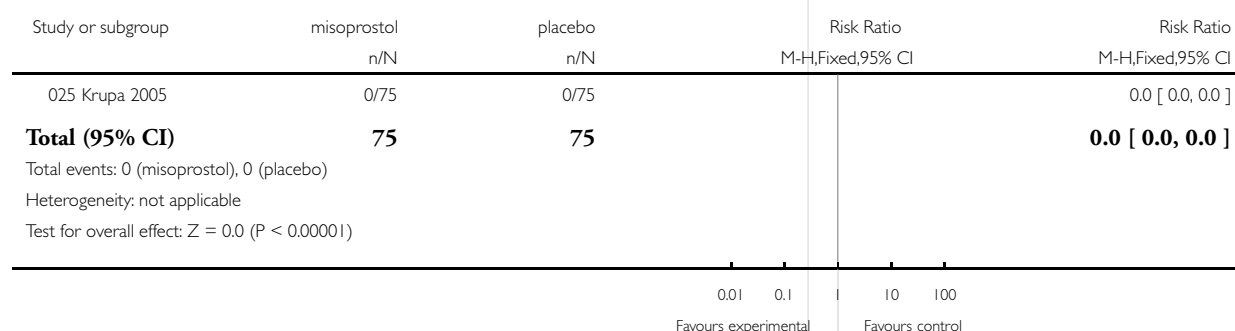


#### Analysis 4.6. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 6 Neonatal encephalopathy.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 6 Neonatal encephalopathy

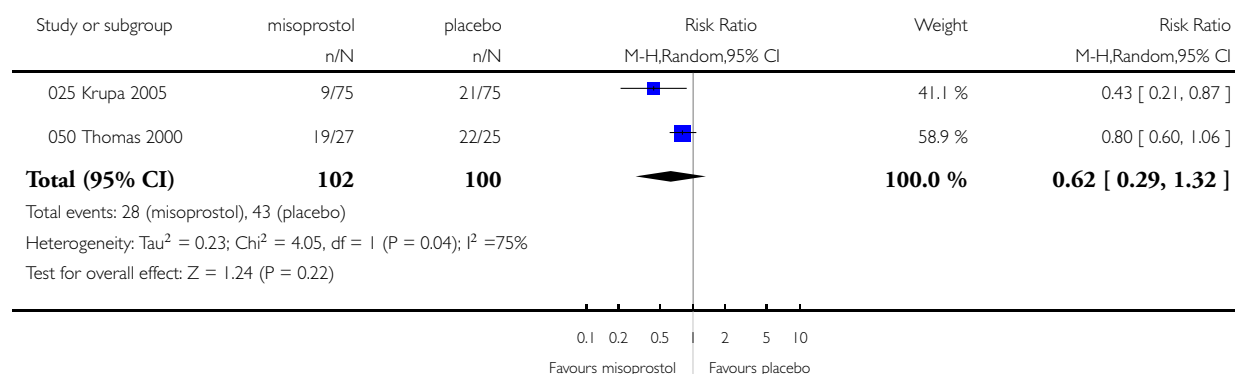


#### Analysis 4.7. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 7 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 7 Oxytocin augmentation

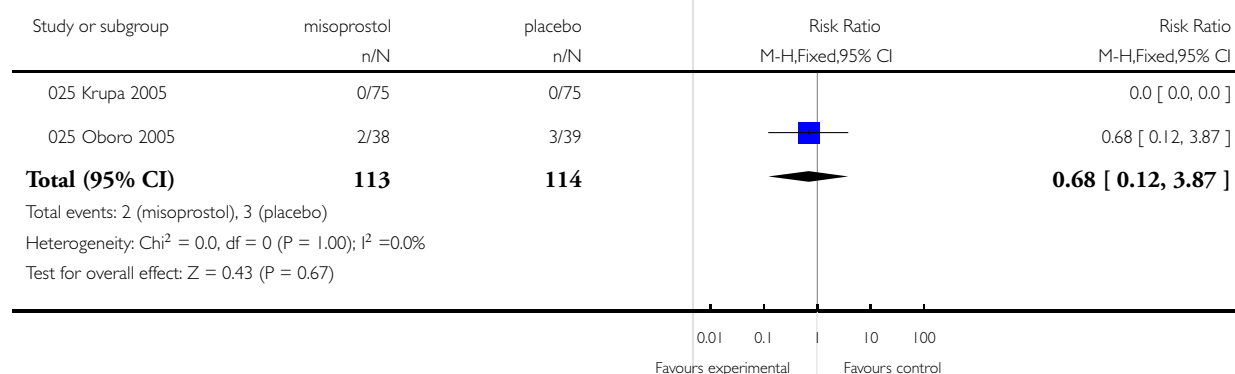


#### Analysis 4.8. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 8 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 8 Serious maternal complications

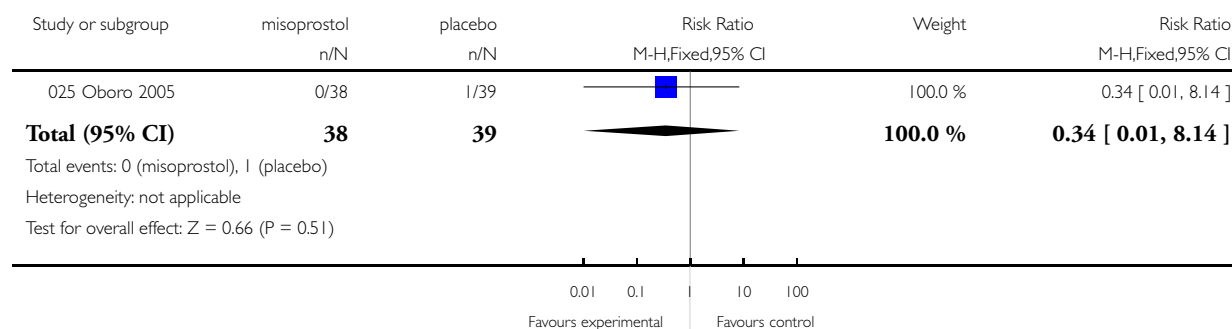


#### Analysis 4.9. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 9 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 9 Perinatal death

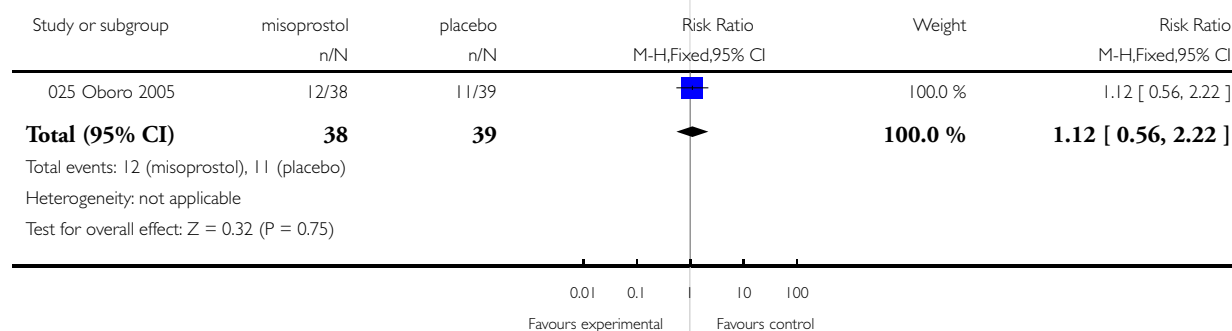


#### Analysis 4.10. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 10 Instrumental delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 10 Instrumental delivery

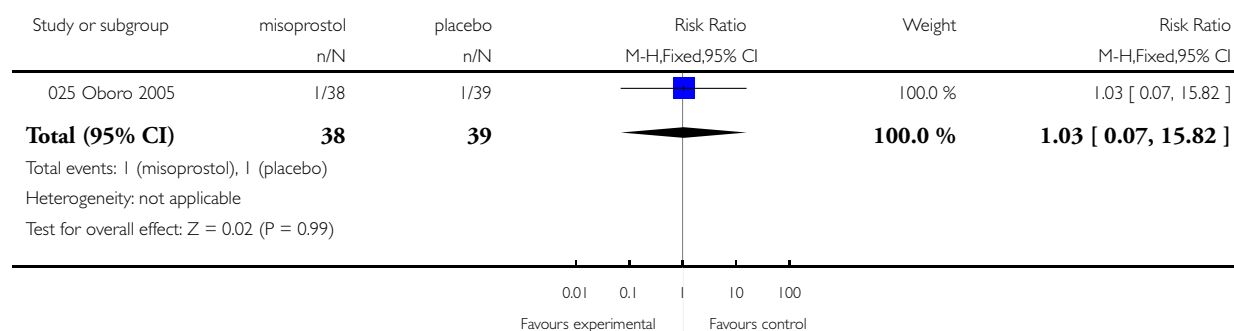


#### Analysis 4.11. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 11 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 11 Postpartum haemorrhage

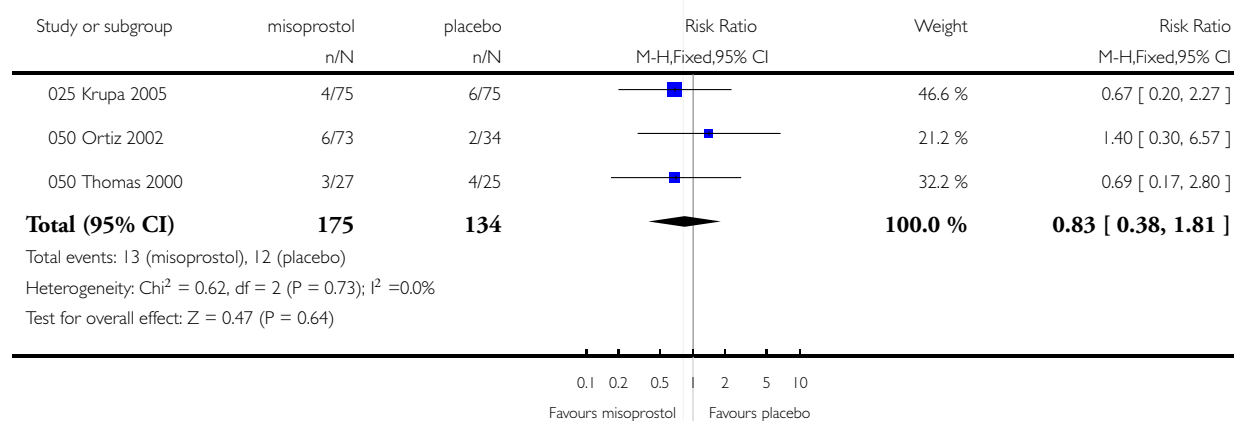


#### Analysis 4.12. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 12 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 12 Meconium-stained liquor

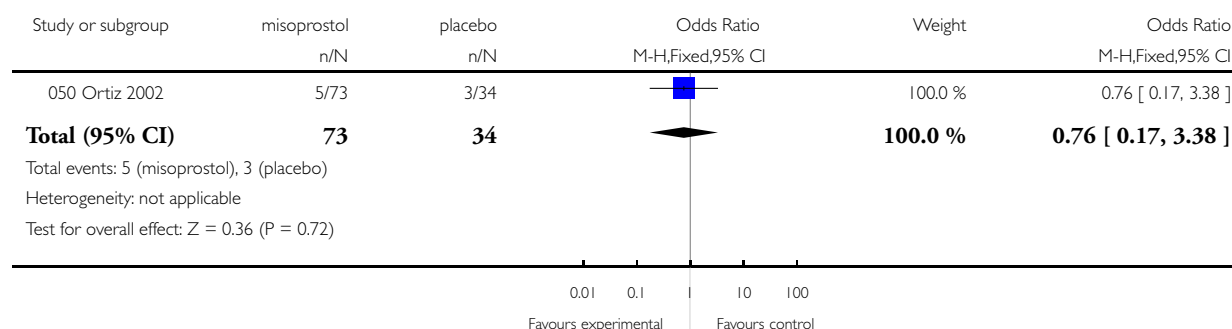


### Analysis 4.13. Comparison 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix, Outcome 13 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 4 Misoprostol versus placebo/no treatment: all women, ruptured membranes, variable or undefined cervix

Outcome: 13 Uterine hyperstimulation without FHR changes

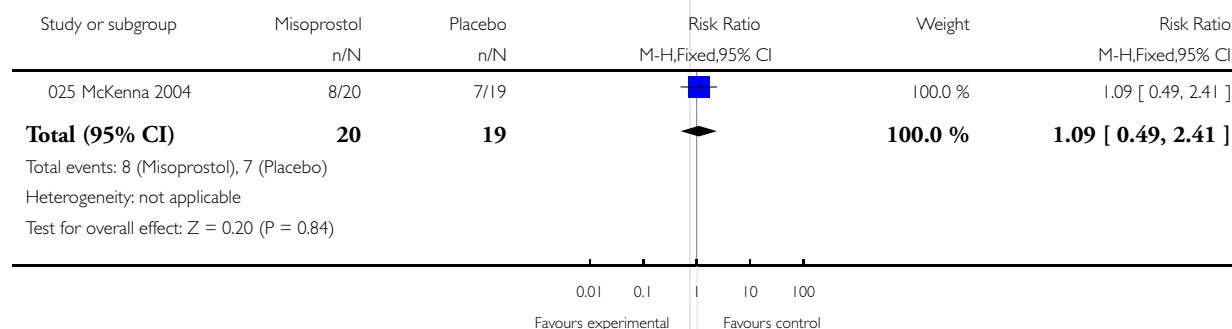


### Analysis 5.1. Comparison 5 Misoprostol versus placebo/no treatment: all primiparae, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 5 Misoprostol versus placebo/no treatment: all primiparae

Outcome: 1 Caesarean section

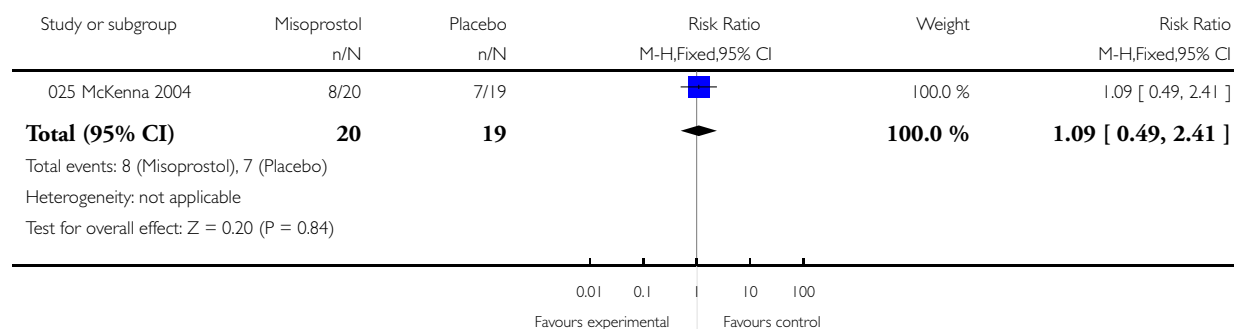


### Analysis 6.1. Comparison 6 Misoprostol versus placebo/no treatment: all primiparae and unfavourable cervix, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 6 Misoprostol versus placebo/no treatment: all primiparae and unfavourable cervix

Outcome: 1 Caesarean section

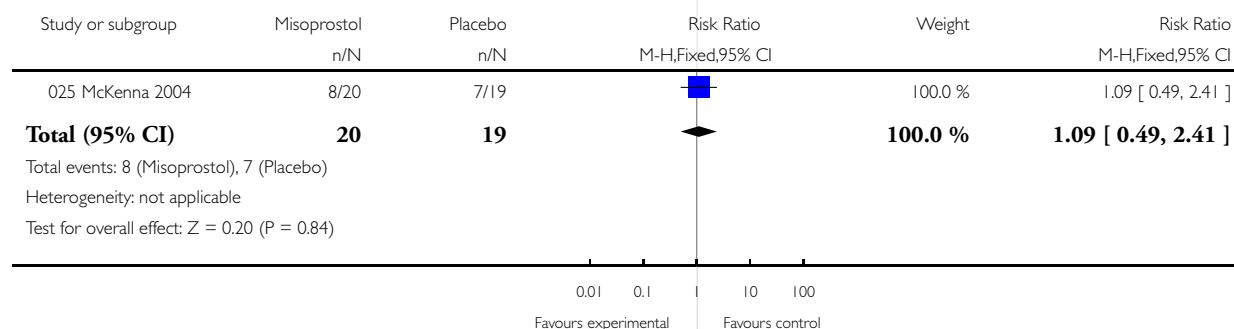


### Analysis 7.1. Comparison 7 Misoprostol versus placebo/no treatment: all primiparae, intact membranes and unfavourable cervix, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 7 Misoprostol versus placebo/no treatment: all primiparae, intact membranes and unfavourable cervix

Outcome: 1 Caesarean section

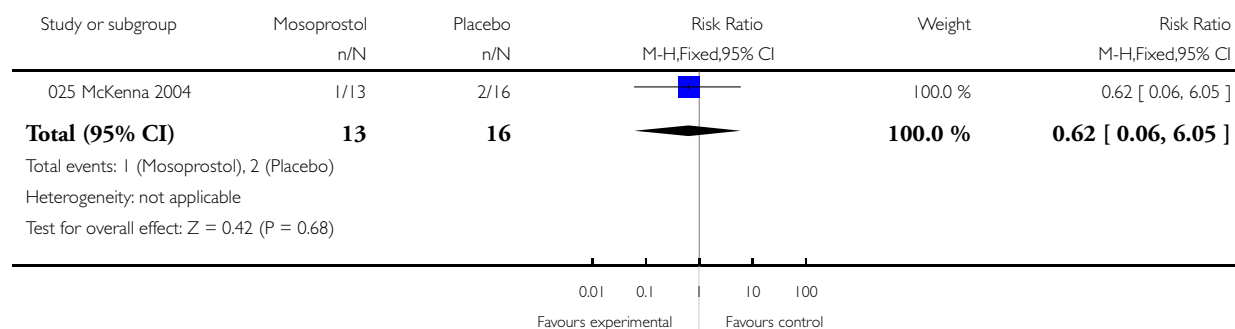


### Analysis 9.1. Comparison 9 Misoprostol versus placebo/no treatment: all multiparous and unfavourable cervix, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 9 Misoprostol versus placebo/no treatment: all multiparous and unfavourable cervix

Outcome: 1 Caesarean section

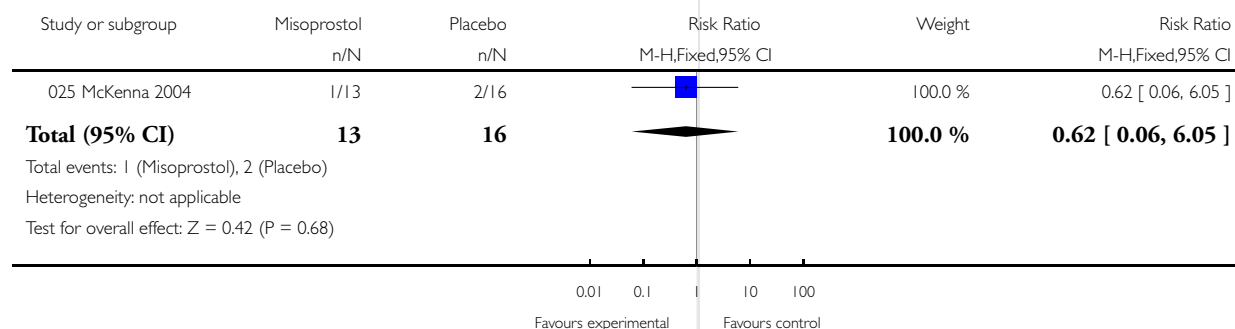


### Analysis 10.1. Comparison 10 Misoprostol versus placebo/no treatment: all multiparous, intact membranes and unfavourable cervix, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 10 Misoprostol versus placebo/no treatment: all multiparous, intact membranes and unfavourable cervix

Outcome: 1 Caesarean section

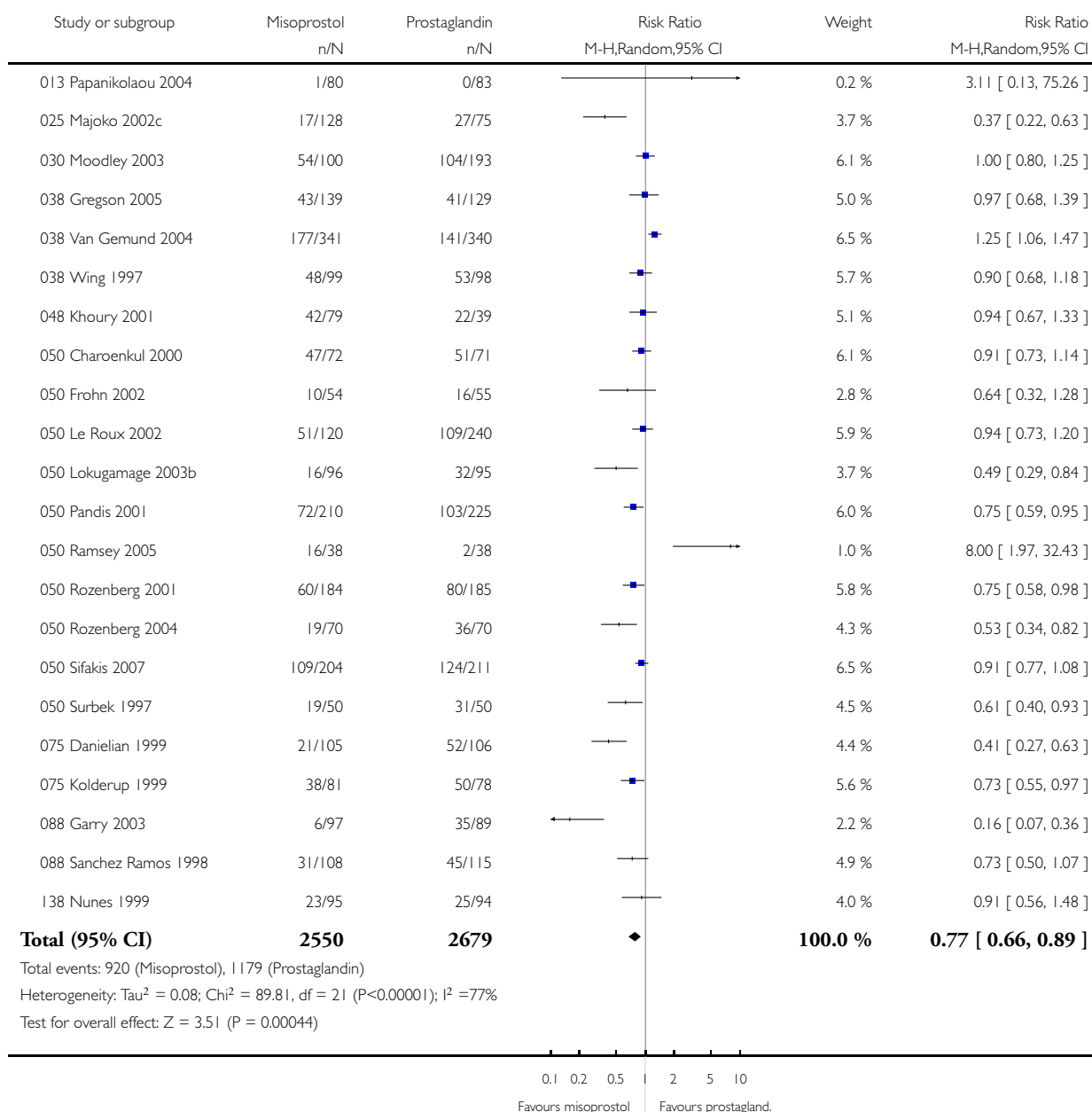


# **Analysis 11.1. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 1 Vaginal delivery not achieved within 24 hours.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 1 Vaginal delivery not achieved within 24 hours



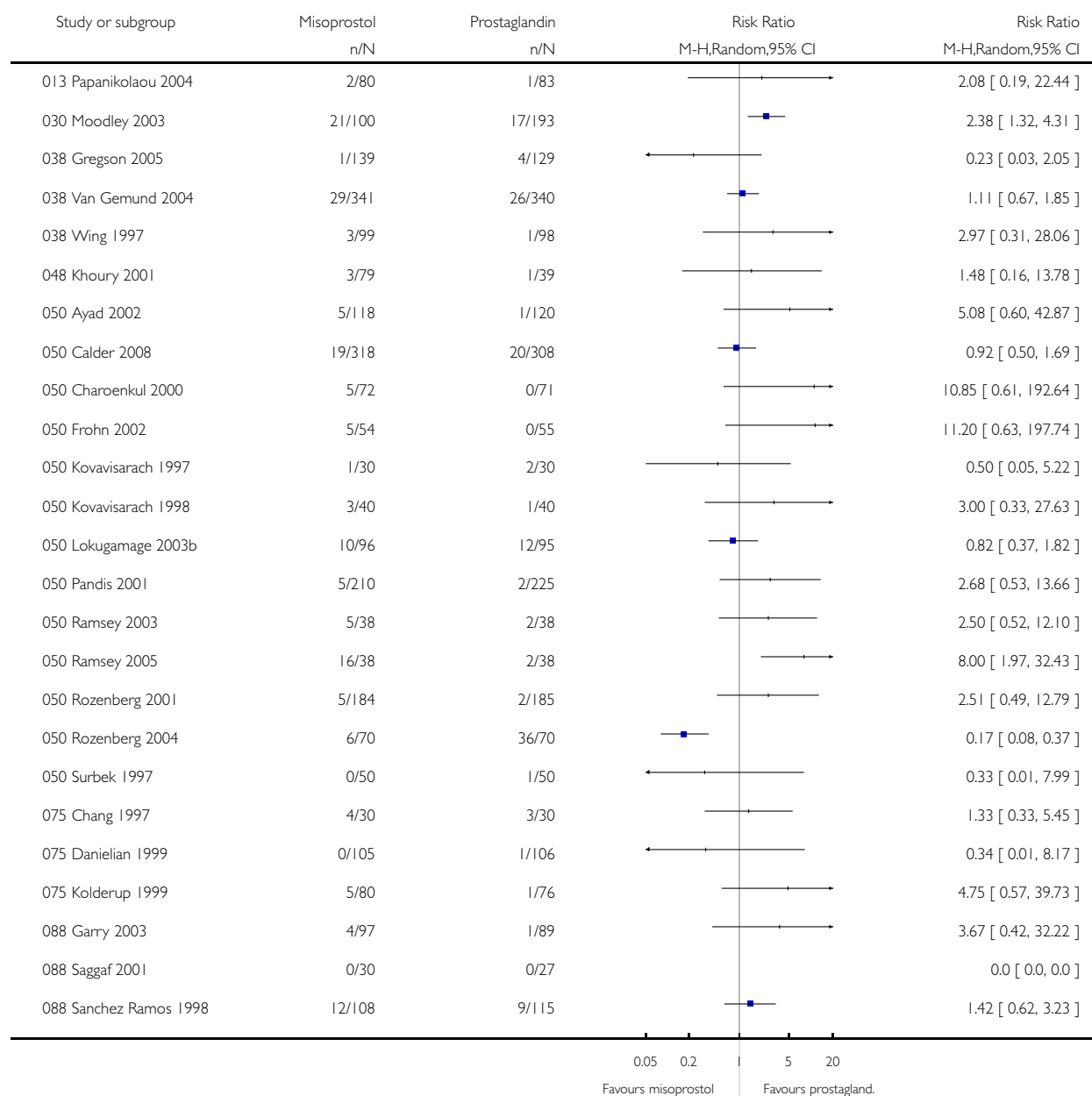


## Analysis 11.2. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 2 Uterine hyperstimulation with FHR changes.

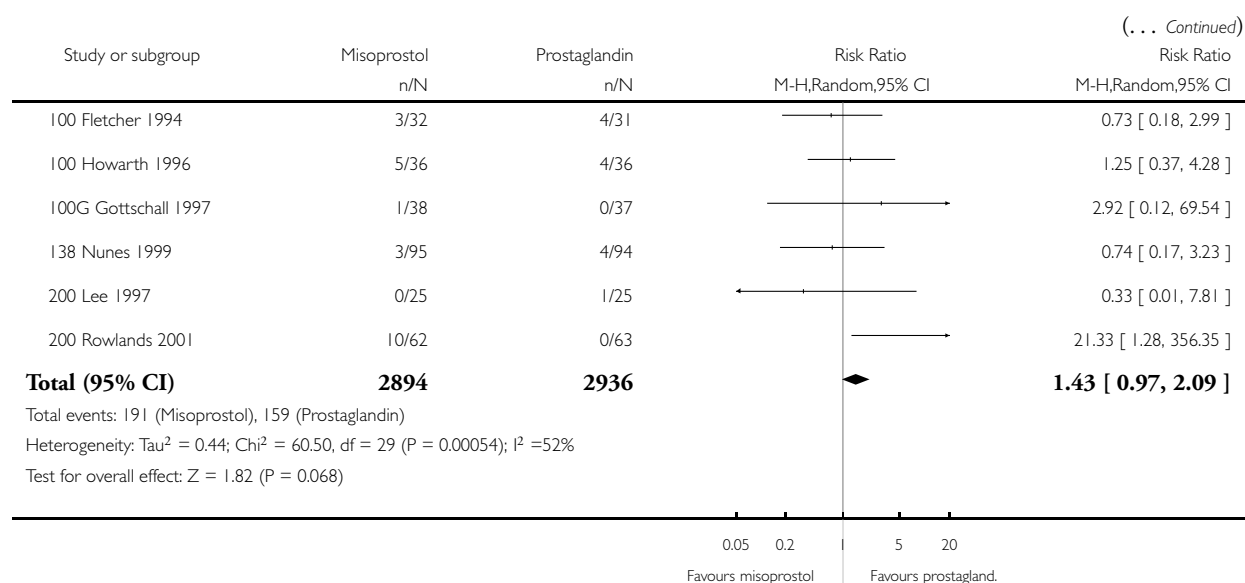
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 2 Uterine hyperstimulation with FHR changes



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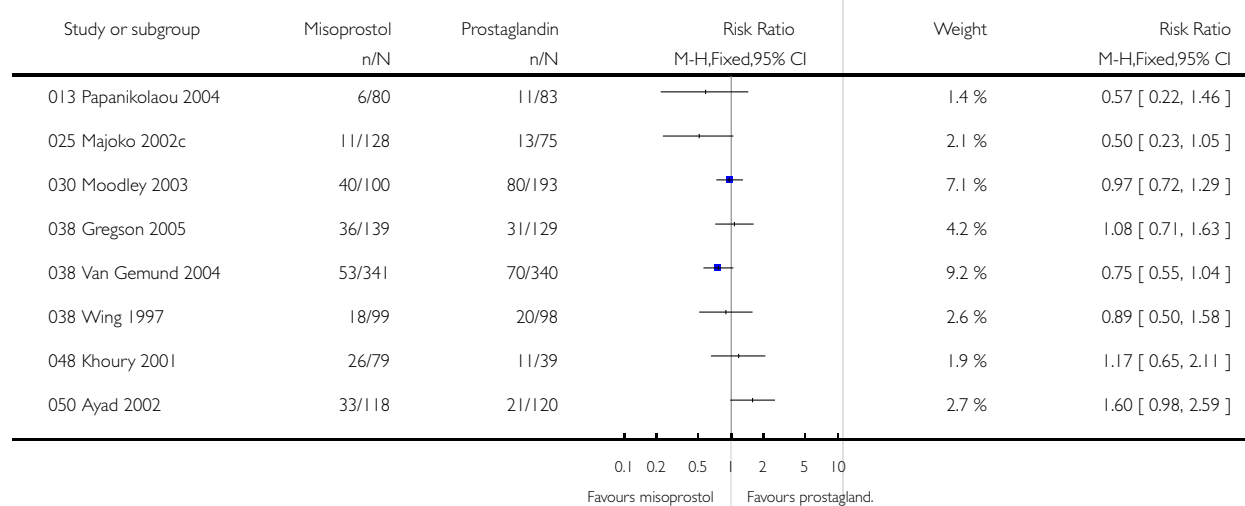


### Analysis 11.3. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 3 Caesarean section.

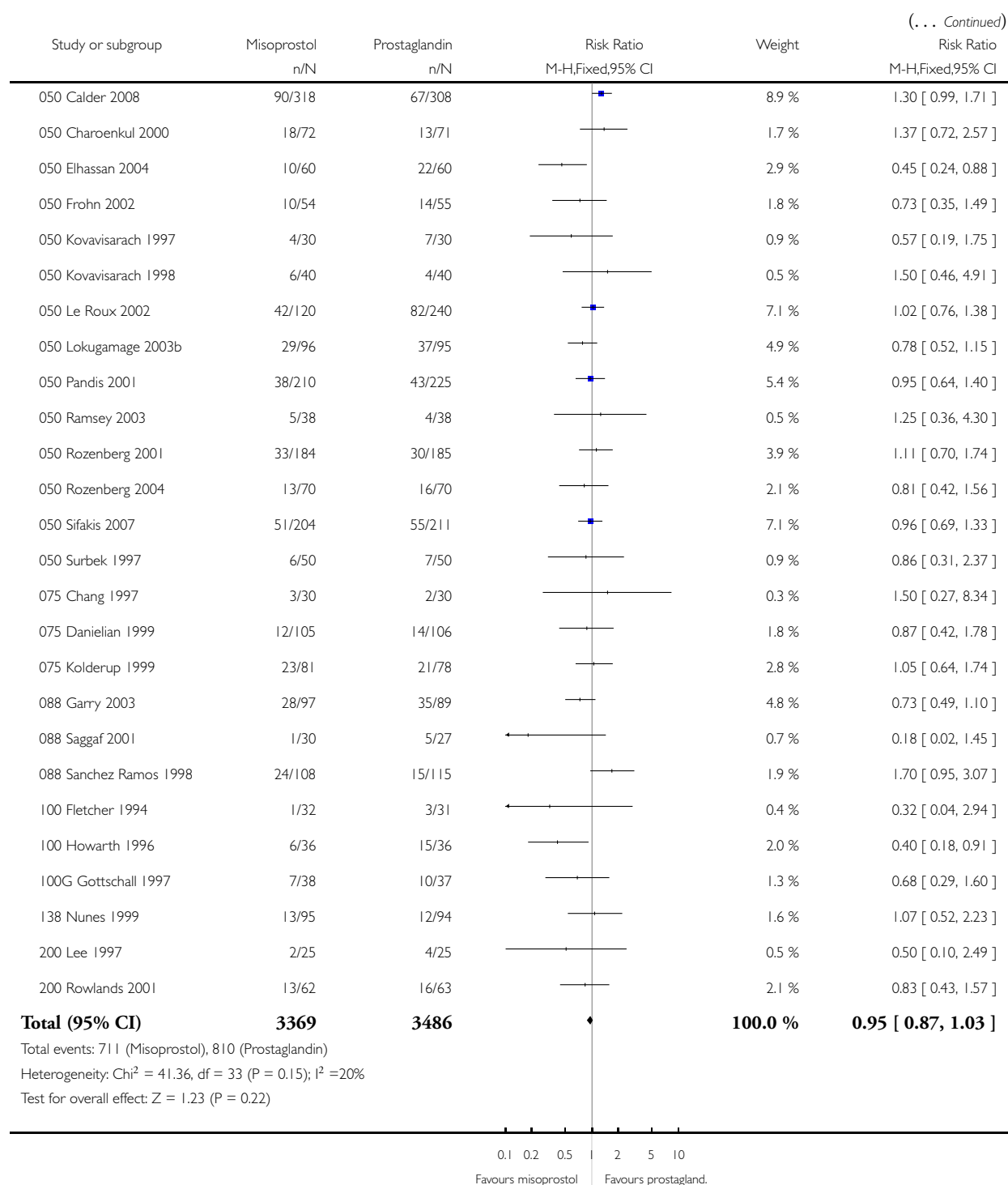
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 3 Caesarean section



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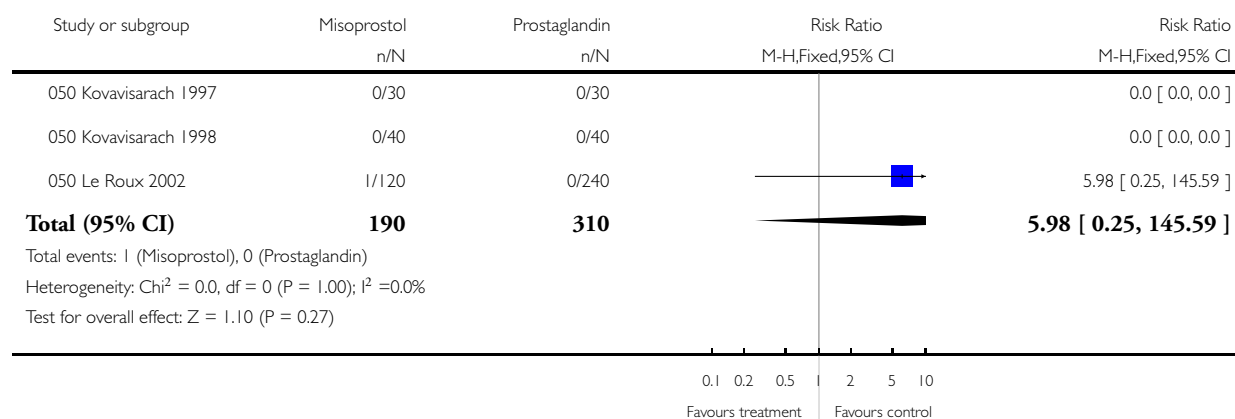


#### Analysis 11.4. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 4 Serious neonatal morbidity or perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 4 Serious neonatal morbidity or perinatal death

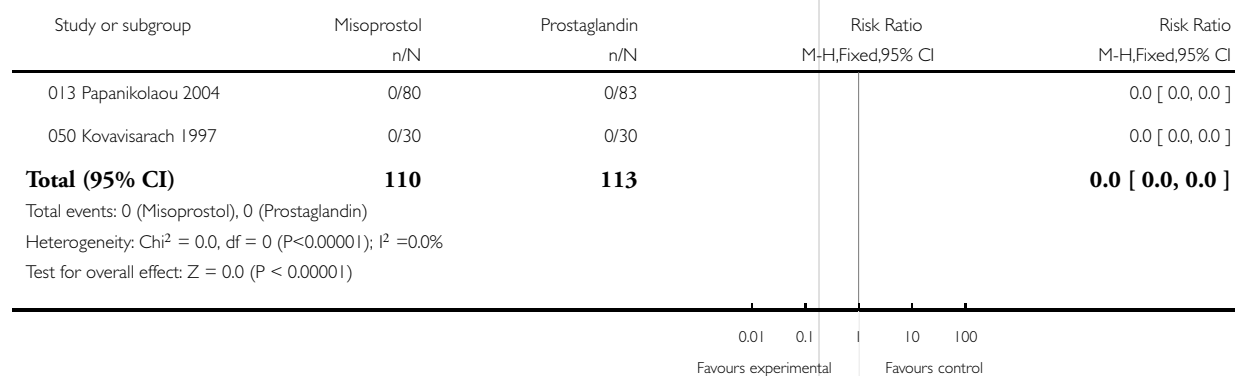


#### Analysis 11.5. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 5 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 5 Serious maternal morbidity or death

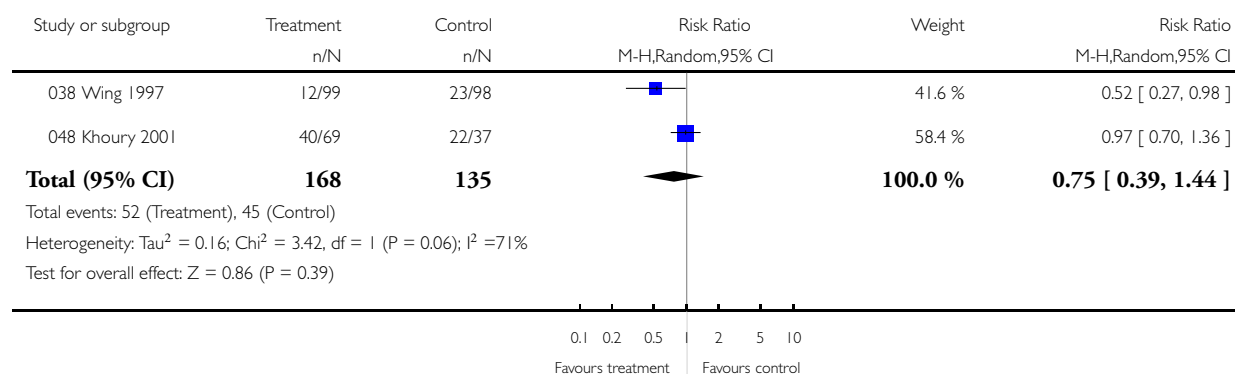


### Analysis 11.6. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 6 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 6 Cervix unfavourable/unchanged after 12-24 hours

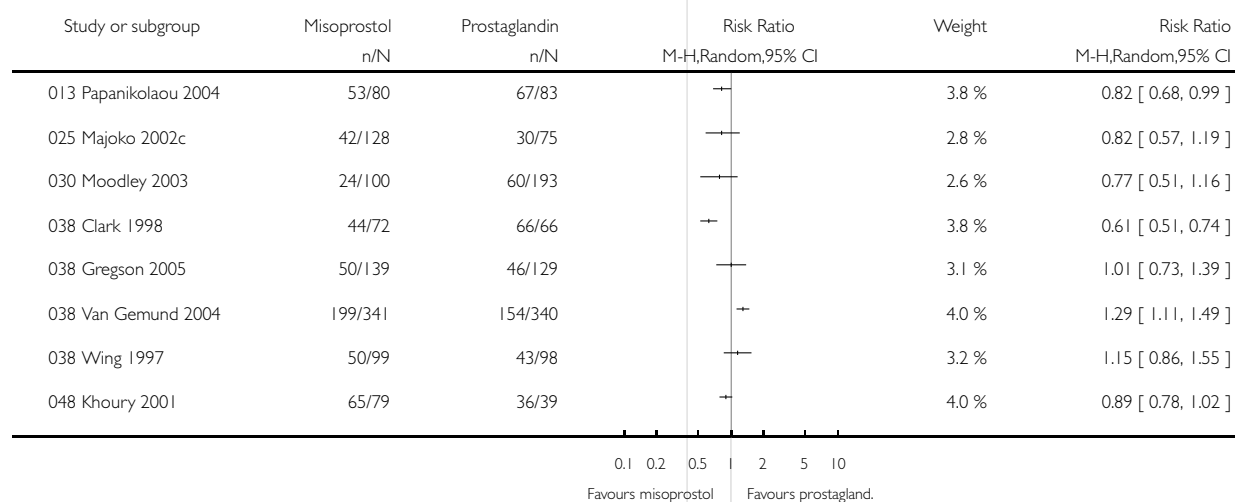


### Analysis 11.7. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 7 Oxytocin augmentation.

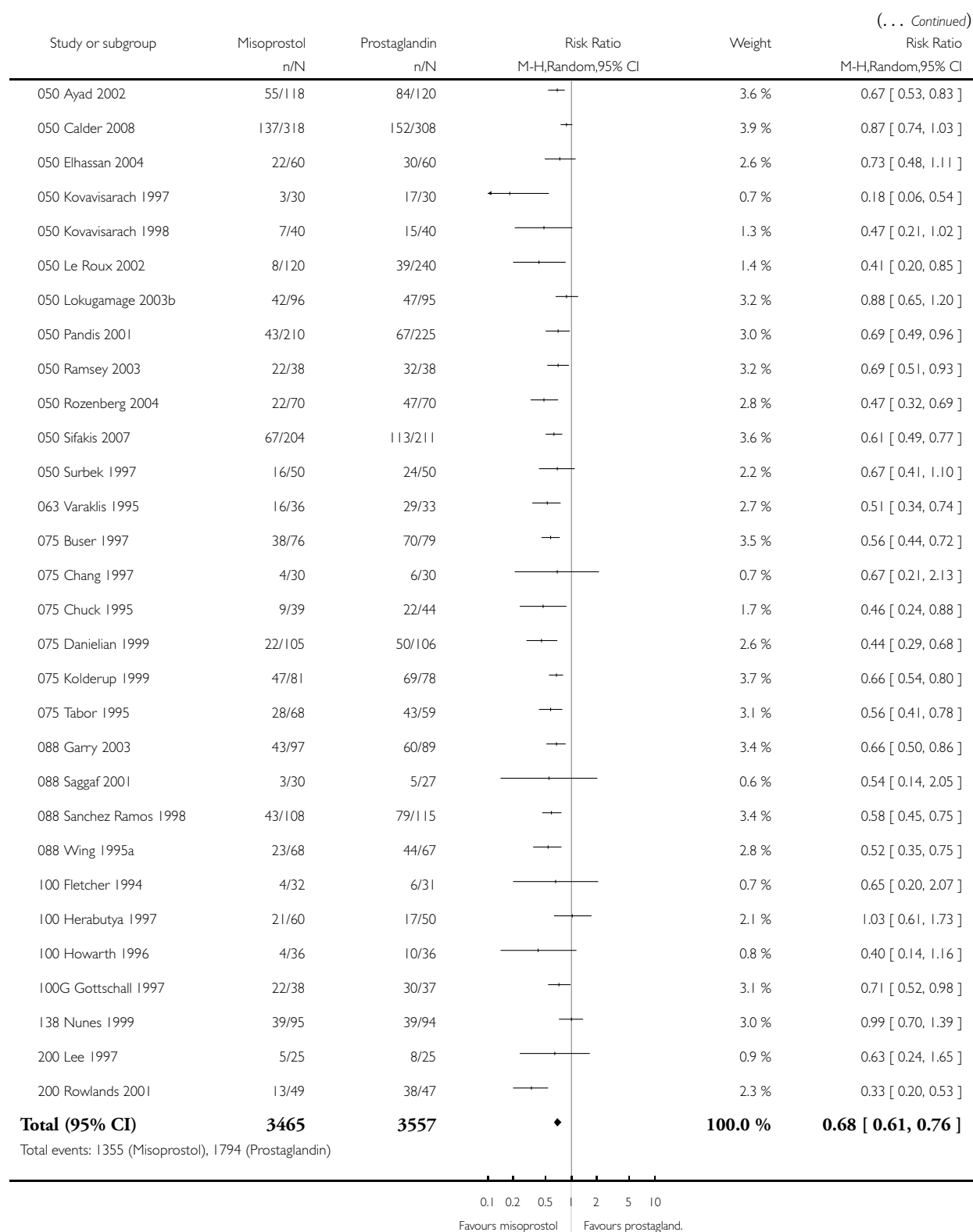
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 7 Oxytocin augmentation



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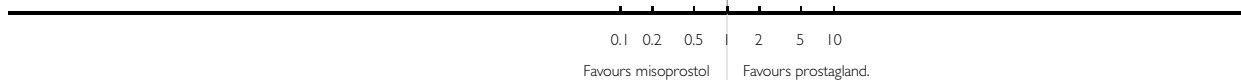


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Study or subgroup	Misoprostol n/N	Prostaglandin n/N	Risk Ratio M-H,Random,95% CI	Weight	Risk Ratio M-H,Random,95% CI
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Heterogeneity:  $\tau^2 = 0.07$ ;  $\chi^2 = 148.59$ ,  $df = 37$  ( $P < 0.00001$ );  $I^2 = 75\%$   
 Test for overall effect:  $Z = 7.01$  ( $P < 0.00001$ )

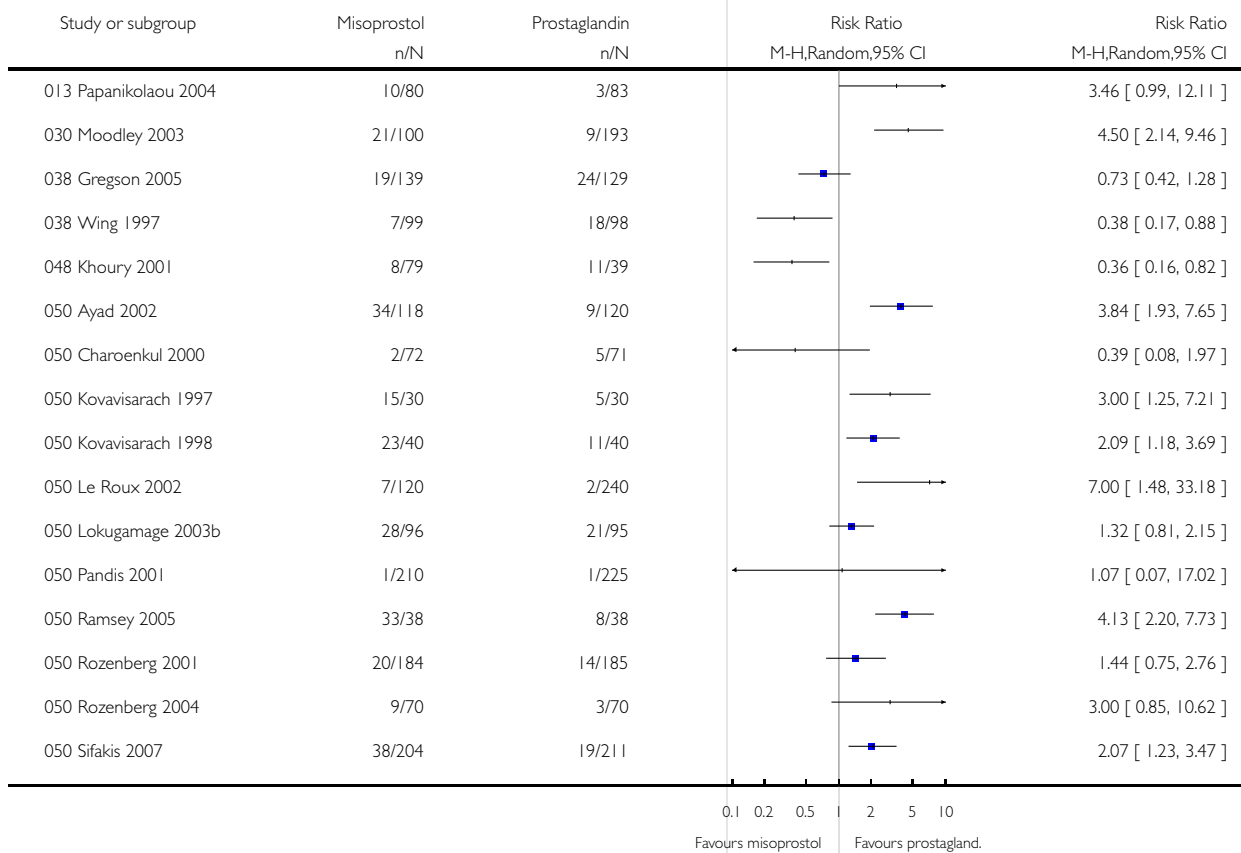


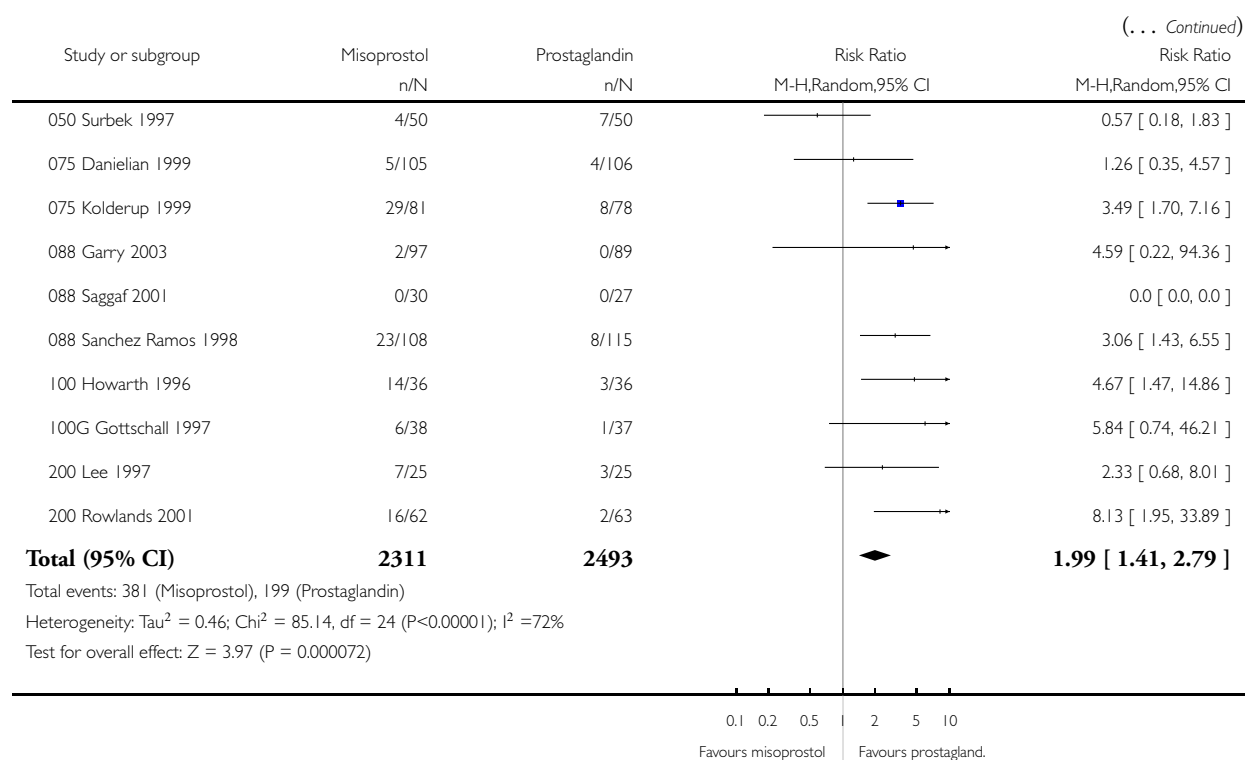
### Analysis 11.8. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 8 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 8 Uterine hyperstimulation without FHR changes





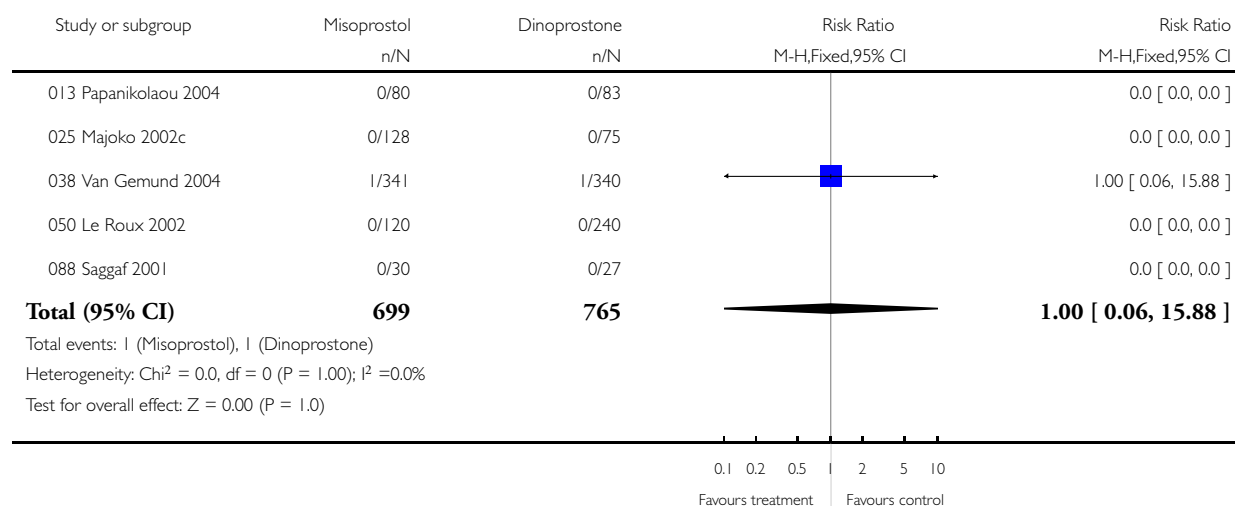


### Analysis 11.9. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 9 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 9 Uterine rupture

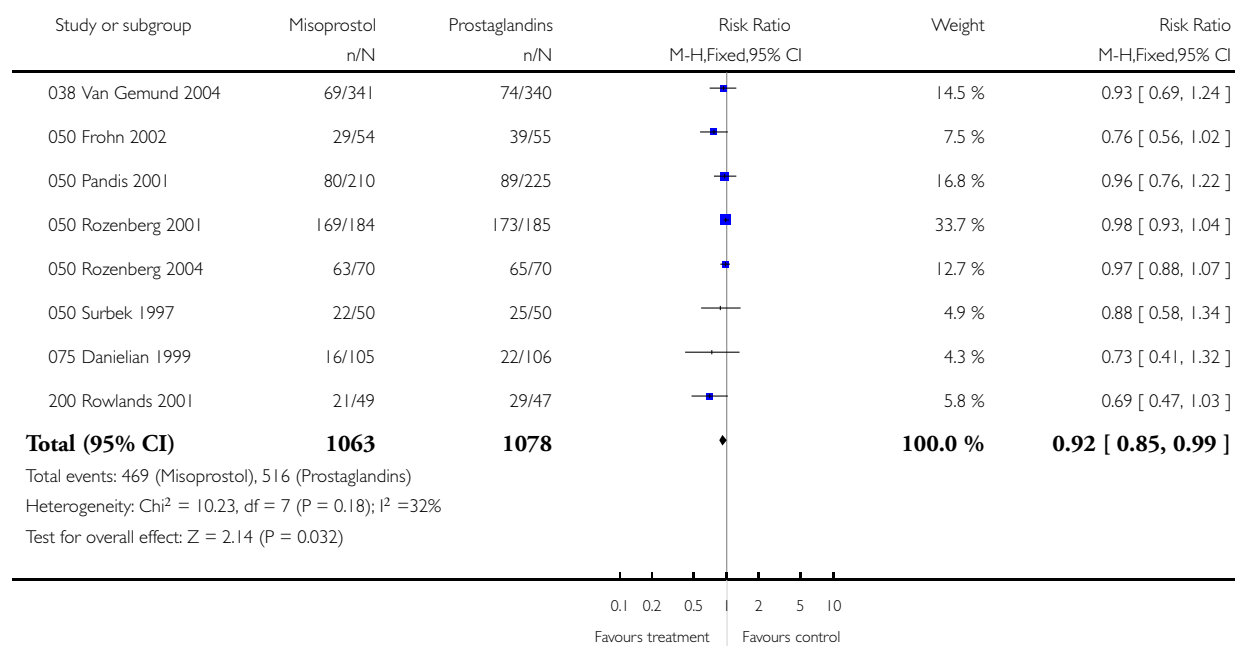


## Analysis 11.10. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 10 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 10 Epidural analgesia

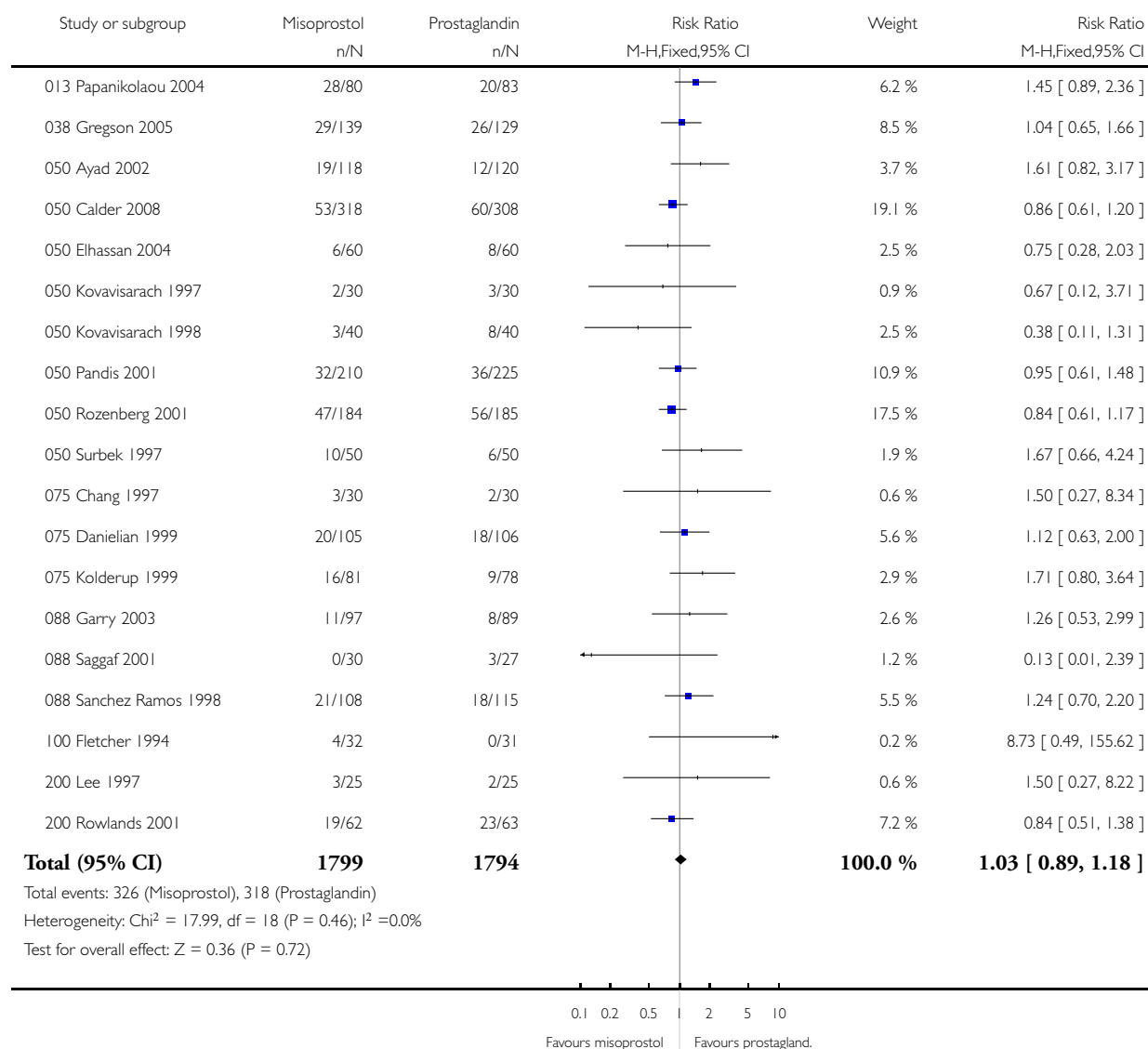


### Analysis 11.11. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 11 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 11 Instrumental vaginal delivery

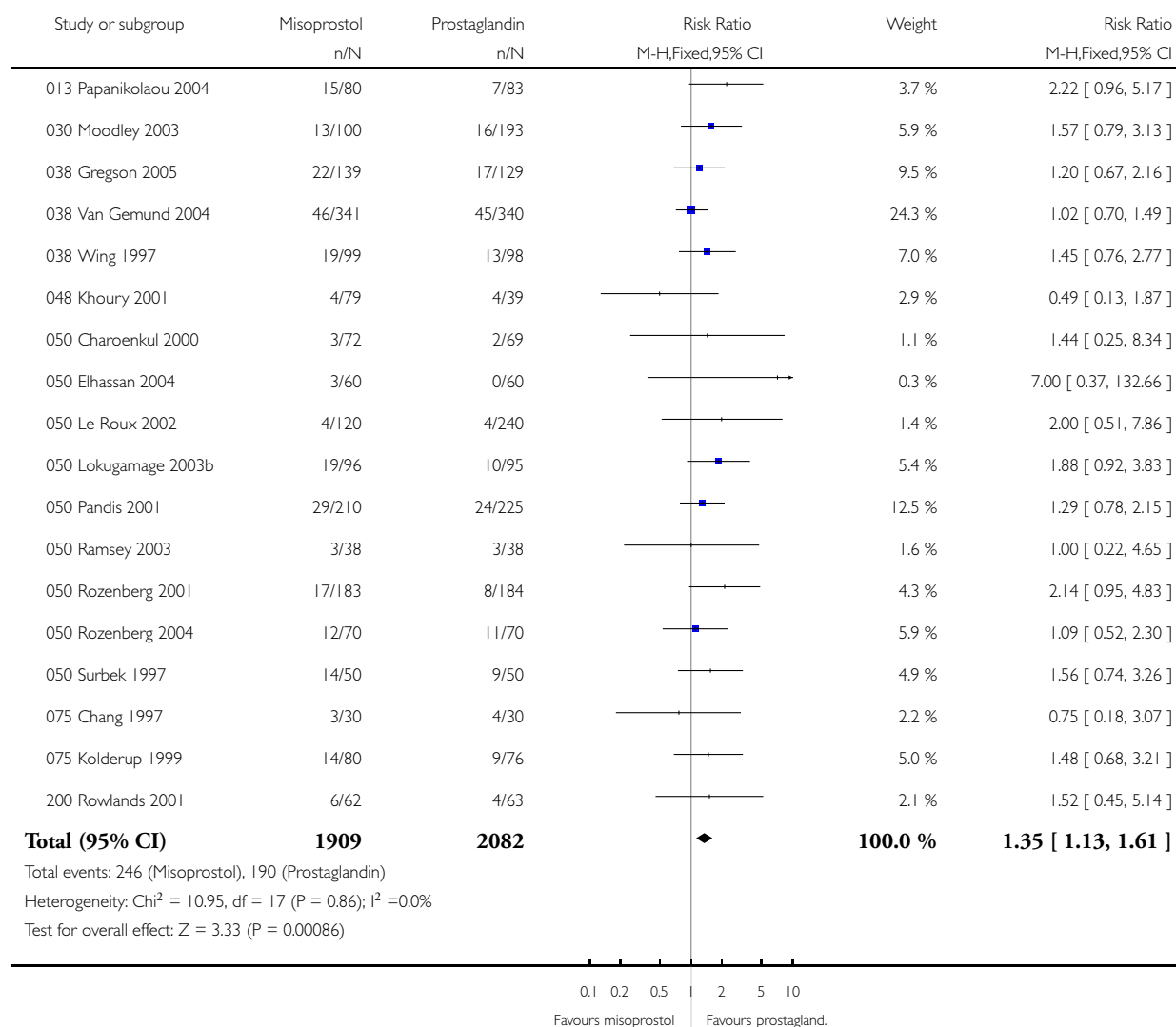


## Analysis 11.12. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 12 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 12 Meconium-stained liquor

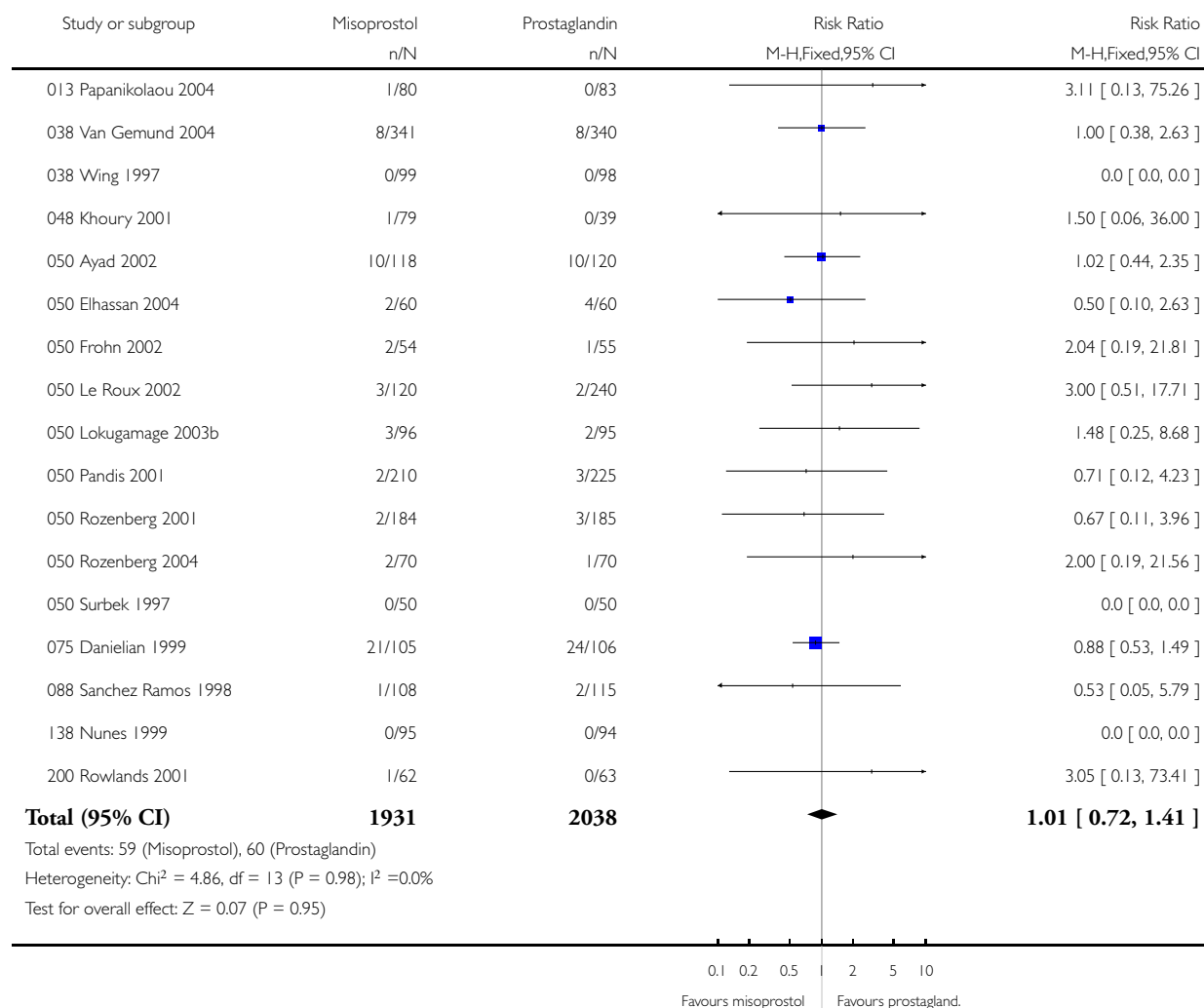


### Analysis 11.13. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 13 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 13 Apgar score < 7 at 5 minutes

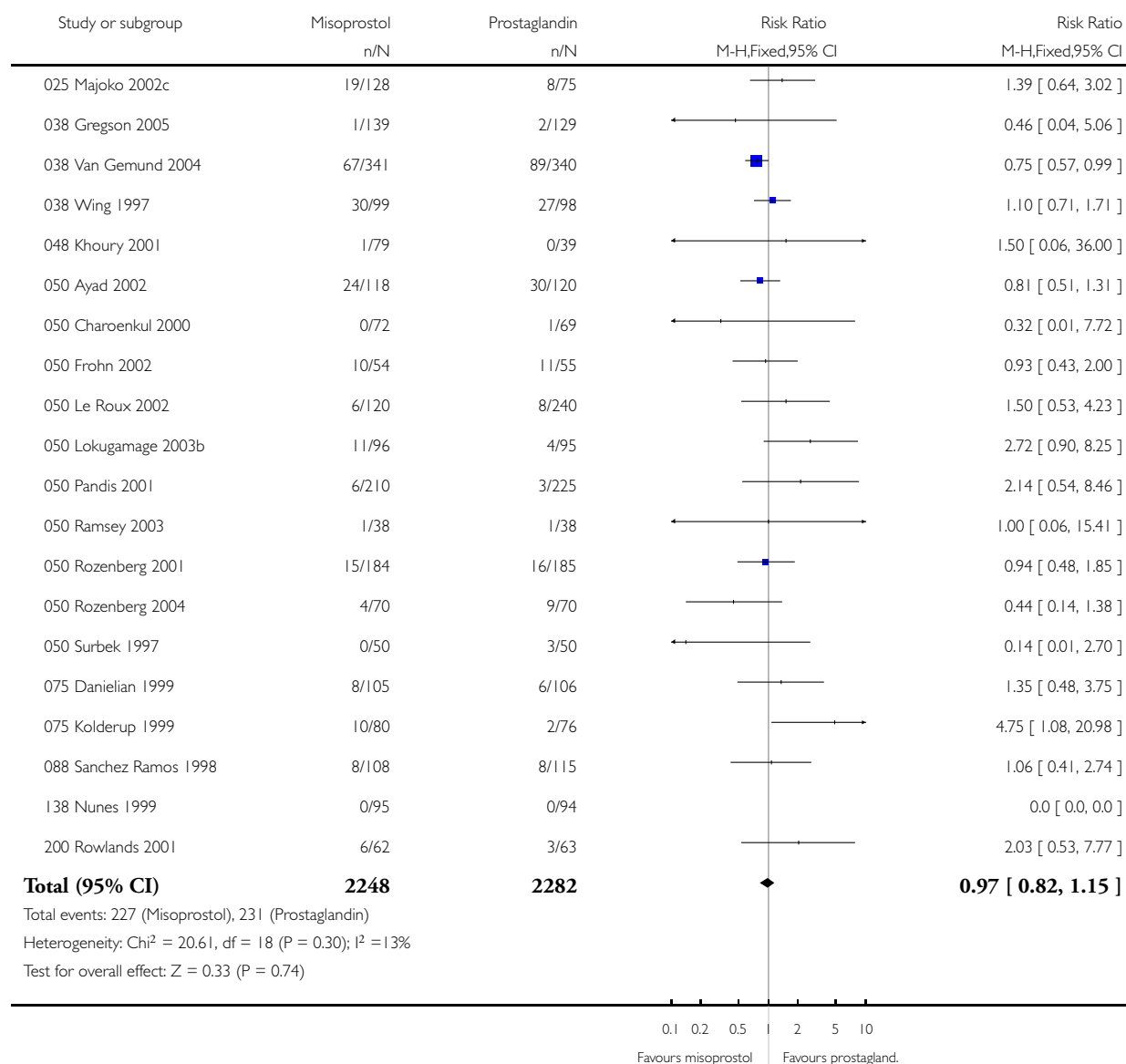


# **Analysis 11.14. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 14 Neonatal intensive care unit admission.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 14 Neonatal intensive care unit admission

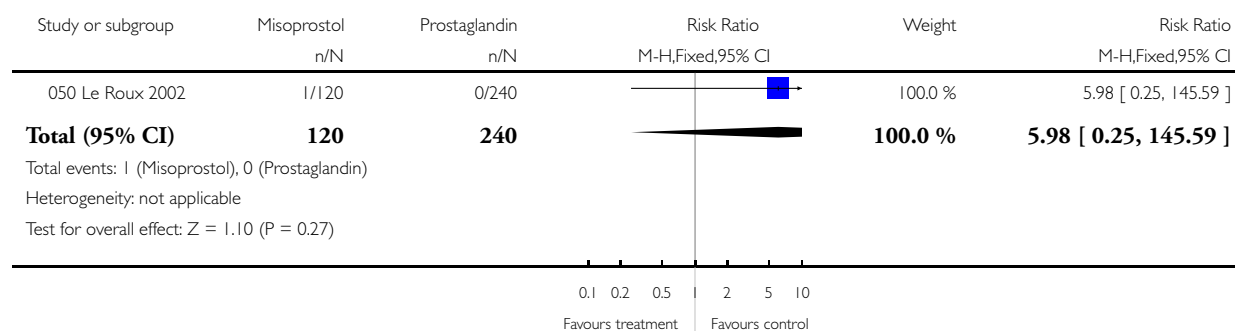


### Analysis 11.15. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 15 Neonatal encephalopathy.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 15 Neonatal encephalopathy

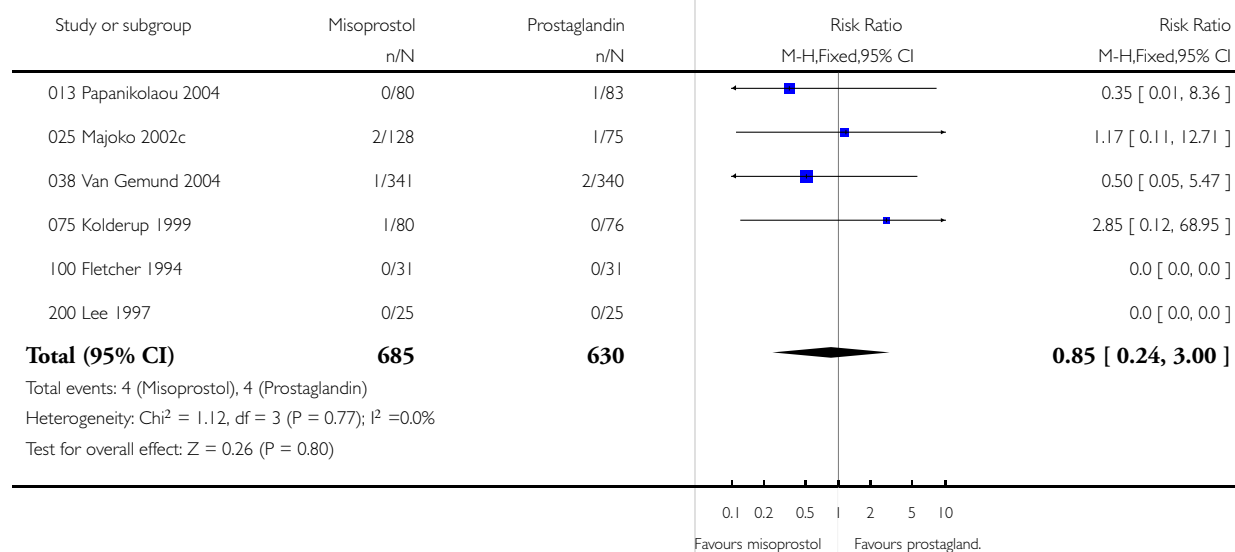


### Analysis 11.16. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 16 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 16 Perinatal death

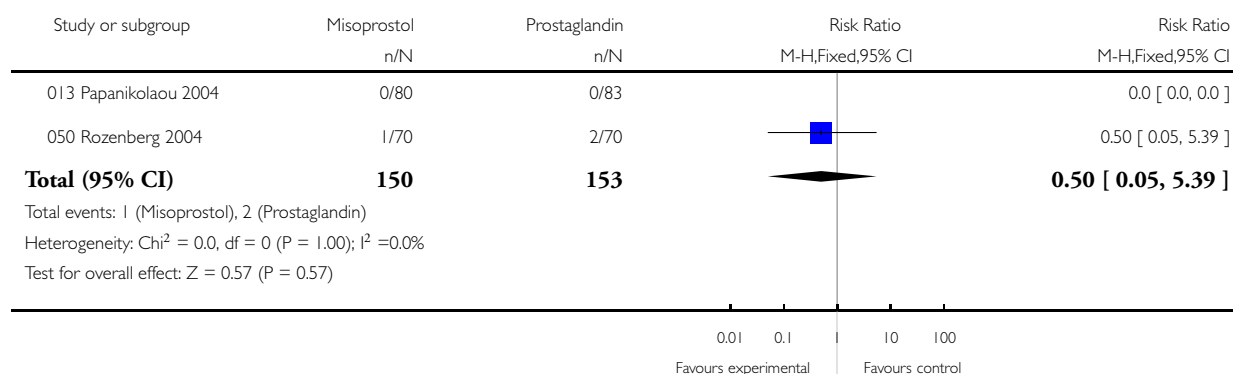


### Analysis 11.17. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 17 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 17 Serious maternal complications

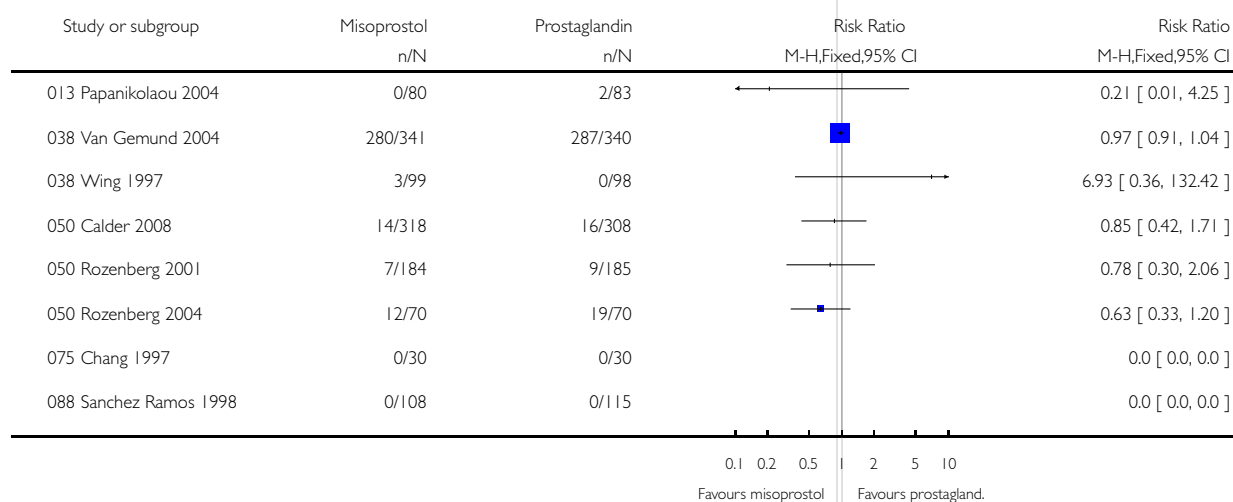


### Analysis 11.18. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 18 Maternal side effects (eg nausea, vomiting, diarrhoea).

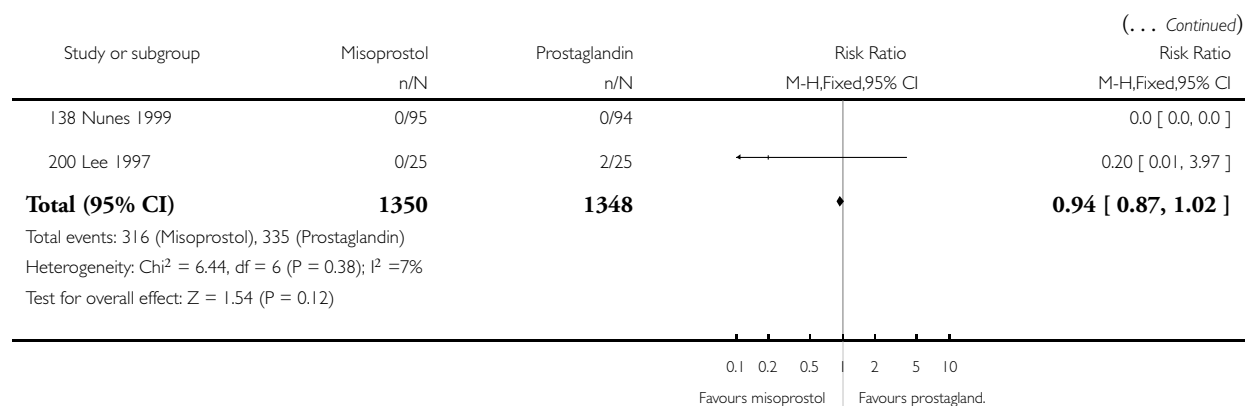
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 18 Maternal side effects (eg nausea, vomiting, diarrhoea)





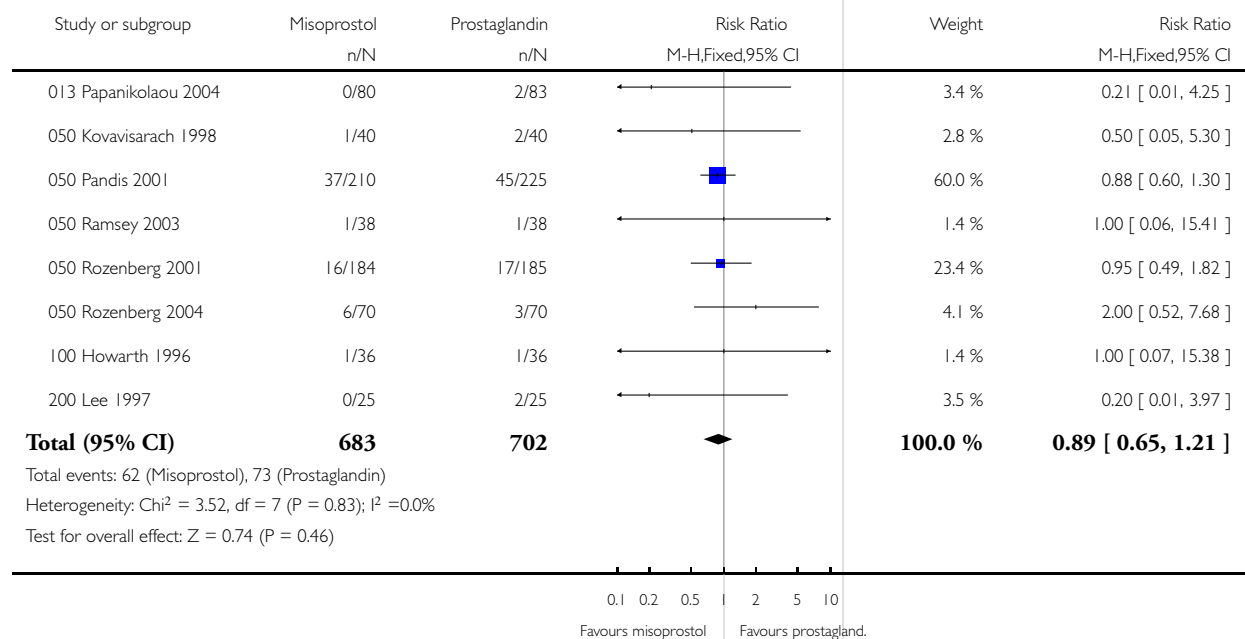


### Analysis 11.19. Comparison 11 Misoprostol versus vaginal prostaglandin: all women, Outcome 19 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 11 Misoprostol versus vaginal prostaglandin: all women

Outcome: 19 Postpartum haemorrhage

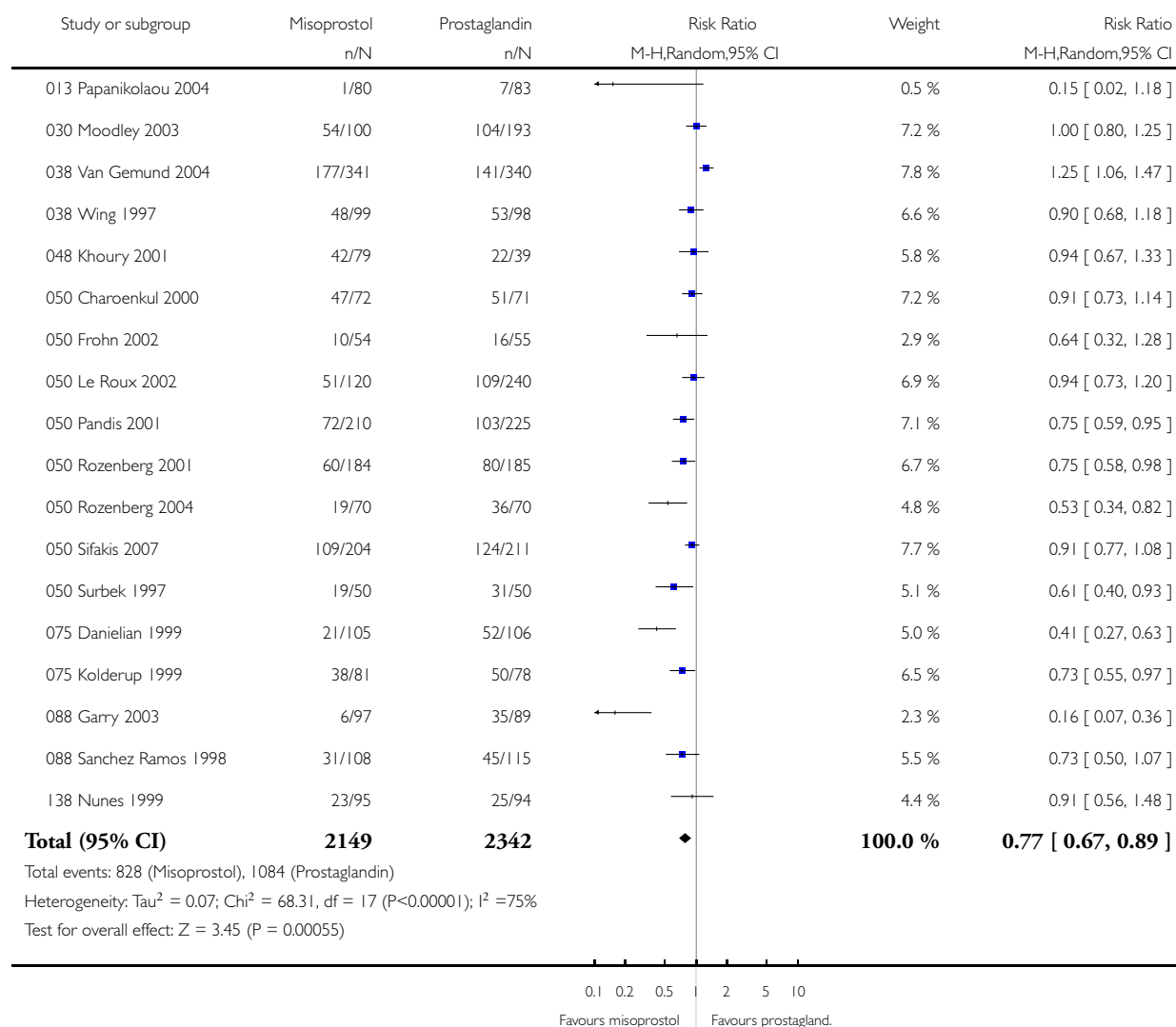


## Analysis 12.1. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

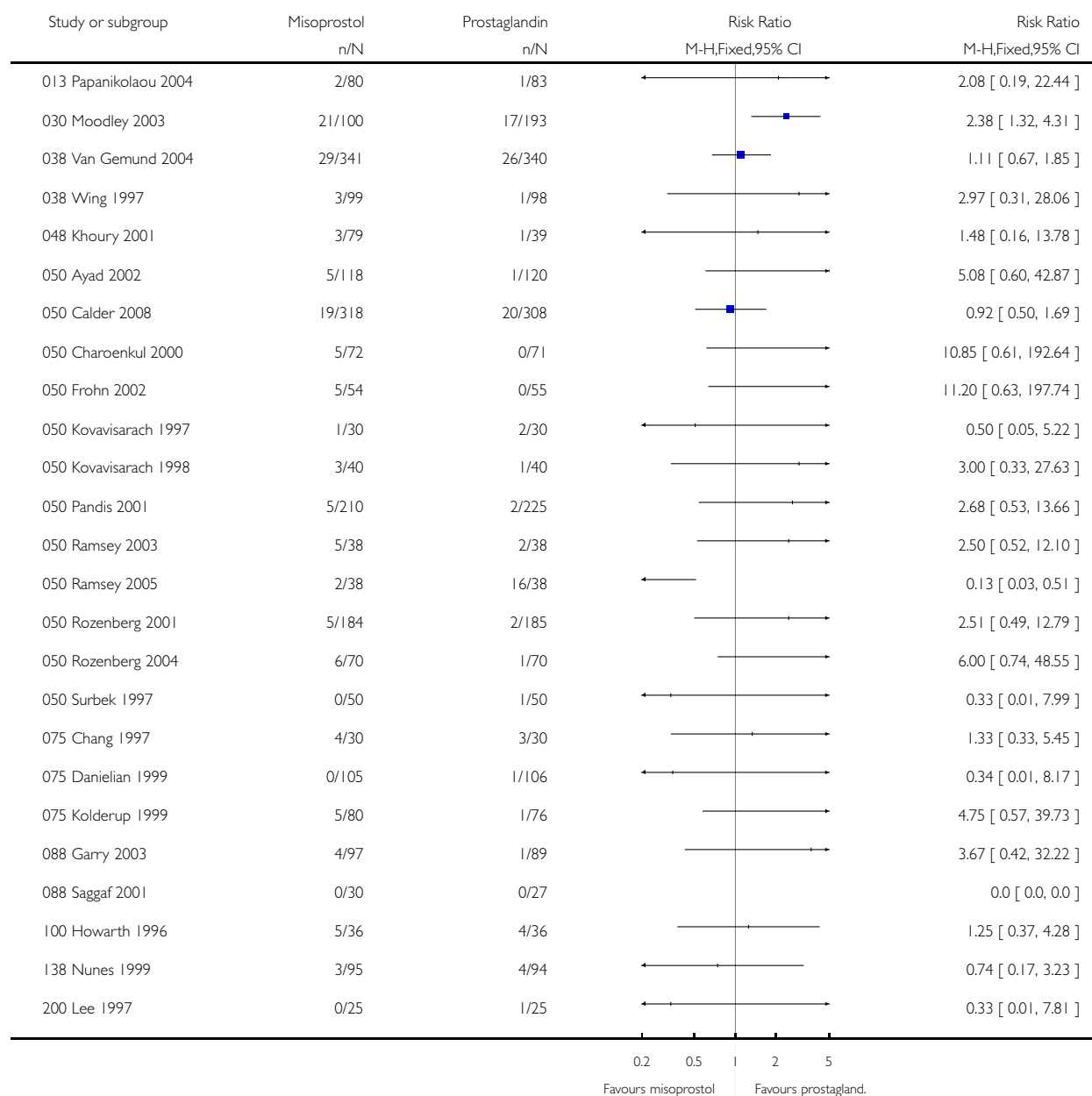


## Analysis 12.2. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

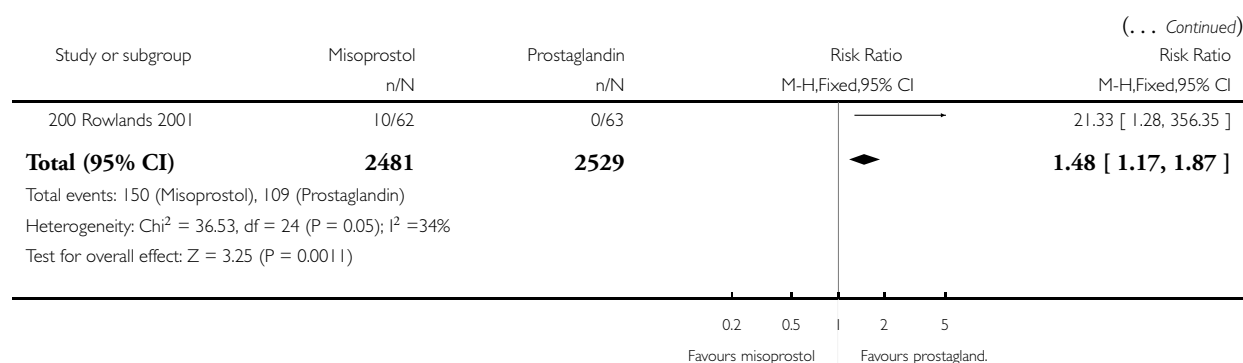
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes



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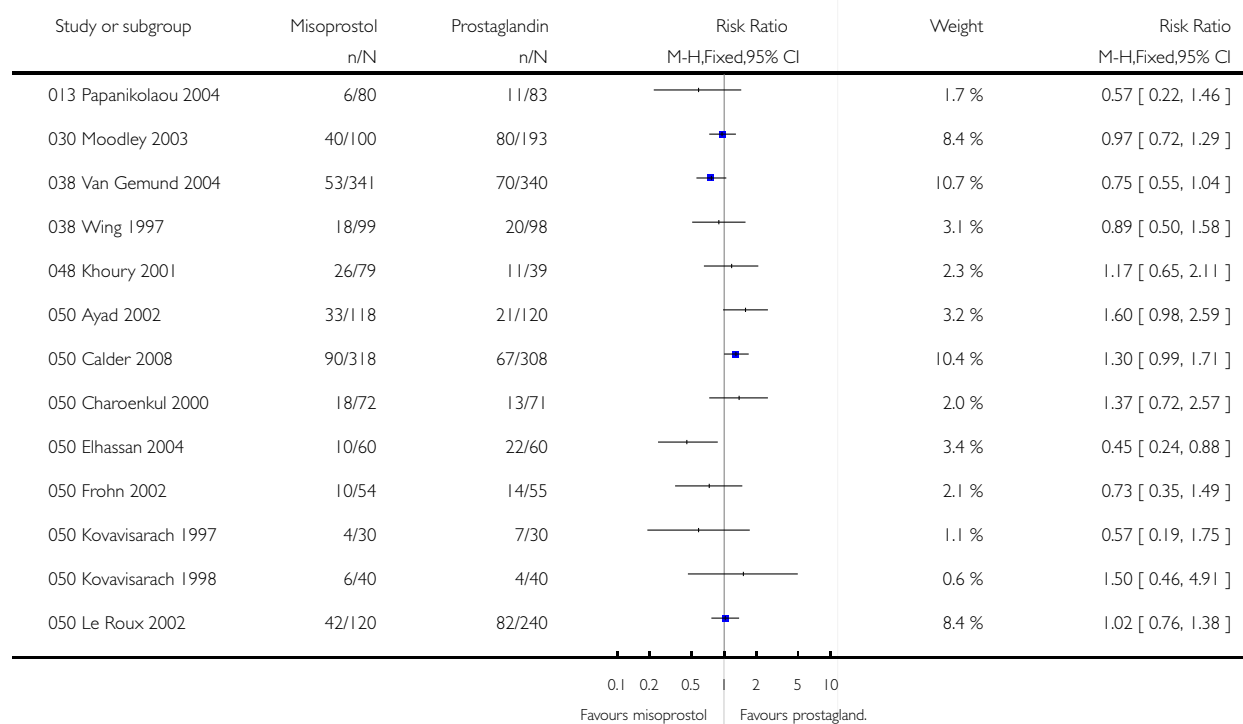


### Analysis 12.3. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 3 Caesarean section.

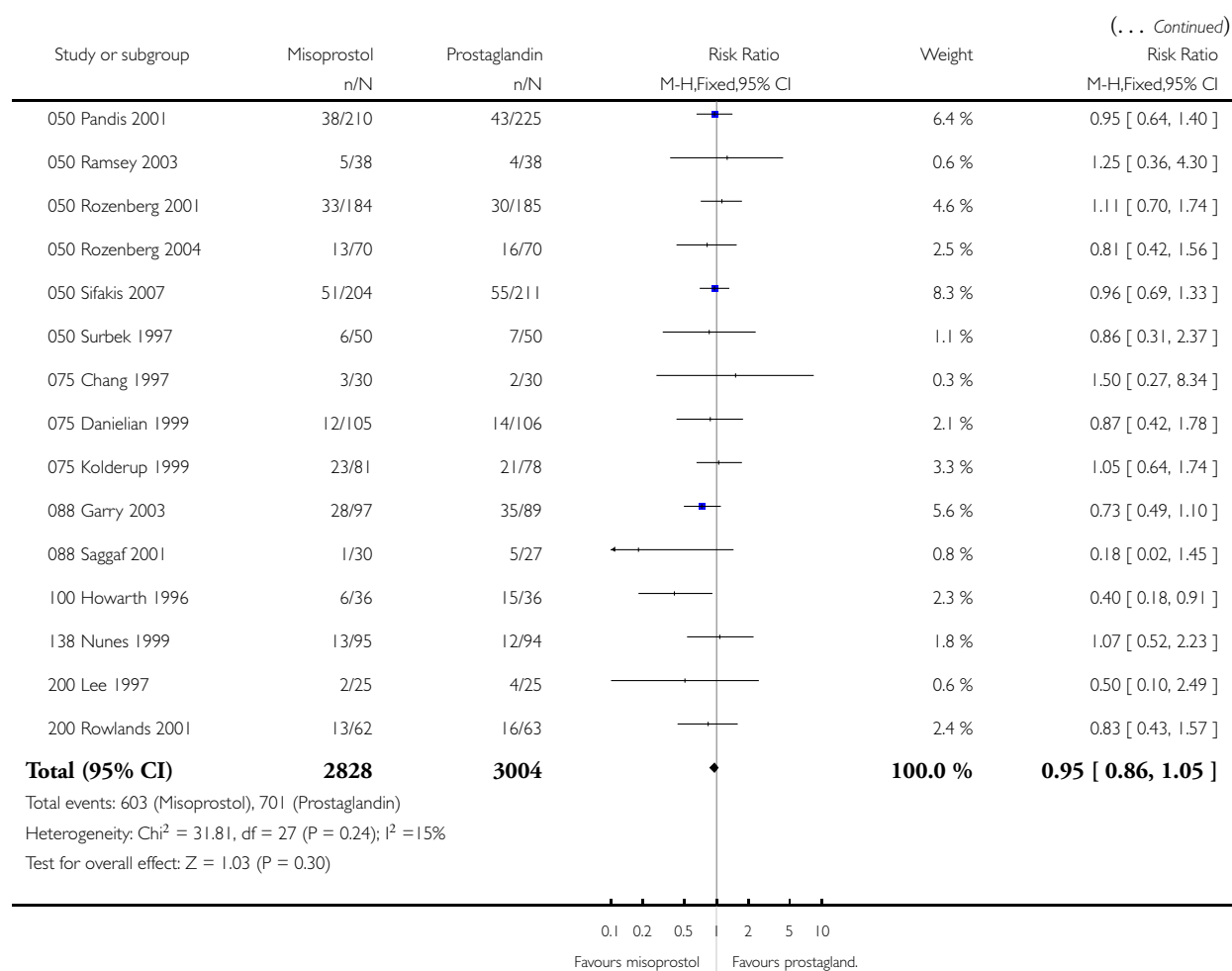
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 3 Caesarean section



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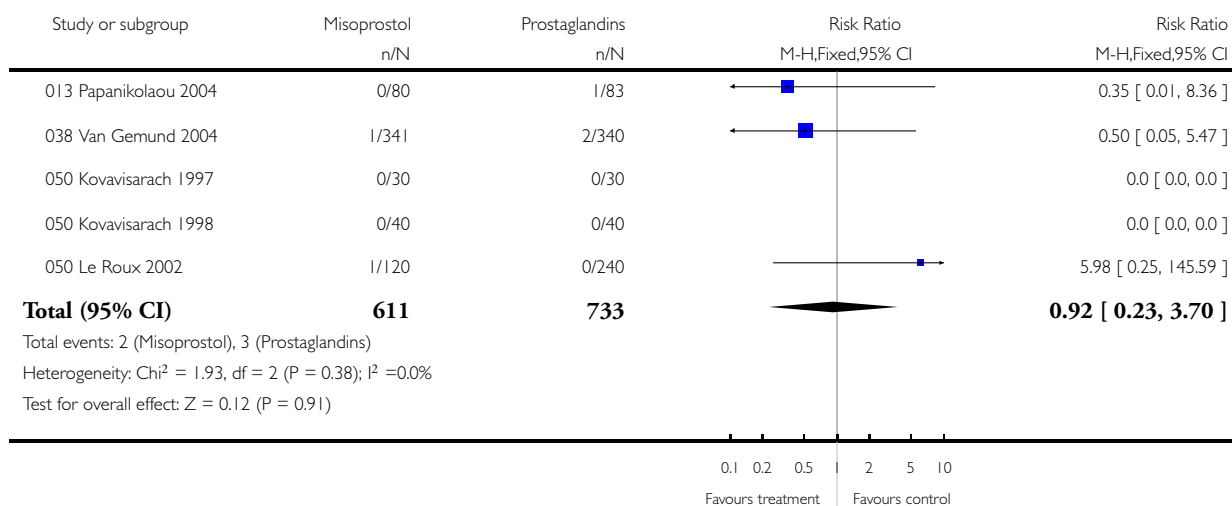


#### Analysis 12.4. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 4 Serious neonatal morbidity or perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 4 Serious neonatal morbidity or perinatal death

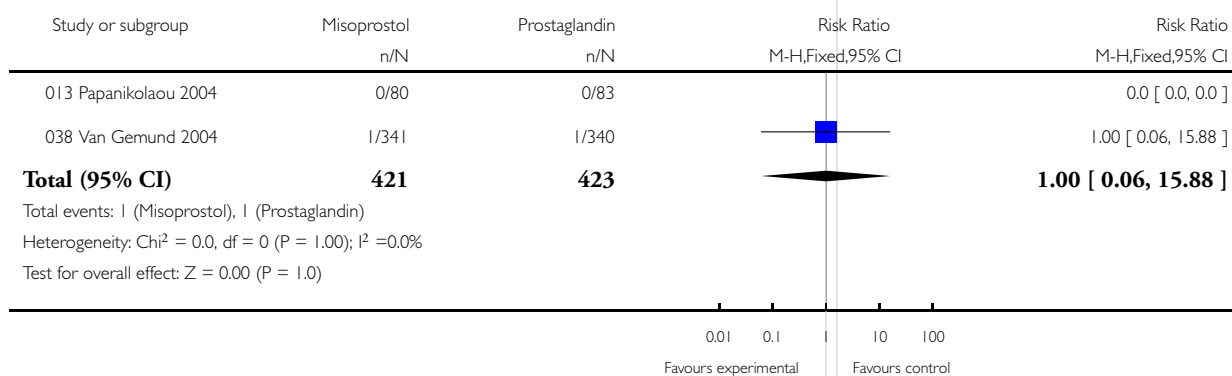


#### Analysis 12.5. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 5 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 5 Uterine rupture

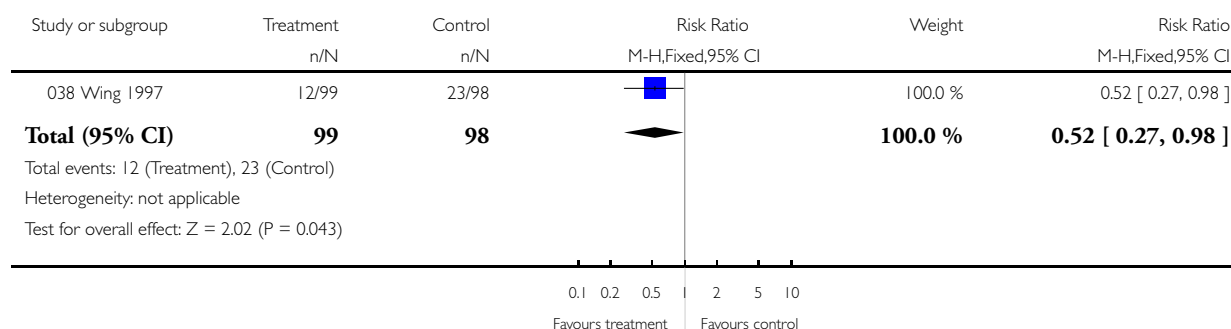


## Analysis 12.6. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 6 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 6 Cervix unfavourable/unchanged after 12-24 hours

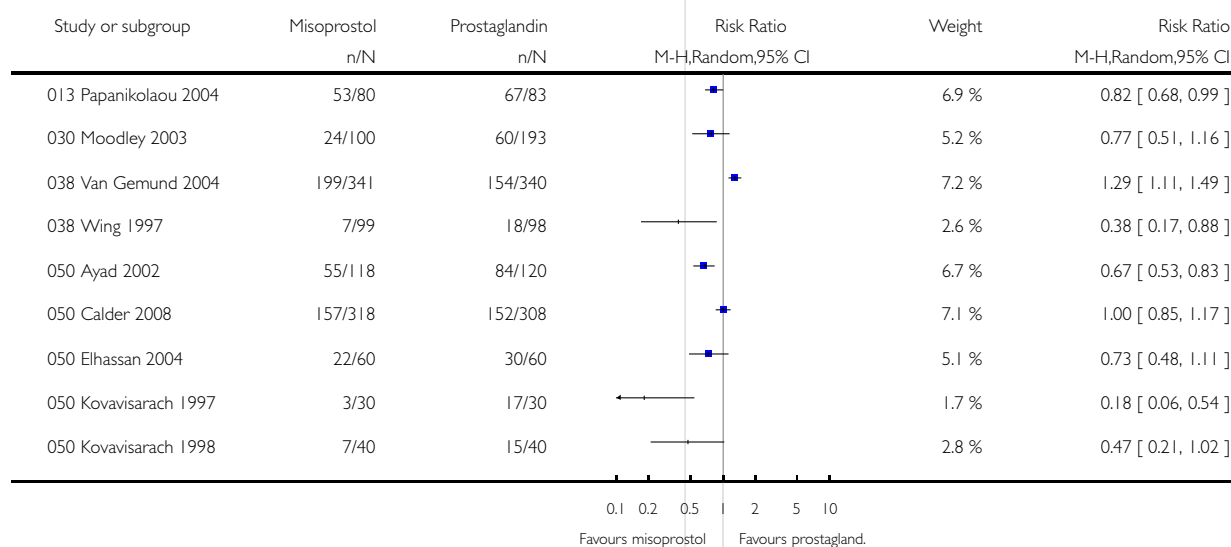


## Analysis 12.7. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 7 Oxytocin augmentation.

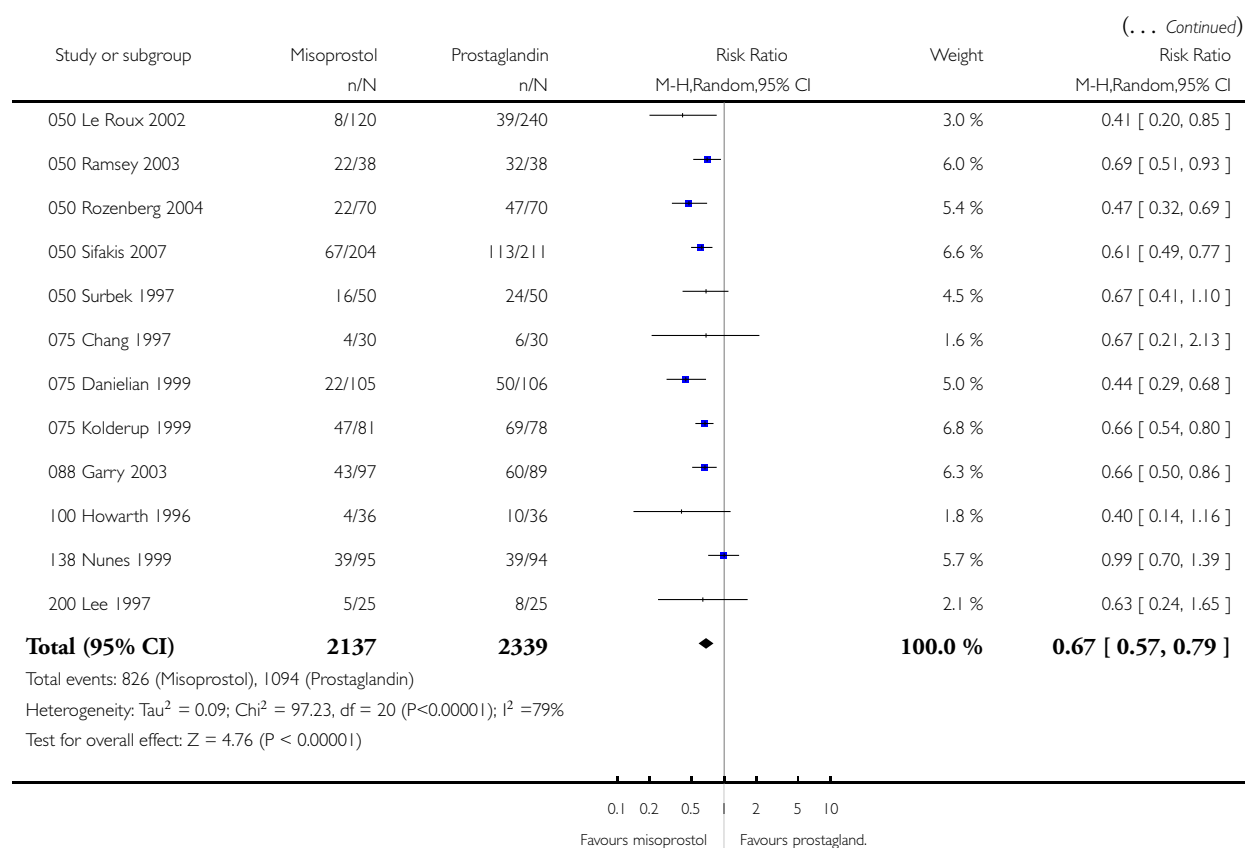
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 7 Oxytocin augmentation



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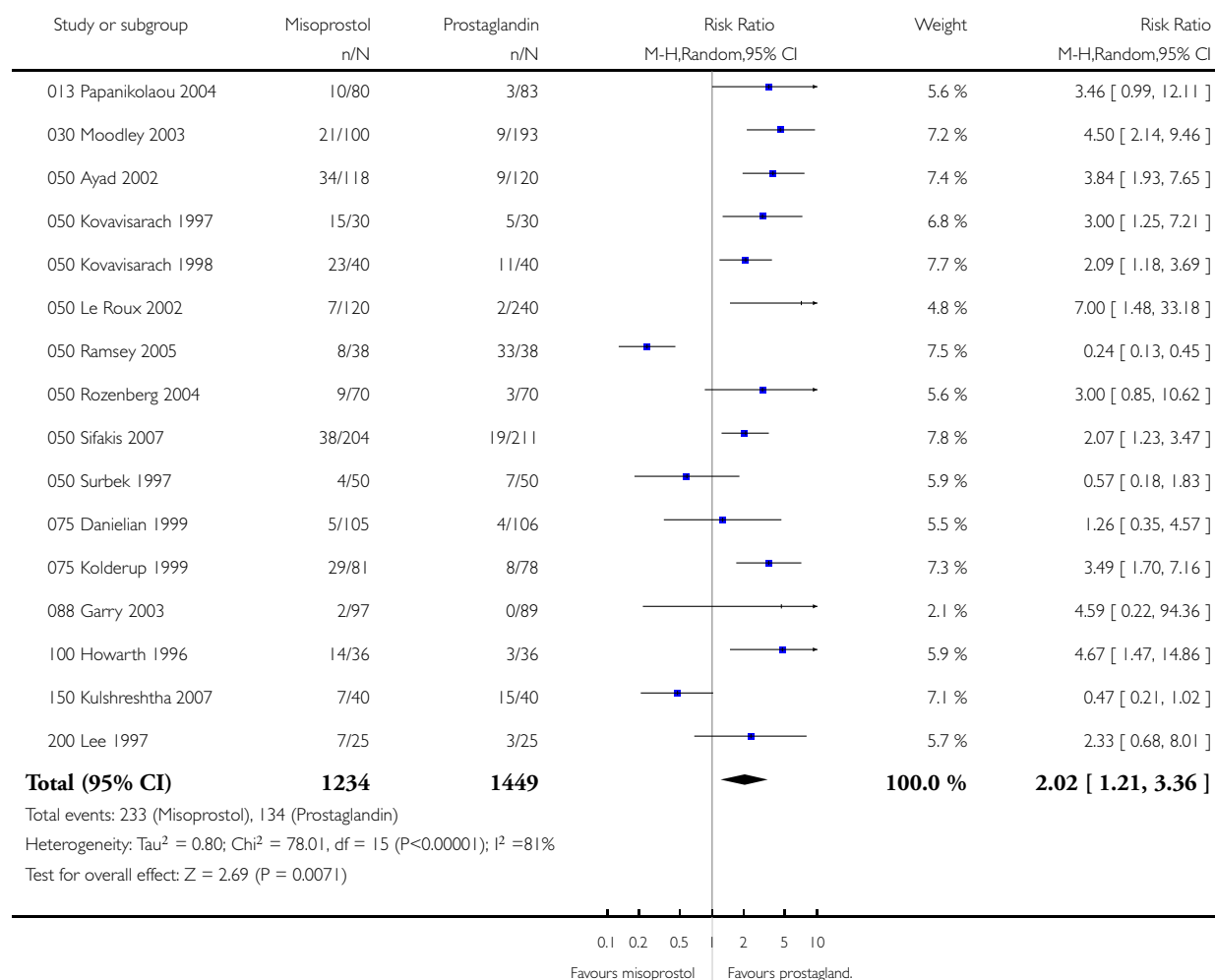


## Analysis 12.8. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 8 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 8 Uterine hyperstimulation without FHR changes

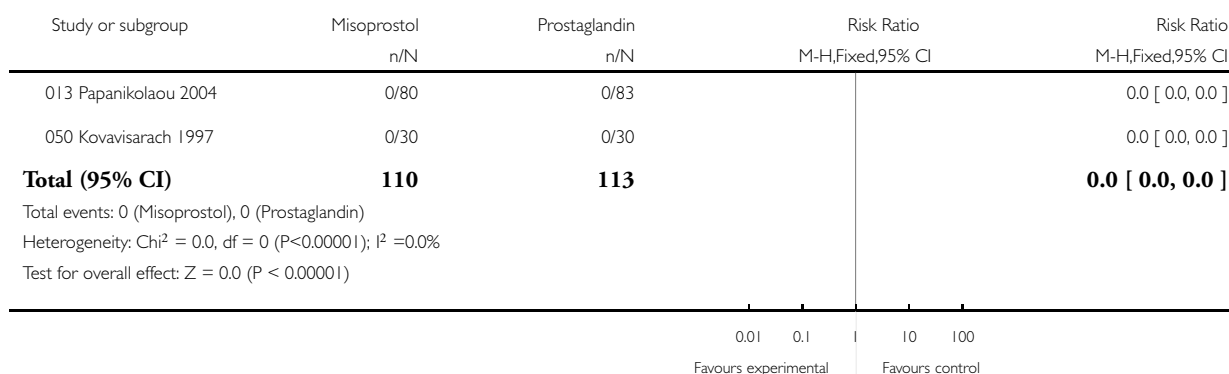


### Analysis 12.9. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 9 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 9 Serious maternal morbidity or death

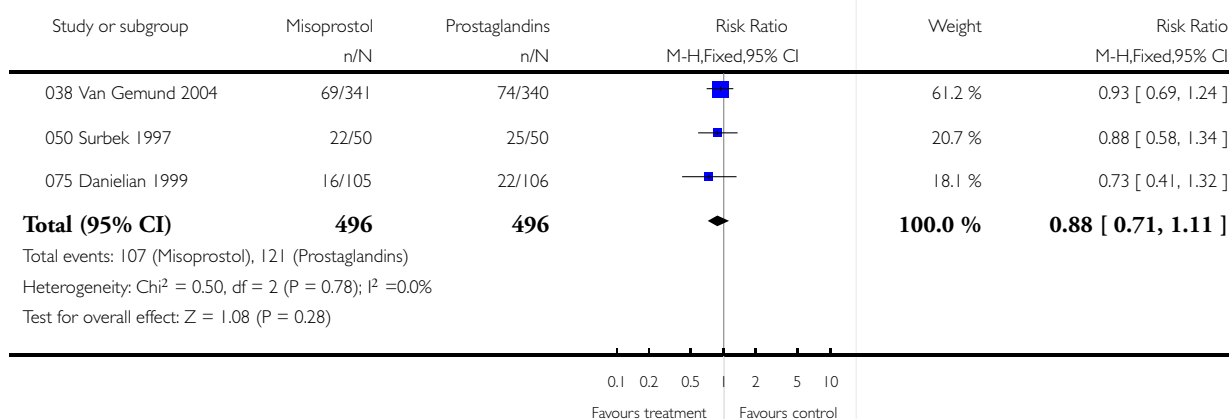


### Analysis 12.10. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 10 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 10 Epidural analgesia

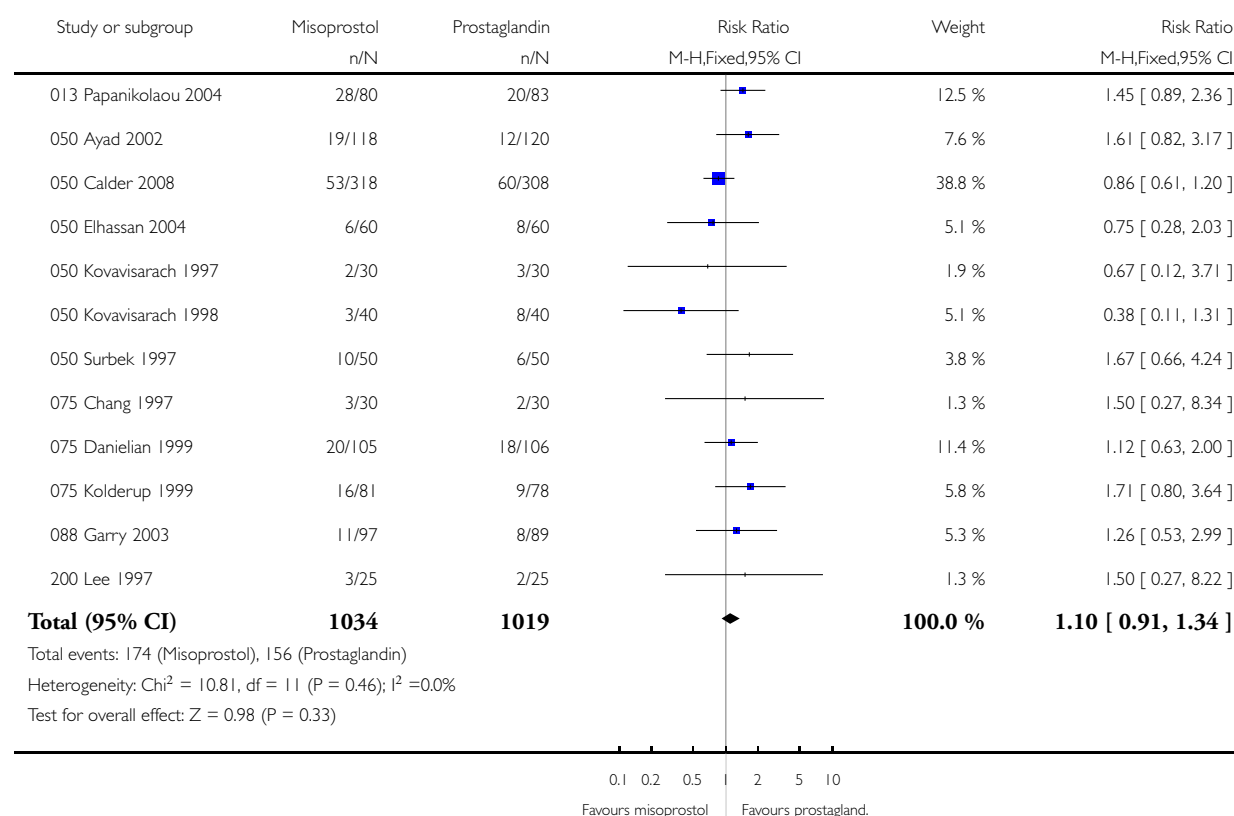


# **Analysis 12.11. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 11 Instrumental vaginal delivery.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 11 Instrumental vaginal delivery

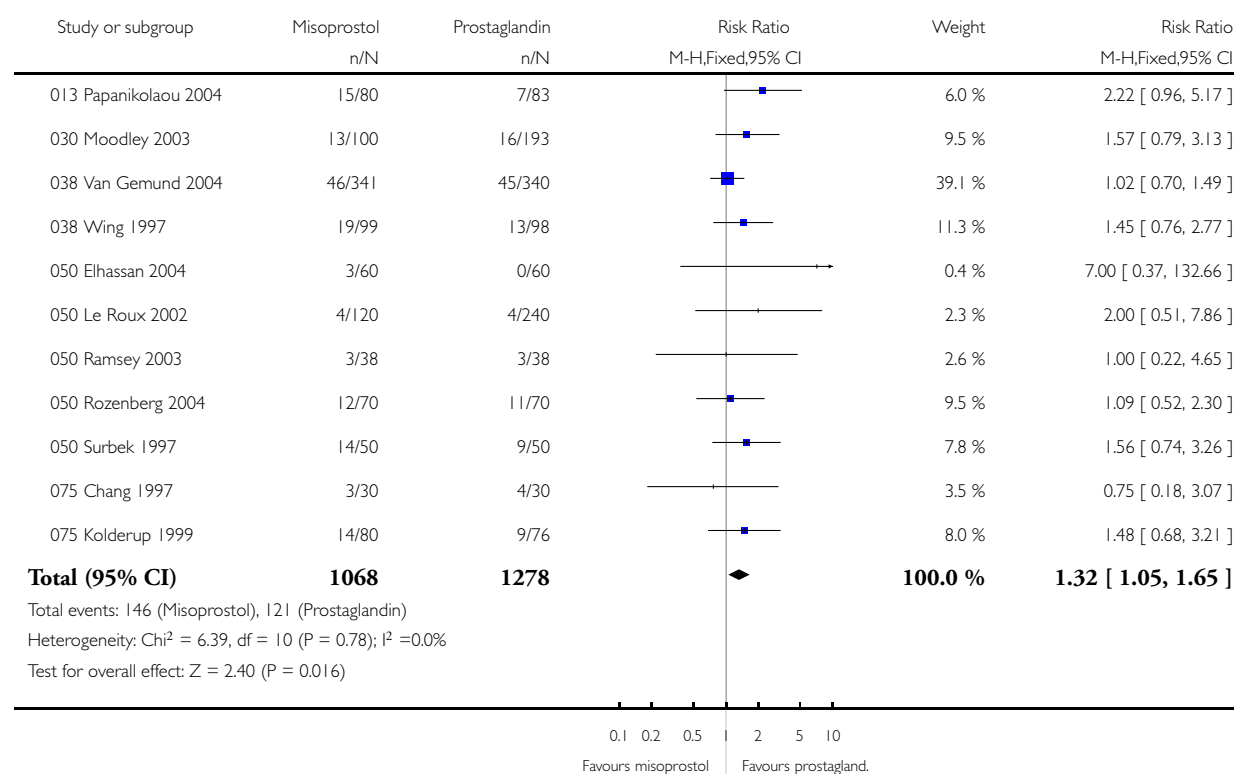


## Analysis 12.12. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 12 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 12 Meconium-stained liquor

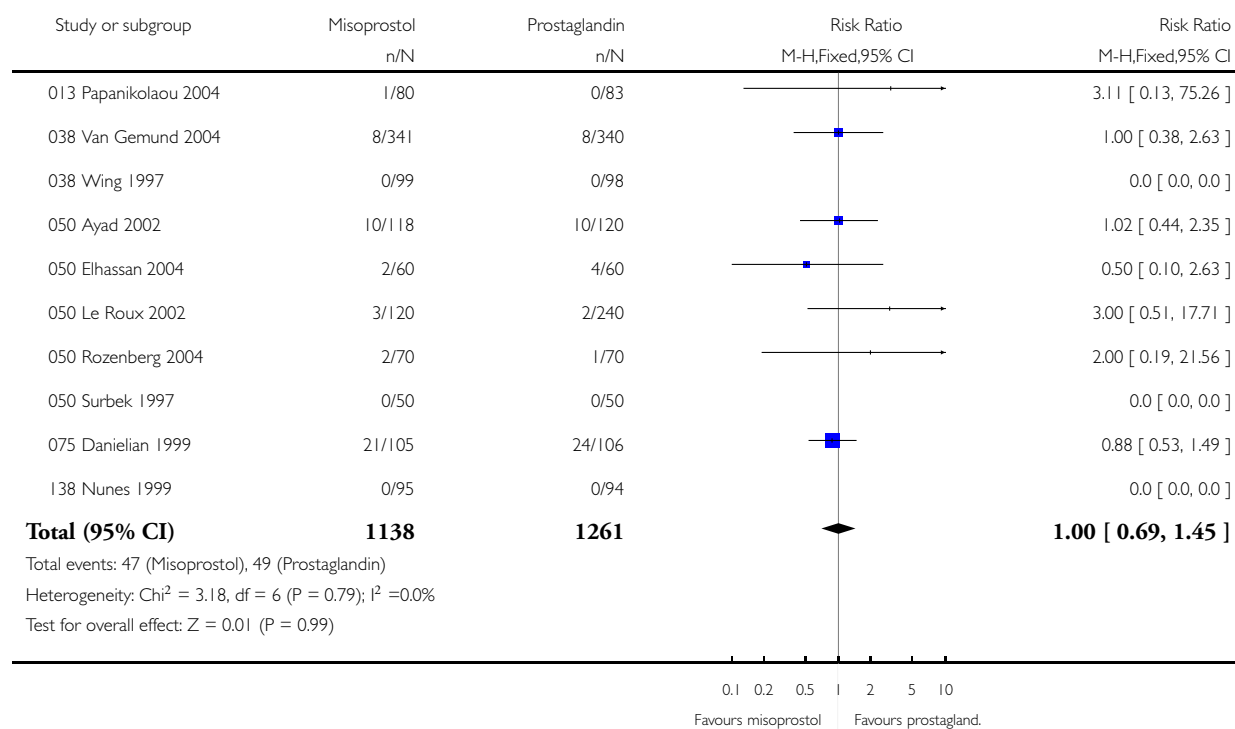


**Analysis 12.13. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 13 Apgar score < 7 at 5 minutes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 13 Apgar score < 7 at 5 minutes

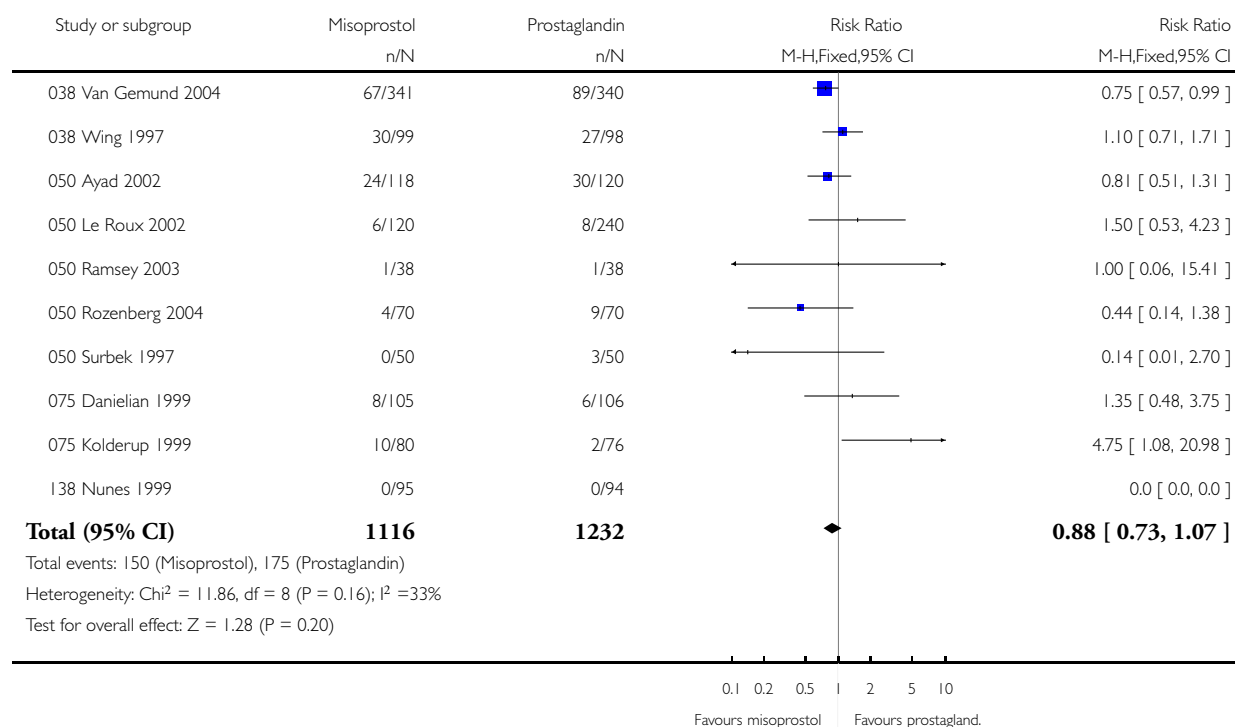


# **Analysis 12.14. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 14 Neonatal intensive care unit admission.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 14 Neonatal intensive care unit admission

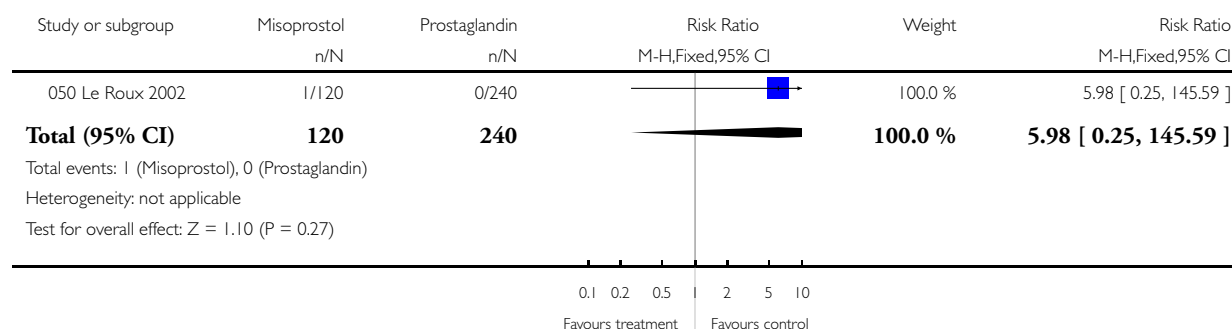


### Analysis 12.15. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 15 Neonatal encephalopathy.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 15 Neonatal encephalopathy

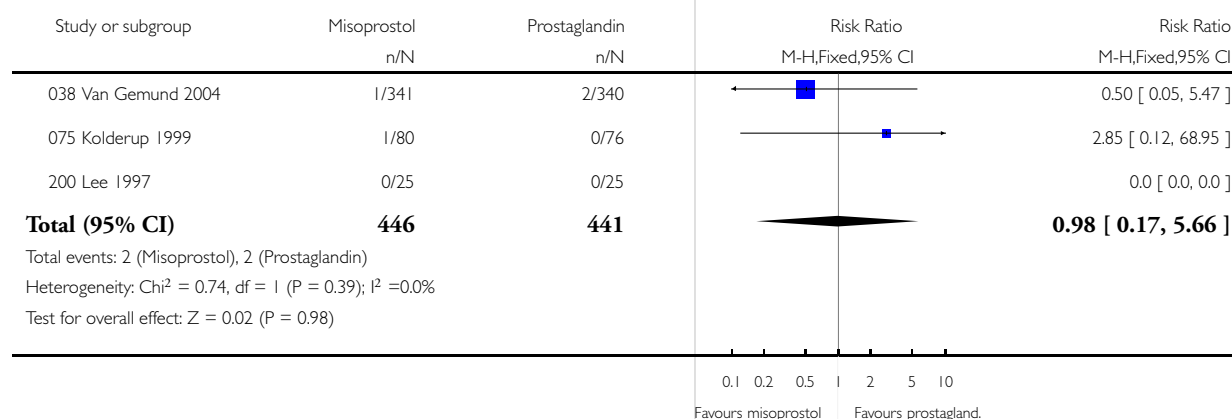


### Analysis 12.16. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 16 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 16 Perinatal death

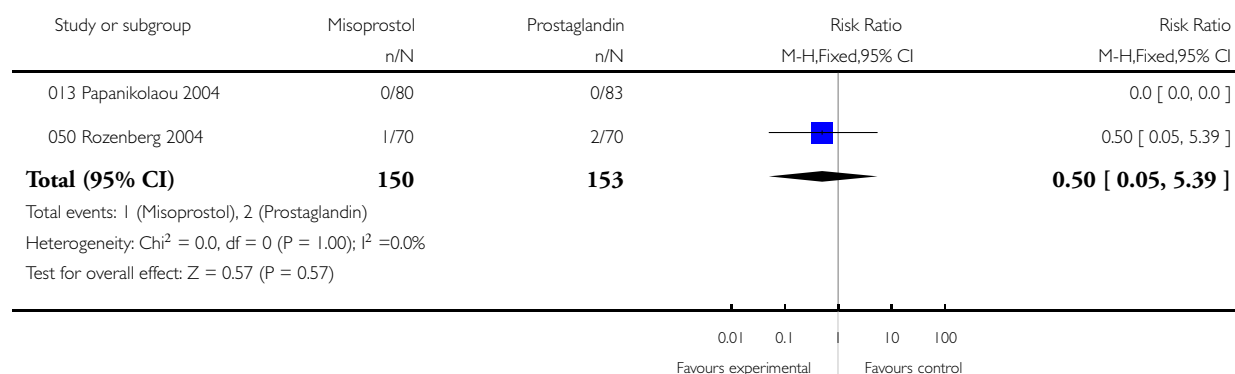


### Analysis 12.17. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 17 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 17 Serious maternal complications

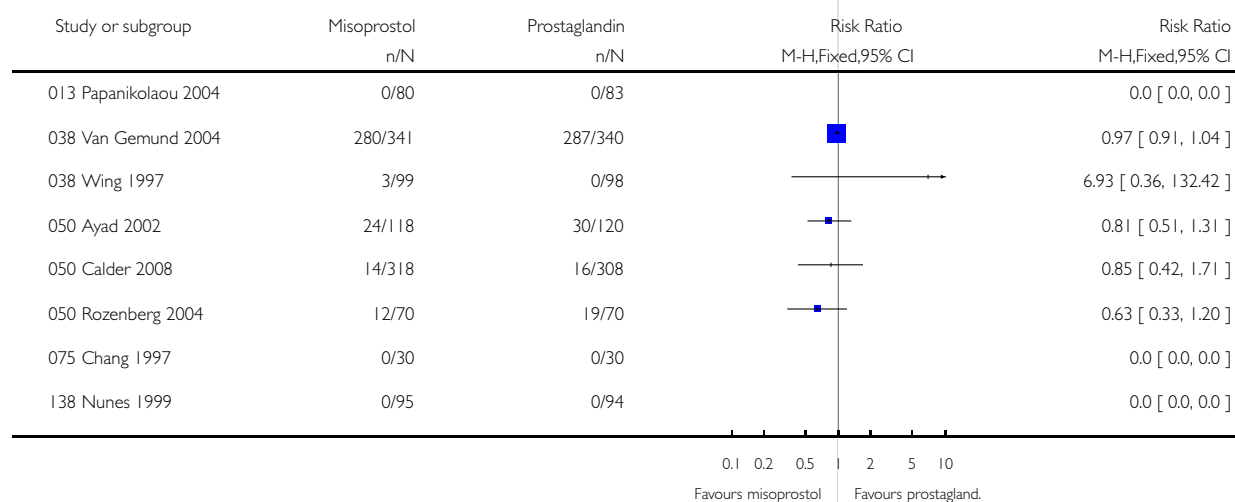


### Analysis 12.18. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 18 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

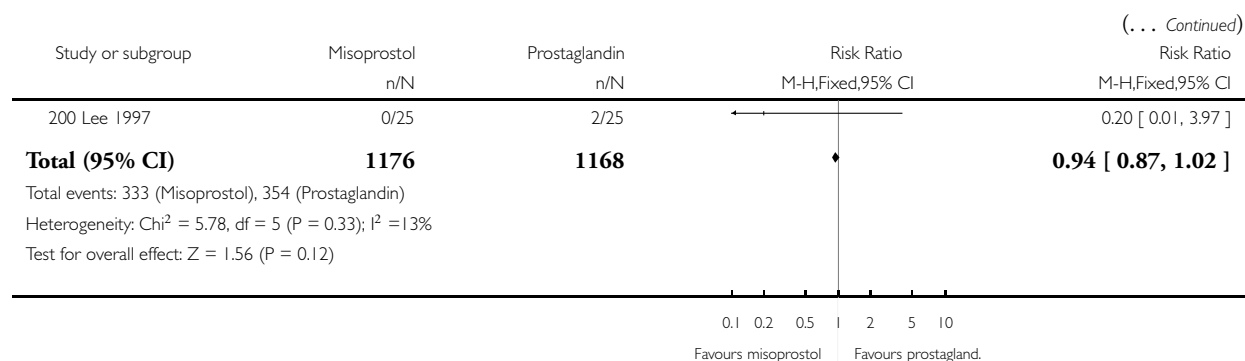
Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 18 Maternal side effects (eg nausea, vomiting, diarrhoea)



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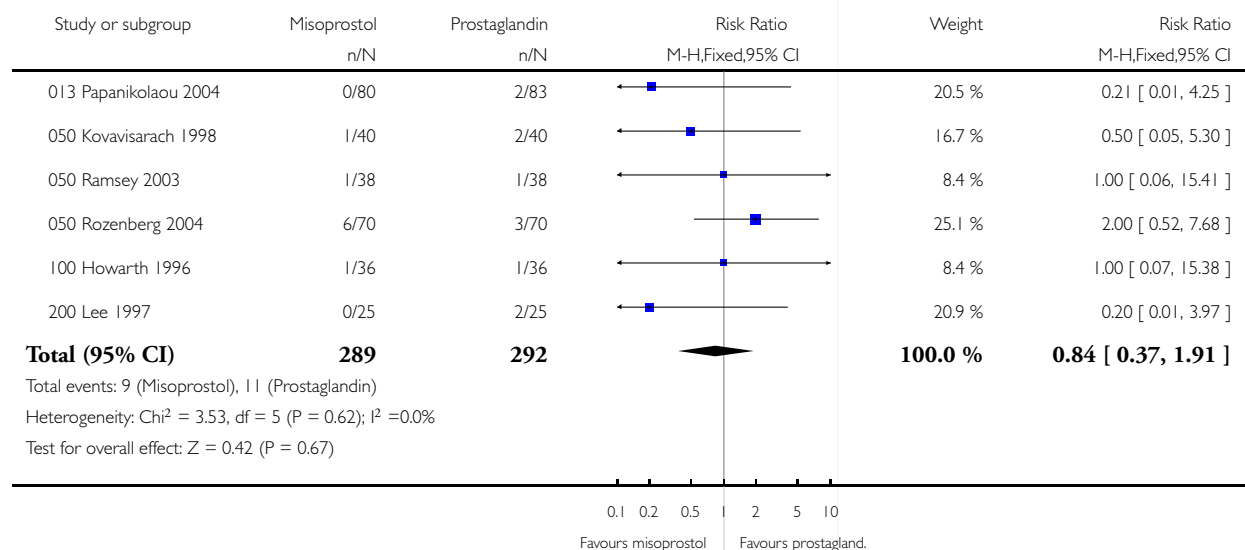


### Analysis 12.19. Comparison 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix, Outcome 19 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 12 Misoprostol versus vaginal prostaglandin: all women, unfavourable cervix

Outcome: 19 Postpartum haemorrhage

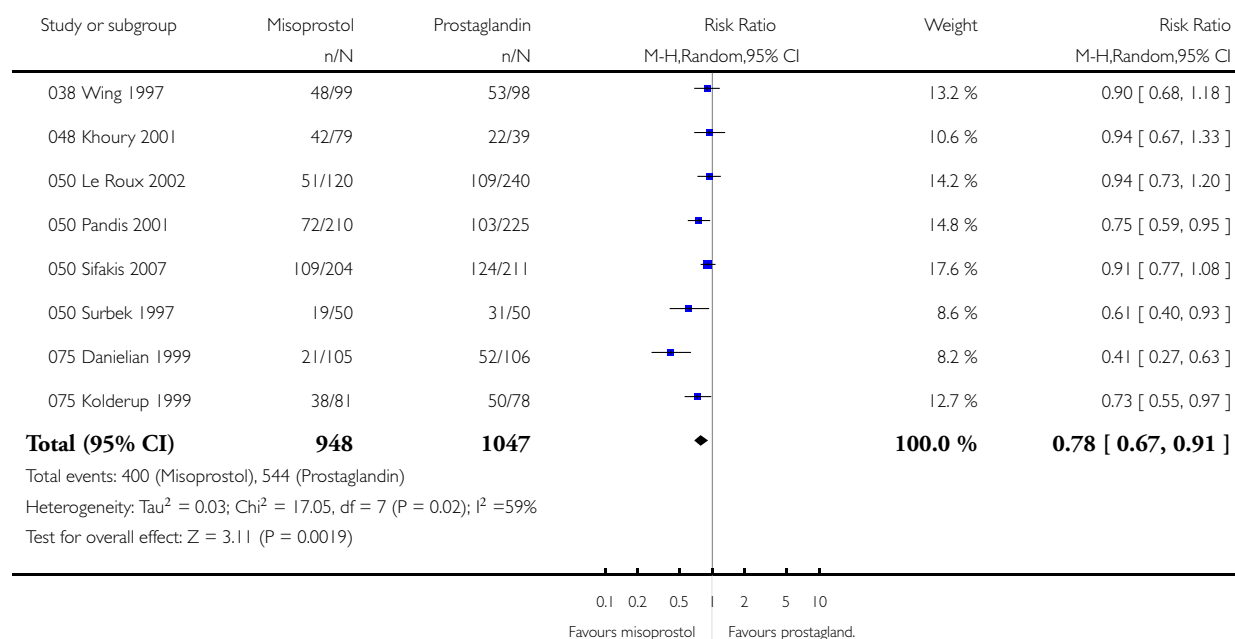


### Analysis 13.1. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

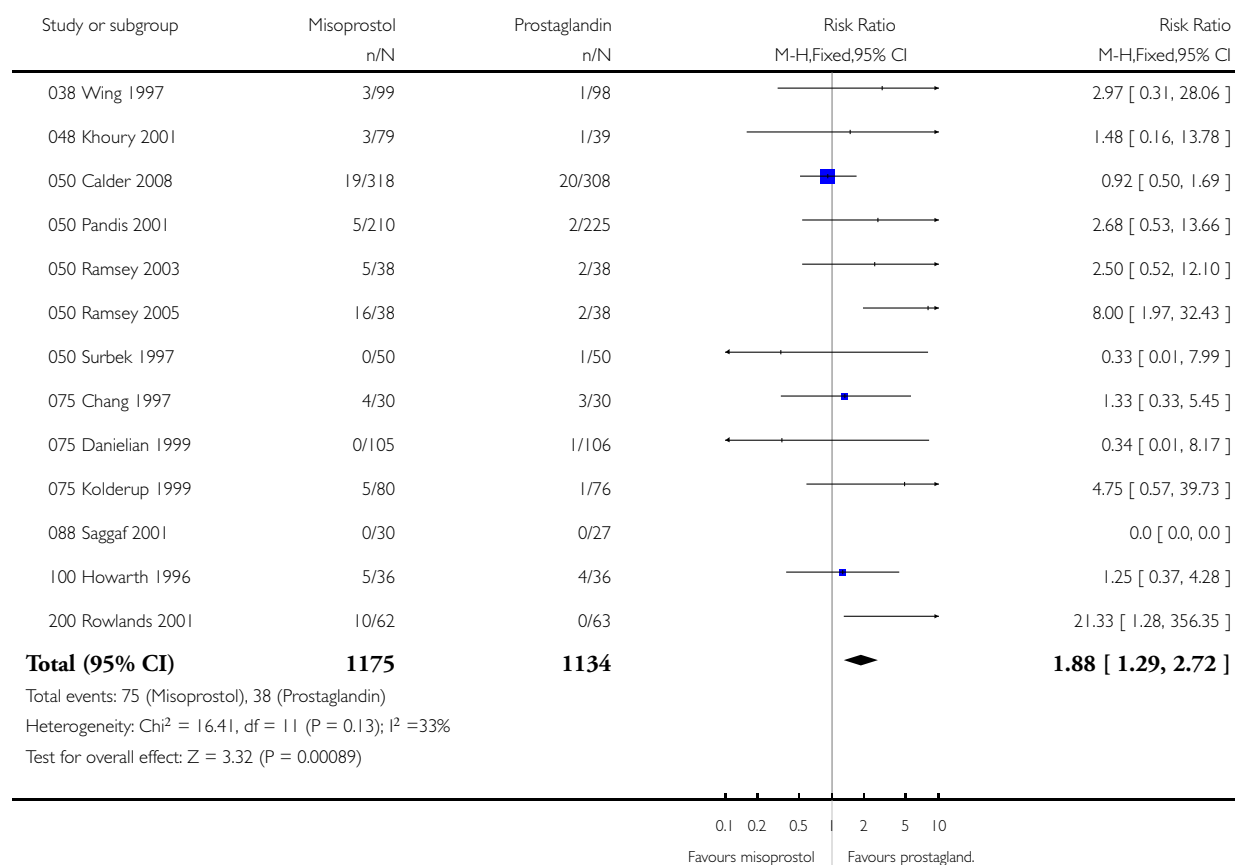


### Analysis 13.2. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

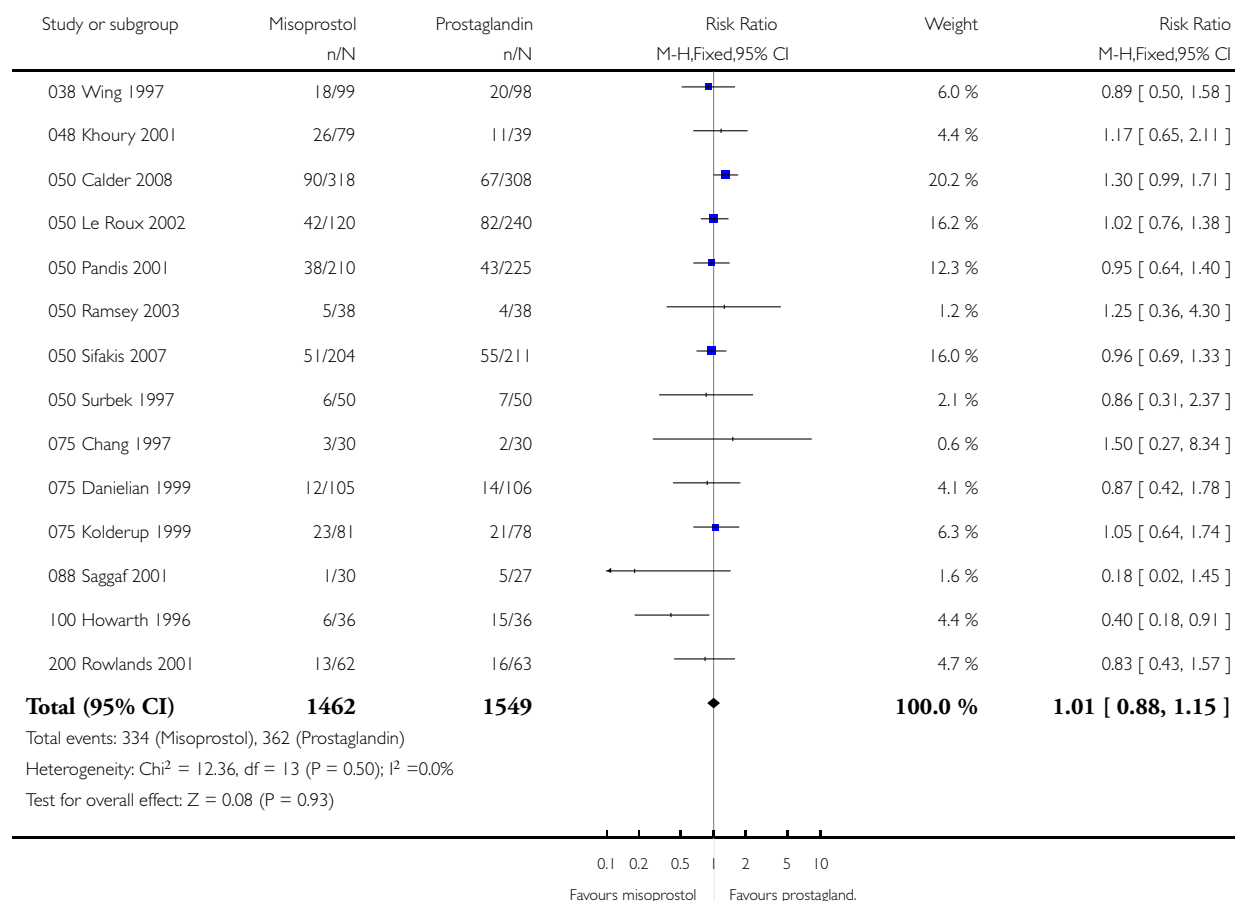


### Analysis 13.3. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 3 Caesarean section

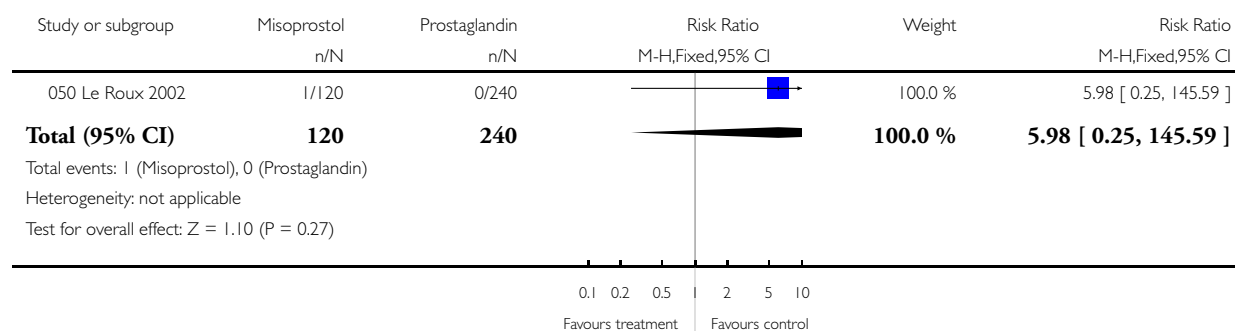


### Analysis 13.4. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 4 Serious neonatal morbidity or perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 4 Serious neonatal morbidity or perinatal death

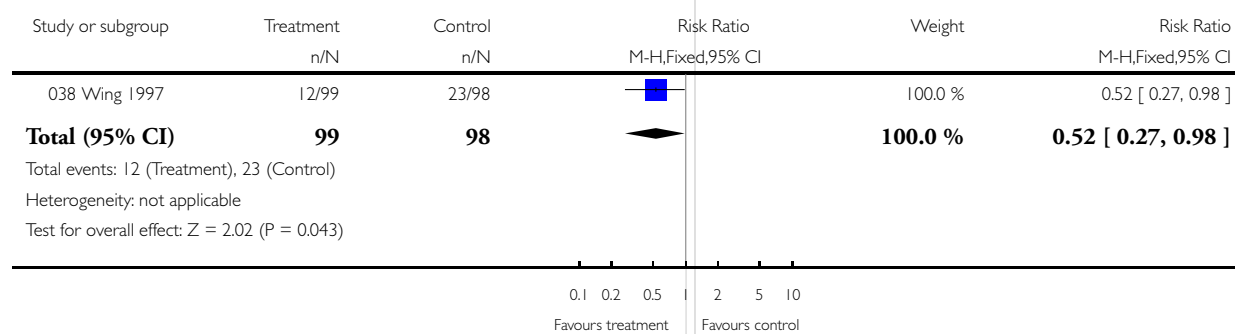


### Analysis 13.5. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 5 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 5 Cervix unfavourable/unchanged after 12-24 hours

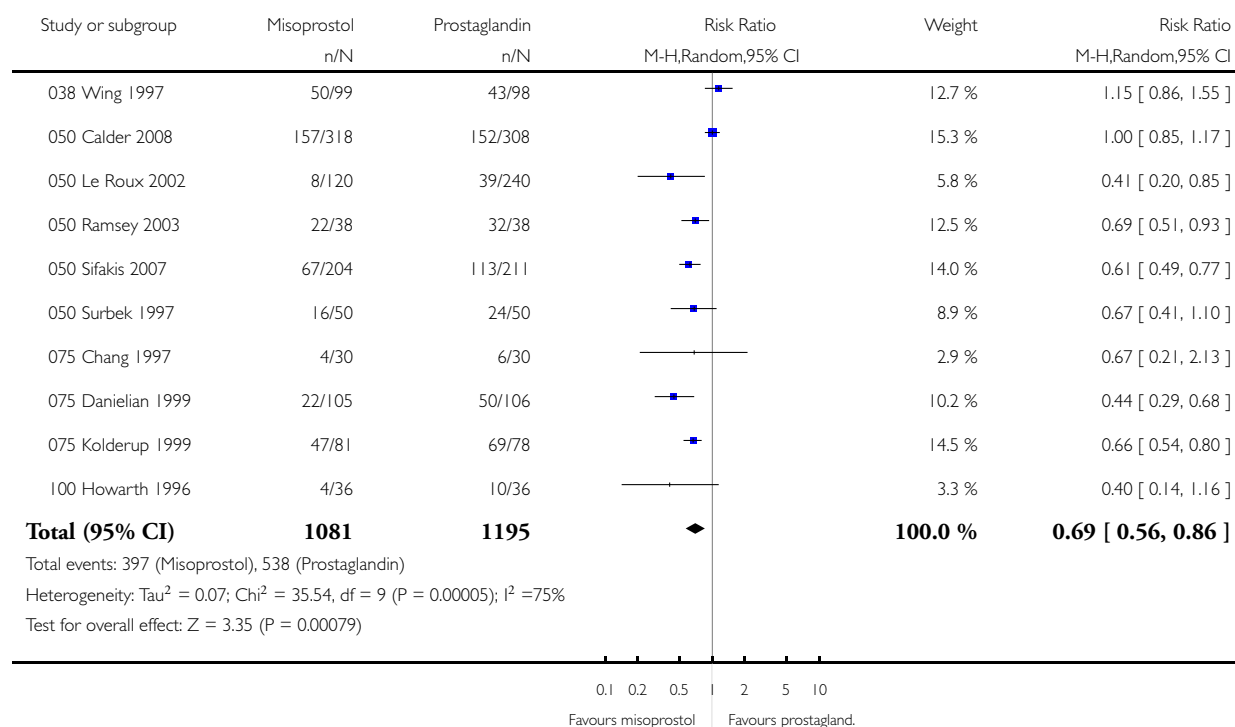


### Analysis 13.6. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 6 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 6 Oxytocin augmentation

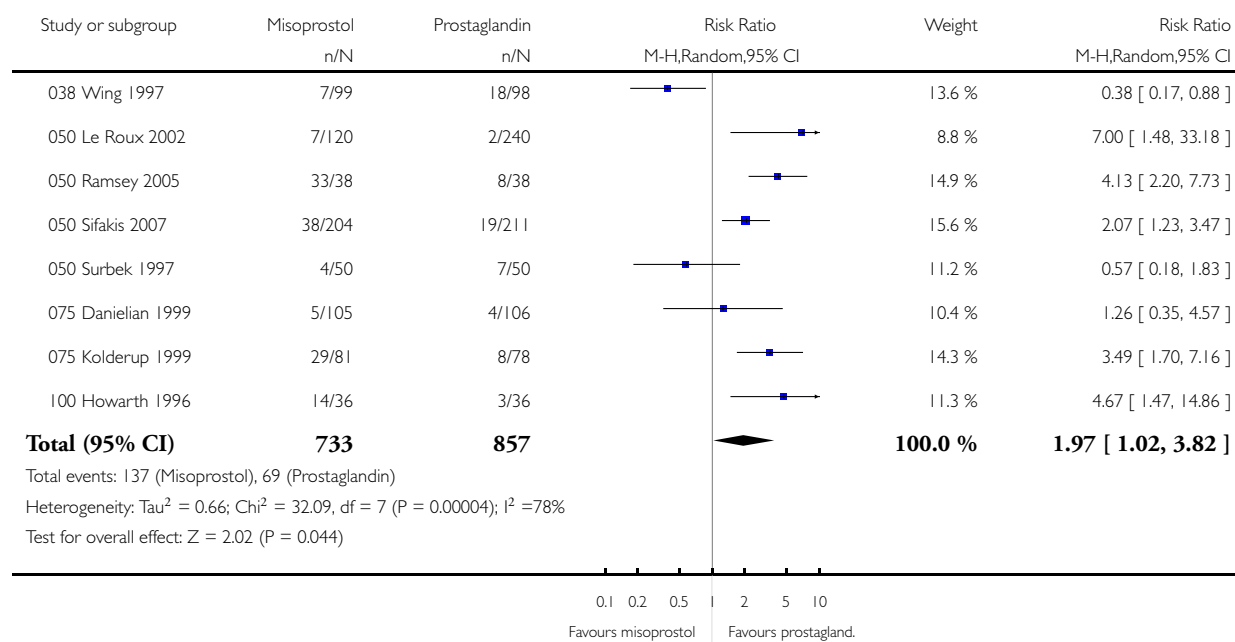


### Analysis 13.7. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 7 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 7 Uterine hyperstimulation without FHR changes

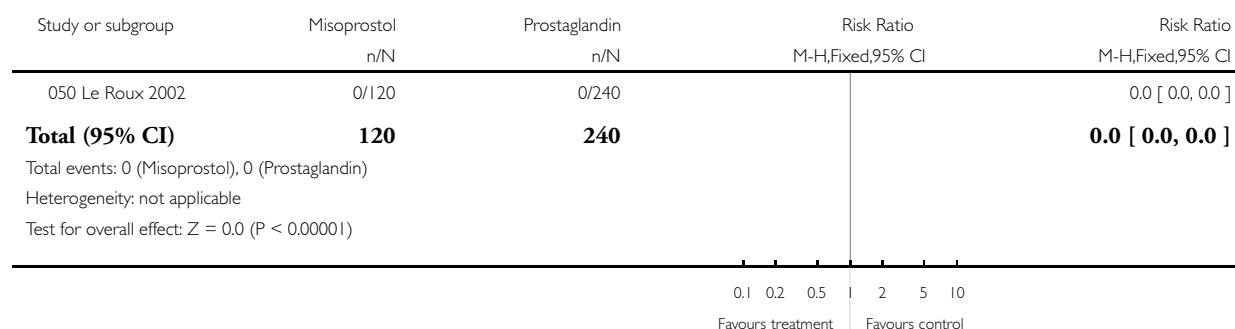


### Analysis 13.8. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 8 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 8 Uterine rupture

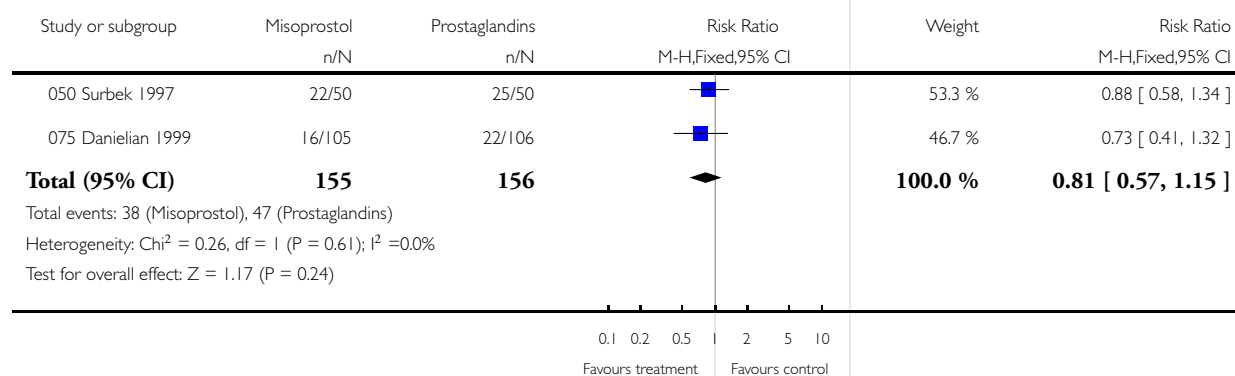


### Analysis 13.9. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 9 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 9 Epidural analgesia



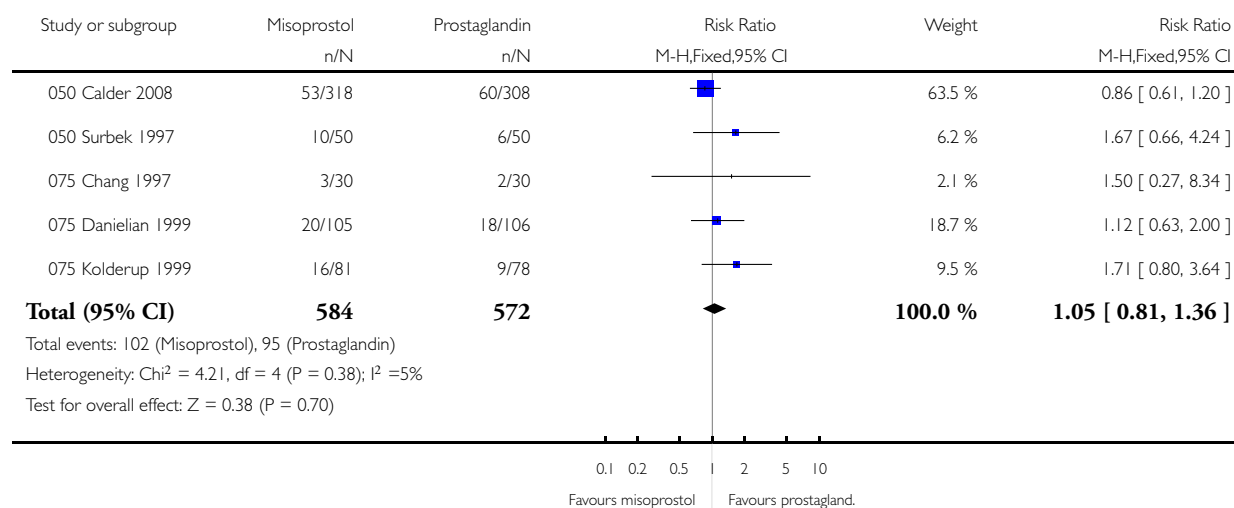


### Analysis 13.10. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 10 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 10 Instrumental vaginal delivery

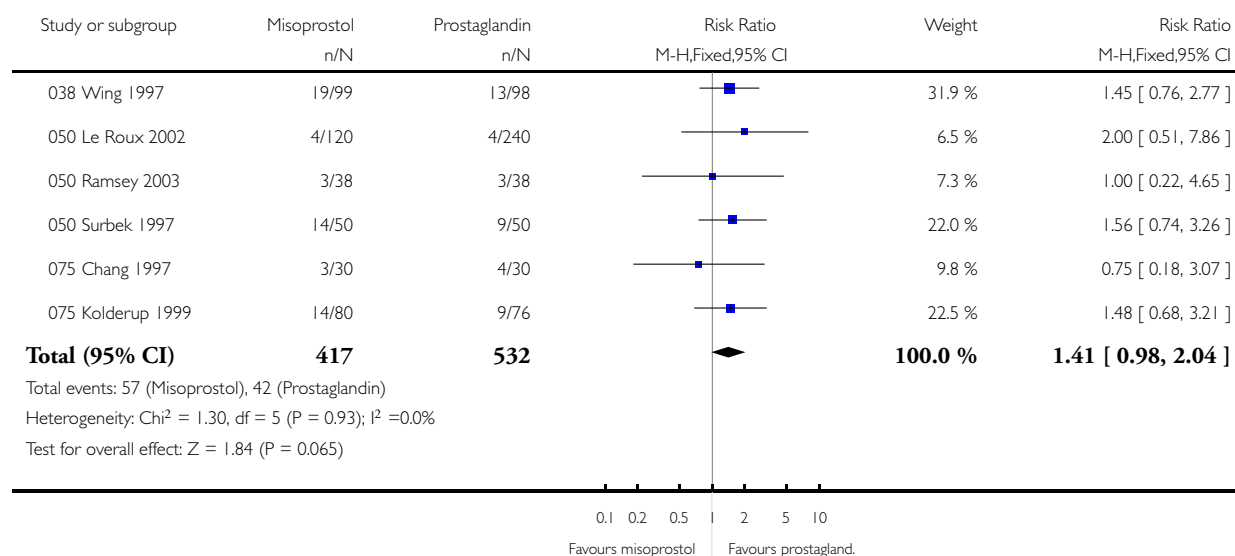


### Analysis 13.11. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 11 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 11 Meconium-stained liquor

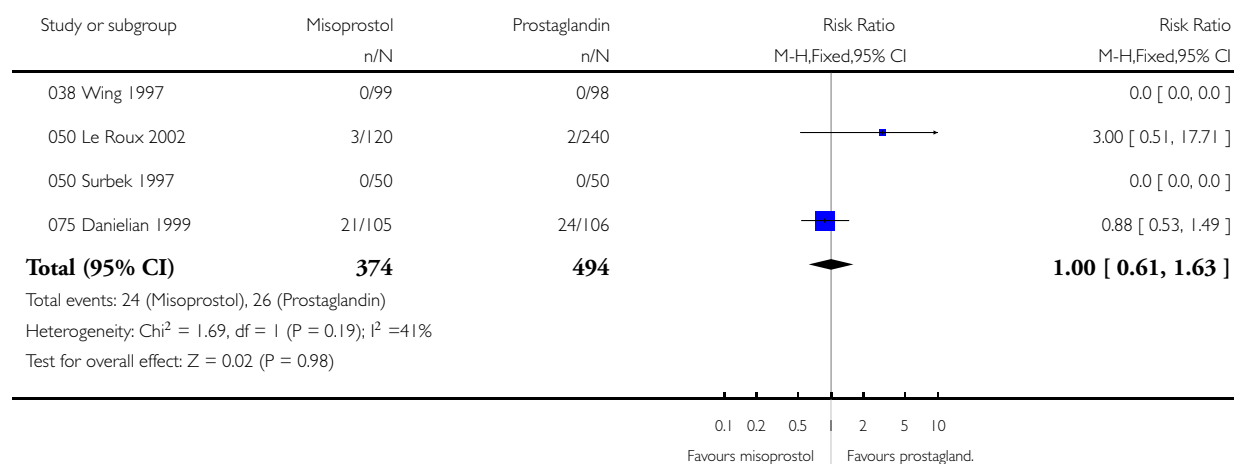


### Analysis 13.12. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 12 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 12 Apgar score < 7 at 5 minutes

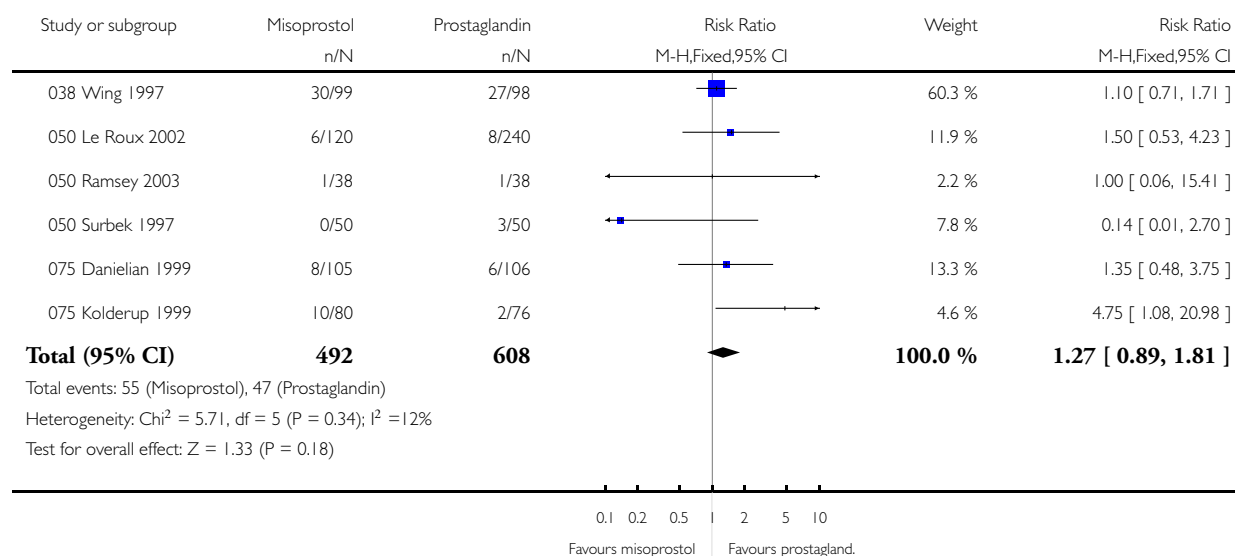


### Analysis 13.13. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 13 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 13 Neonatal intensive care unit admission

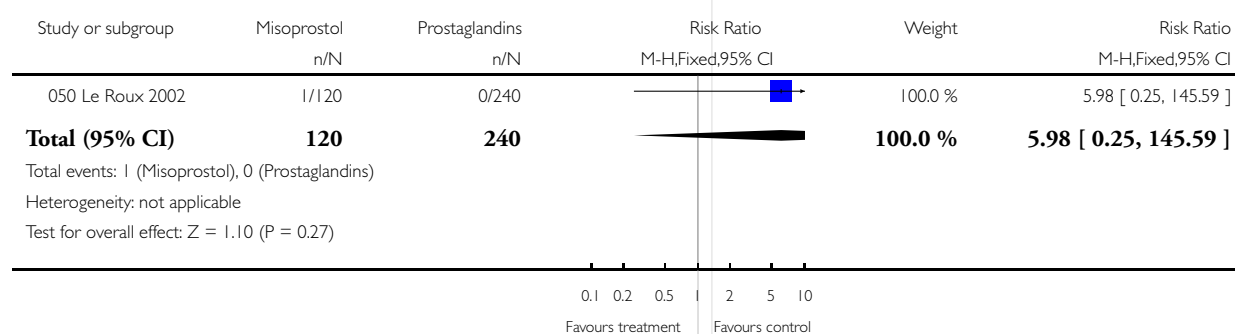


### Analysis 13.14. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 14 Neonatal encephalopathy.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 14 Neonatal encephalopathy

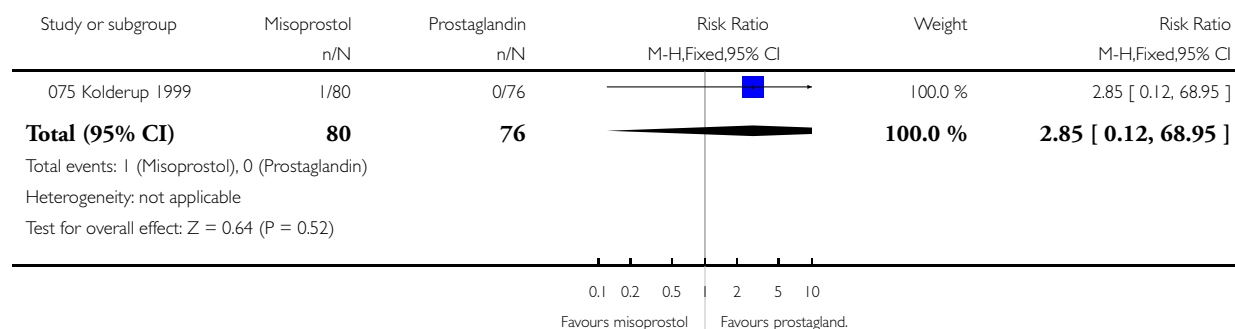


### Analysis 13.15. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 15 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 15 Perinatal death

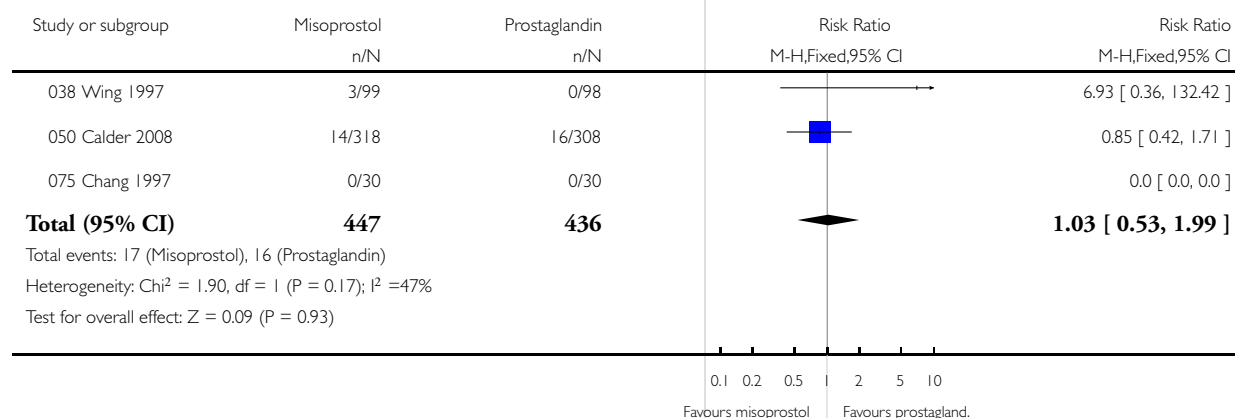


### Analysis 13.16. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 16 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 16 Maternal side effects (eg nausea, vomiting, diarrhoea)

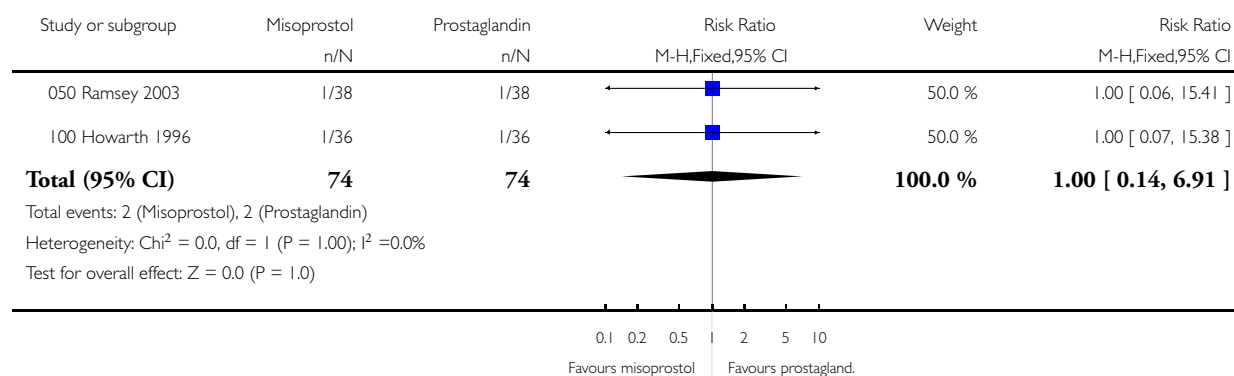


### Analysis 13.17. Comparison 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix, Outcome 17 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 13 Misoprostol versus vaginal prostaglandin: all women, intact membranes, unfavourable cervix

Outcome: 17 Postpartum haemorrhage

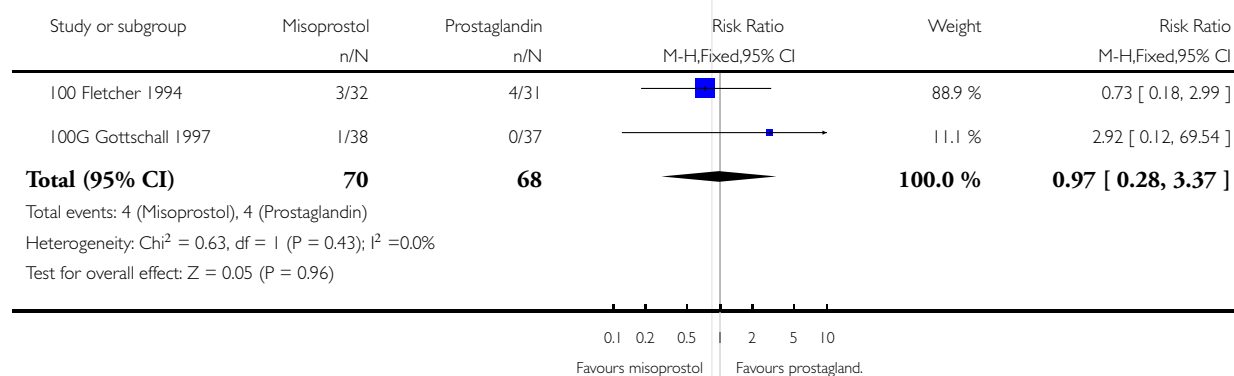


### Analysis 14.1. Comparison 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix, Outcome 1 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix

Outcome: 1 Uterine hyperstimulation with FHR changes

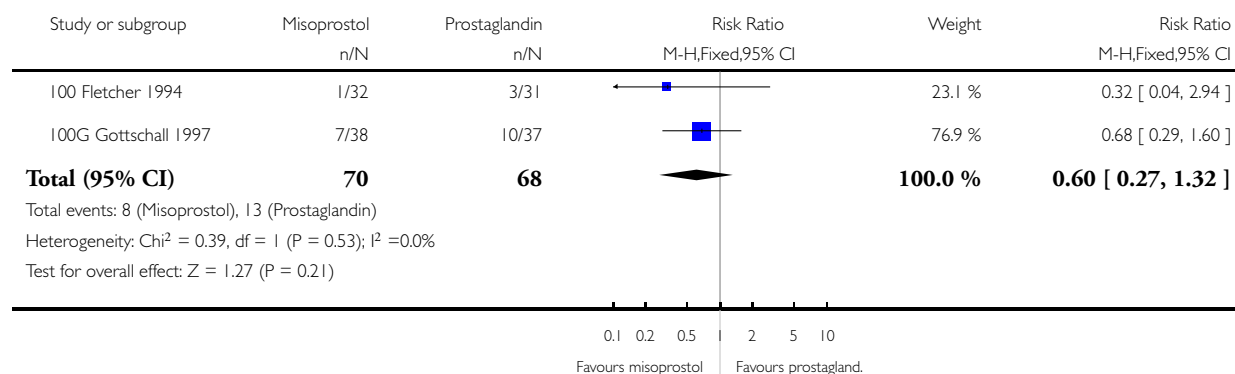


## Analysis 14.2. Comparison 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix

Outcome: 2 Caesarean section

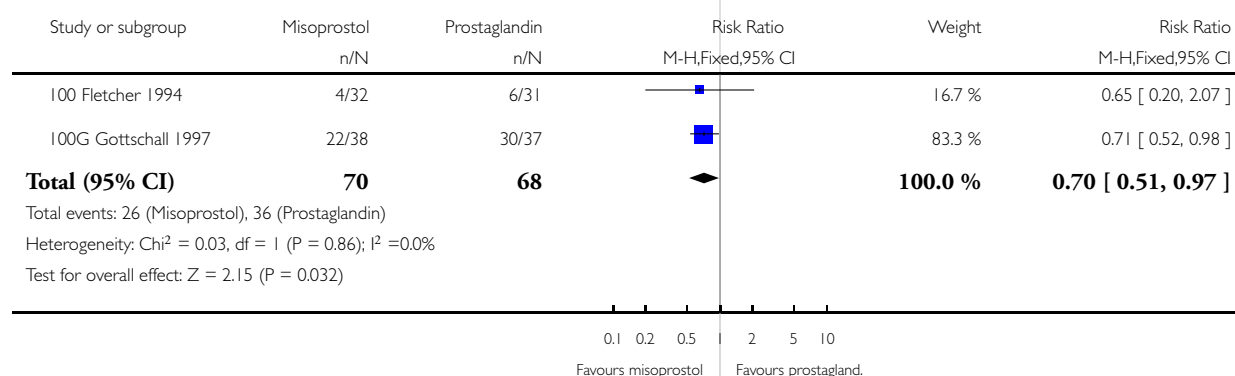


## Analysis 14.3. Comparison 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix, Outcome 3 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix

Outcome: 3 Oxytocin augmentation

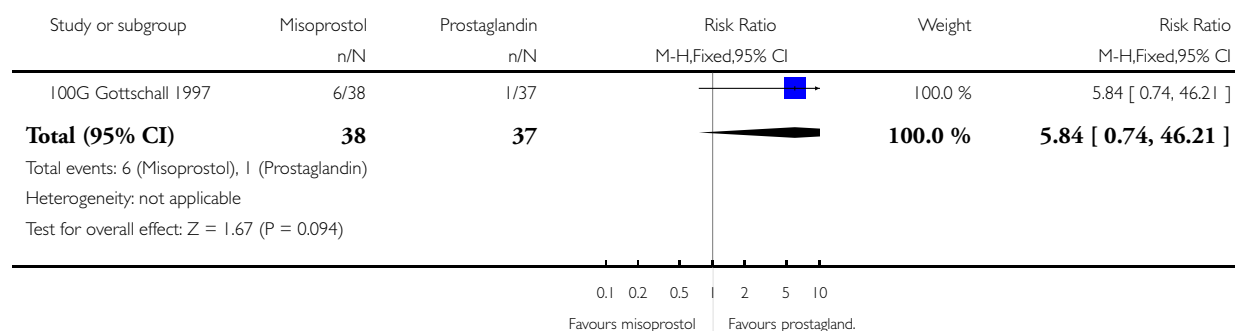


#### Analysis 14.4. Comparison 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix, Outcome 4 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix

Outcome: 4 Uterine hyperstimulation without FHR changes

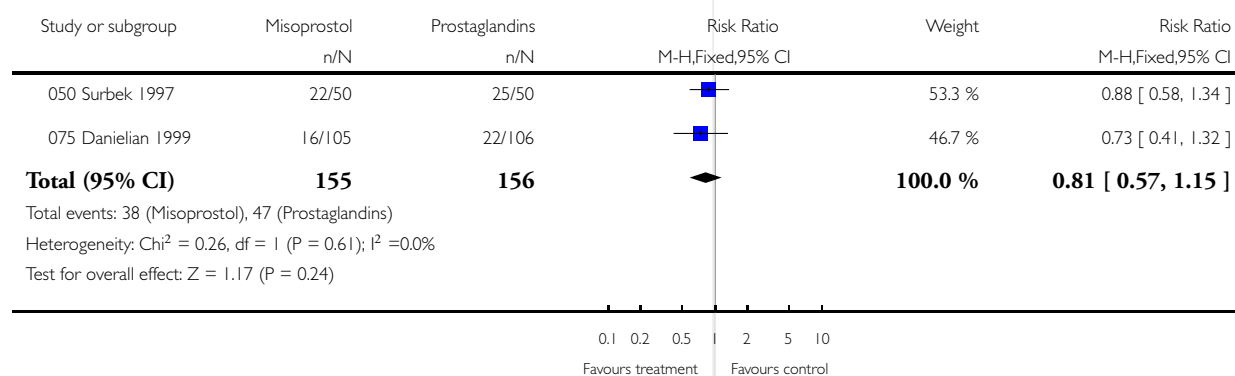


#### Analysis 14.5. Comparison 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix, Outcome 5 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix

Outcome: 5 Epidural analgesia



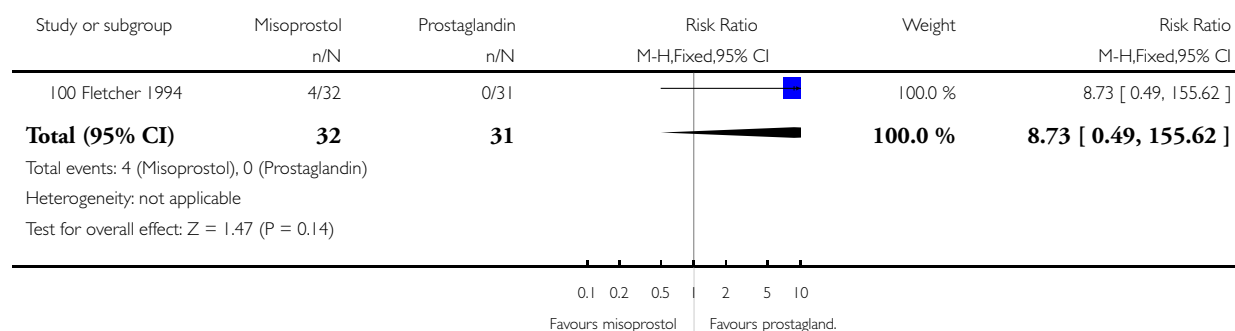


#### Analysis 14.6. Comparison 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix, Outcome 6 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix

Outcome: 6 Instrumental vaginal delivery

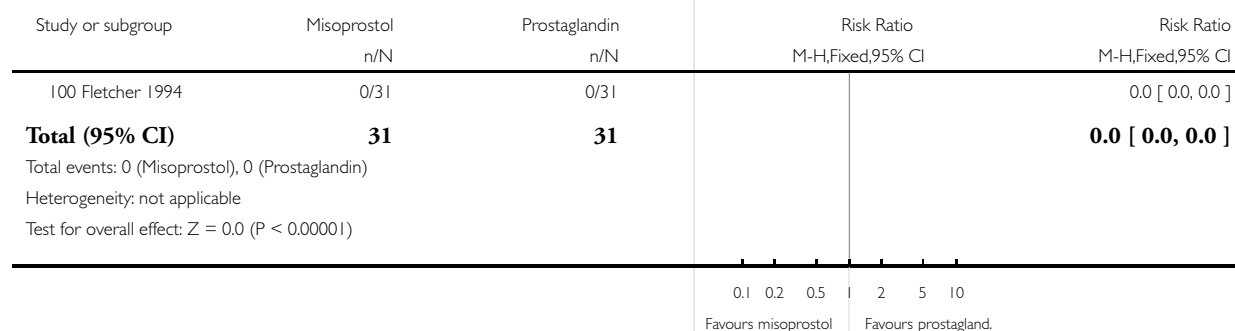


#### Analysis 14.7. Comparison 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix, Outcome 7 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 14 Misoprostol versus vaginal prostaglandin: all women, intact membranes, variable or undefined cervix

Outcome: 7 Perinatal death

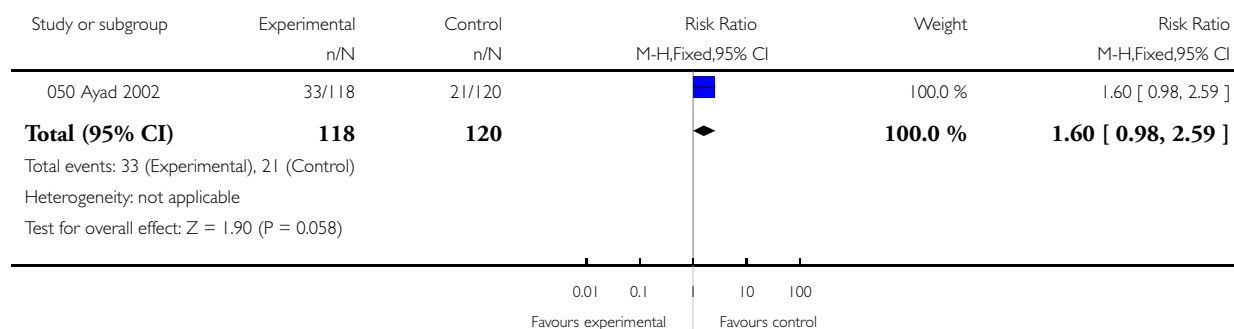


### Analysis 15.1. Comparison 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix

Outcome: 1 Caesarean section

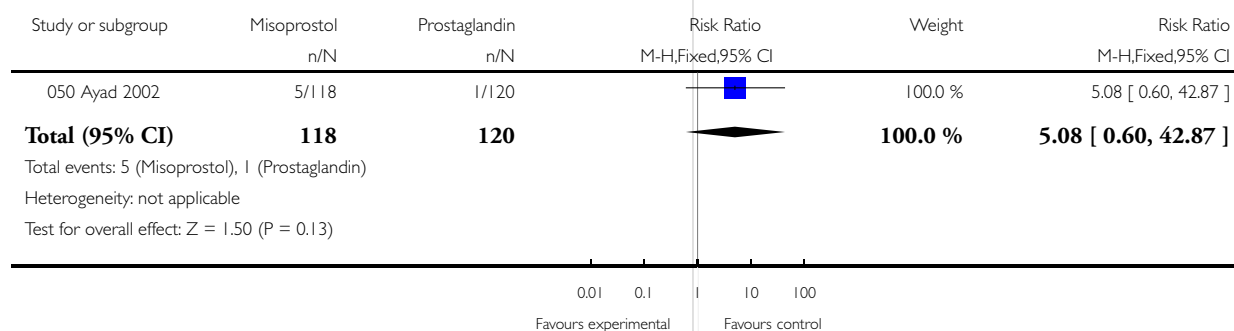


### Analysis 15.2. Comparison 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

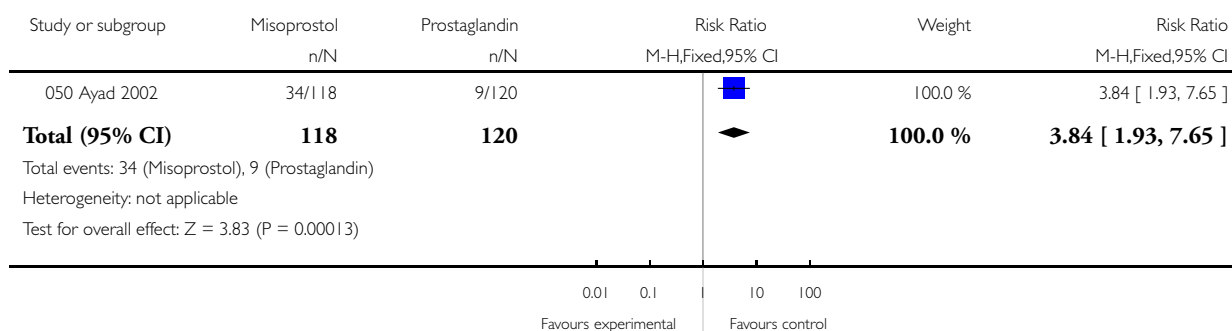


### Analysis 15.3. Comparison 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix, Outcome 3 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix

Outcome: 3 Uterine hyperstimulation without FHR changes

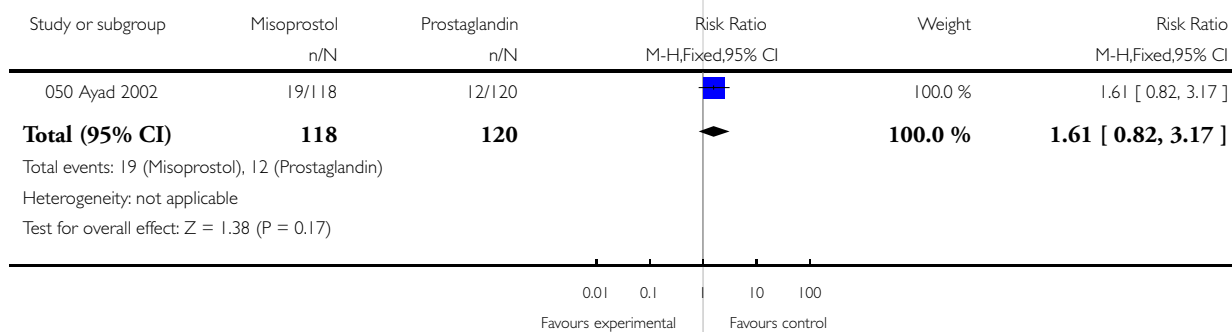


### Analysis 15.4. Comparison 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix, Outcome 4 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix

Outcome: 4 Instrumental vaginal delivery

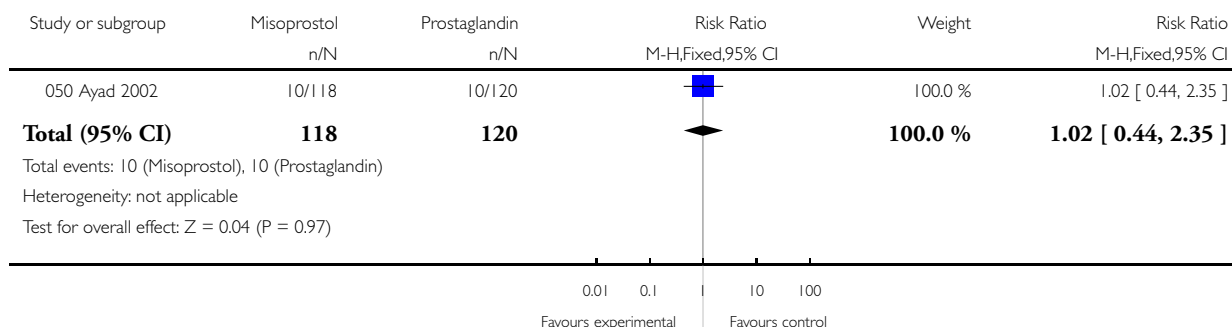


### Analysis 15.5. Comparison 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix, Outcome 5 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix

Outcome: 5 Apgar score < 7 at 5 minutes

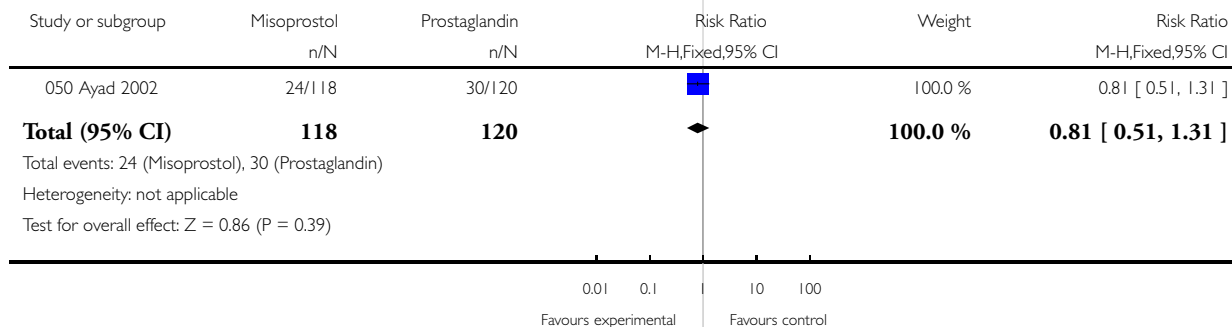


### Analysis 15.6. Comparison 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix, Outcome 6 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix

Outcome: 6 Neonatal intensive care unit admission

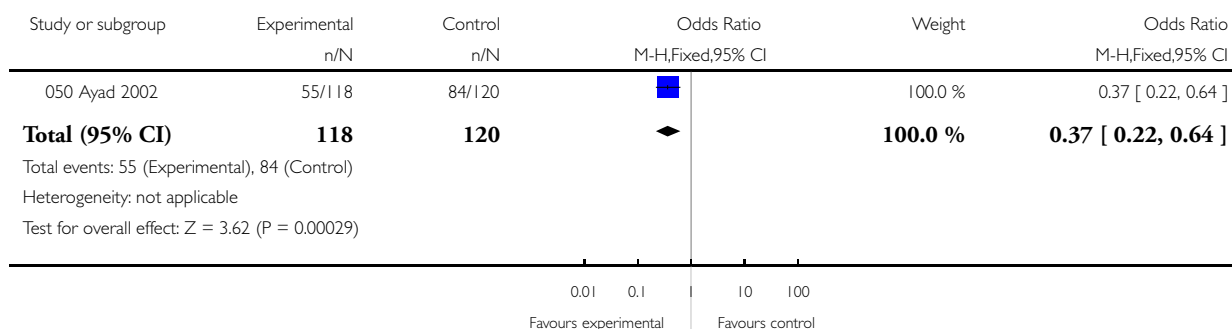


### Analysis 15.7. Comparison 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix, Outcome 7 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 15 Misoprostol versus vaginal prostaglandin: all women with ruptured membranes and unfavourable cervix

Outcome: 7 Oxytocin augmentation

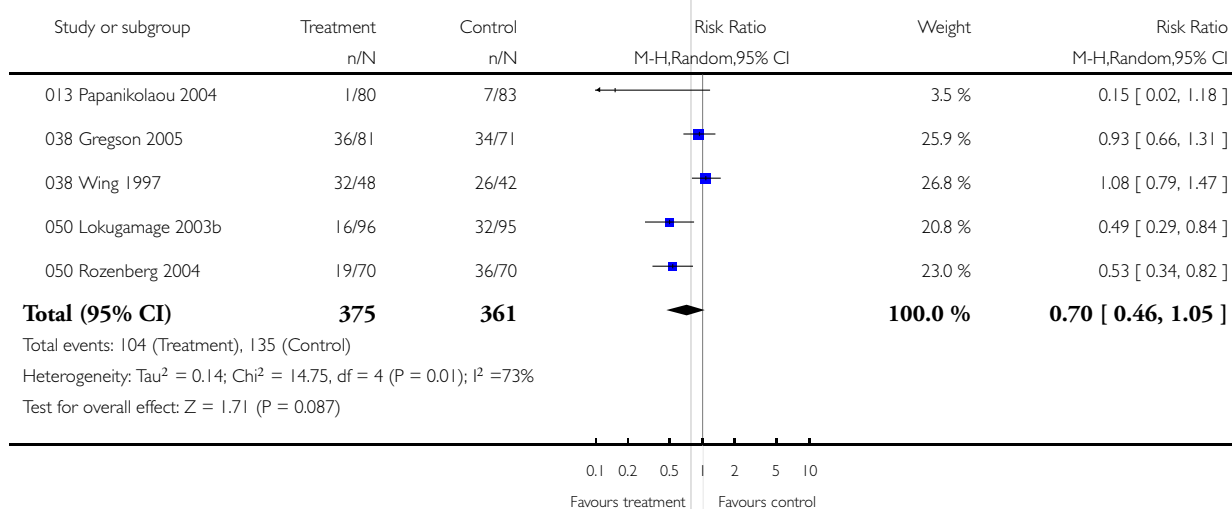


### Analysis 16.1. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 1 Vaginal delivery not achieved within 24 hours

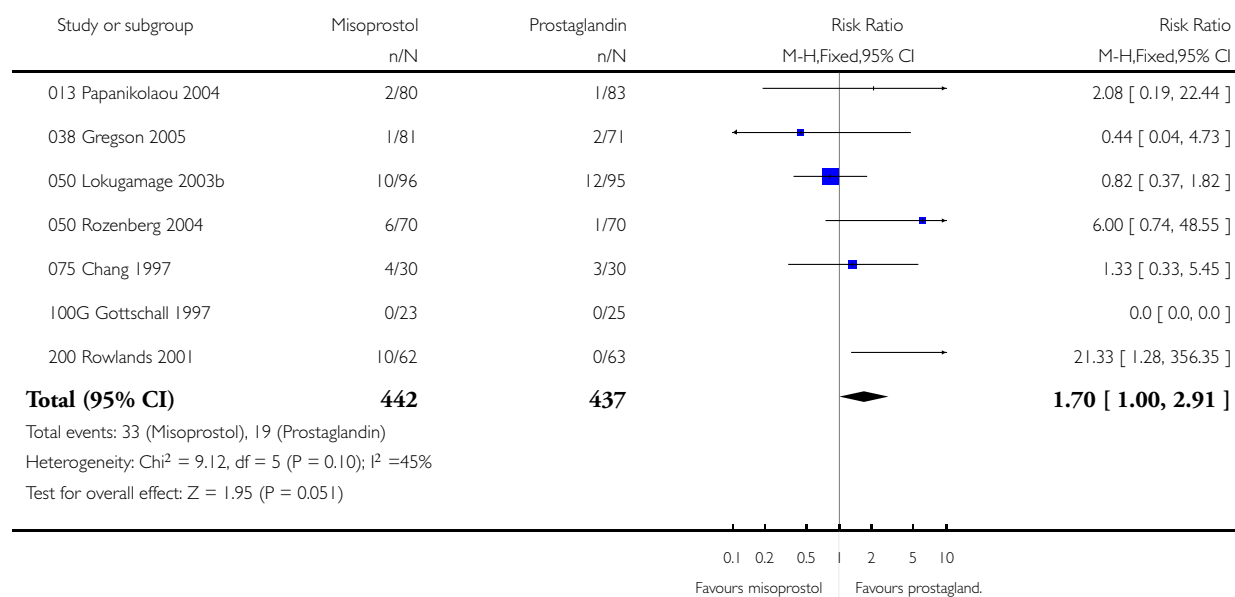


## Analysis 16.2. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 2 Uterine hyperstimulation with FHR changes

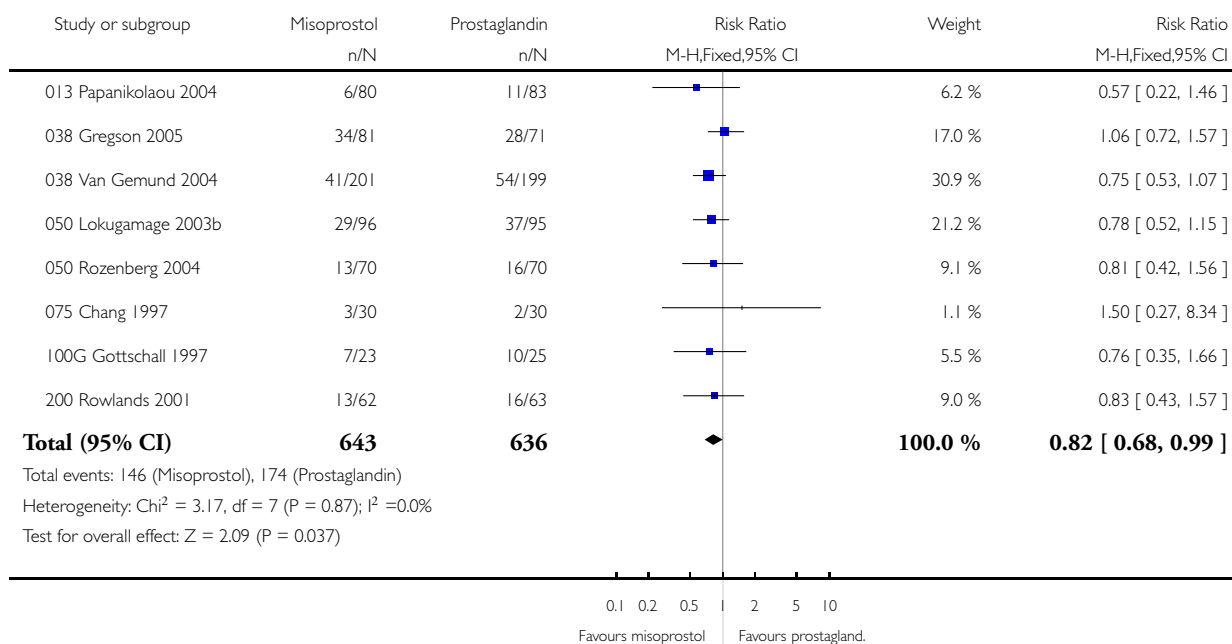


### Analysis 16.3. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 3 Caesarean section

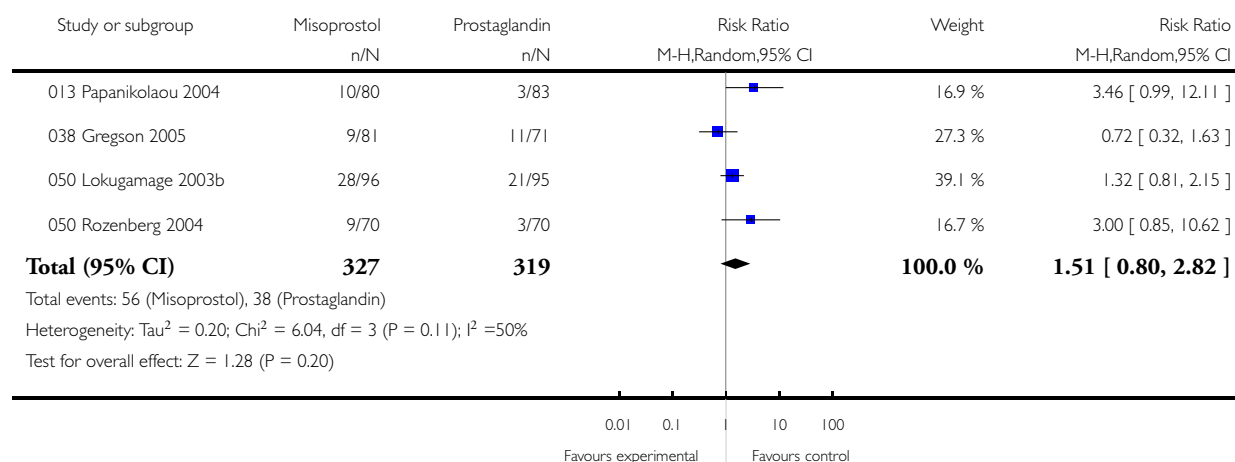


#### Analysis 16.4. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 4 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 4 Uterine hyperstimulation without FHR changes

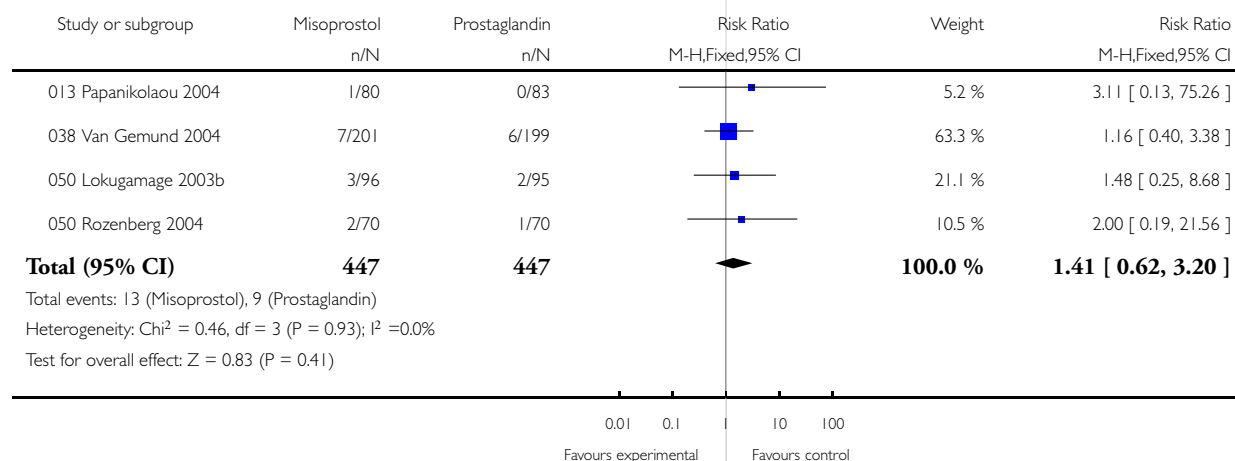


#### Analysis 16.5. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 5 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 5 Apgar score < 7 at 5 minutes



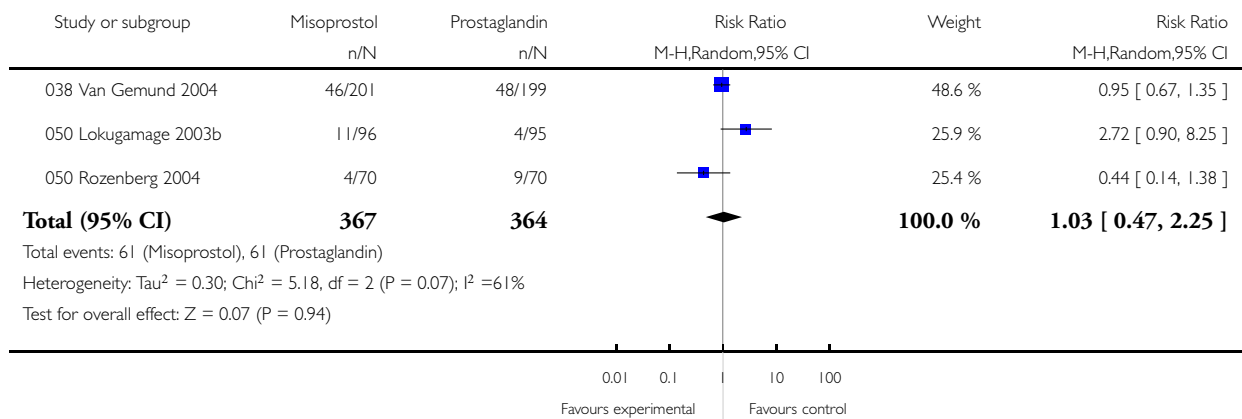


### Analysis 16.6. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 6 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 6 Neonatal intensive care unit admission

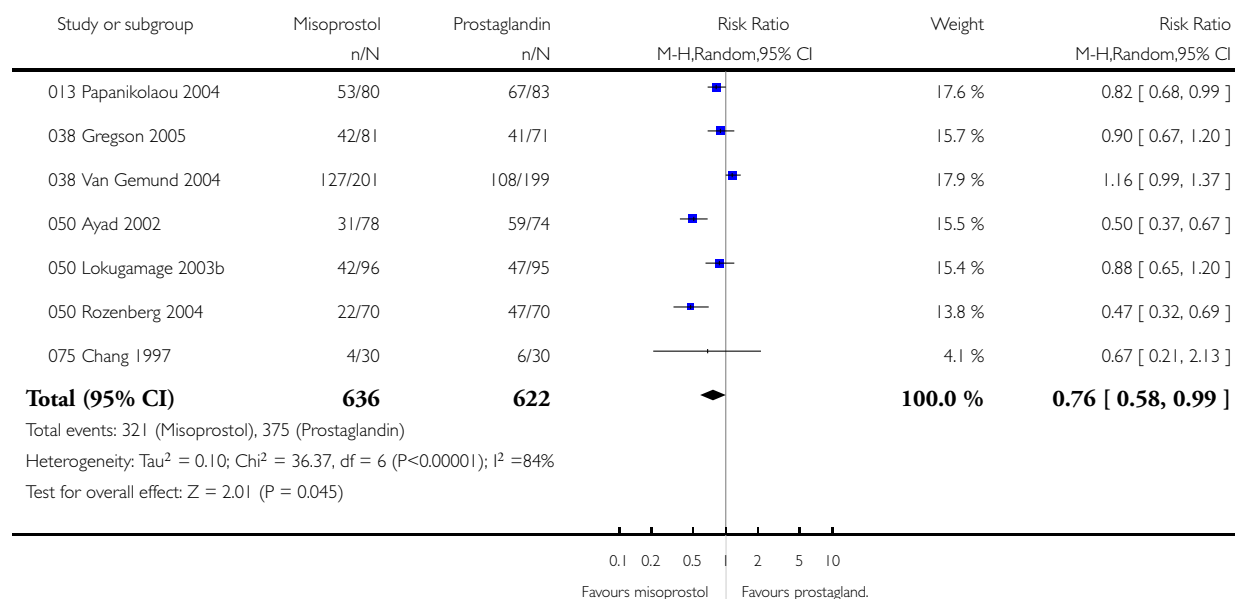


### Analysis 16.7. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 7 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 7 Oxytocin augmentation

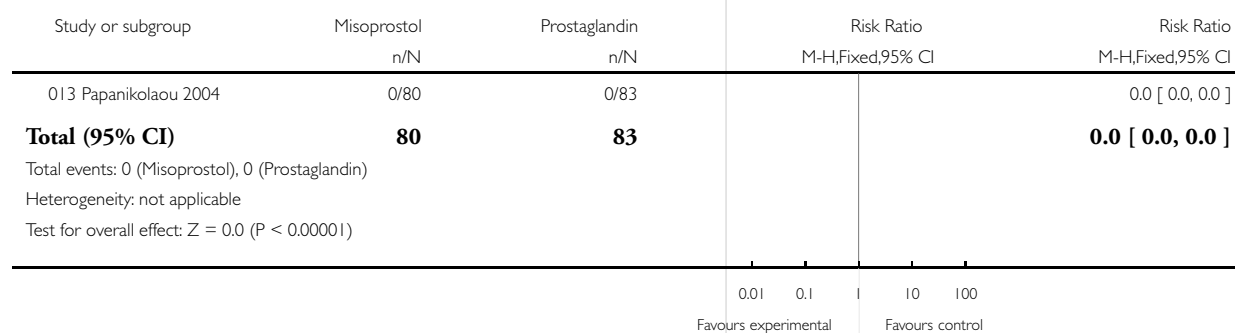


### Analysis 16.8. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 8 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 8 Serious neonatal morbidity

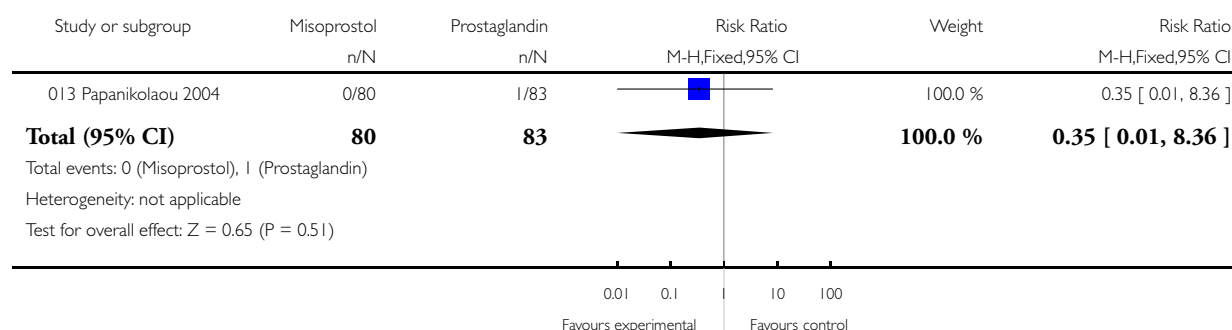


### Analysis 16.9. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 9 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 9 Perinatal death

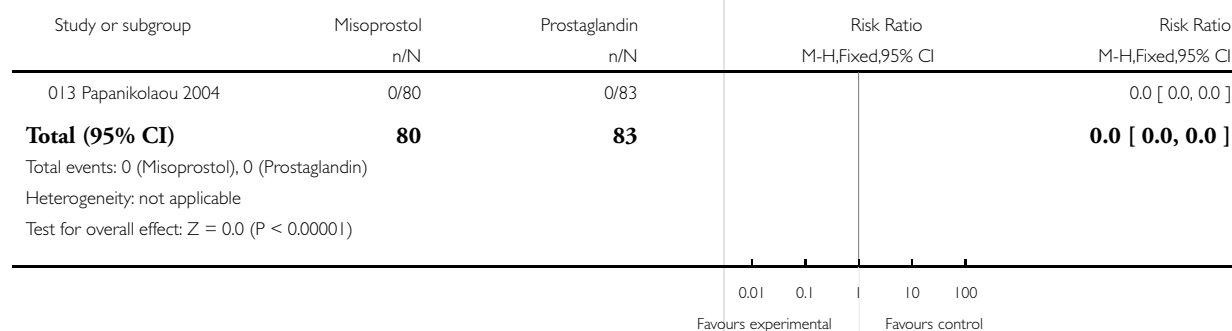


### Analysis 16.10. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 10 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 10 Serious maternal morbidity or death

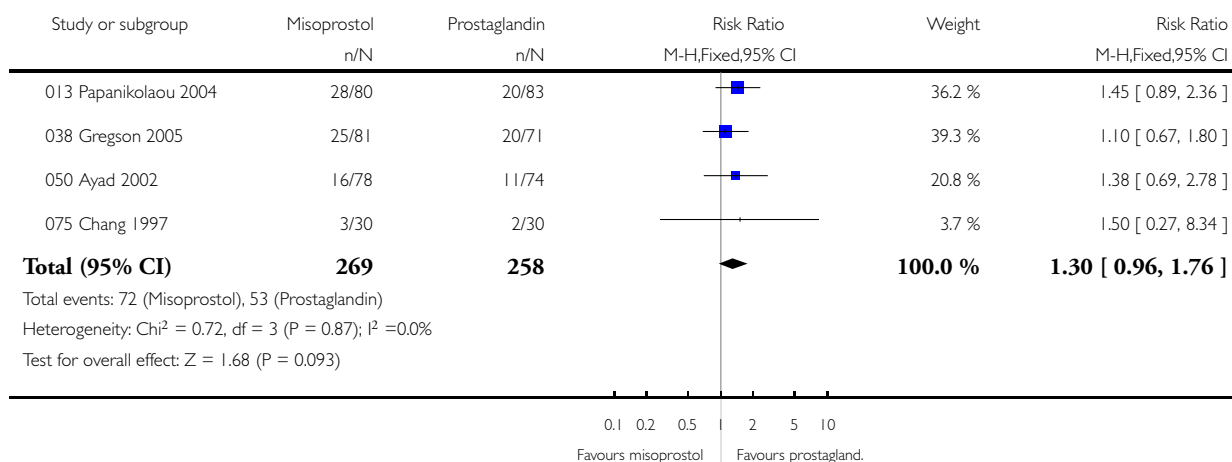


### Analysis 16.11. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 11 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 11 Instrumental vaginal delivery

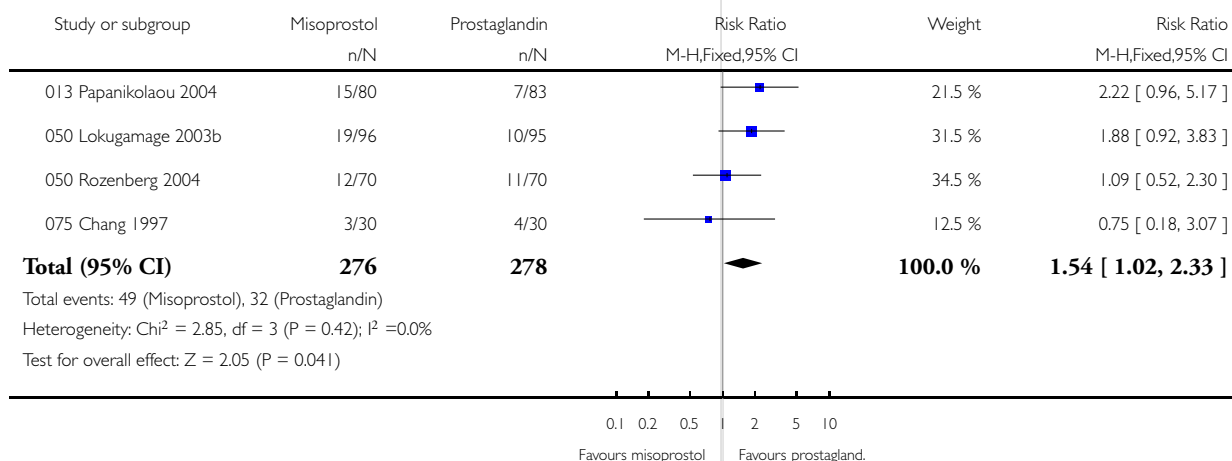


### Analysis 16.12. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 12 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 12 Meconium-stained liquor

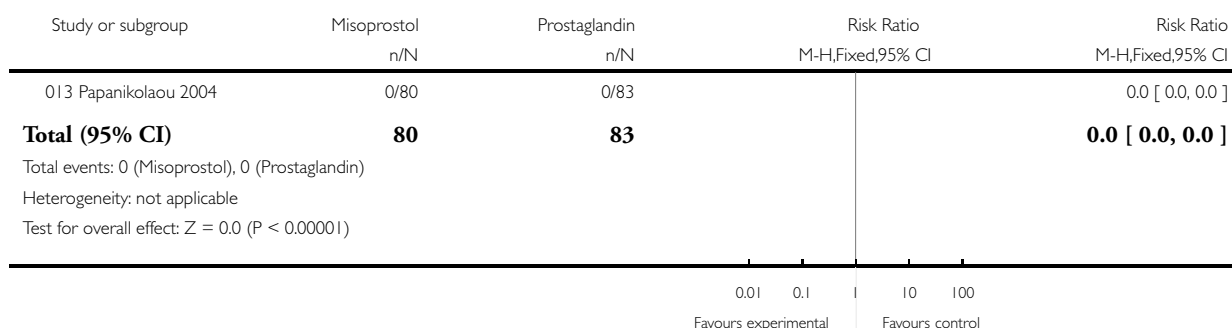


### Analysis 16.13. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 13 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 13 Uterine rupture

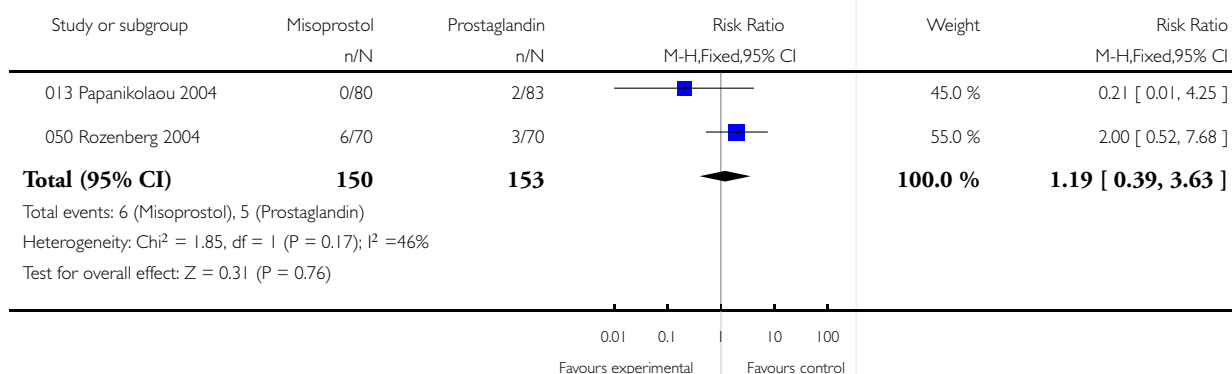


### Analysis 16.14. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 14 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 14 Postpartum haemorrhage

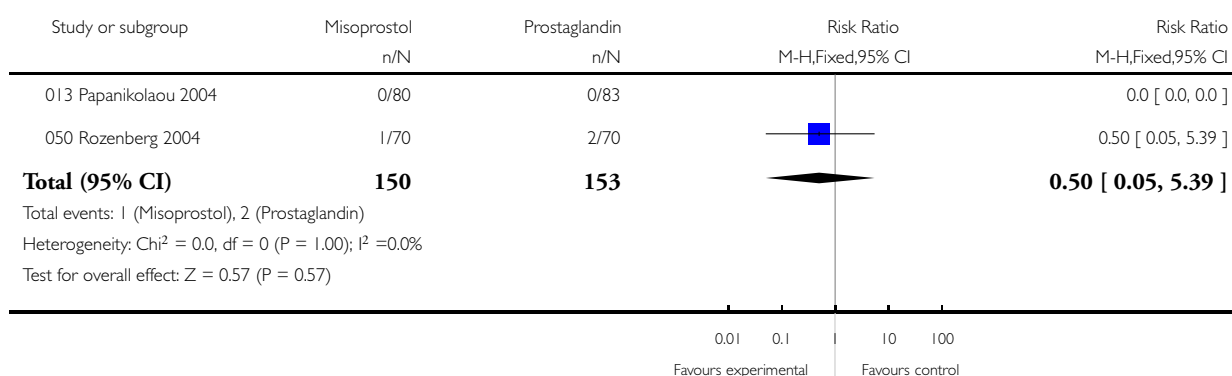


### Analysis 16.15. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 15 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 15 Serious maternal complications

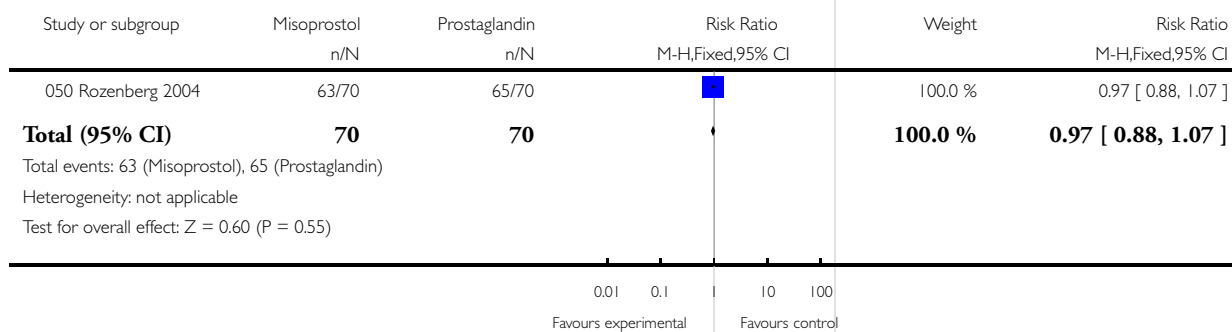


### Analysis 16.16. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 16 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 16 Epidural analgesia

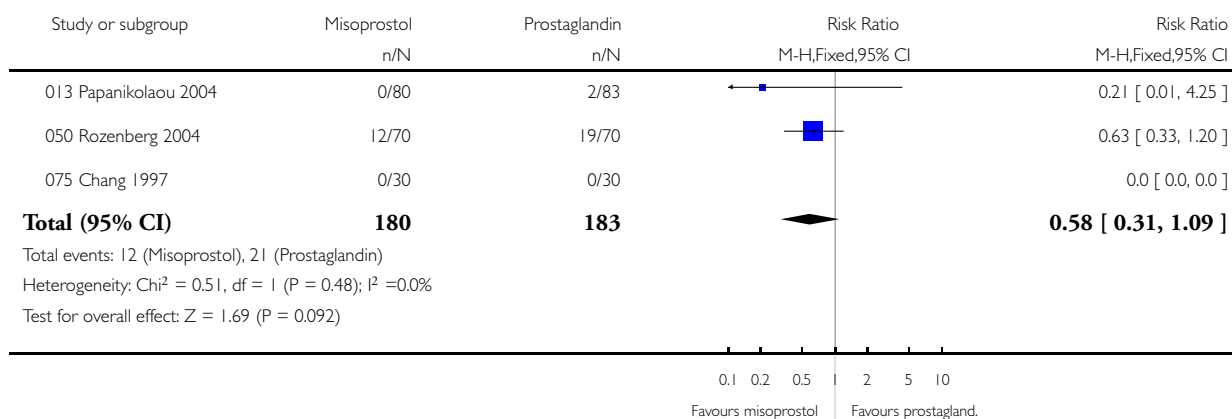


### Analysis 16.17. Comparison 16 Misoprostol versus vaginal prostaglandin: all primiparae, Outcome 17 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 16 Misoprostol versus vaginal prostaglandin: all primiparae

Outcome: 17 Maternal side effects (eg nausea, vomiting, diarrhoea)

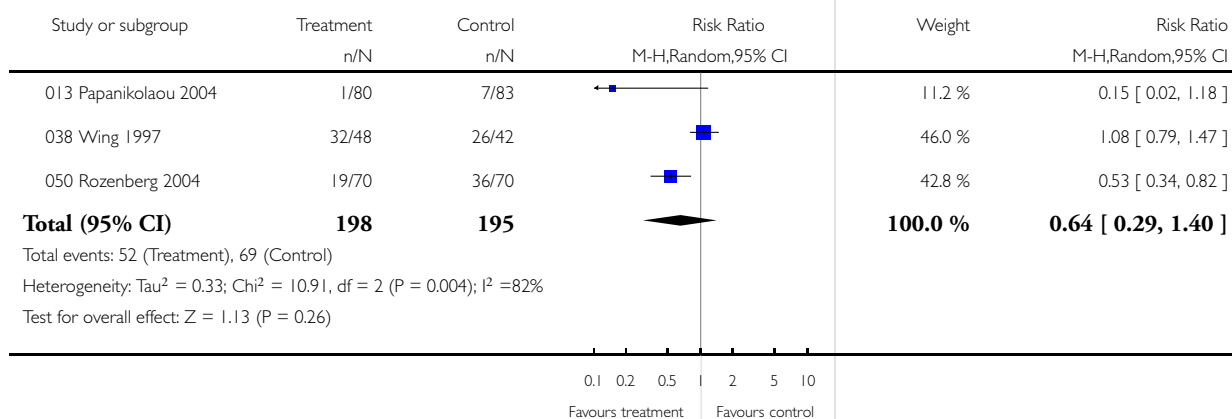


### Analysis 17.1. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

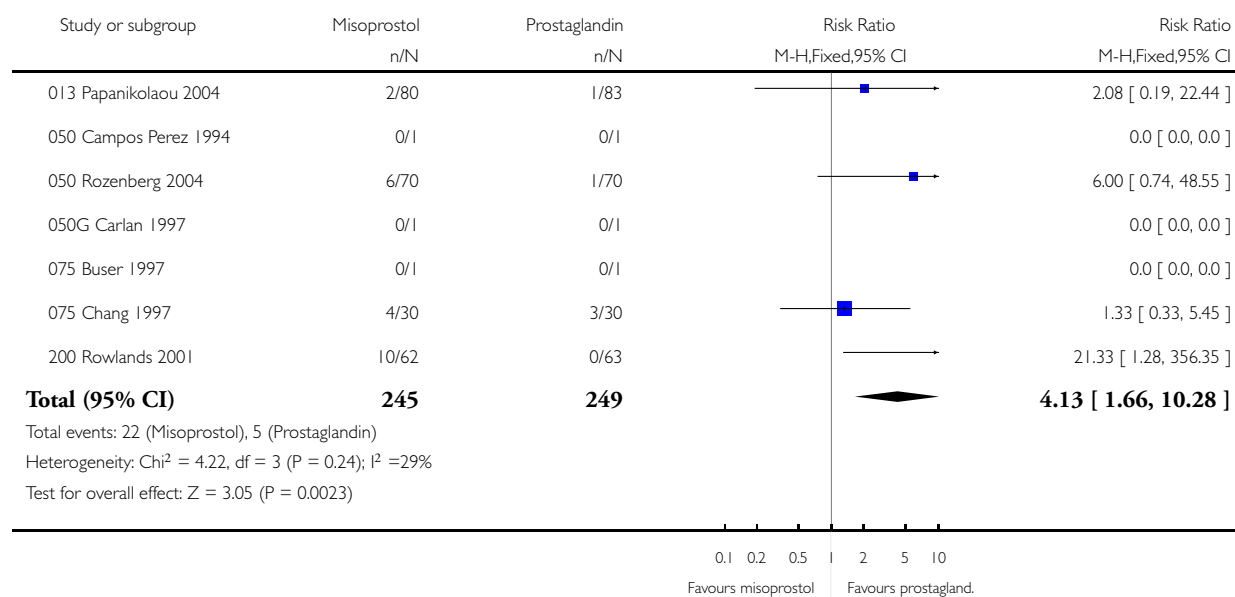


## Analysis 17.2. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes



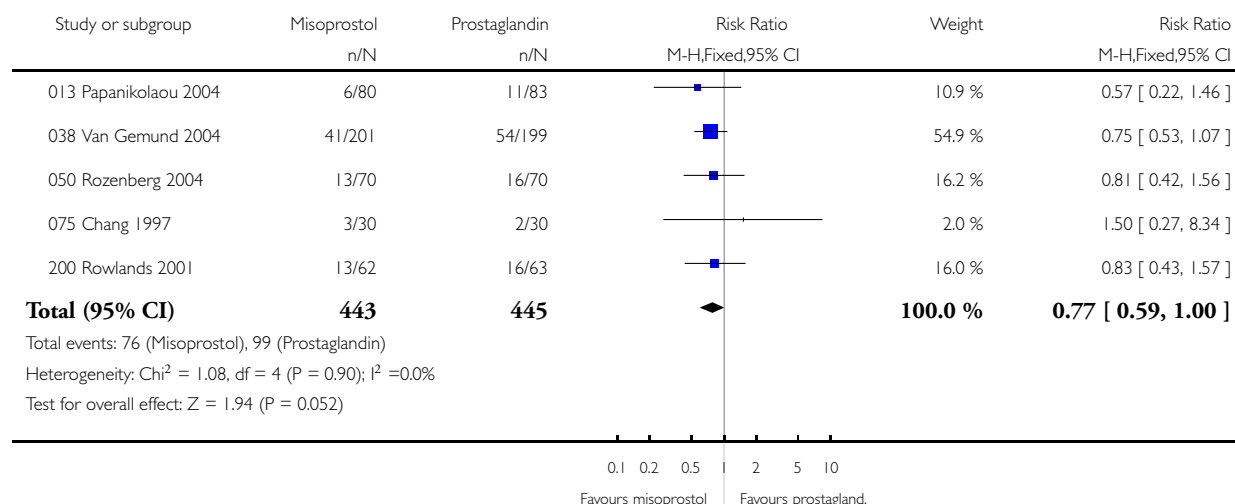


### Analysis 17.3. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 3 Caesarean section

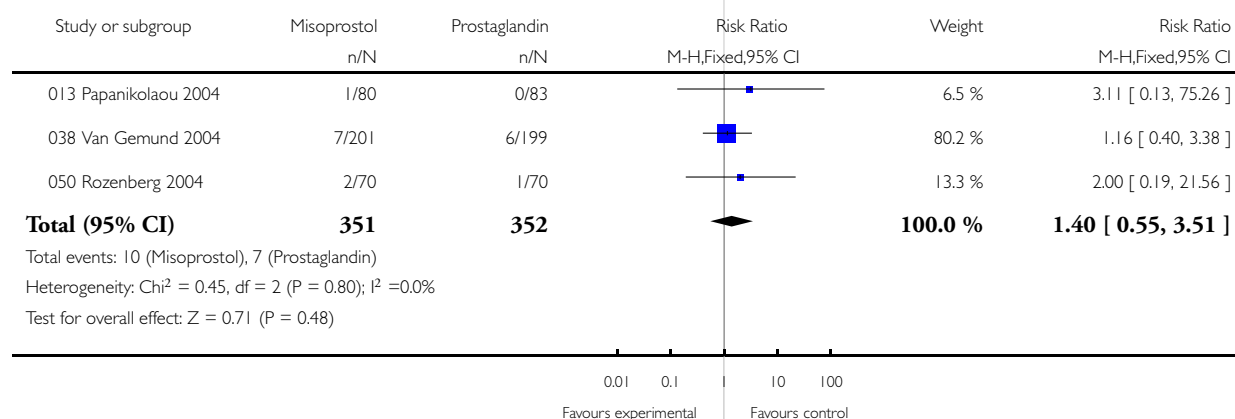


### Analysis 17.4. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 4 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 4 Apgar score < 7 at 5 minutes

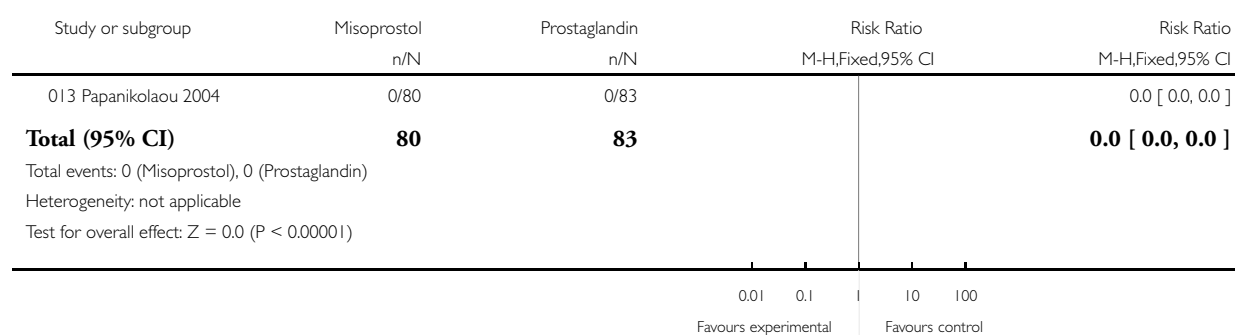


### Analysis 17.5. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 5 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 5 Serious neonatal morbidity

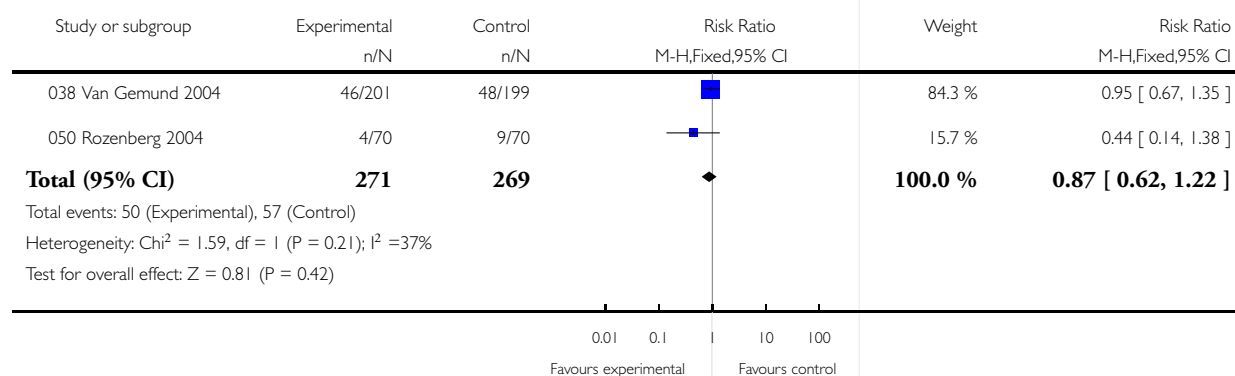


### Analysis 17.6. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 6 Neonatal intensive care admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 6 Neonatal intensive care admission

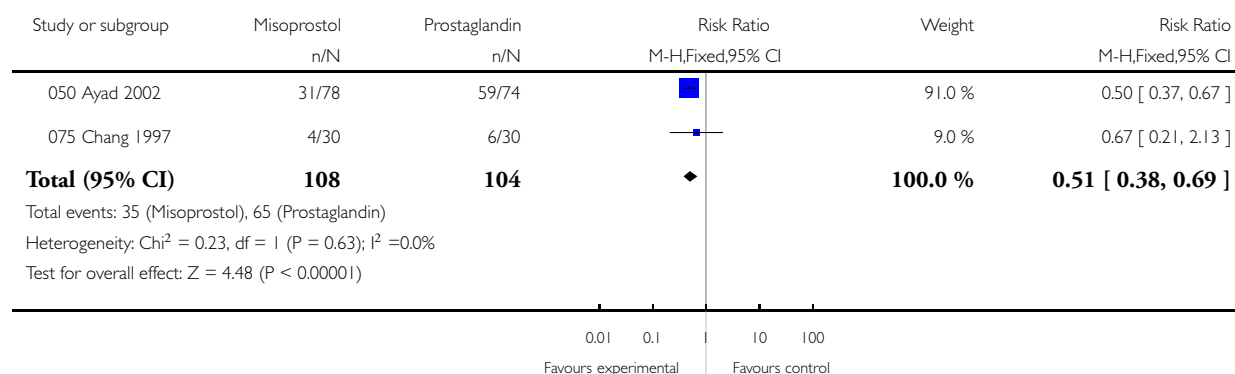


### Analysis 17.7. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 7 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 7 Oxytocin augmentation

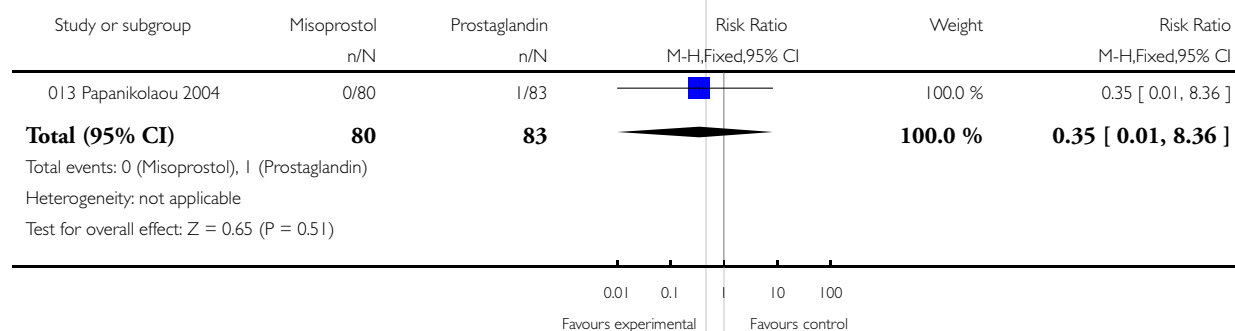


### Analysis 17.8. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 8 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 8 Perinatal death

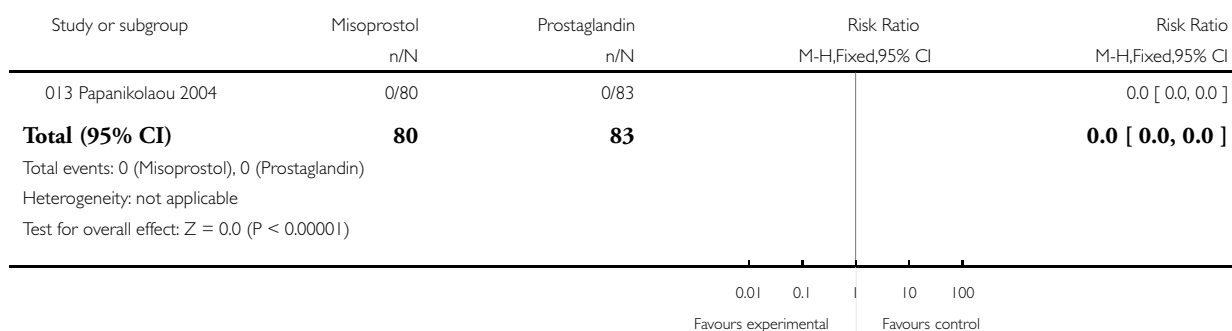


### Analysis 17.9. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 9 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 9 Serious maternal morbidity or death

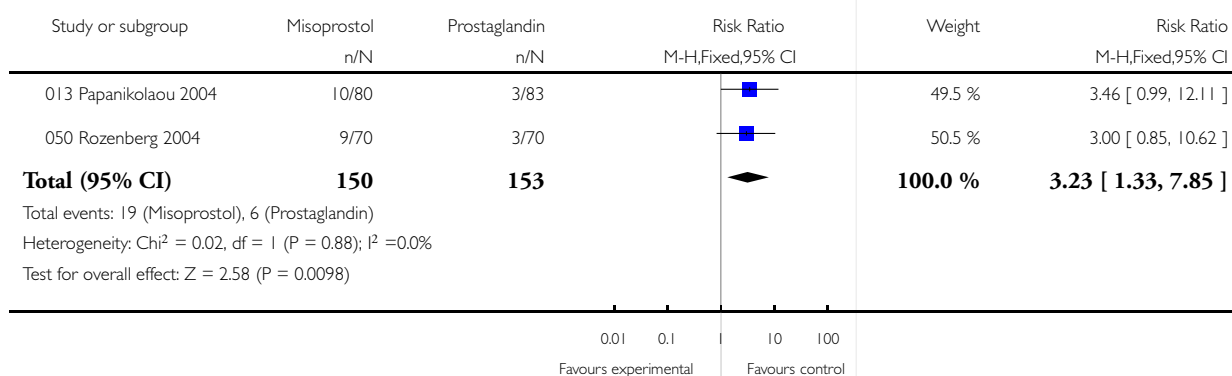


### Analysis 17.10. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 10 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 10 Uterine hyperstimulation without FHR changes

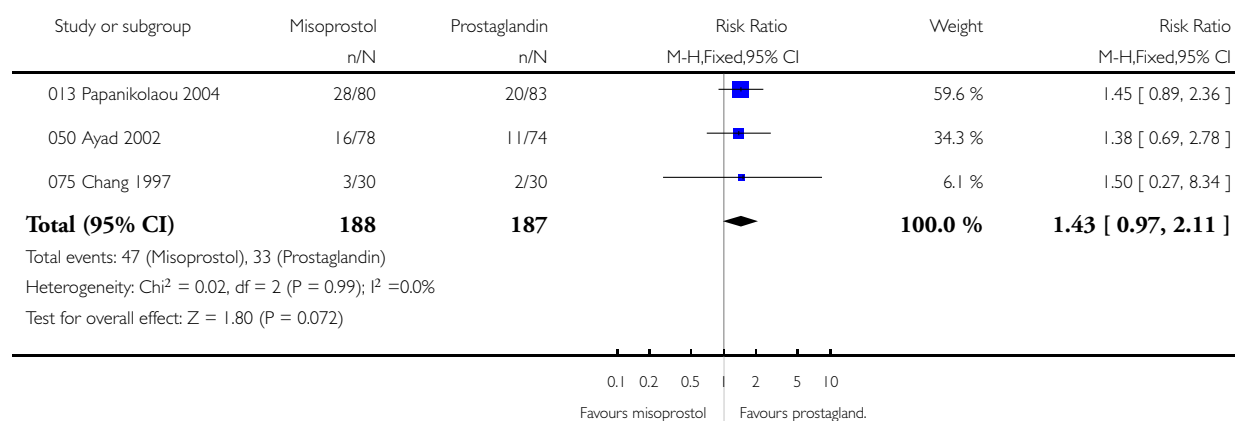


### Analysis 17.11. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 11 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 11 Instrumental vaginal delivery

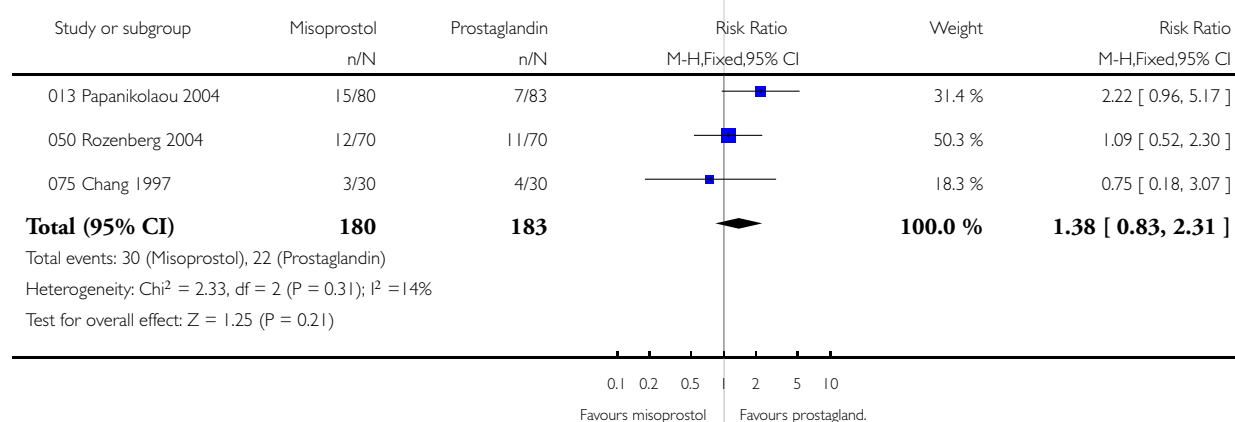


### Analysis 17.12. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 12 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 12 Meconium-stained liquor

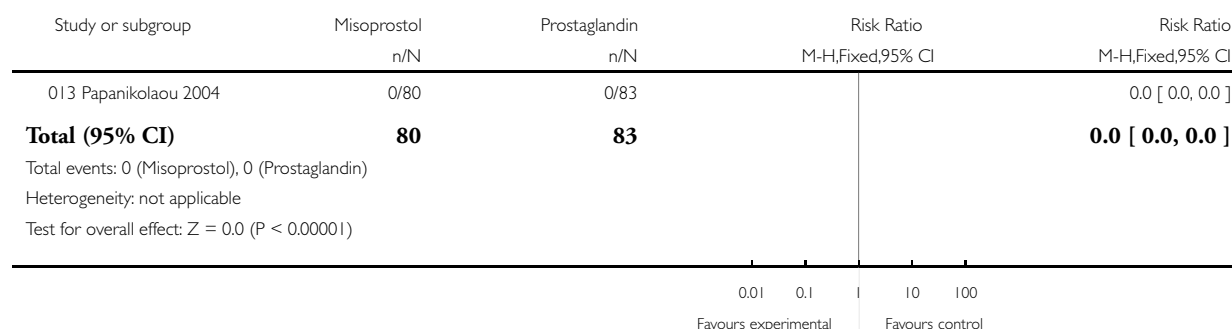


### Analysis 17.13. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 13 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 13 Uterine rupture

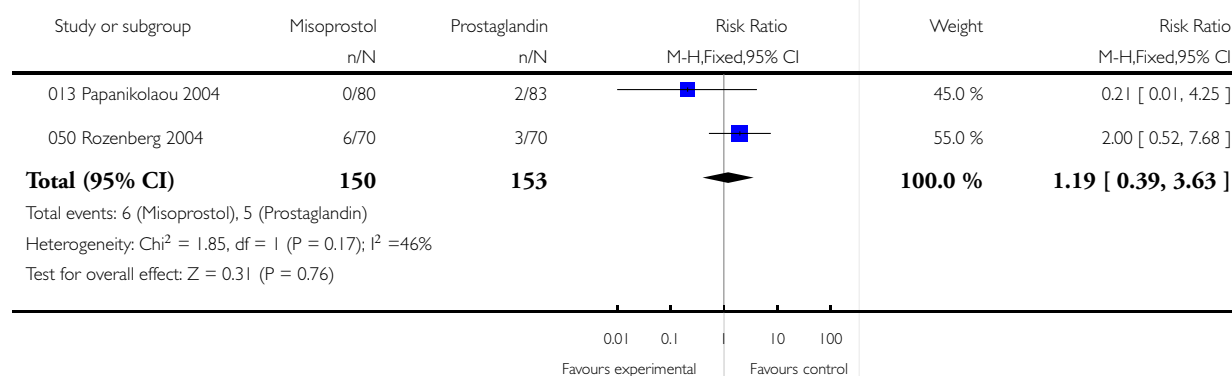


### Analysis 17.14. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 14 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 14 Postpartum haemorrhage

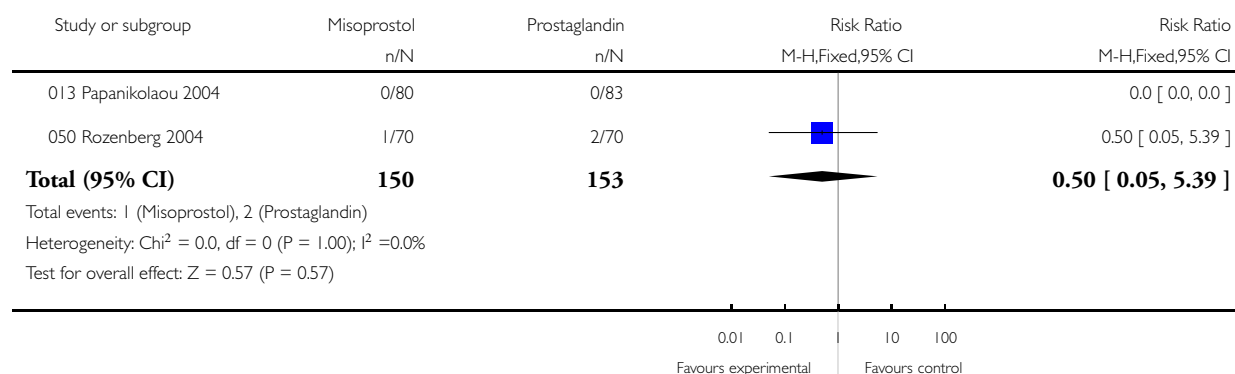


### Analysis 17.15. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 15 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 15 Serious maternal complications

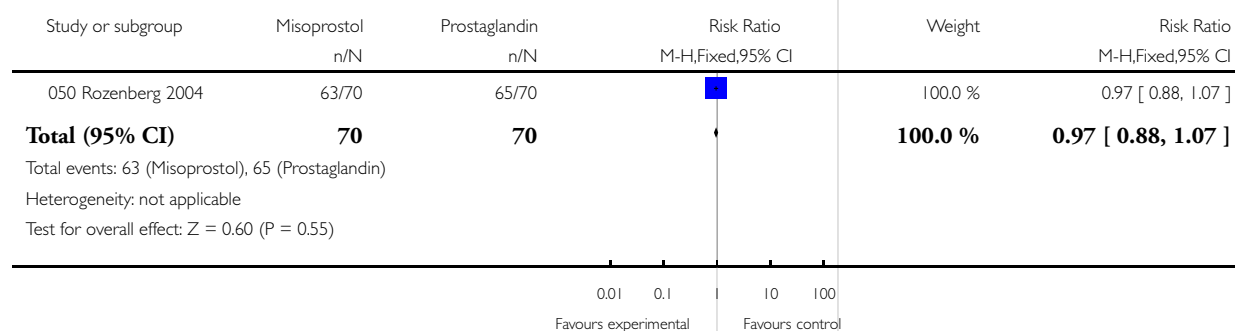


### Analysis 17.16. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 16 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 16 Epidural analgesia

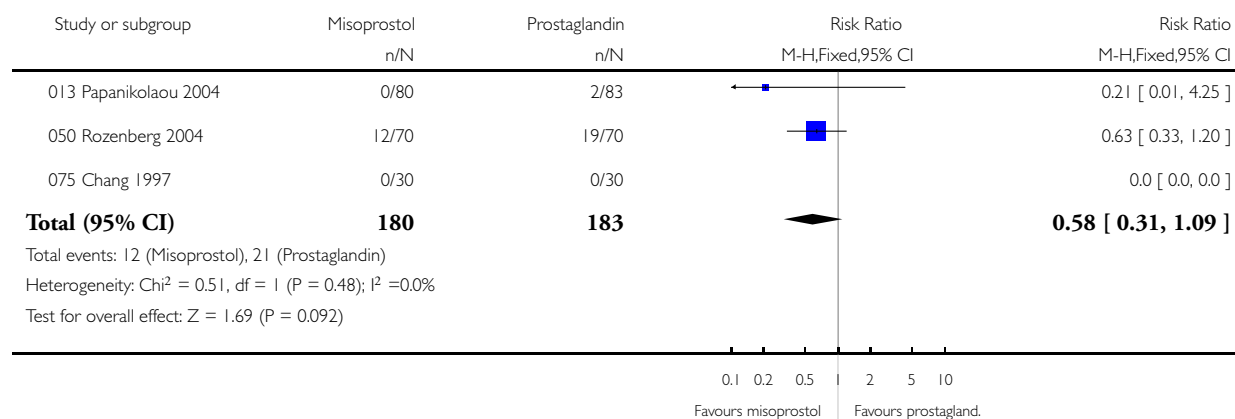


### Analysis 17.17. Comparison 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix, Outcome 17 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 17 Misoprostol versus vaginal prostaglandin: all primiparae, unfavourable cervix

Outcome: 17 Maternal side effects (eg nausea, vomiting, diarrhoea)

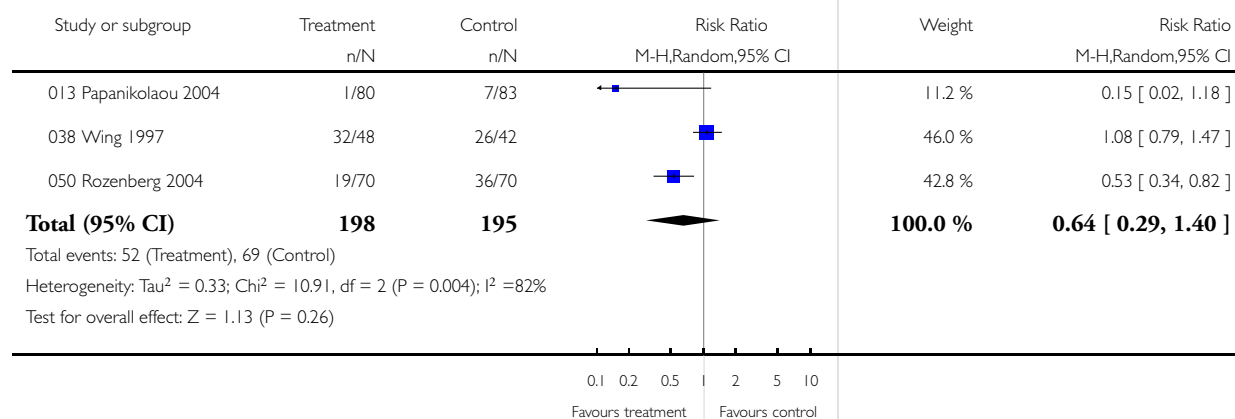


### Analysis 18.1. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours



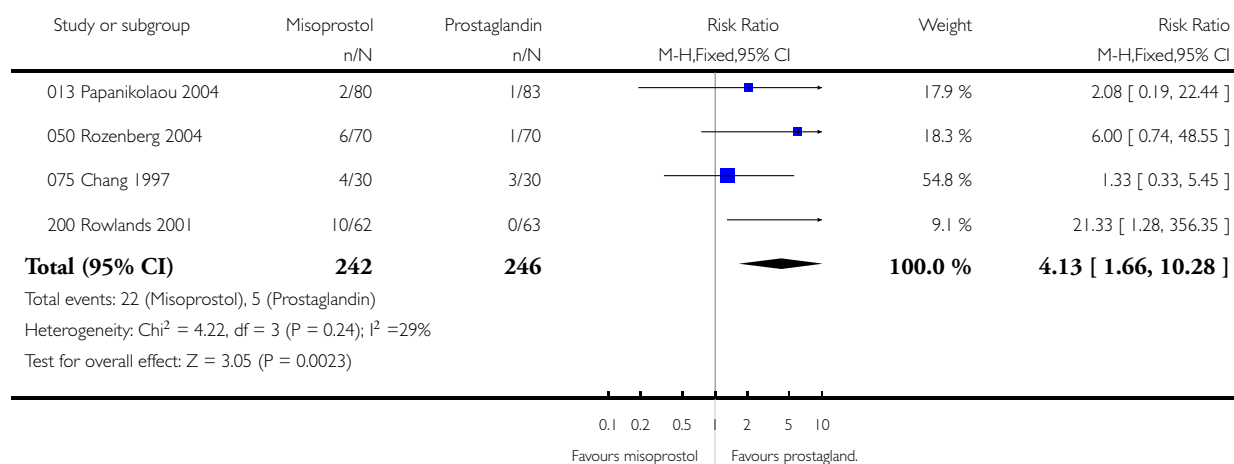


## Analysis 18.2. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

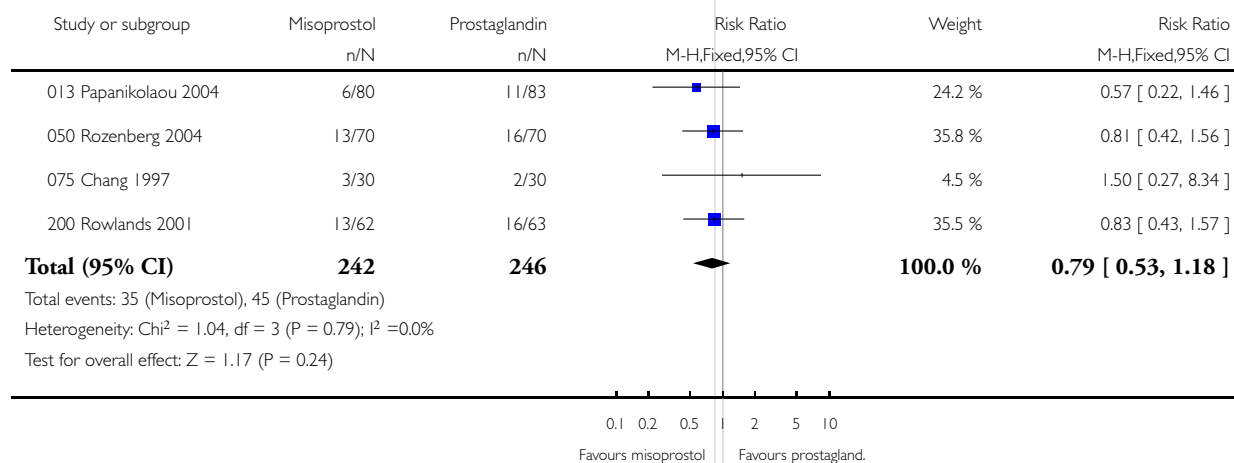


## Analysis 18.3. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 3 Caesarean section

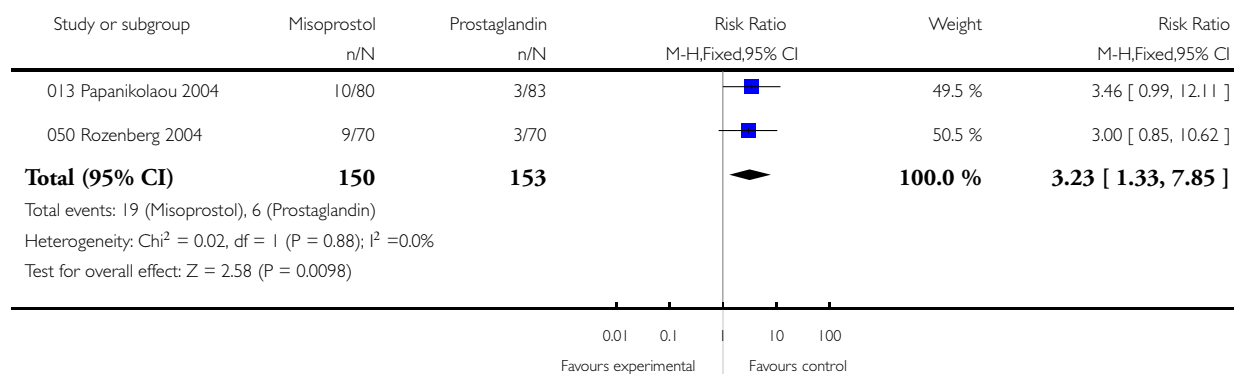


#### Analysis 18.4. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 4 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 4 Uterine hyperstimulation without FHR changes

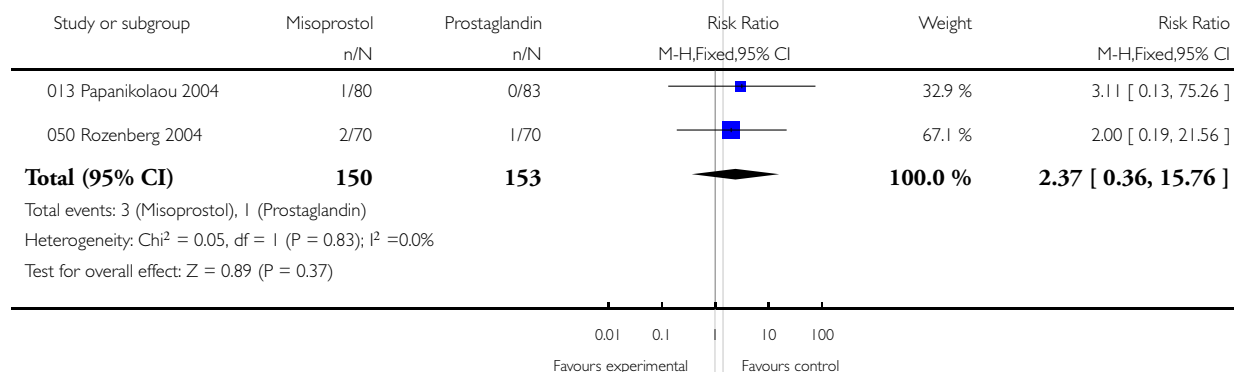


#### Analysis 18.5. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 5 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 5 Apgar score < 7 at 5 minutes

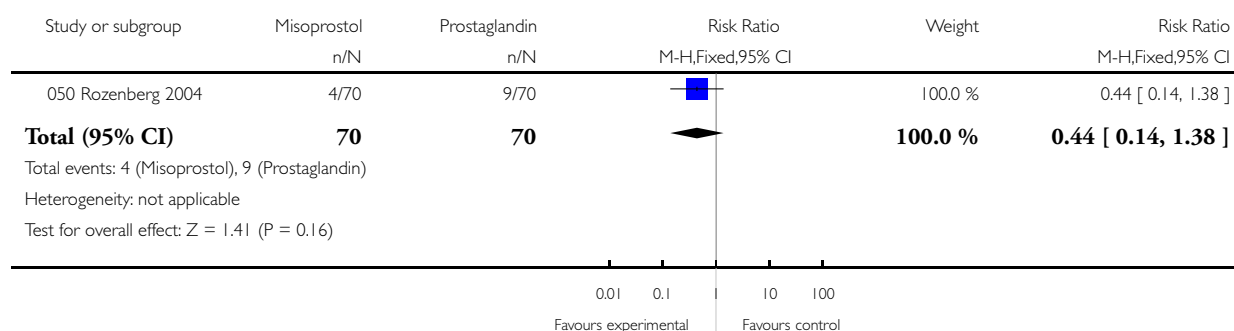


**Analysis 18.6. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 6 Neonatal intensive care unit admission.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 6 Neonatal intensive care unit admission

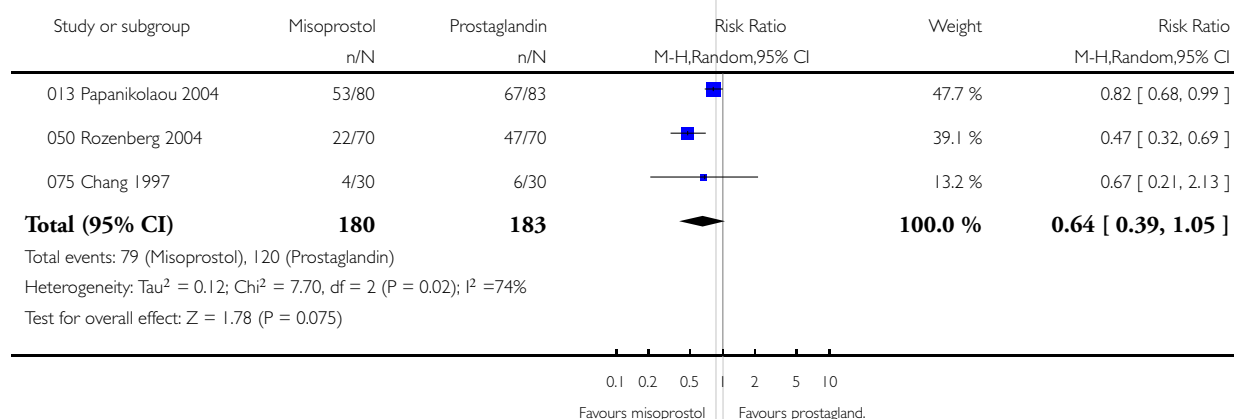


**Analysis 18.7. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 7 Oxytocin augmentation.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 7 Oxytocin augmentation

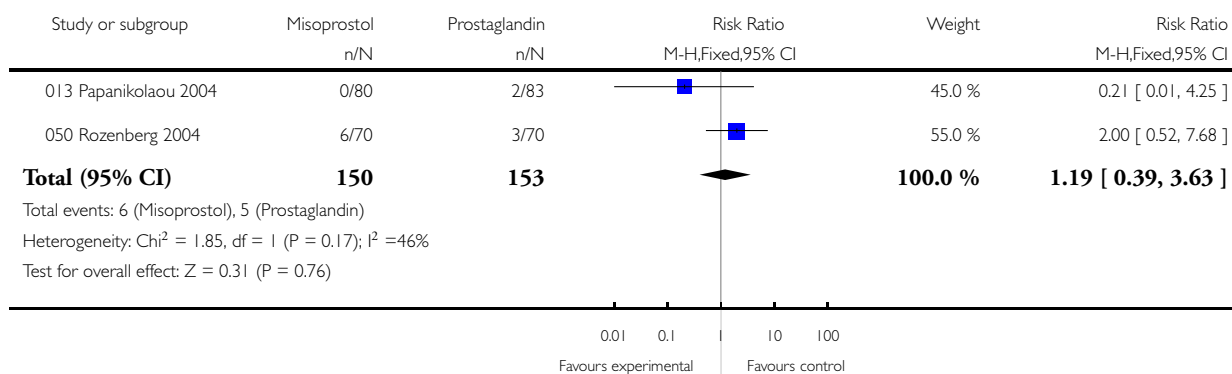


### Analysis 18.8. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 8 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 8 Postpartum haemorrhage

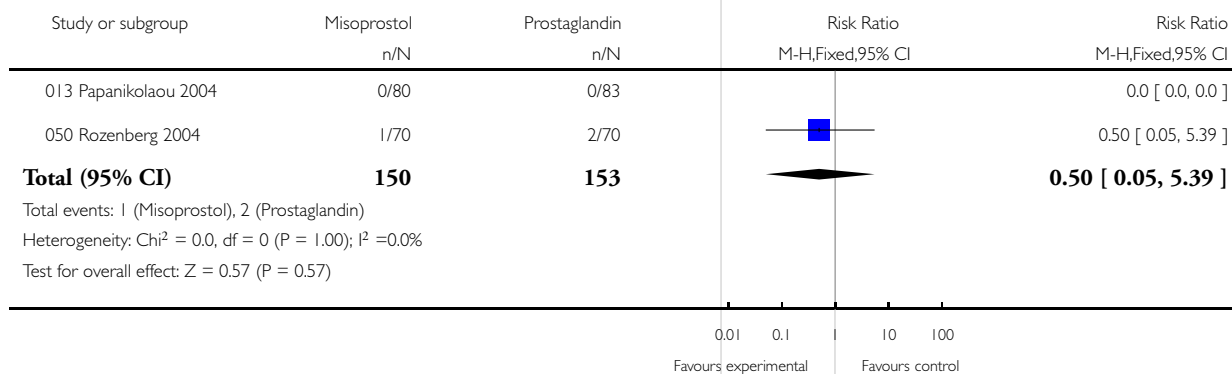


### Analysis 18.9. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 9 Serious maternal complication.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 9 Serious maternal complication

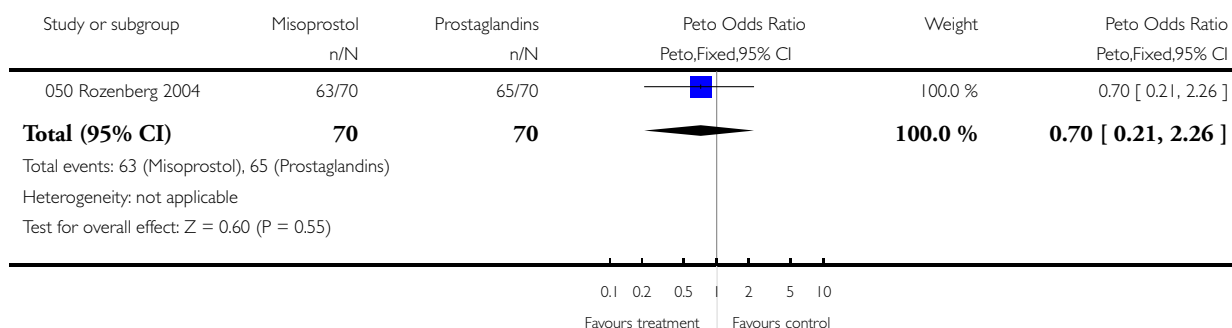


### Analysis 18.10. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 10 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 10 Epidural analgesia

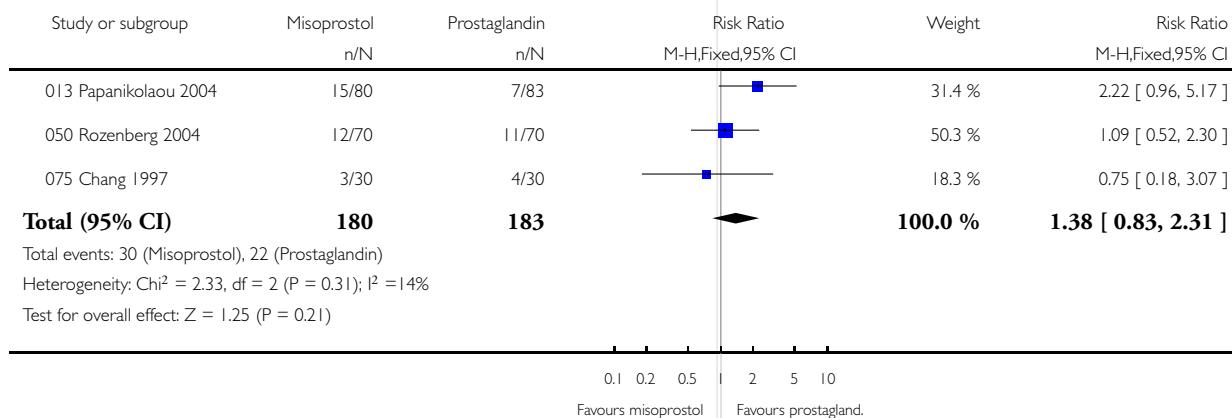


### Analysis 18.11. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 11 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 11 Meconium-stained liquor



### Analysis 18.12. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 12 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 12 Serious maternal morbidity or death

Study or subgroup	Misoprostol n/N	Prostaglandin n/N	Risk Ratio M-H,Fixed,95% CI	Risk Ratio M-H,Fixed,95% CI
013 Papanikolaou 2004	0/80	0/83		0.0 [ 0.0, 0.0 ]
<b>Total (95% CI)</b>	<b>80</b>	<b>83</b>		<b>0.0 [ 0.0, 0.0 ]</b>
Total events: 0 (Misoprostol), 0 (Prostaglandin)				
Heterogeneity: not applicable				
Test for overall effect: Z = 0.0 (P < 0.00001)				
			0.01 0.1	10 100
			Favours experimental	Favours control

### Analysis 18.13. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 13 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 13 Serious neonatal morbidity

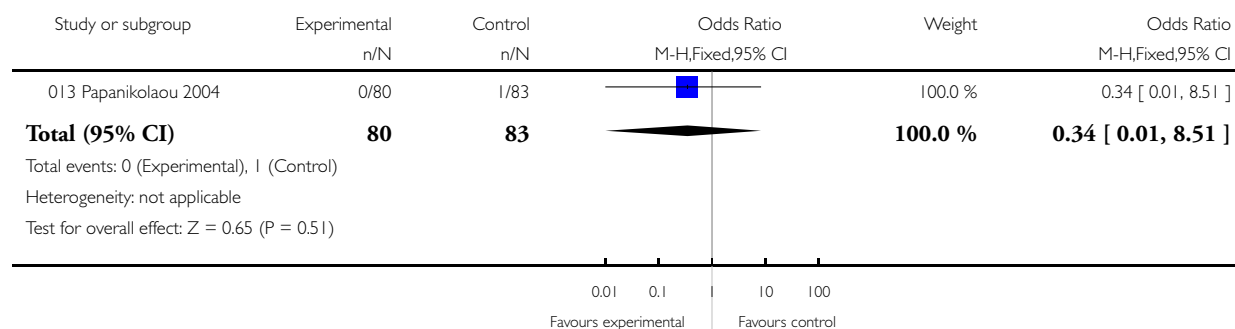
Study or subgroup	Misoprostol n/N	Prostaglandin n/N	Risk Ratio M-H,Fixed,95% CI	Risk Ratio M-H,Fixed,95% CI
013 Papanikolaou 2004	0/80	0/83		0.0 [ 0.0, 0.0 ]
<b>Total (95% CI)</b>	<b>80</b>	<b>83</b>		<b>0.0 [ 0.0, 0.0 ]</b>
Total events: 0 (Misoprostol), 0 (Prostaglandin)				
Heterogeneity: not applicable				
Test for overall effect: Z = 0.0 (P < 0.00001)				
			0.01 0.1	10 100
			Favours experimental	Favours control

### Analysis 18.14. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 14 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 14 Perinatal death

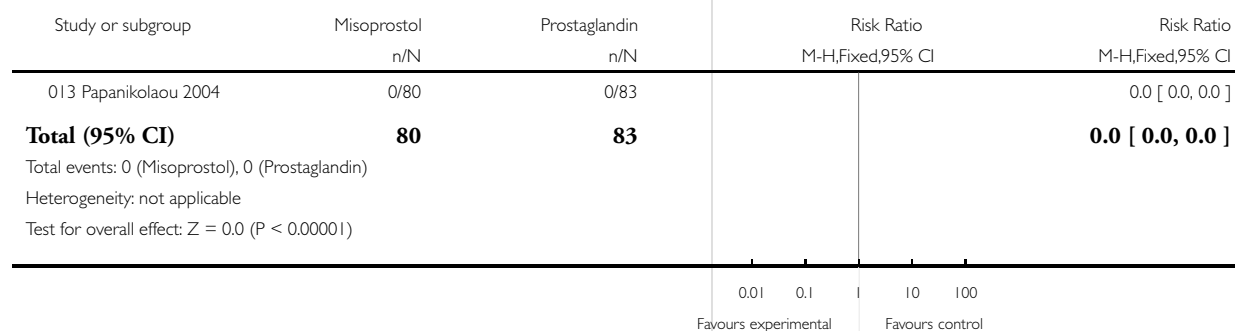


### Analysis 18.15. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 15 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 15 Uterine rupture

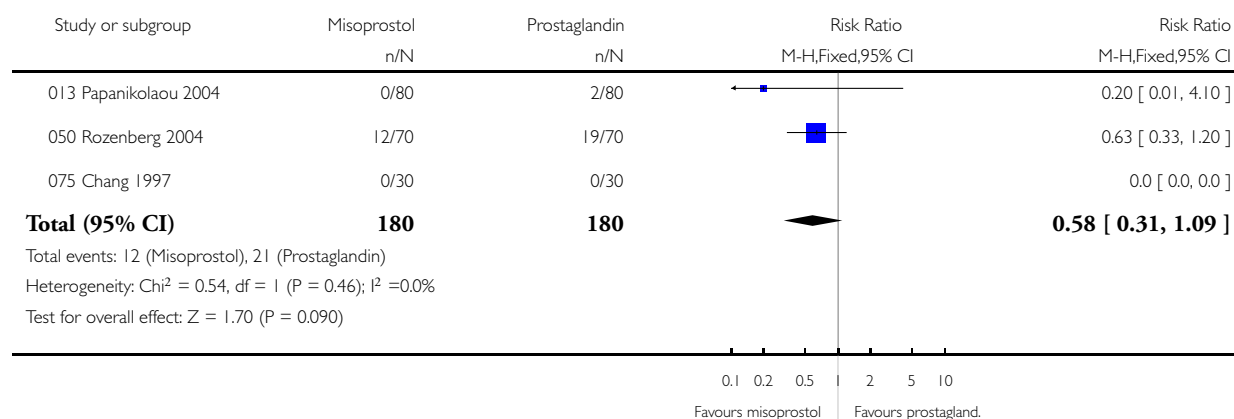


### Analysis 18.16. Comparison 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix, Outcome 16 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 18 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, unfavourable cervix

Outcome: 16 Maternal side effects (eg nausea, vomiting, diarrhoea)

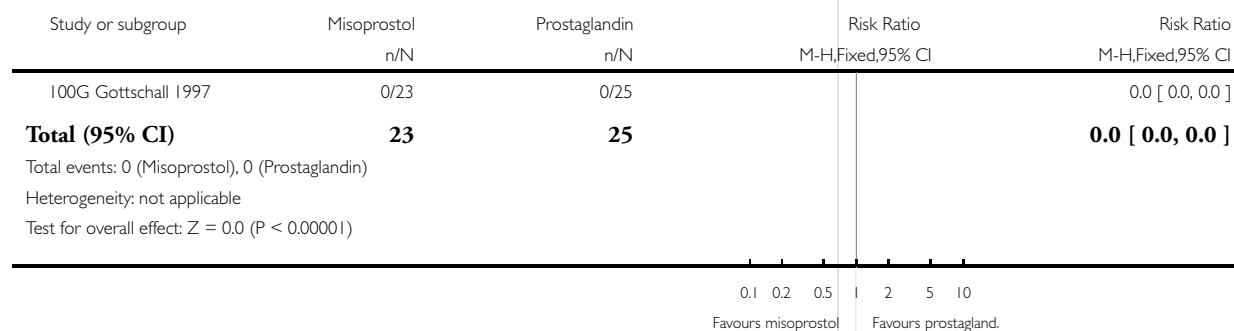


### Analysis 19.1. Comparison 19 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, variable or undefined cervix, Outcome 1 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 19 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, variable or undefined cervix

Outcome: 1 Uterine hyperstimulation with FHR changes



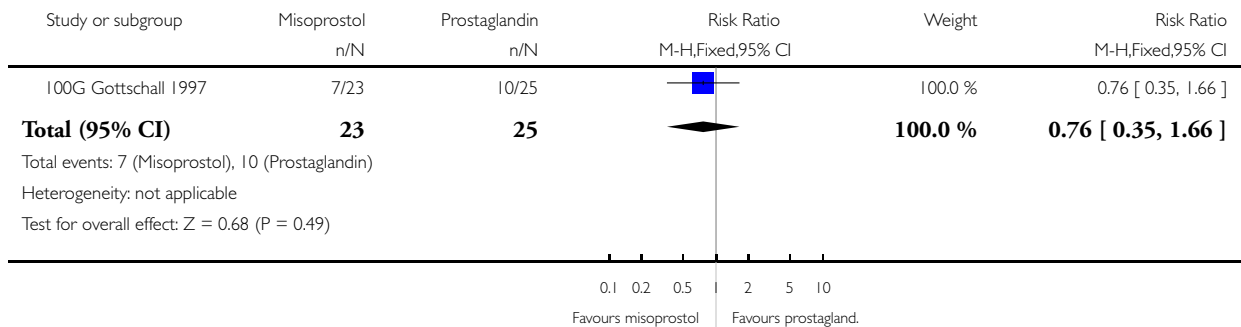


## Analysis 19.2. Comparison 19 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, variable or undefined cervix, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 19 Misoprostol versus vaginal prostaglandin: all primiparae, intact membranes, variable or undefined cervix

Outcome: 2 Caesarean section

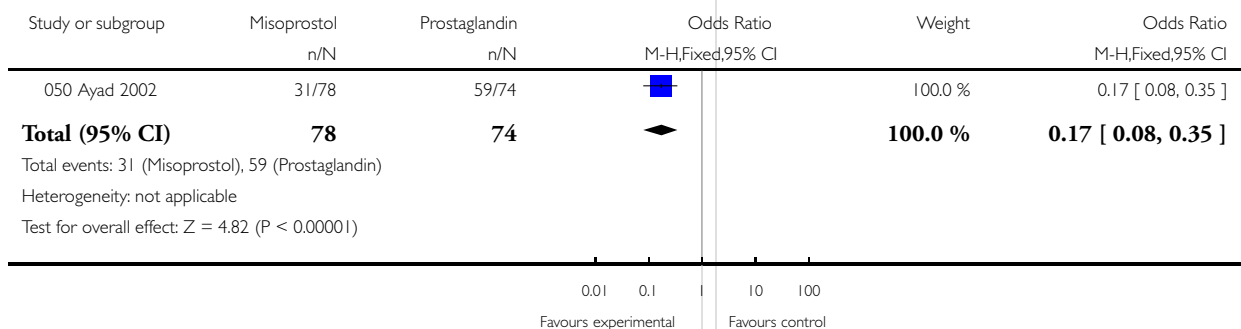


## Analysis 20.1. Comparison 20 Misoprostol versus vaginal prostaglandin: all primiparae with ruptured membranes and unfavourable cervix, Outcome 1 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 20 Misoprostol versus vaginal prostaglandin: all primiparae with ruptured membranes and unfavourable cervix

Outcome: 1 Oxytocin augmentation

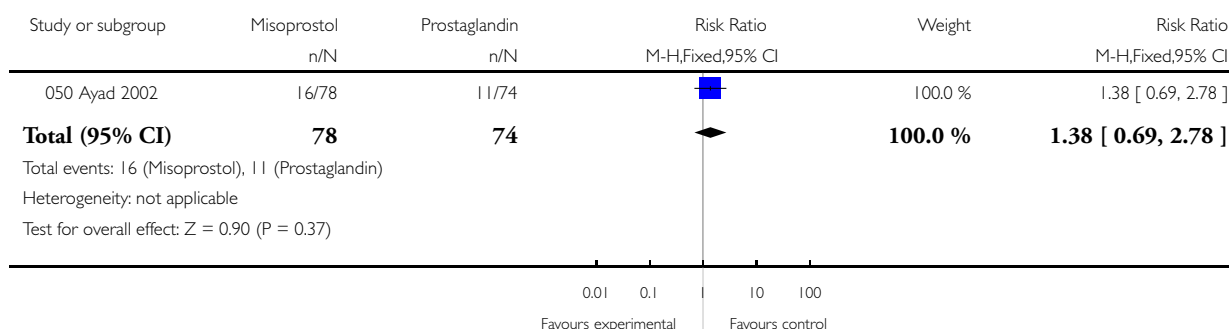


## Analysis 20.2. Comparison 20 Misoprostol versus vaginal prostaglandin: all primiparae with ruptured membranes and unfavourable cervix, Outcome 2 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 20 Misoprostol versus vaginal prostaglandin: all primiparae with ruptured membranes and unfavourable cervix

Outcome: 2 Instrumental vaginal delivery

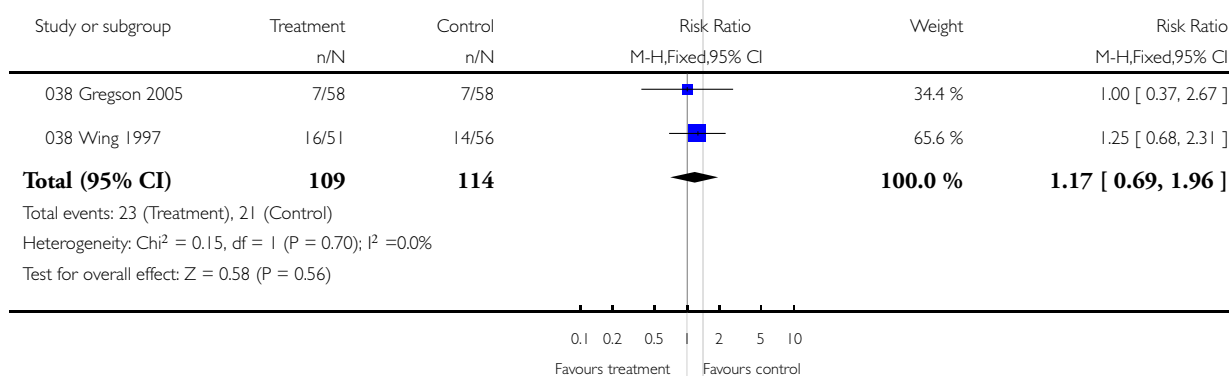


## Analysis 21.1. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 1 Vaginal delivery not achieved within 24 hours

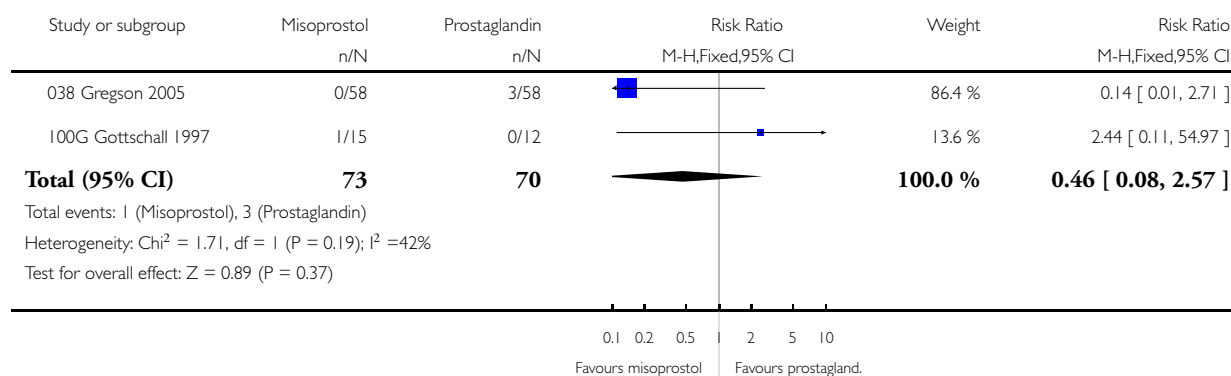


## Analysis 21.2. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 2 Uterine hyperstimulation with FHR changes

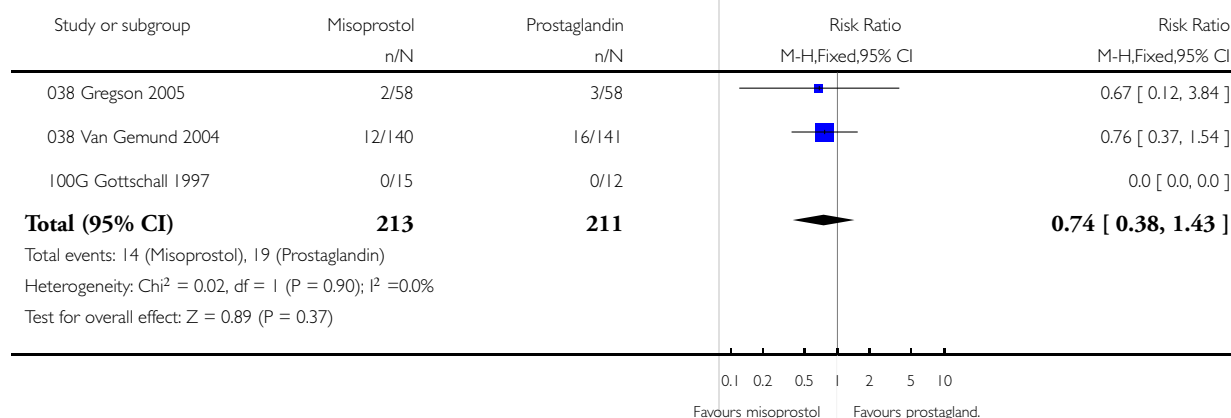


## Analysis 21.3. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 3 Caesarean section

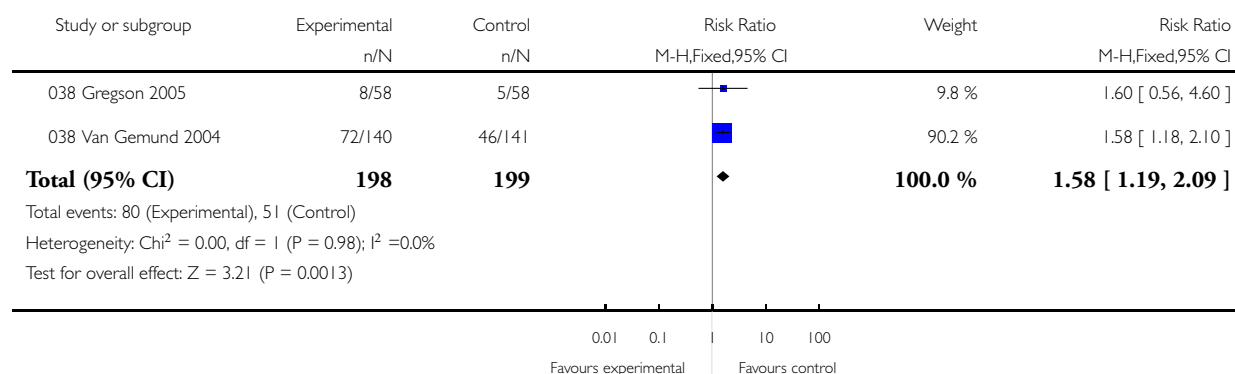


#### Analysis 21.4. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 4 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 4 Oxytocin augmentation

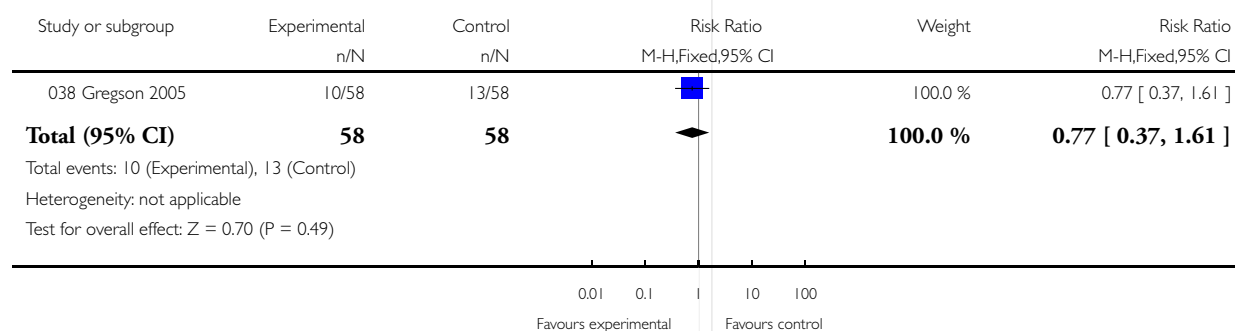


#### Analysis 21.5. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 5 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 5 Uterine hyperstimulation without FHR changes

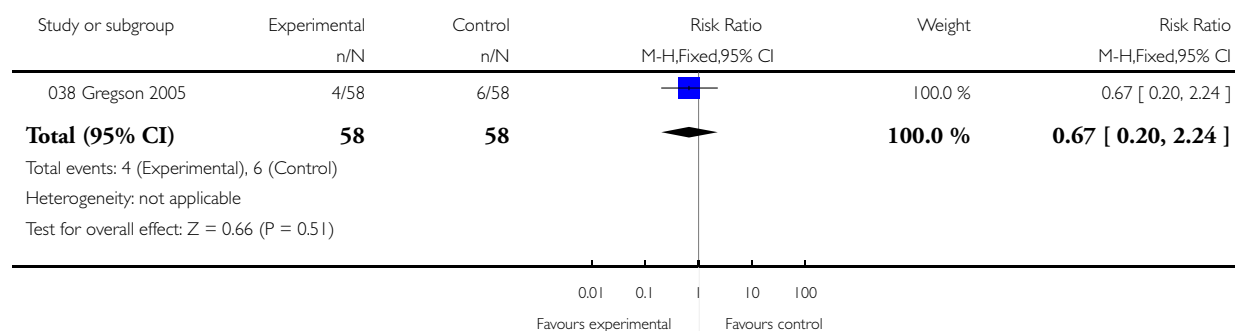


### Analysis 21.6. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 6 Instrumental delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 6 Instrumental delivery

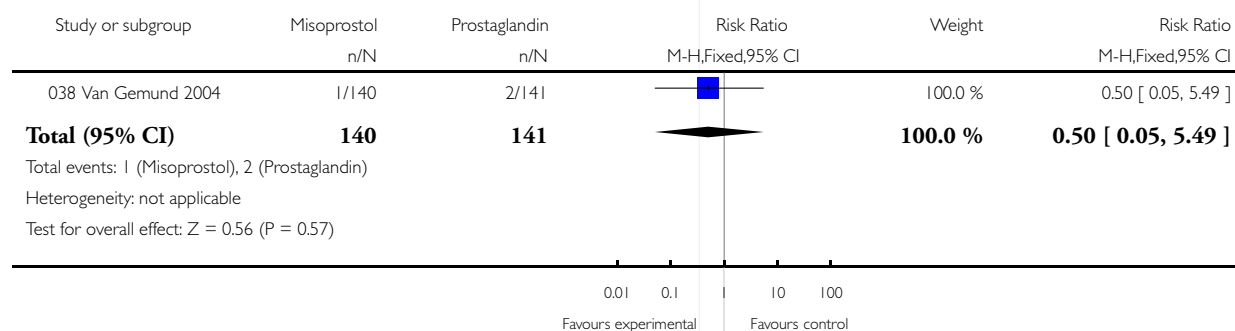


### Analysis 21.7. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 7 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 7 Apgar score < 7 at 5 minutes

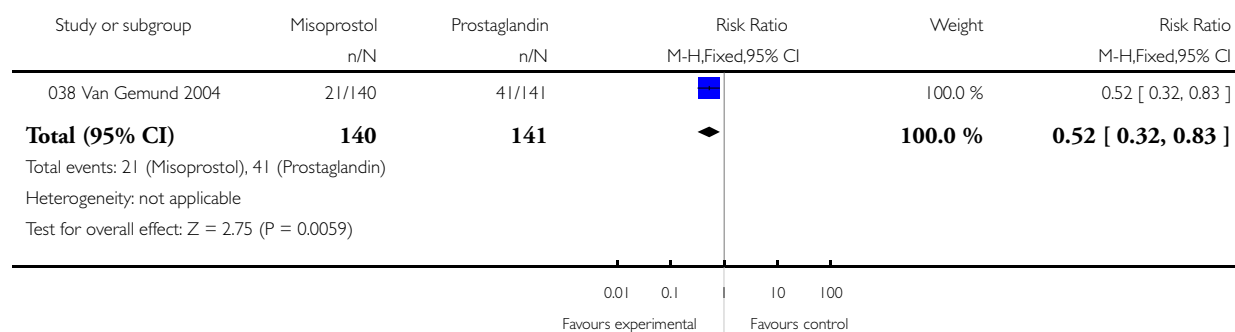


### Analysis 21.8. Comparison 21 Misoprostol versus vaginal prostaglandin: all multiparae, Outcome 8 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 21 Misoprostol versus vaginal prostaglandin: all multiparae

Outcome: 8 Neonatal intensive care unit admission

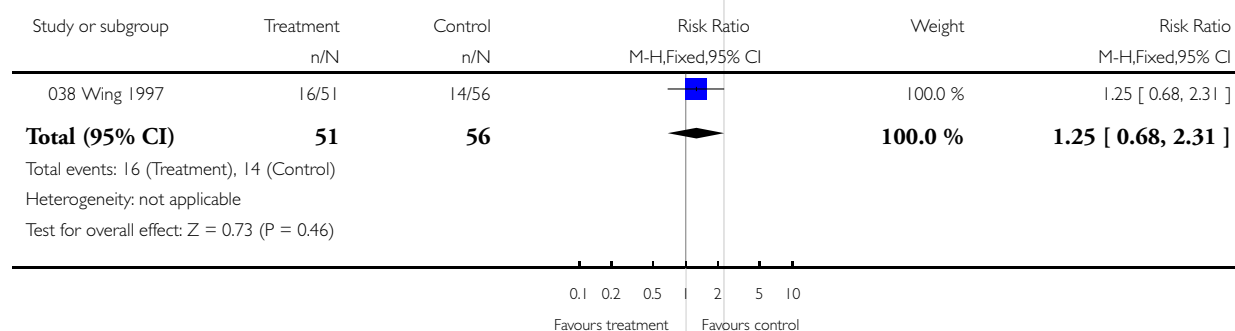


### Analysis 22.1. Comparison 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

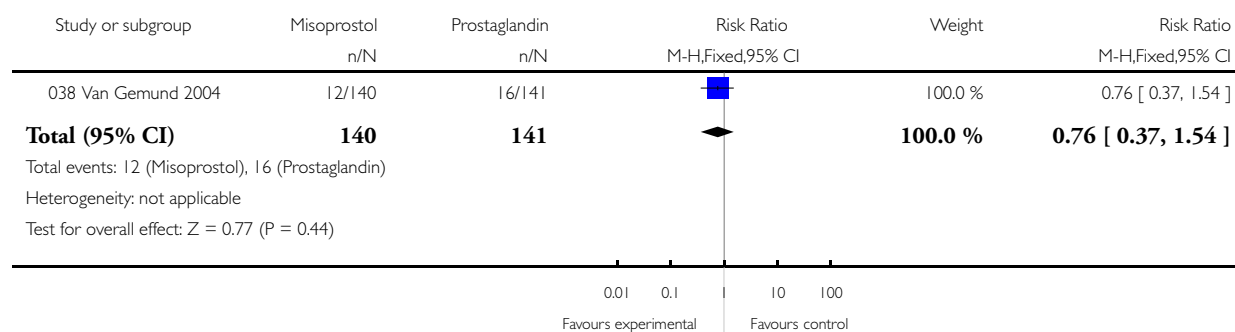


## Analysis 22.2. Comparison 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix

Outcome: 2 Caesarean section

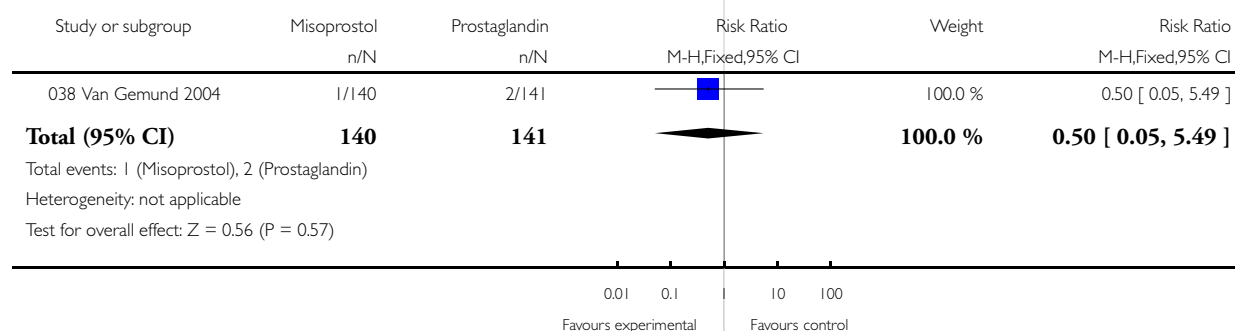


## Analysis 22.3. Comparison 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix, Outcome 3 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix

Outcome: 3 Apgar score < 7 at 5 minutes

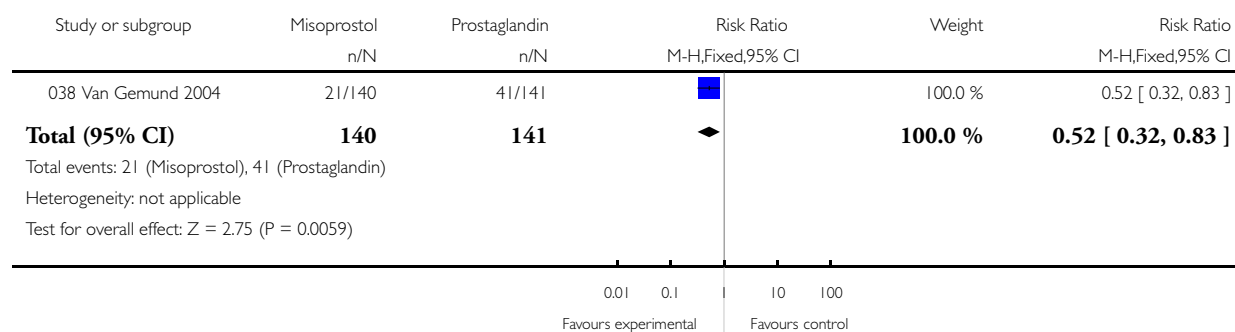


#### Analysis 22.4. Comparison 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix, Outcome 4 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix

Outcome: 4 Neonatal intensive care unit admission

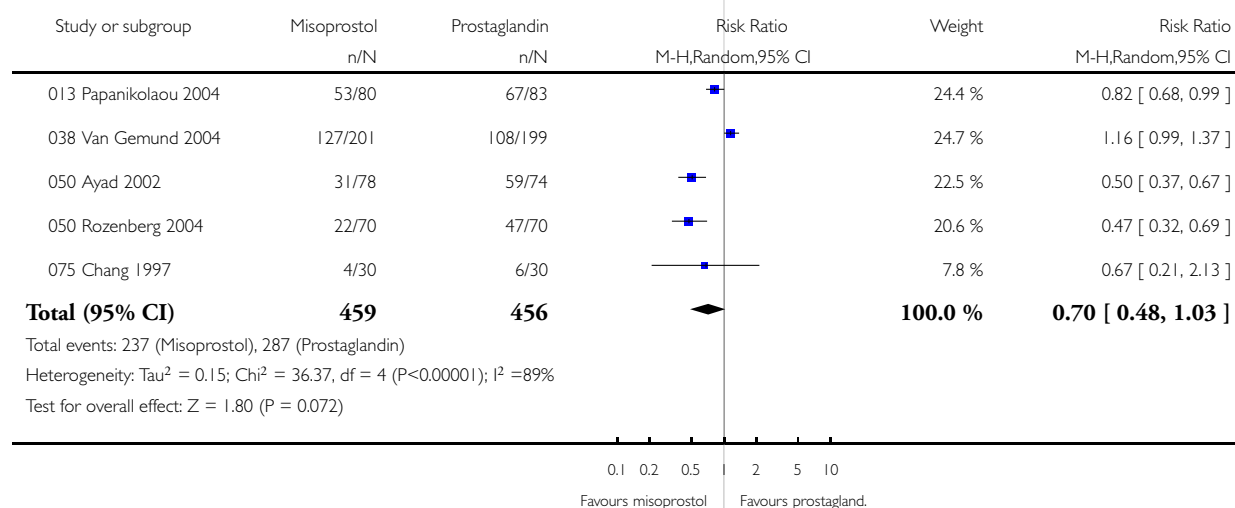


#### Analysis 22.5. Comparison 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix, Outcome 5 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 22 Misoprostol versus vaginal prostaglandin: all multiparae, unfavourable cervix

Outcome: 5 Oxytocin augmentation



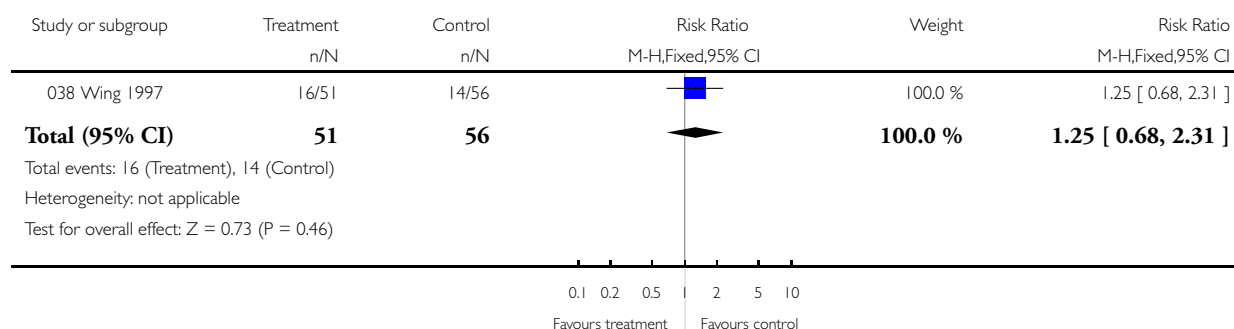


### Analysis 23.1. Comparison 23 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, unfavourable cervix, Outcome 1 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 23 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, unfavourable cervix

Outcome: 1 Vaginal delivery not achieved in 24 hours

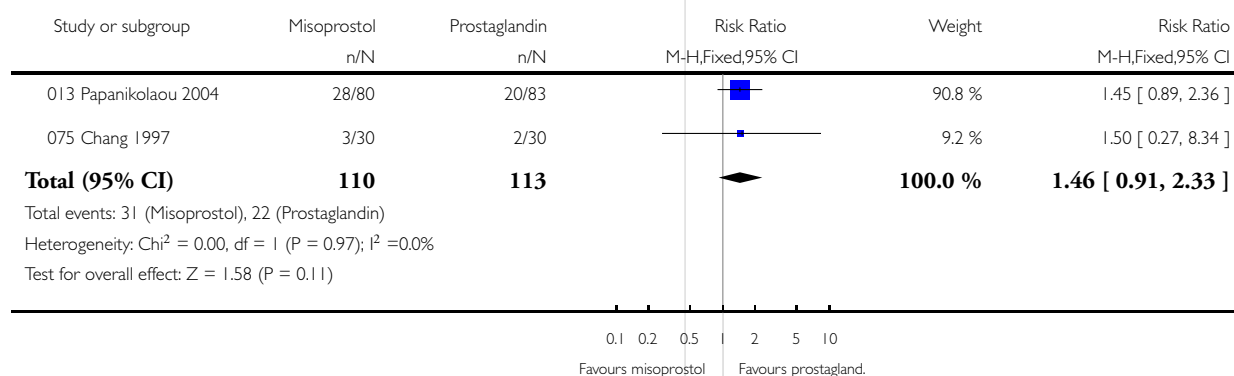


### Analysis 23.2. Comparison 23 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, unfavourable cervix, Outcome 2 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 23 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, unfavourable cervix

Outcome: 2 Instrumental vaginal delivery

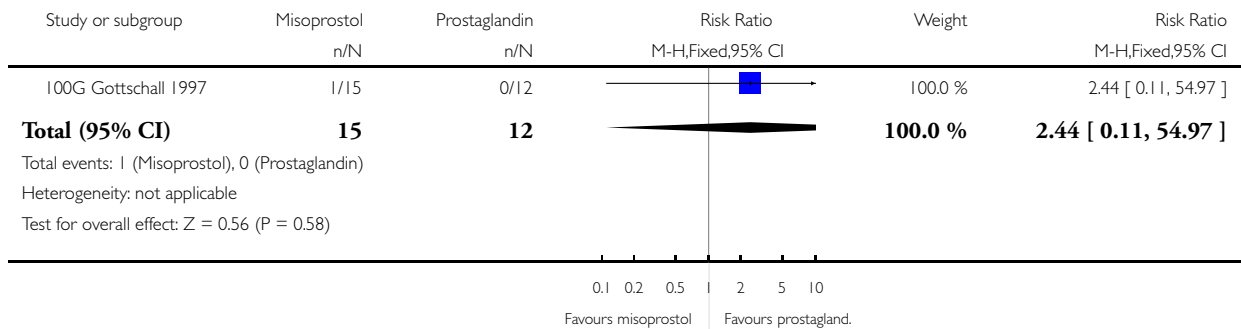


**Analysis 24.1. Comparison 24 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, variable or undefined cervix, Outcome 1 Uterine hyperstimulation with FHR changes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 24 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, variable or undefined cervix

Outcome: 1 Uterine hyperstimulation with FHR changes

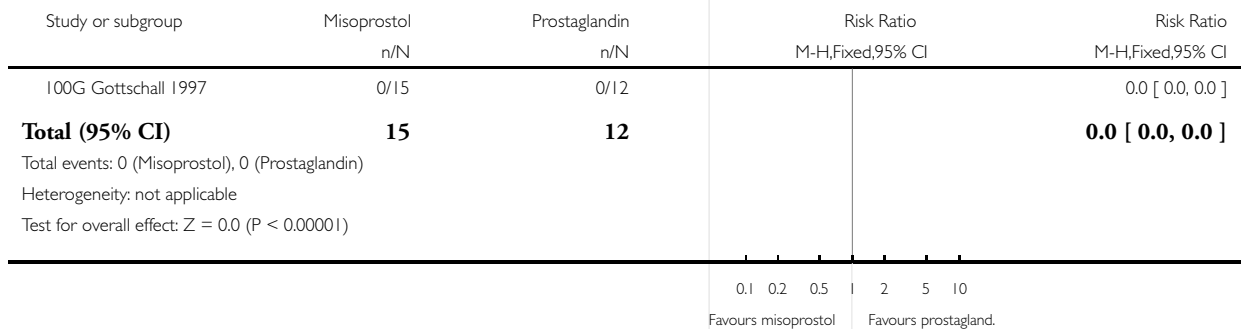


**Analysis 24.2. Comparison 24 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, variable or undefined cervix, Outcome 2 Caesarean section.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 24 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, variable or undefined cervix

Outcome: 2 Caesarean section

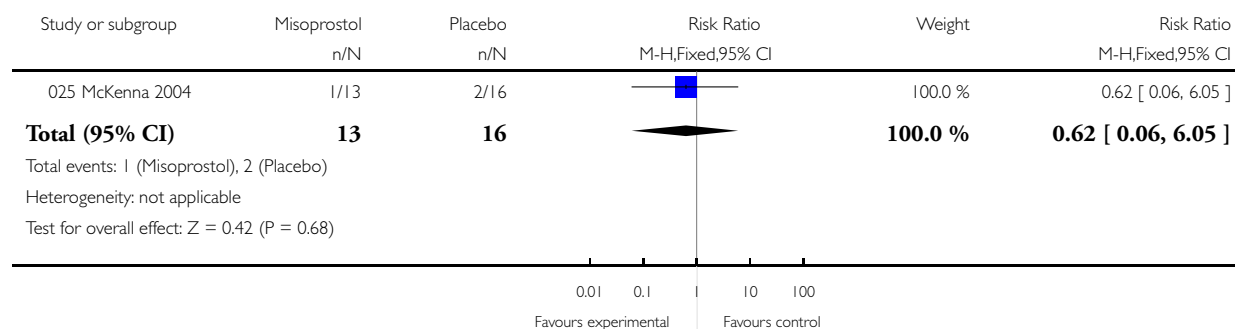


### Analysis 24.3. Comparison 24 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, variable or undefined cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 24 Misoprostol versus vaginal prostaglandin: all multiparae, intact membranes, variable or undefined cervix

Outcome: 3 Caesarean section

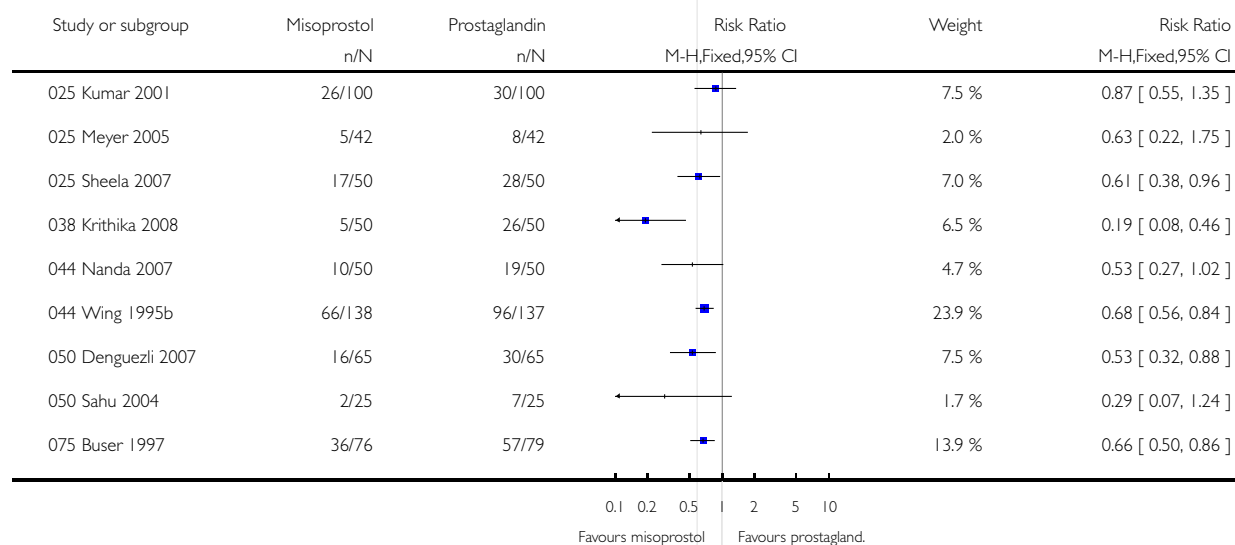


### Analysis 25.1. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 1 Vaginal delivery not achieved within 24 hours.

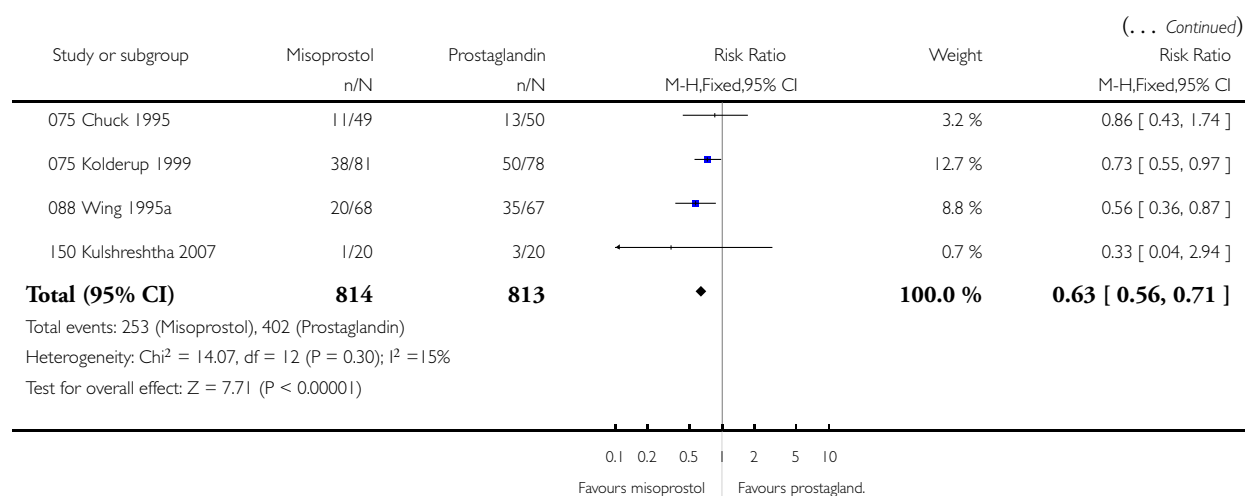
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 1 Vaginal delivery not achieved within 24 hours



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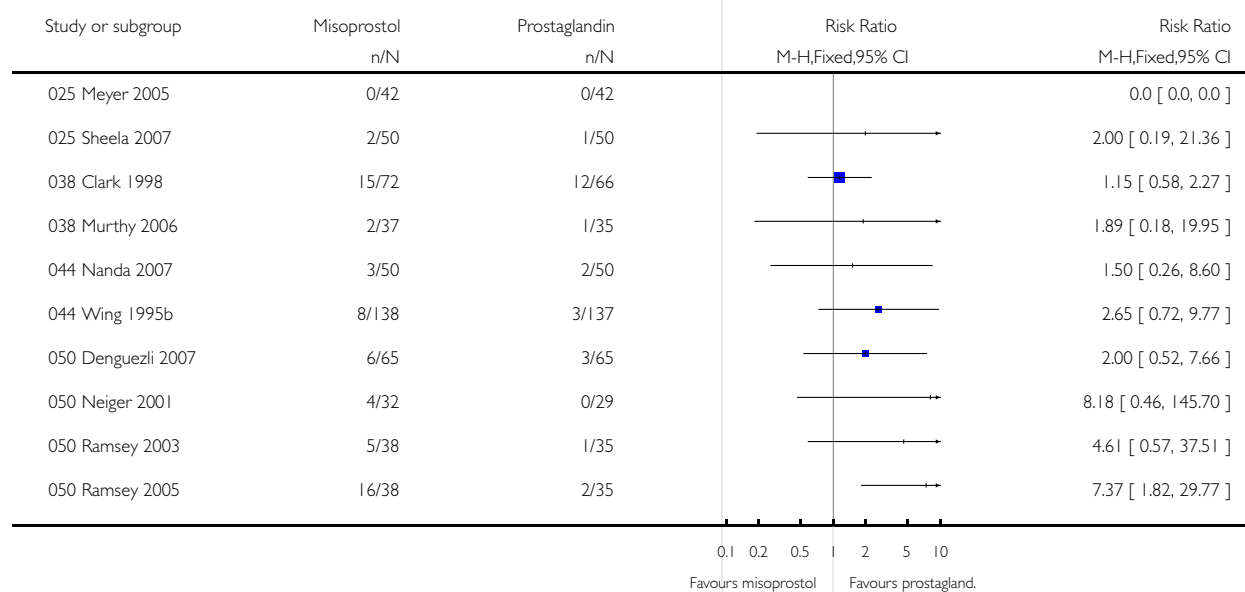


## Analysis 25.2. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 2 Uterine hyperstimulation with FHR changes.

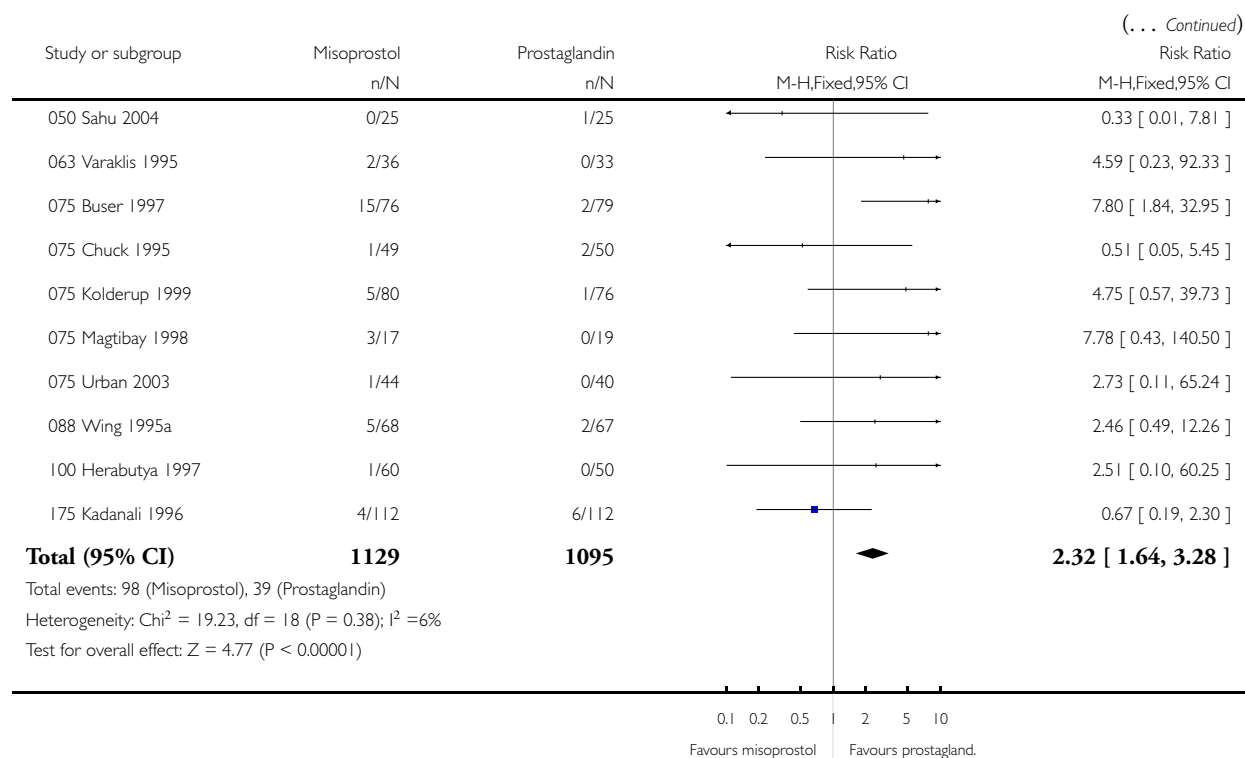
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 2 Uterine hyperstimulation with FHR changes



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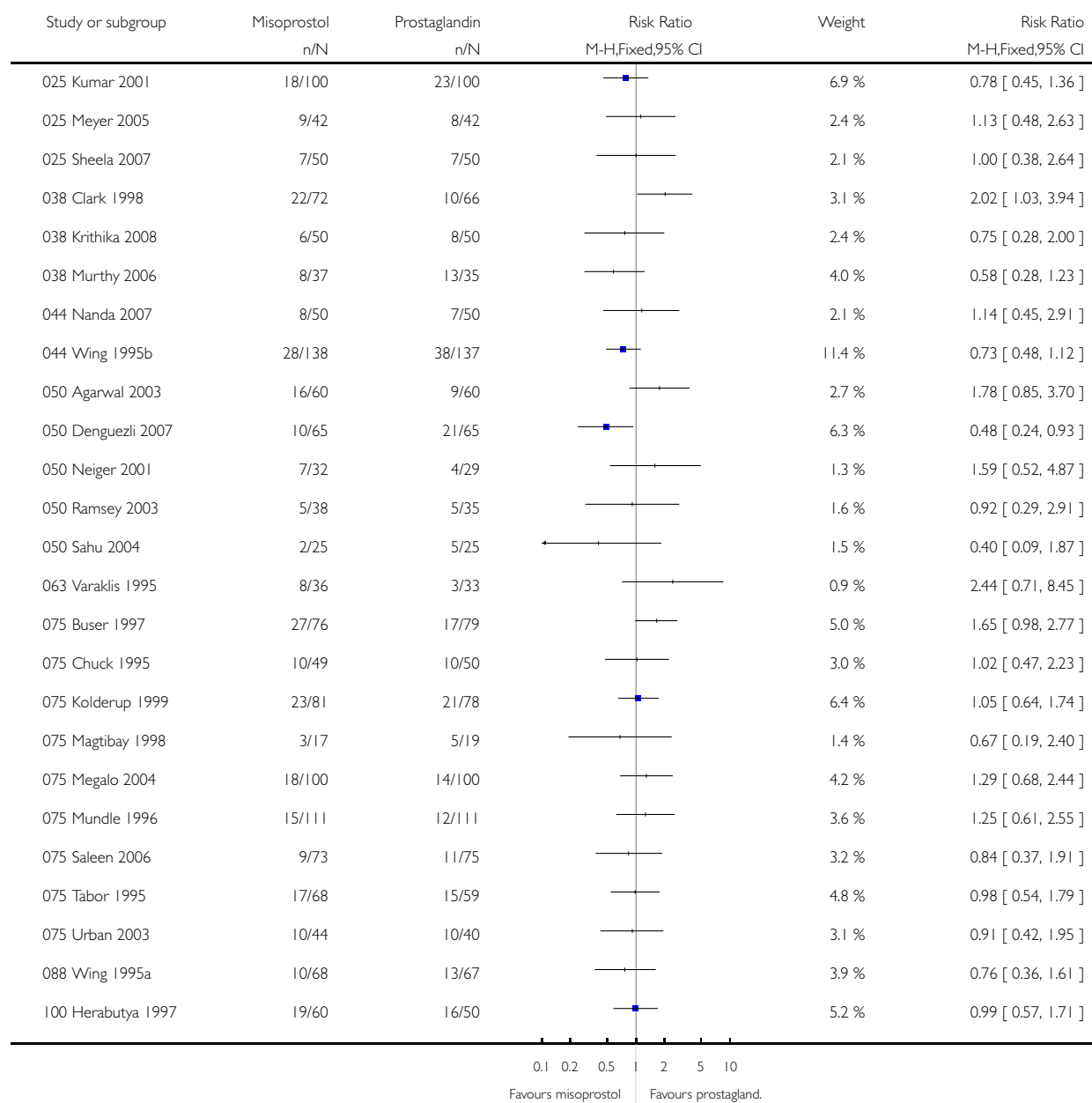


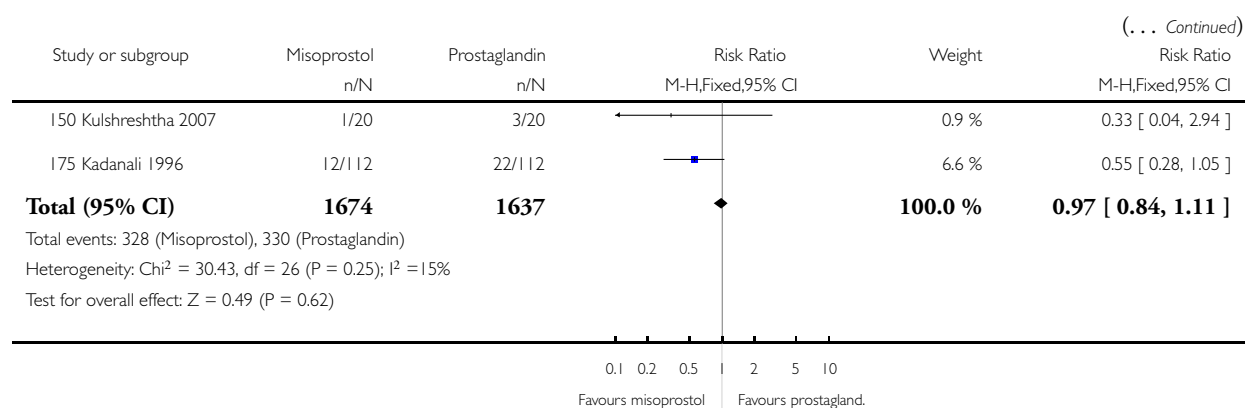
### Analysis 25.3. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 3 Caesarean section



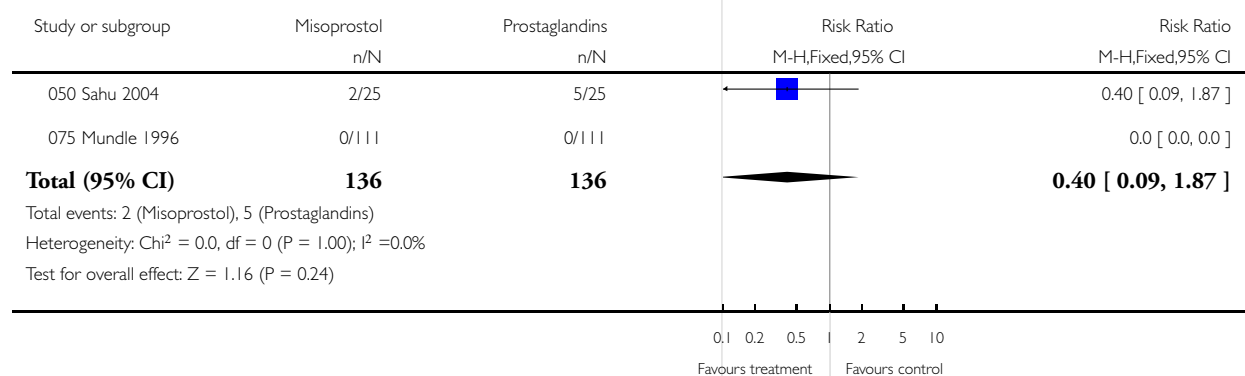


#### Analysis 25.4. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 4 Serious neonatal morbidity/perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 4 Serious neonatal morbidity/perinatal death

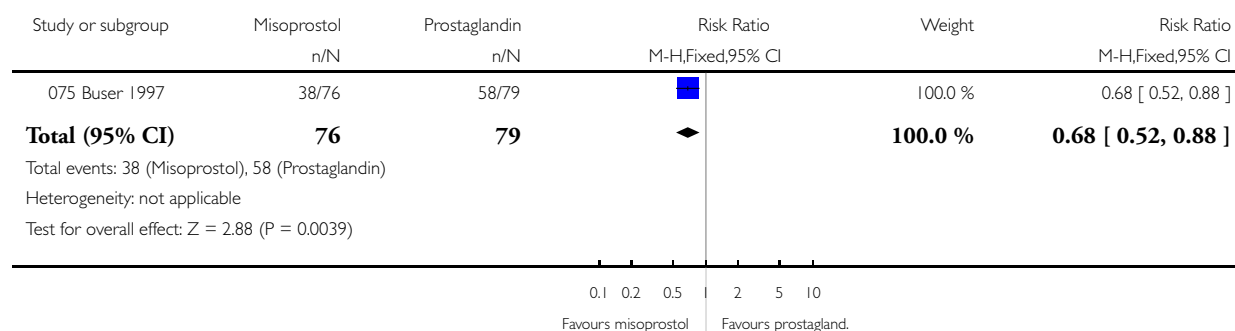


## Analysis 25.5. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 5 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 5 Cervix unfavourable/unchanged after 12-24 hours

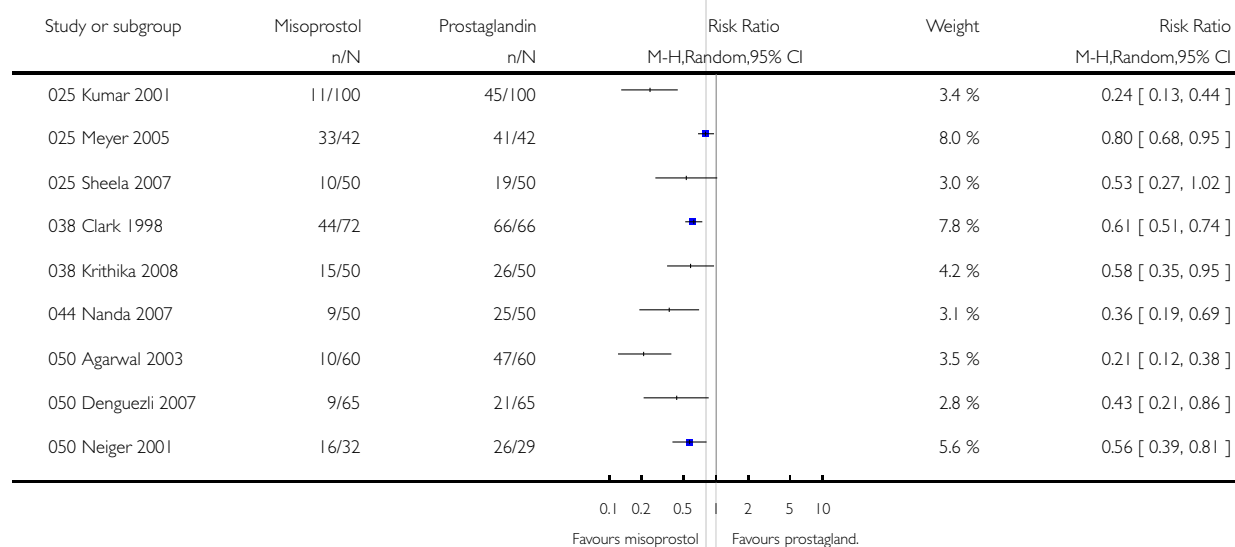


## Analysis 25.6. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 6 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

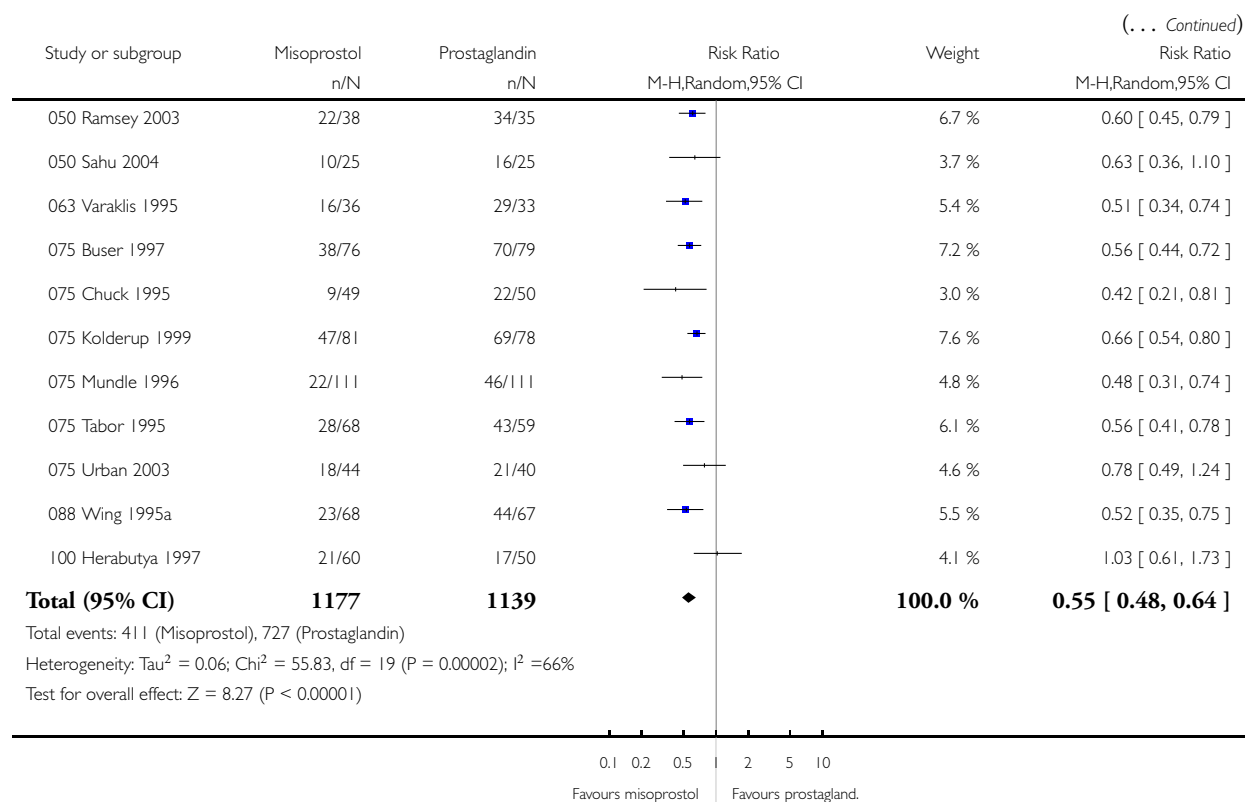
Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 6 Oxytocin augmentation



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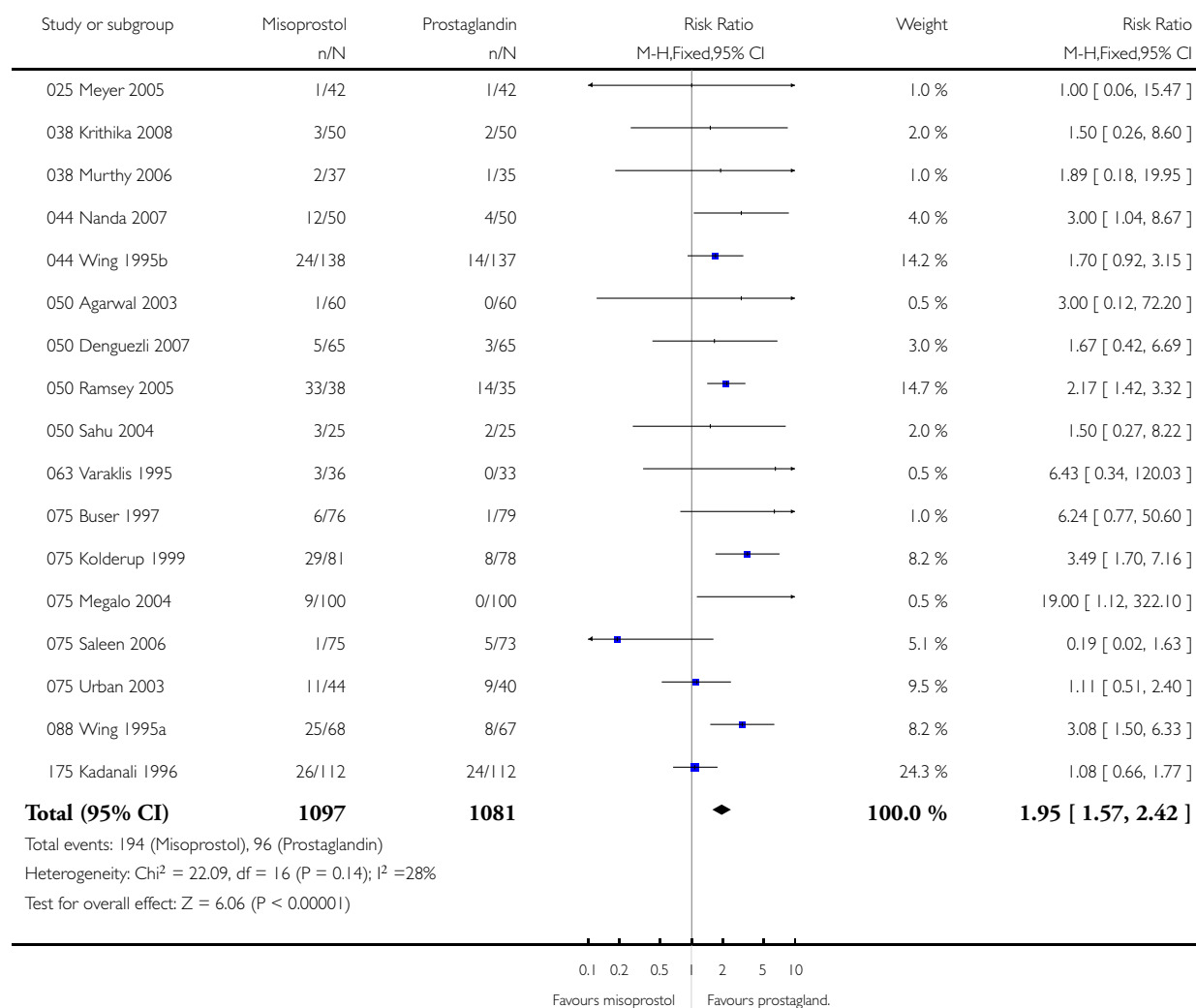


## Analysis 25.7. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 7 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 7 Uterine hyperstimulation without FHR changes

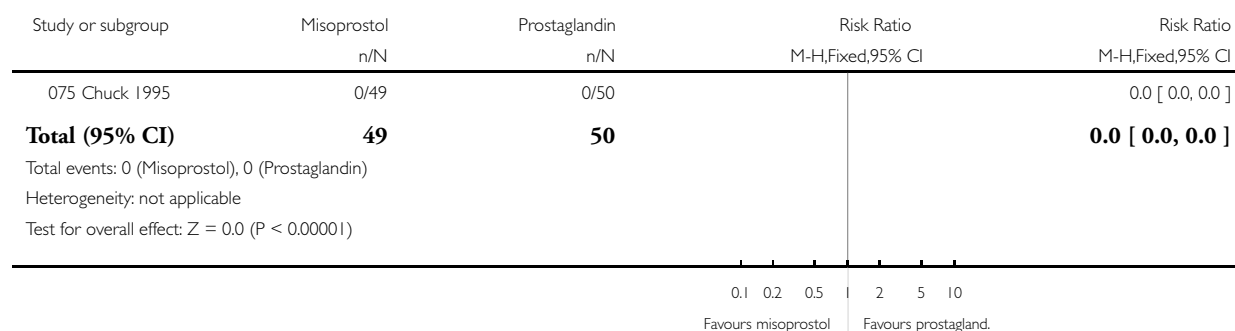


### Analysis 25.8. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 8 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 8 Uterine rupture

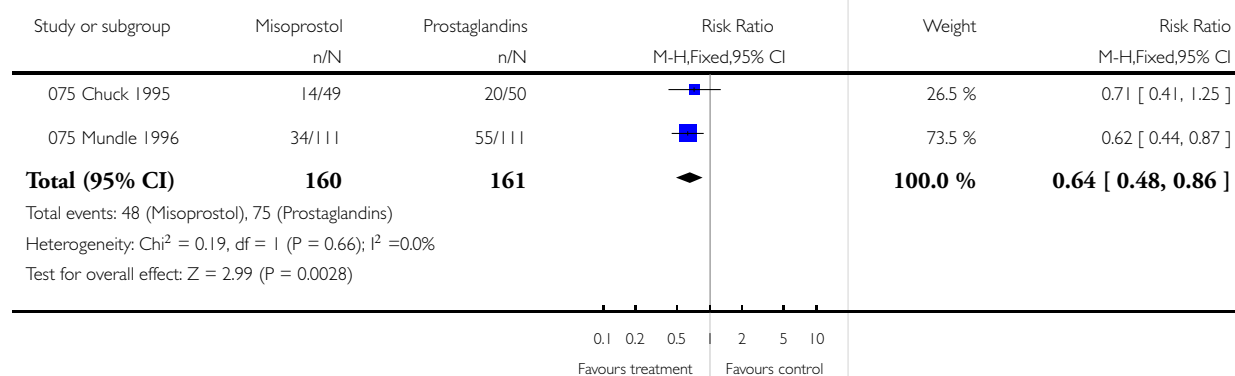


### Analysis 25.9. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 9 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 9 Epidural analgesia

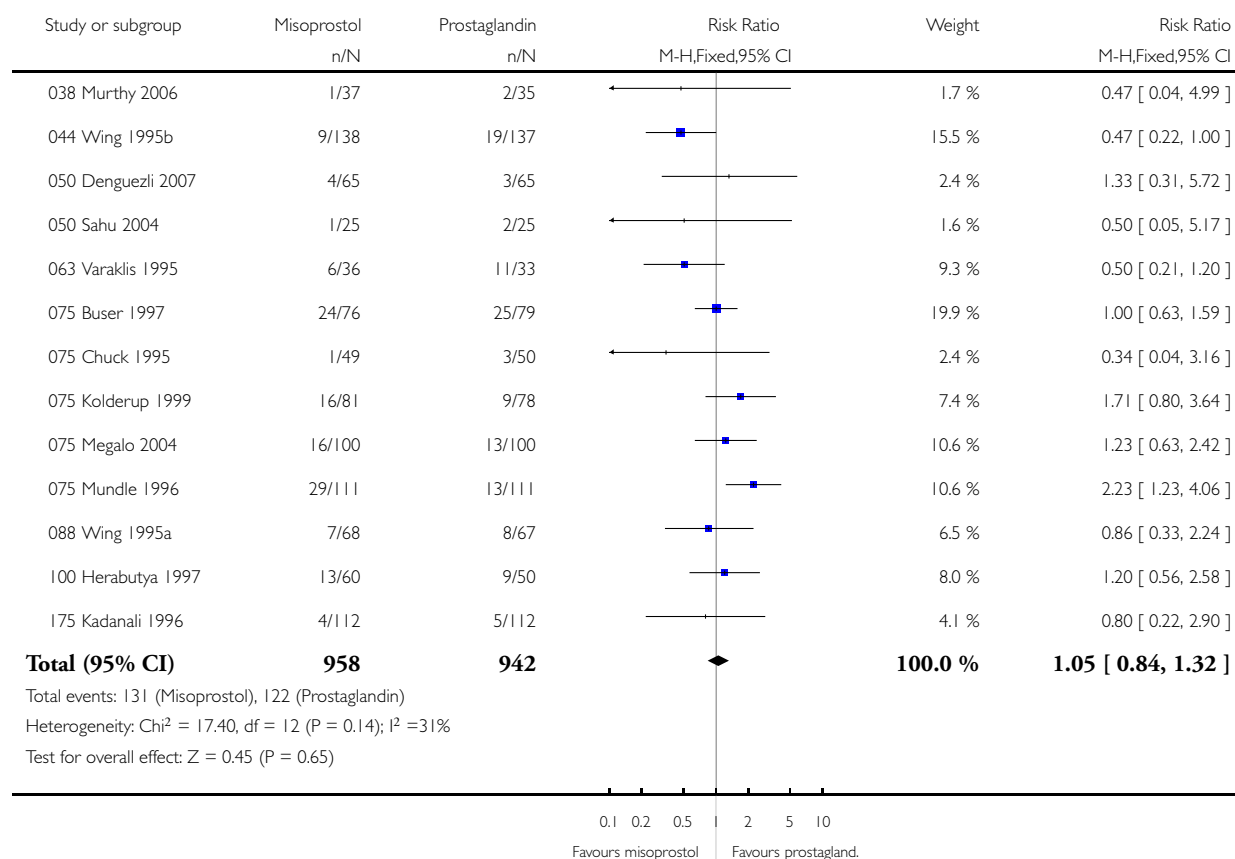


# **Analysis 25.10. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 10 Instrumental vaginal delivery.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 10 Instrumental vaginal delivery

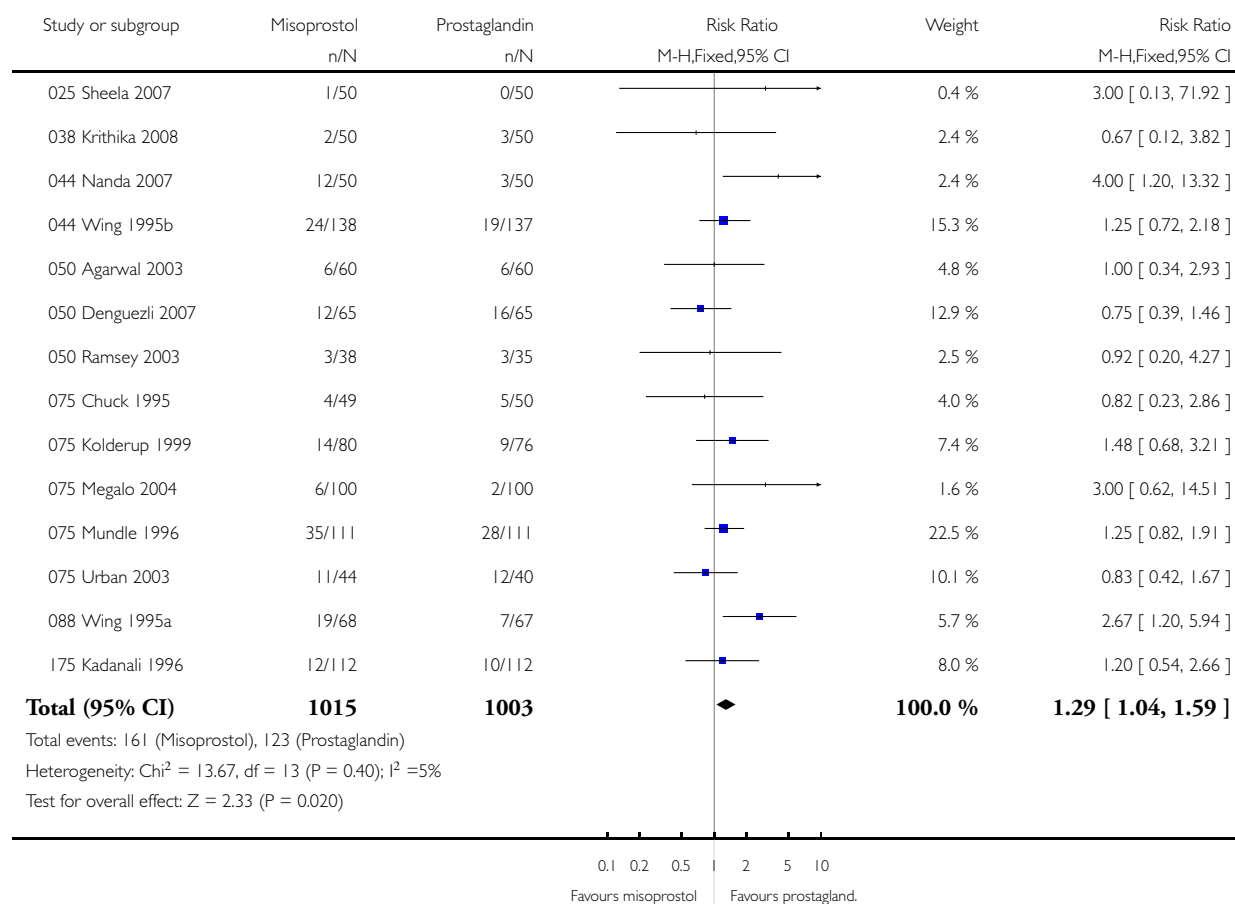


## Analysis 25.11. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 11 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 11 Meconium-stained liquor

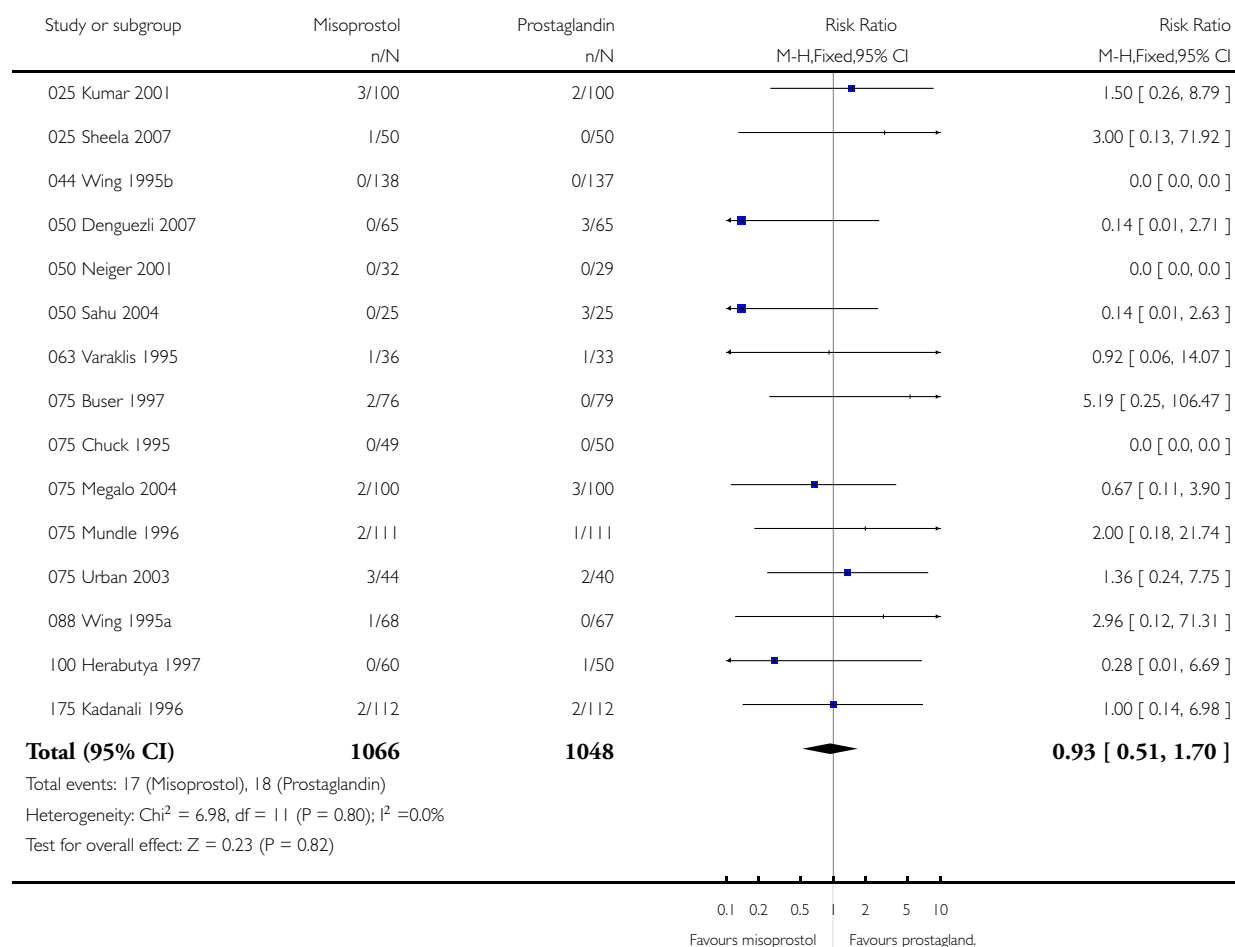


# **Analysis 25.12. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 12 Apgar score < 7 at 5 minutes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 12 Apgar score < 7 at 5 minutes

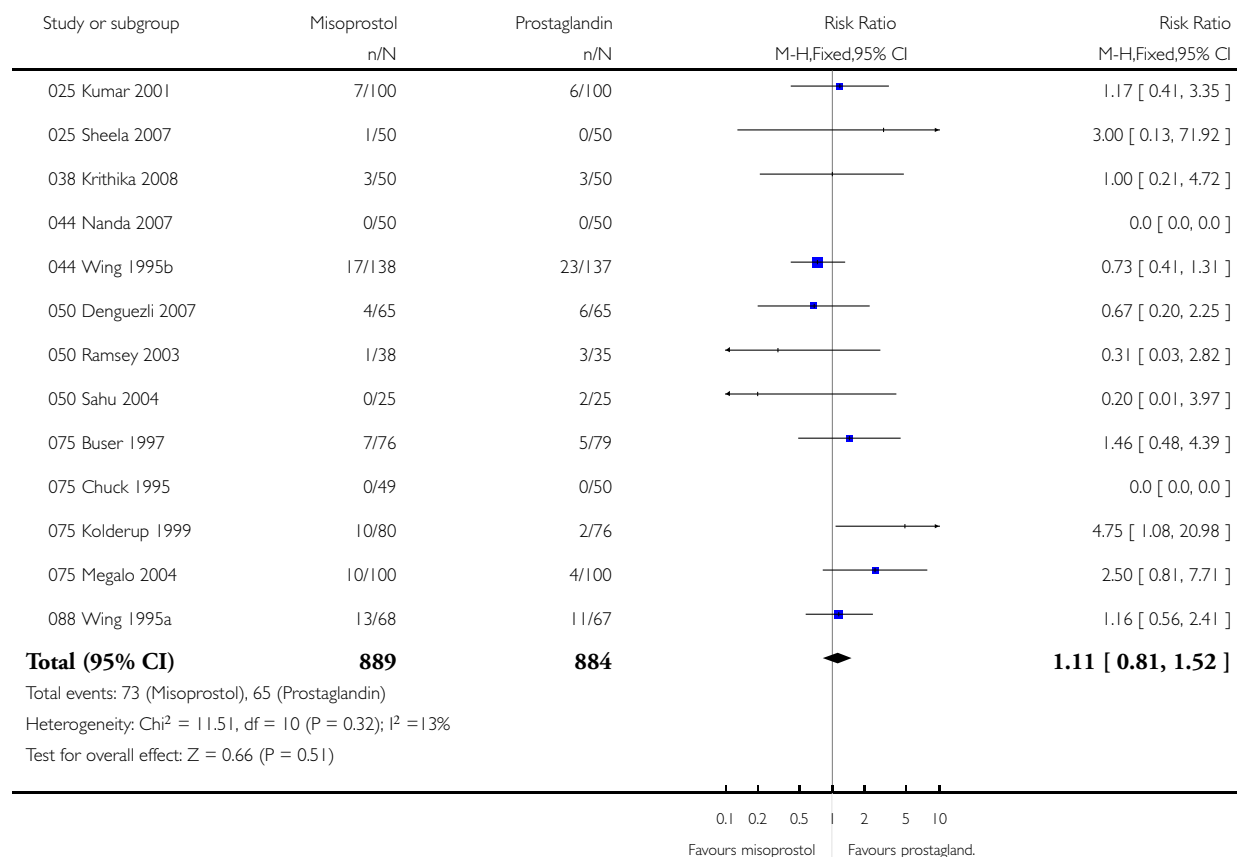


### Analysis 25.13. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 13 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 13 Neonatal intensive care unit admission

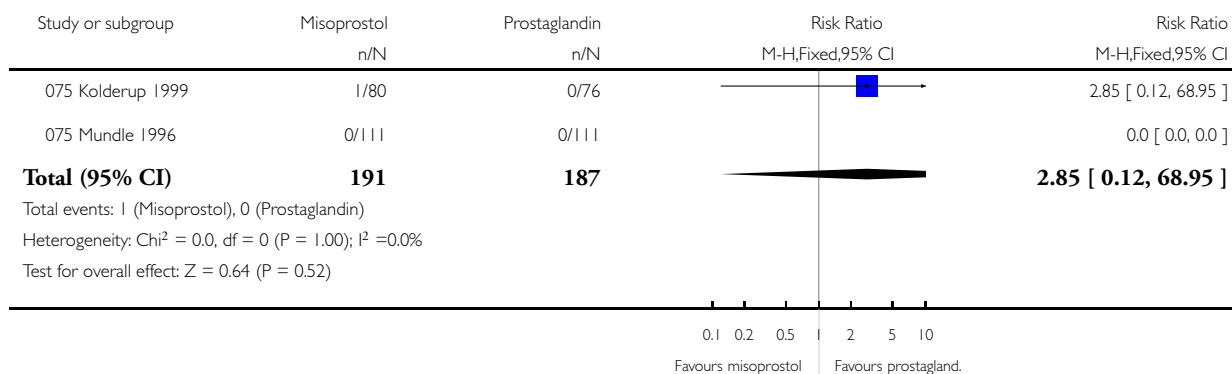


### Analysis 25.14. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 14 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 14 Perinatal death

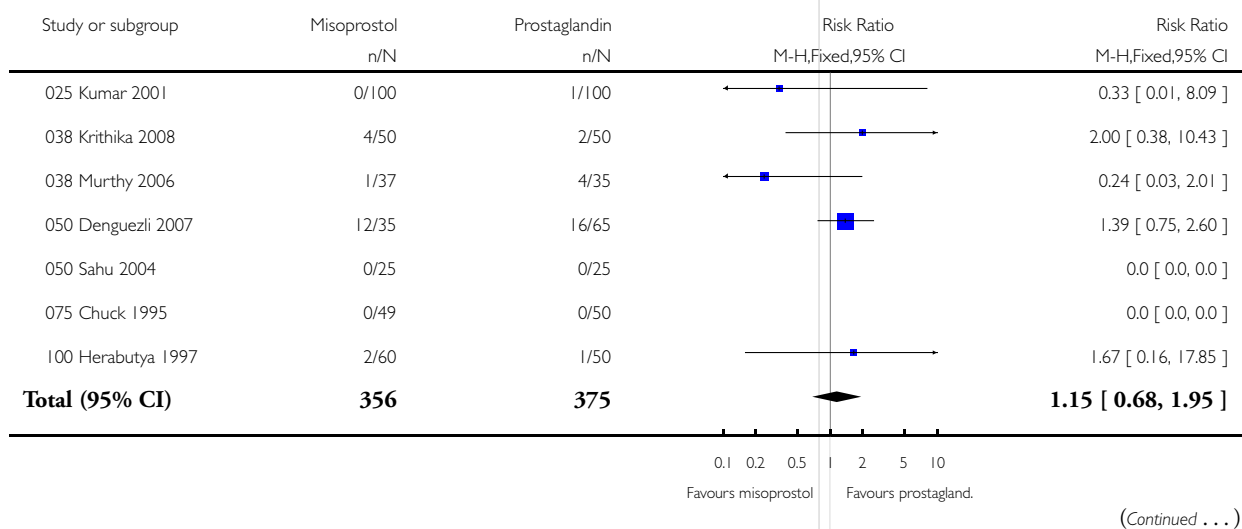


### Analysis 25.15. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 15 Maternal side effects (eg nausea, vomiting, diarrhoea).

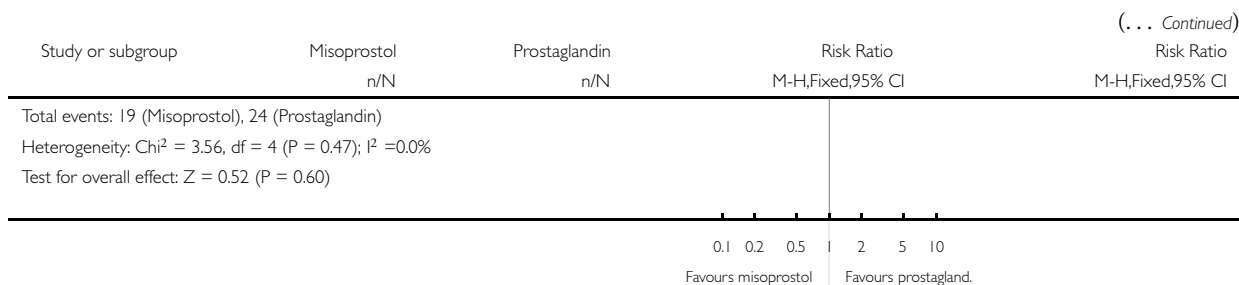
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 15 Maternal side effects (eg nausea, vomiting, diarrhoea)





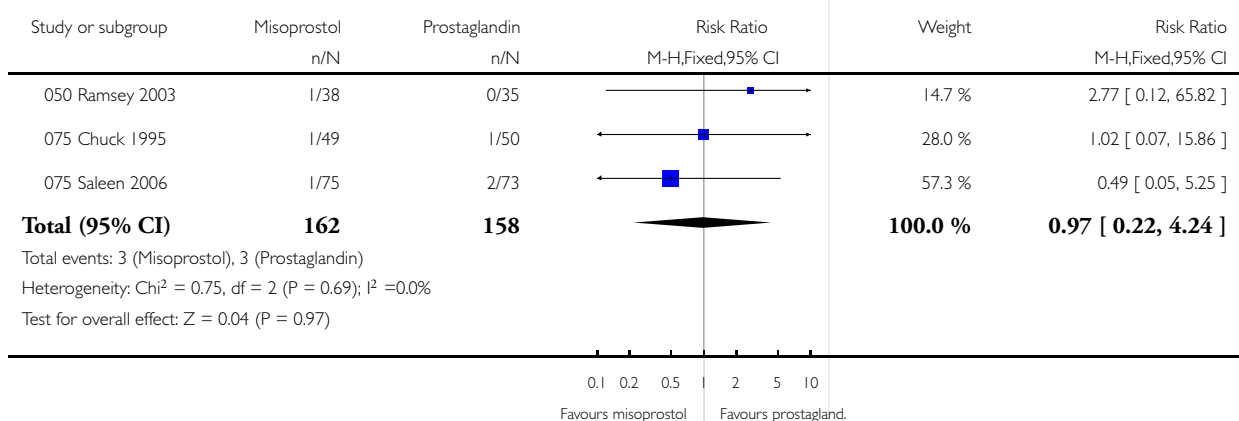


### Analysis 25.16. Comparison 25 Misoprostol versus intracervical prostaglandin: all women, Outcome 16 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 25 Misoprostol versus intracervical prostaglandin: all women

Outcome: 16 Postpartum haemorrhage

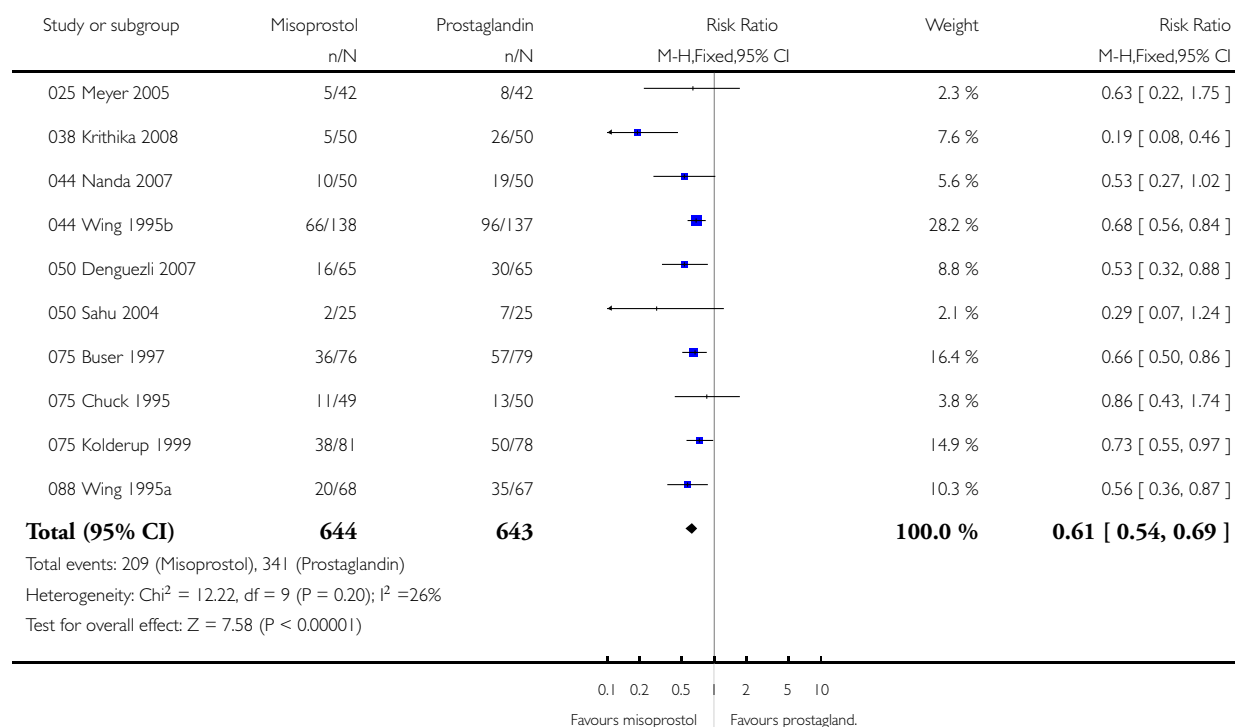


# **Analysis 26.1. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

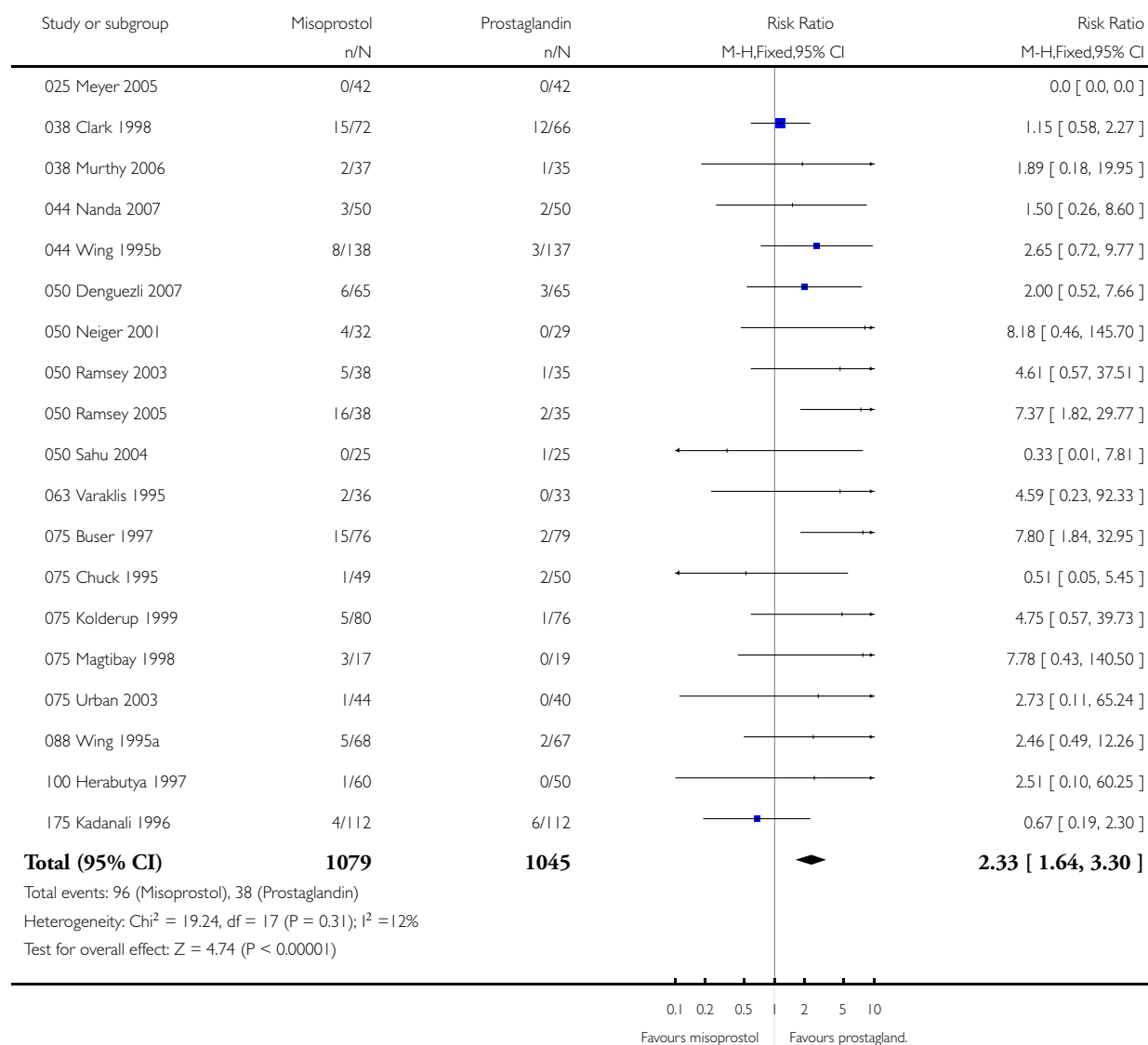


## Analysis 26.2. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

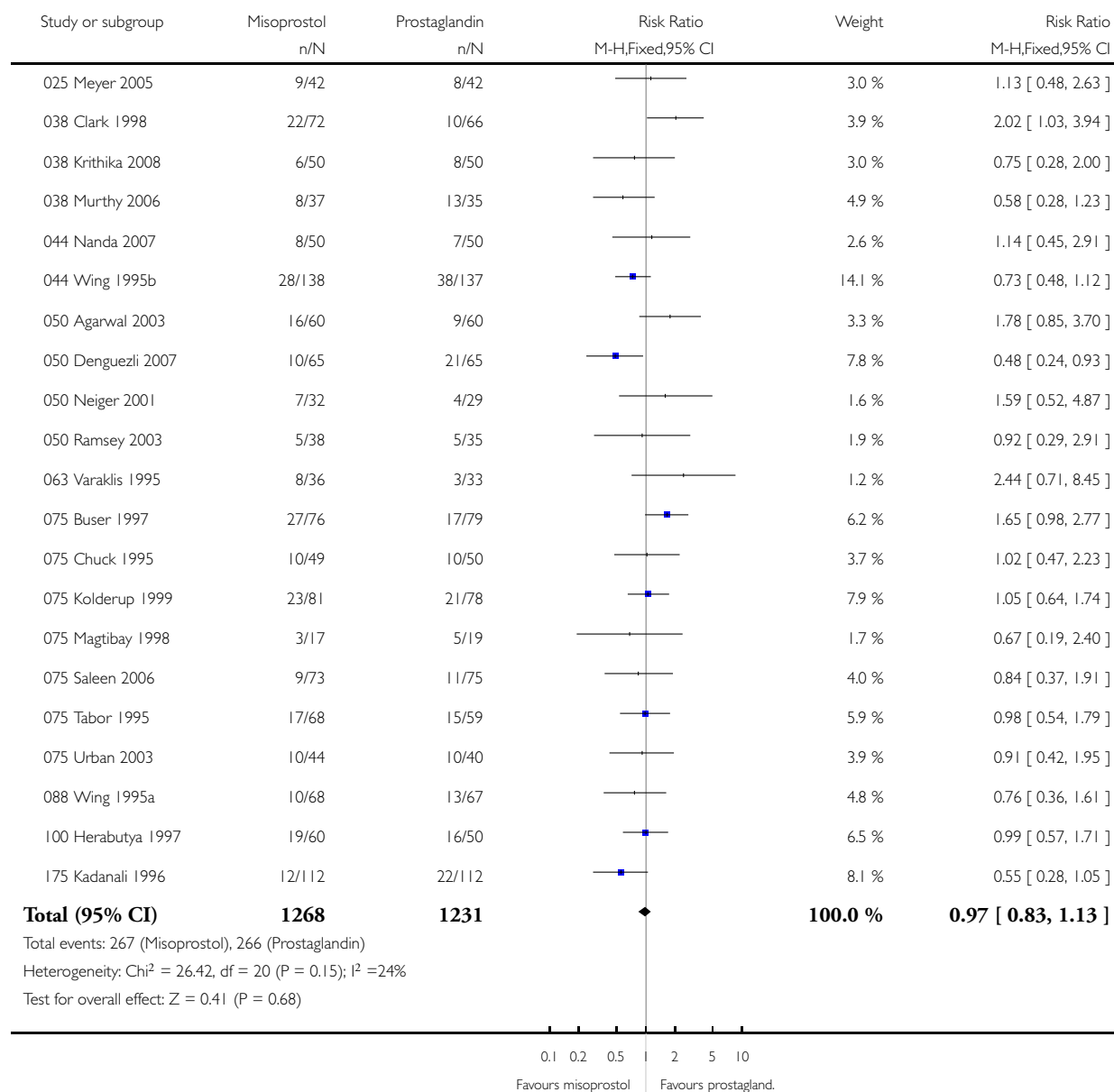


### Analysis 26.3. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 3 Caesarean section

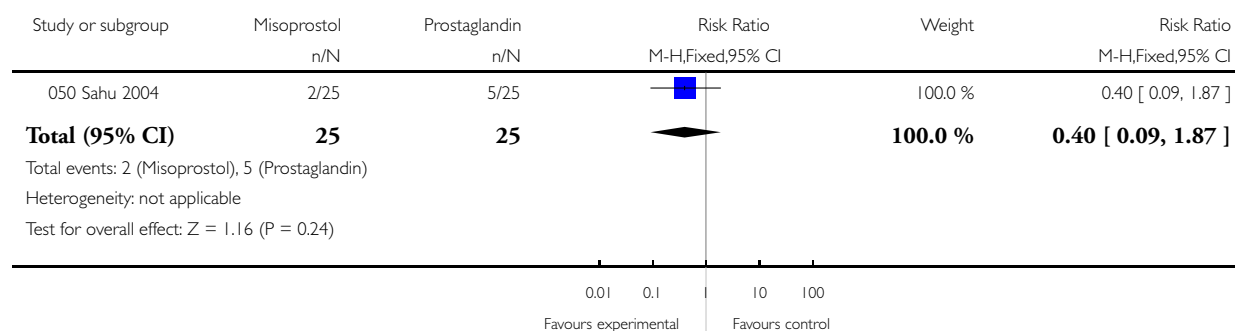


#### Analysis 26.4. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 4 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 4 Serious neonatal morbidity

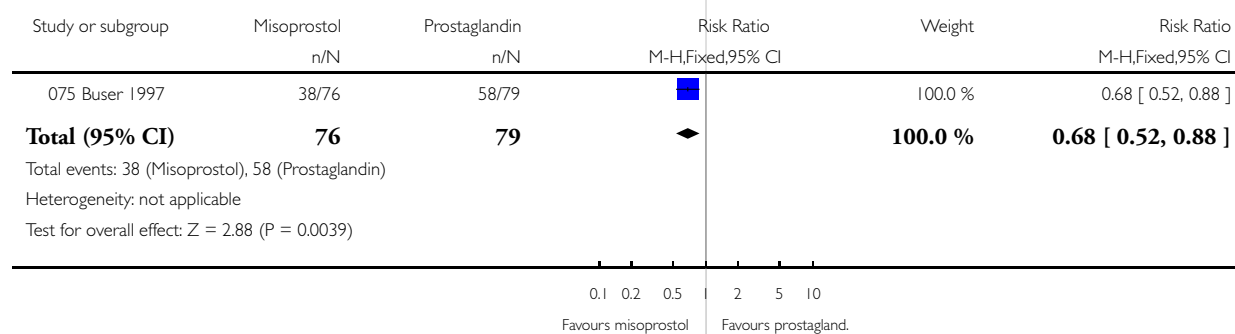


#### Analysis 26.5. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 5 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 5 Cervix unfavourable/unchanged after 12-24 hours

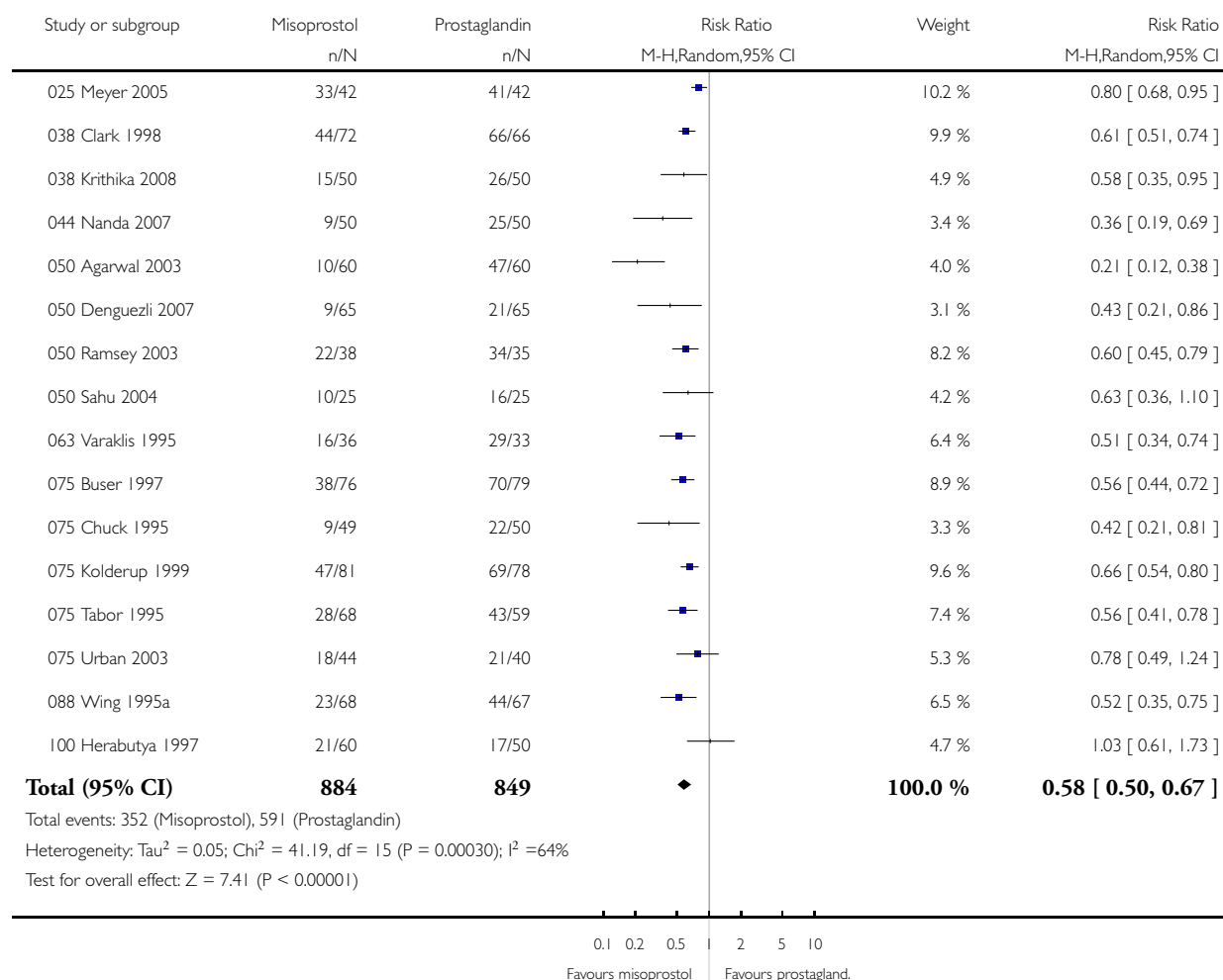


## Analysis 26.6. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 6 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 6 Oxytocin augmentation

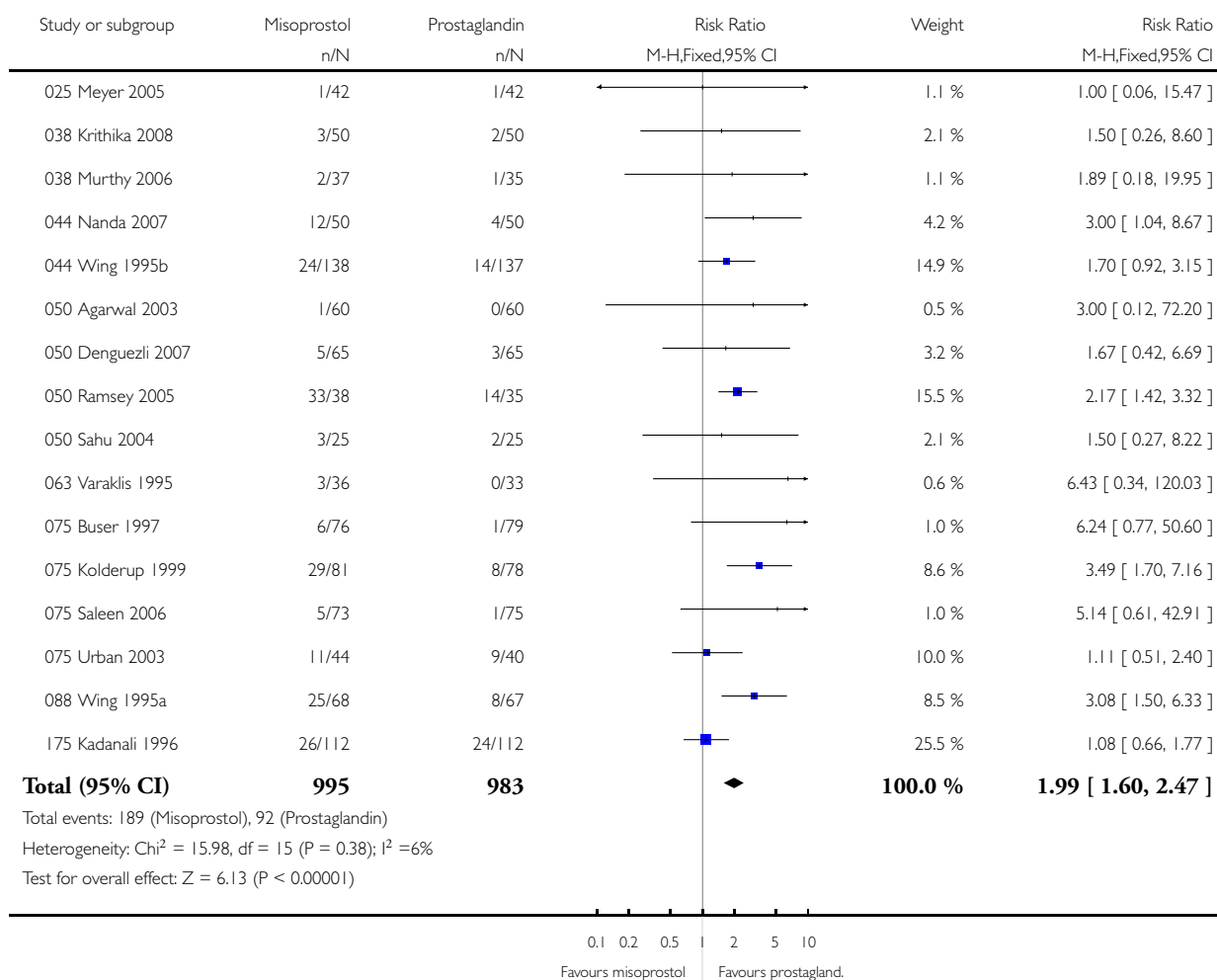


# **Analysis 26.7. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 7 Uterine hyperstimulation without FHR changes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 7 Uterine hyperstimulation without FHR changes

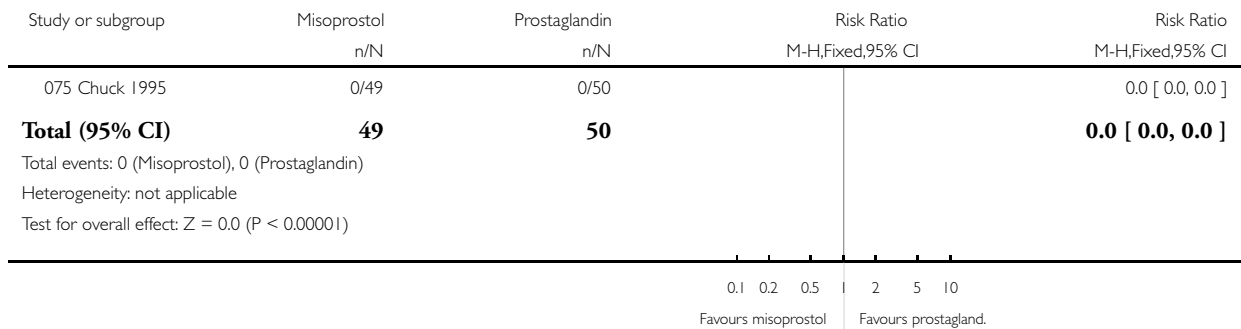


### Analysis 26.8. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 8 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 8 Uterine rupture

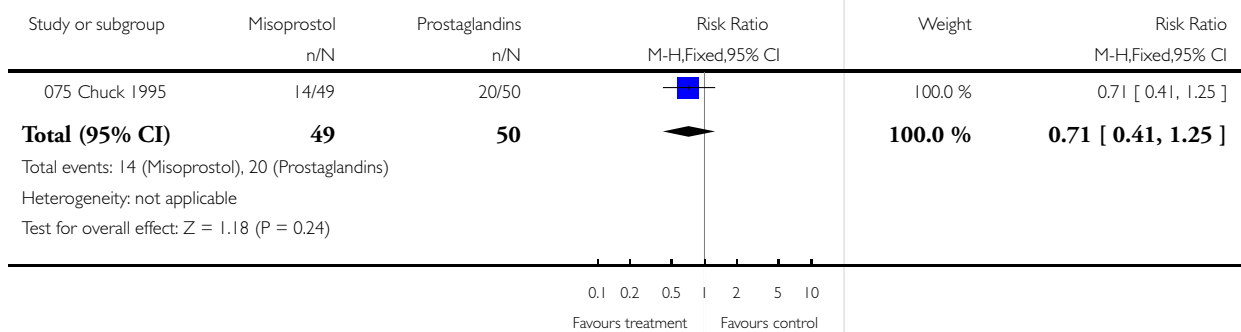


### Analysis 26.9. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 9 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 9 Epidural analgesia



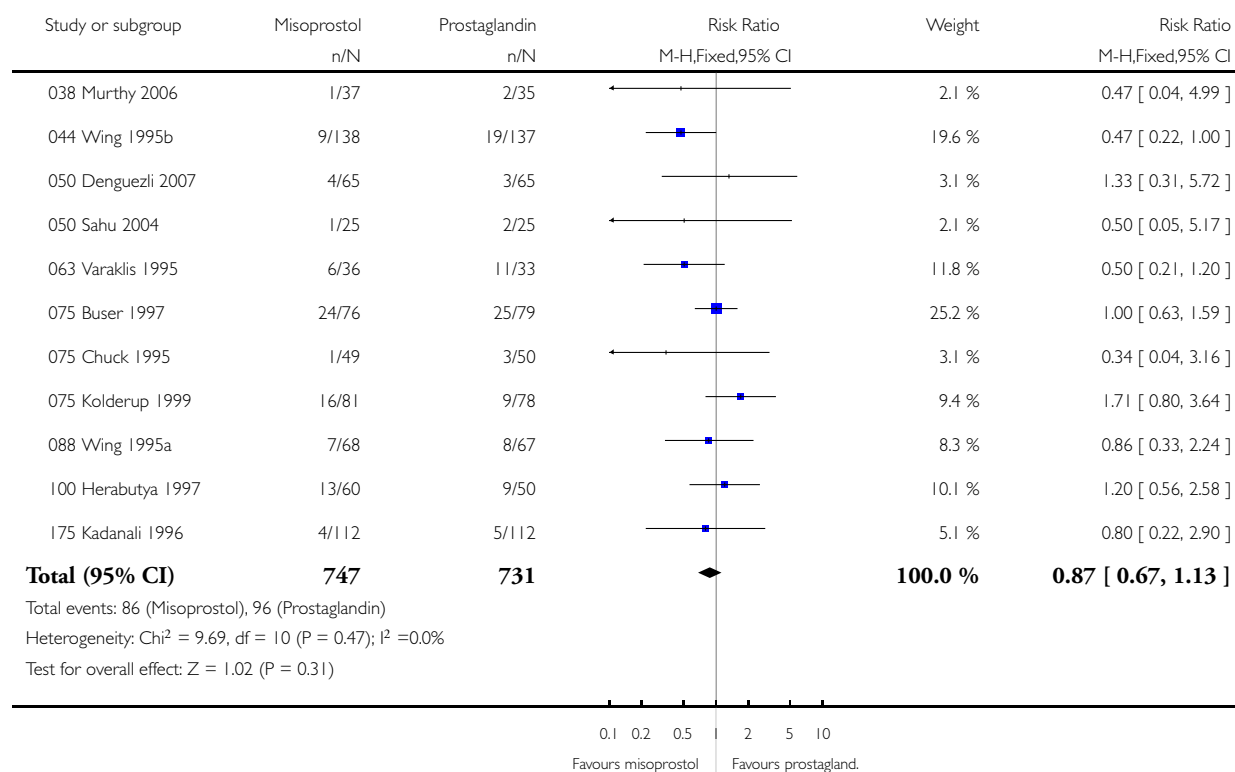


# **Analysis 26.10. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 10 Instrumental vaginal delivery.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 10 Instrumental vaginal delivery

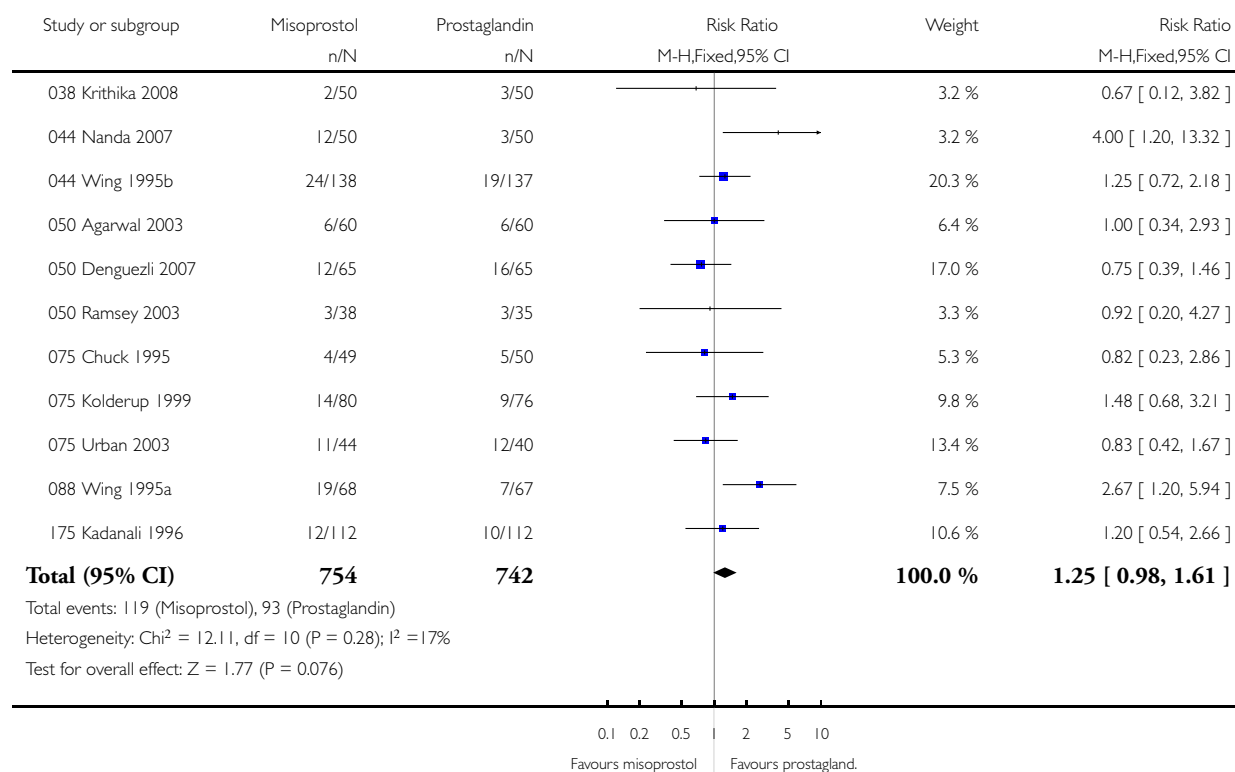


## Analysis 26.11. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 11 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 11 Meconium-stained liquor

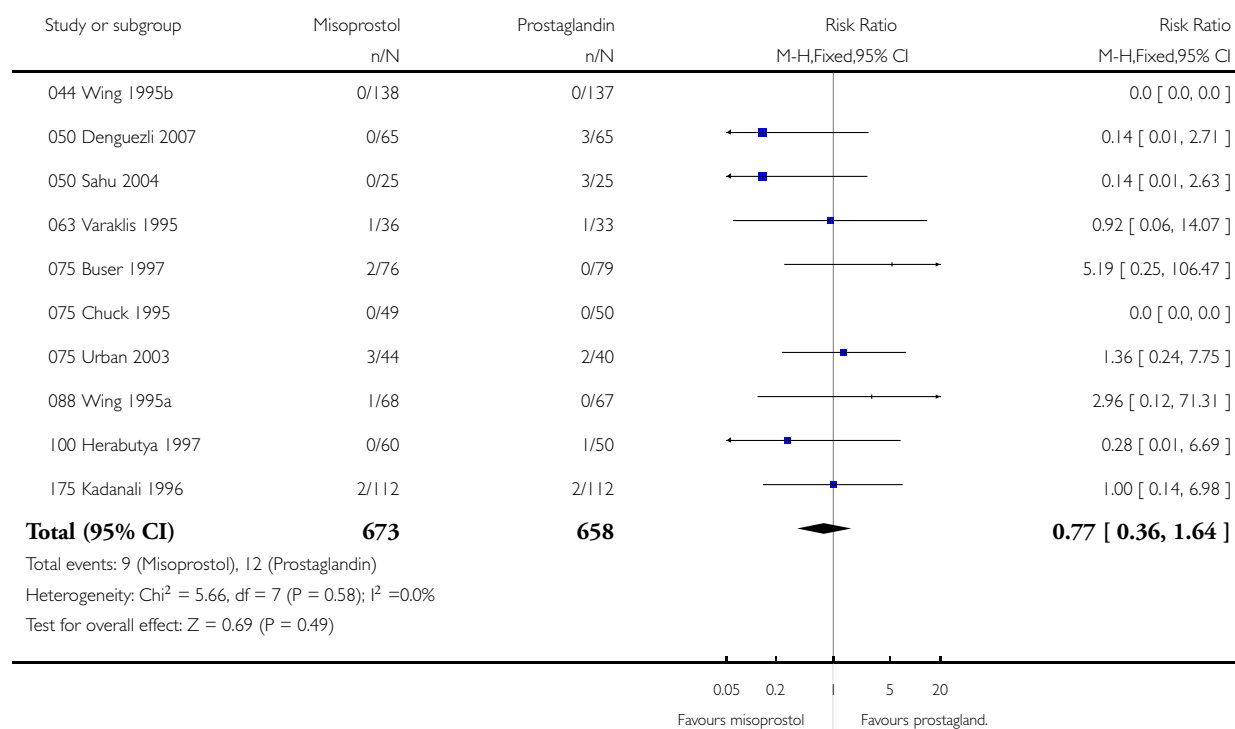


## Analysis 26.12. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 12 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 12 Apgar score < 7 at 5 minutes

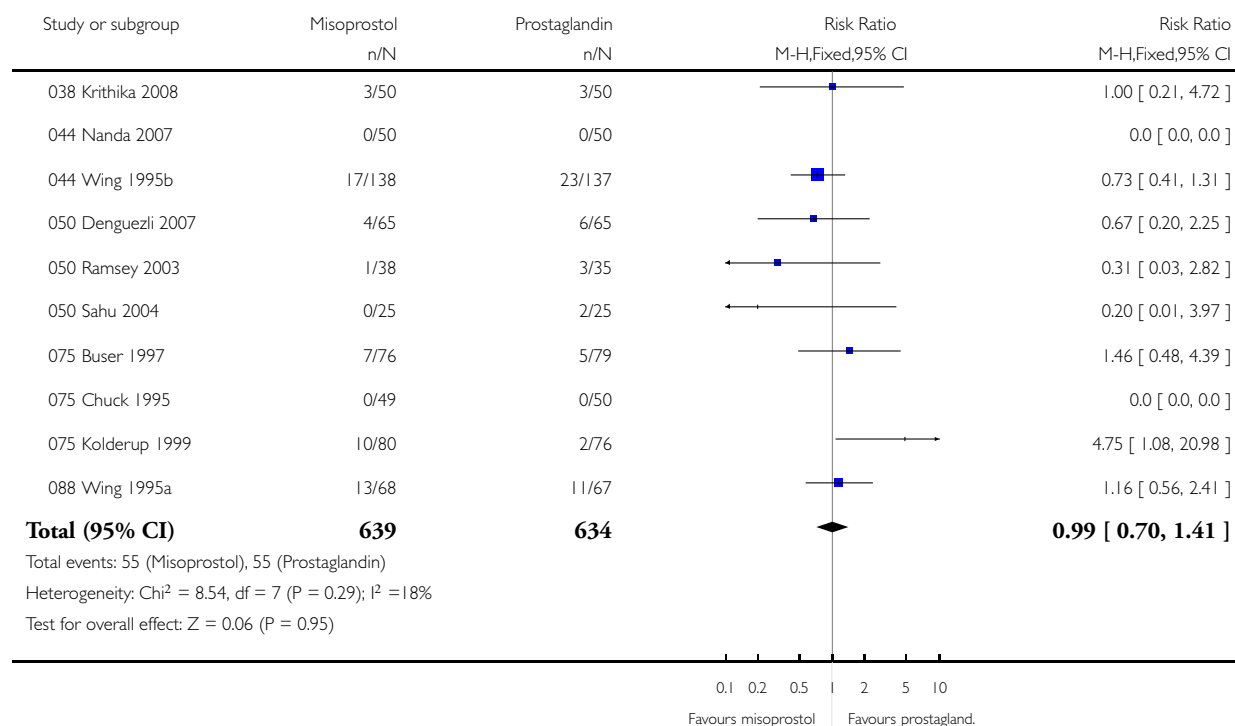


### Analysis 26.13. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 13 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 13 Neonatal intensive care unit admission

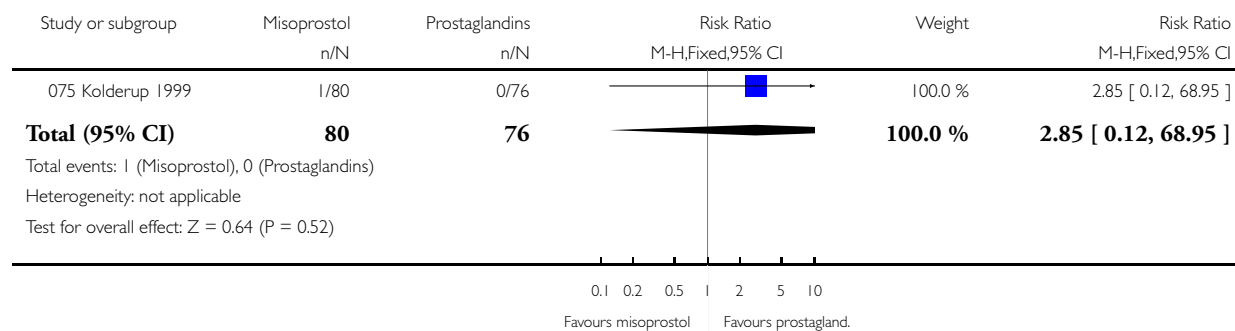


### Analysis 26.14. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 14 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 14 Perinatal death

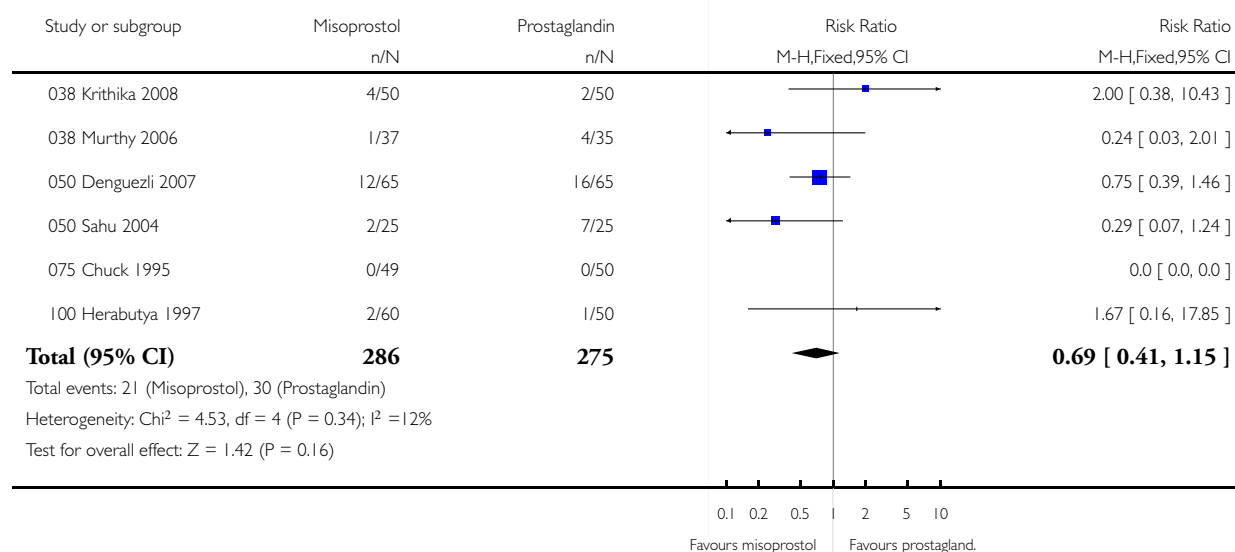


### Analysis 26.15. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 15 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 15 Maternal side effects (eg nausea, vomiting, diarrhoea)

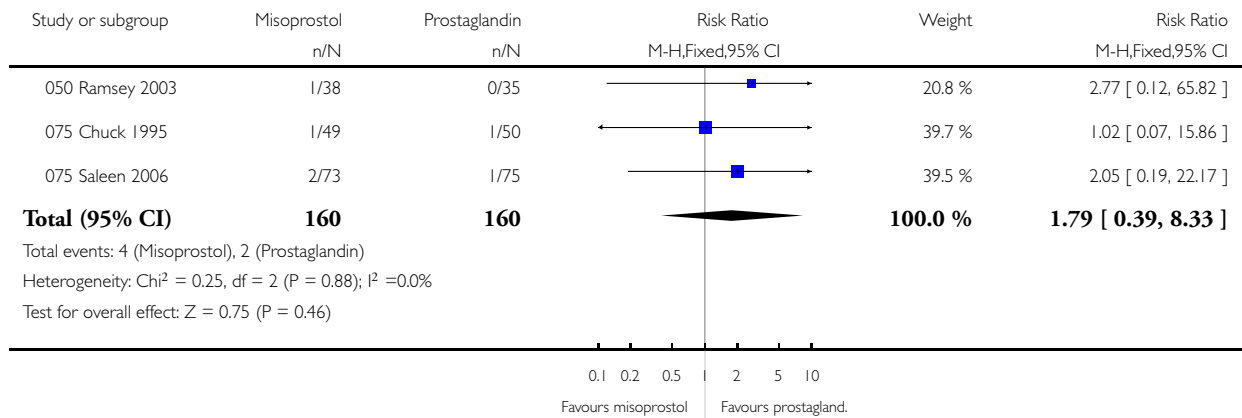


# **Analysis 26.16. Comparison 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix, Outcome 16 Postpartum haemorrhage.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 26 Misoprostol versus intracervical prostaglandin: all women with unfavourable cervix

Outcome: 16 Postpartum haemorrhage

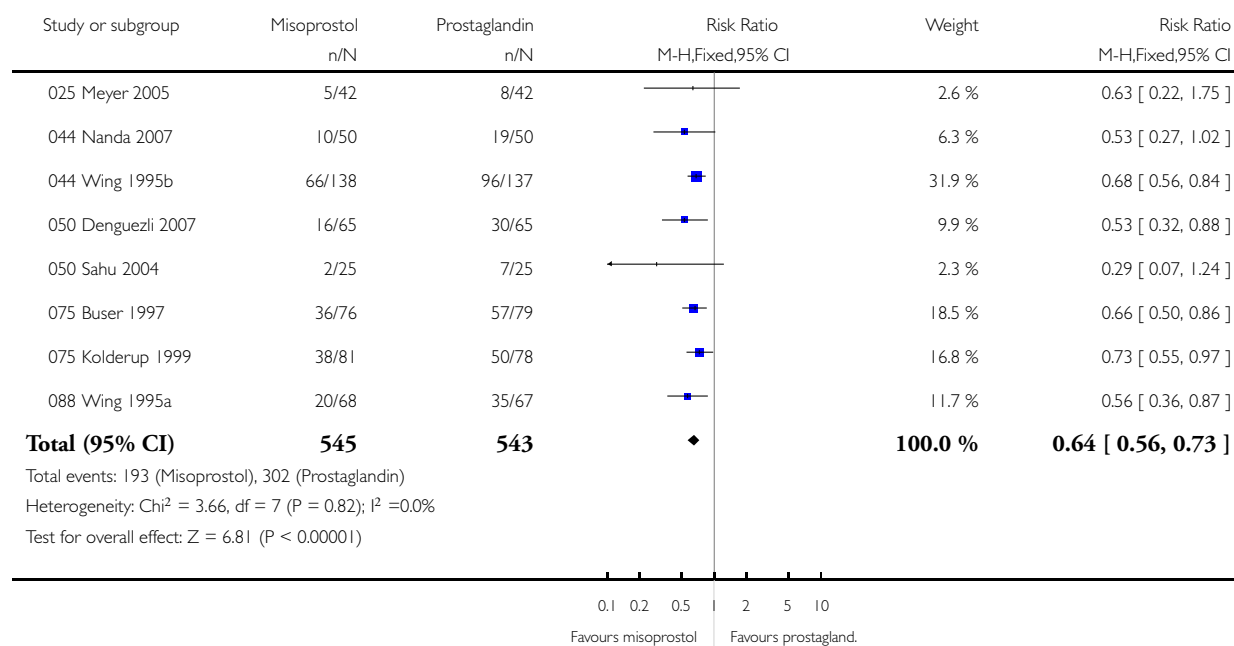


# **Analysis 27.1. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

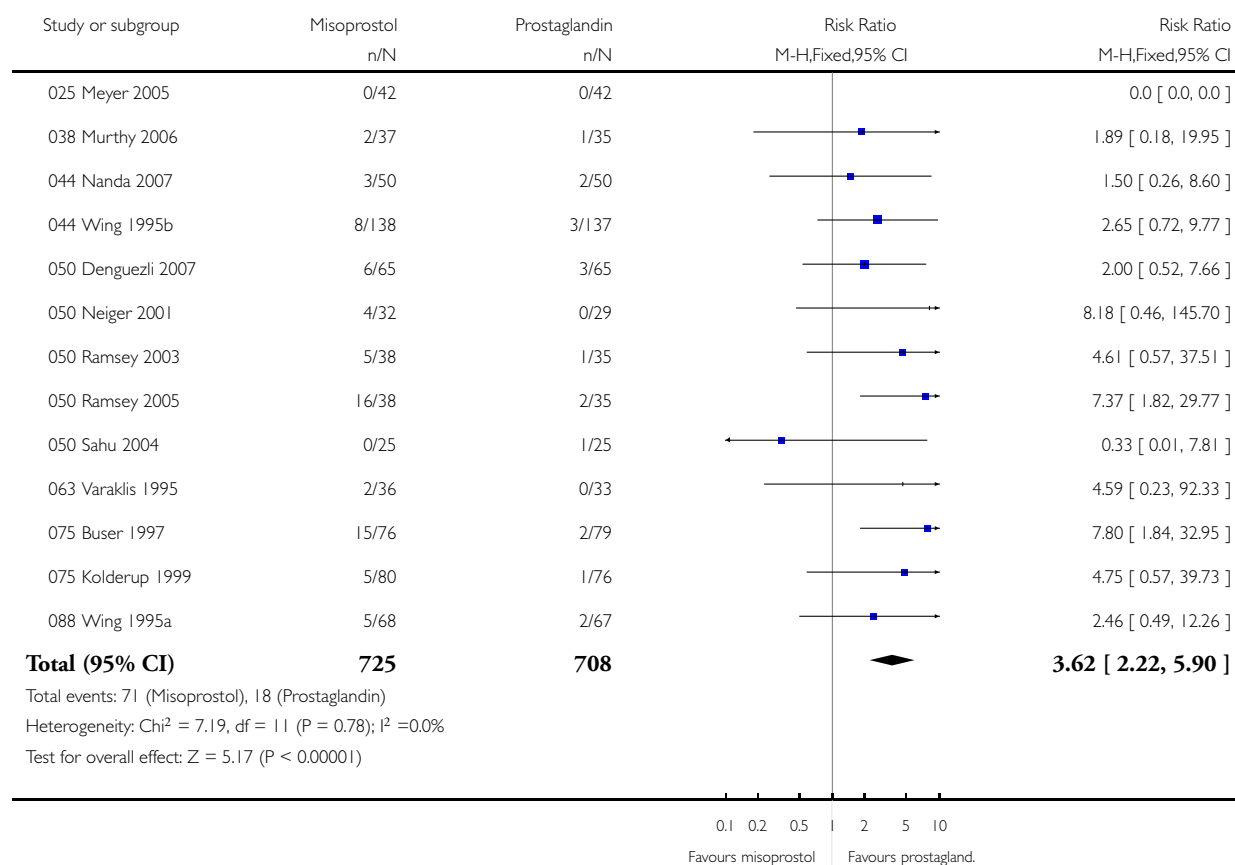


## Analysis 27.2. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes



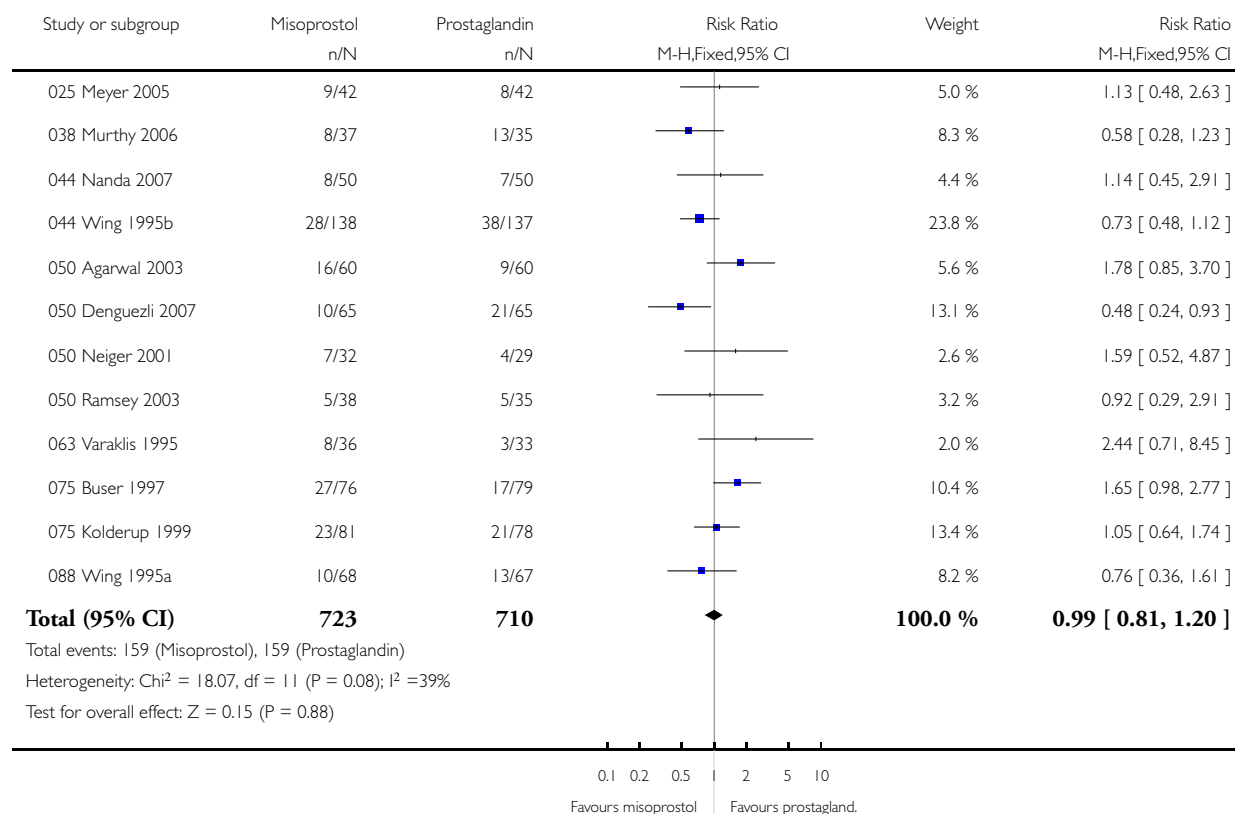


### Analysis 27.3. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 3 Caesarean section

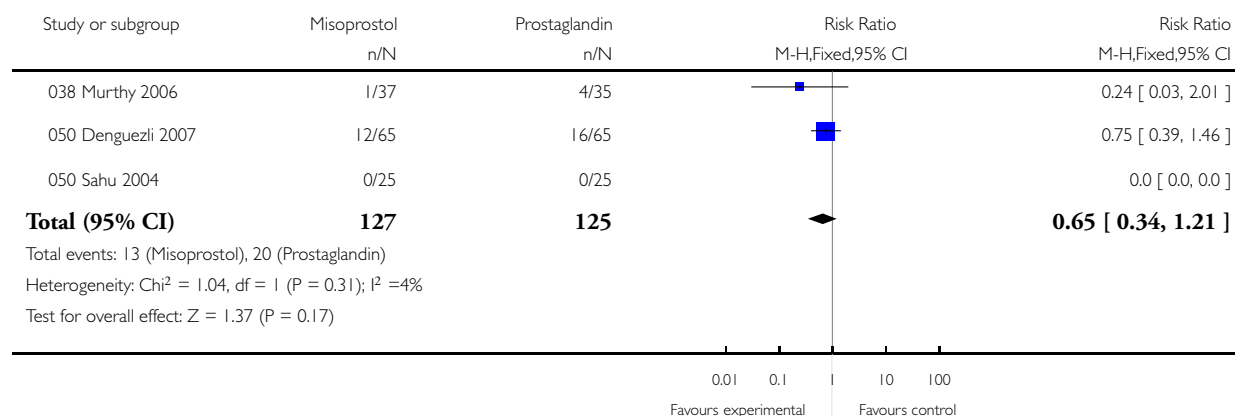


### Analysis 27.4. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 4 Maternal side effects.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 4 Maternal side effects

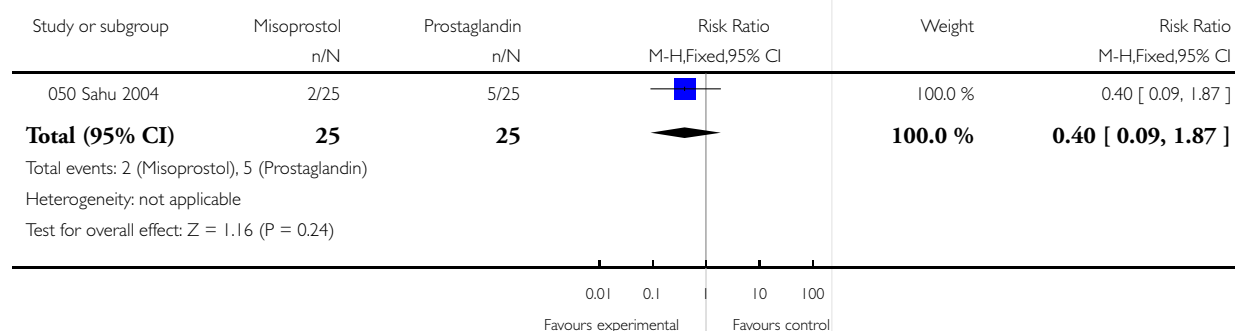


### Analysis 27.5. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 5 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 5 Serious neonatal morbidity

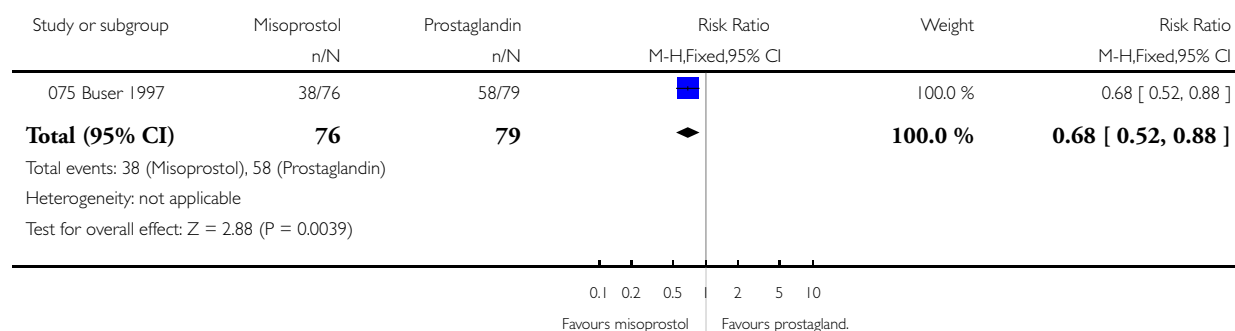


### Analysis 27.6. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 6 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 6 Cervix unfavourable/unchanged after 12-24 hours

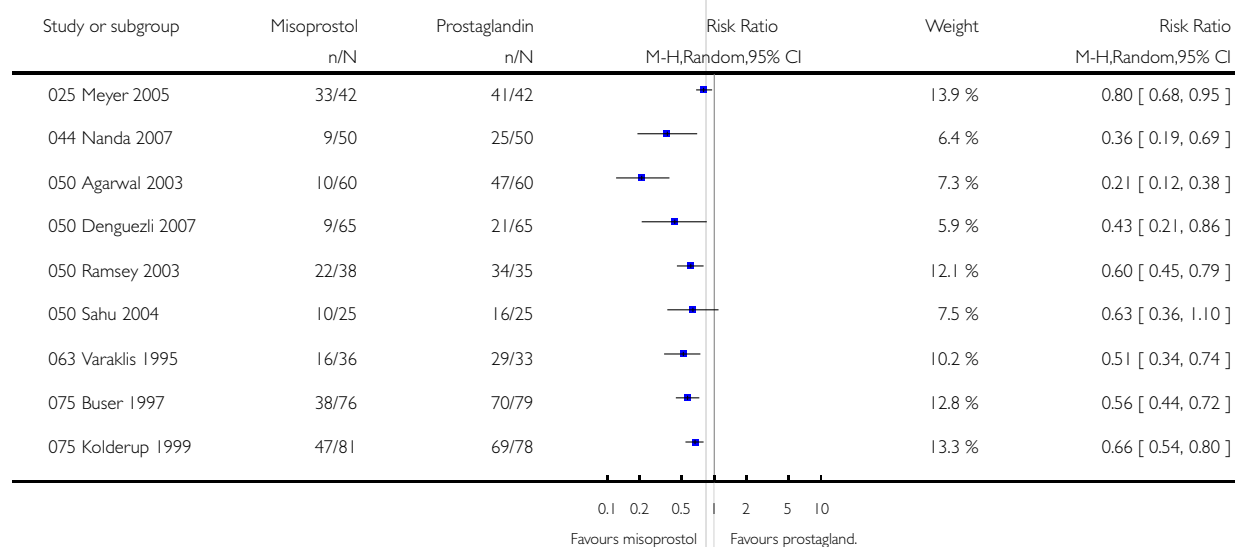


### Analysis 27.7. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 7 Oxytocin augmentation.

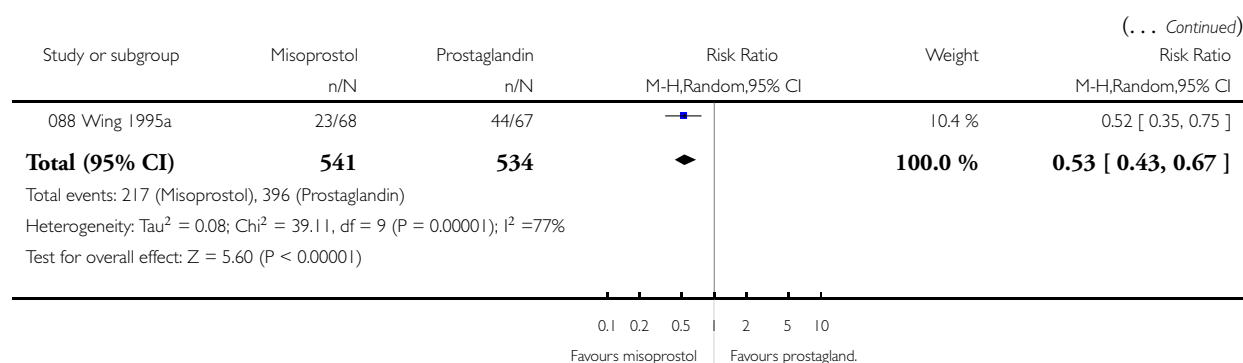
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 7 Oxytocin augmentation



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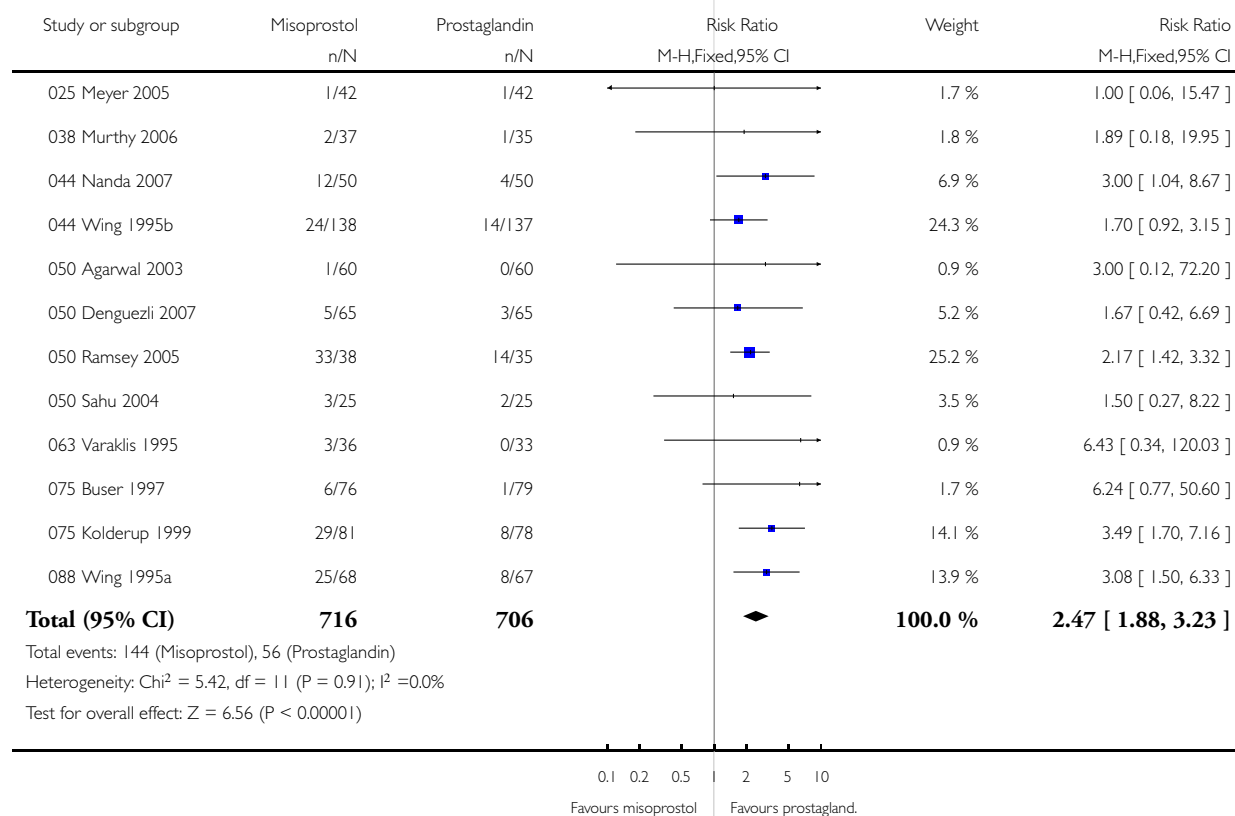


### Analysis 27.8. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 8 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 8 Uterine hyperstimulation without FHR changes

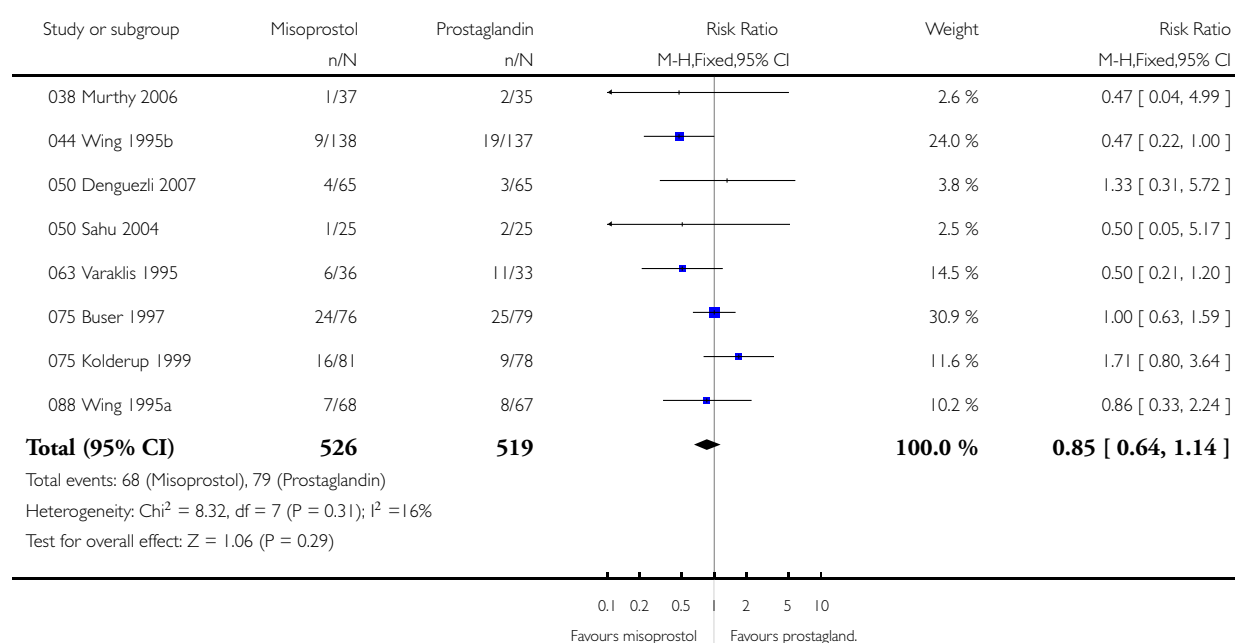


### Analysis 27.9. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 9 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 9 Instrumental vaginal delivery

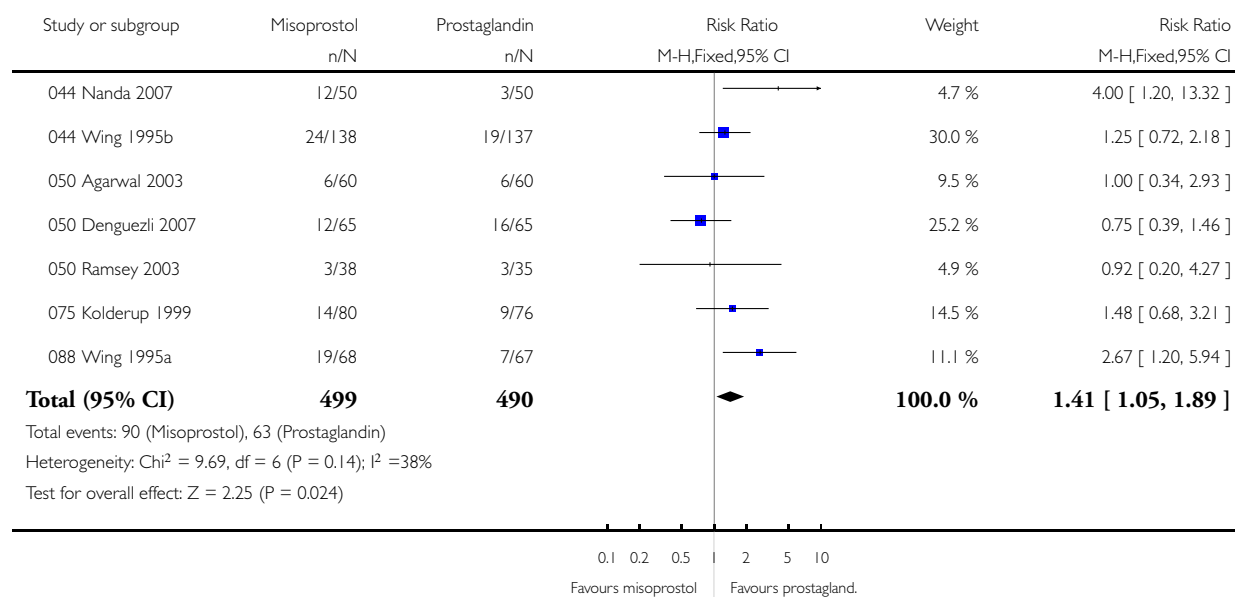


# **Analysis 27.10. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 10 Meconium-stained liquor.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 10 Meconium-stained liquor

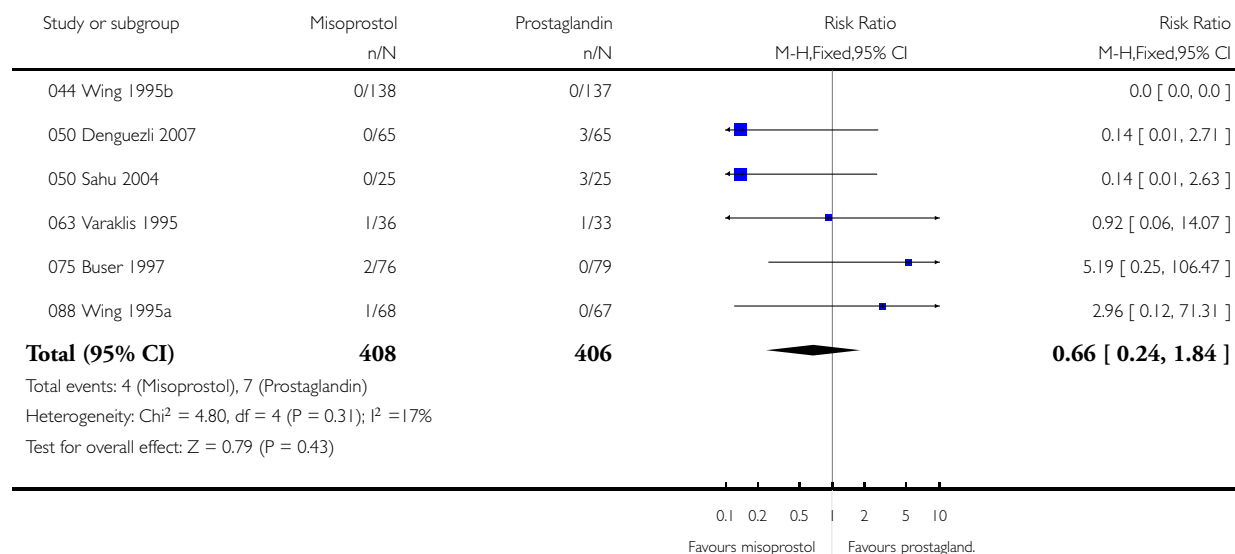


# **Analysis 27.11. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 11 Apgar score < 7 at 5 minutes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 11 Apgar score < 7 at 5 minutes

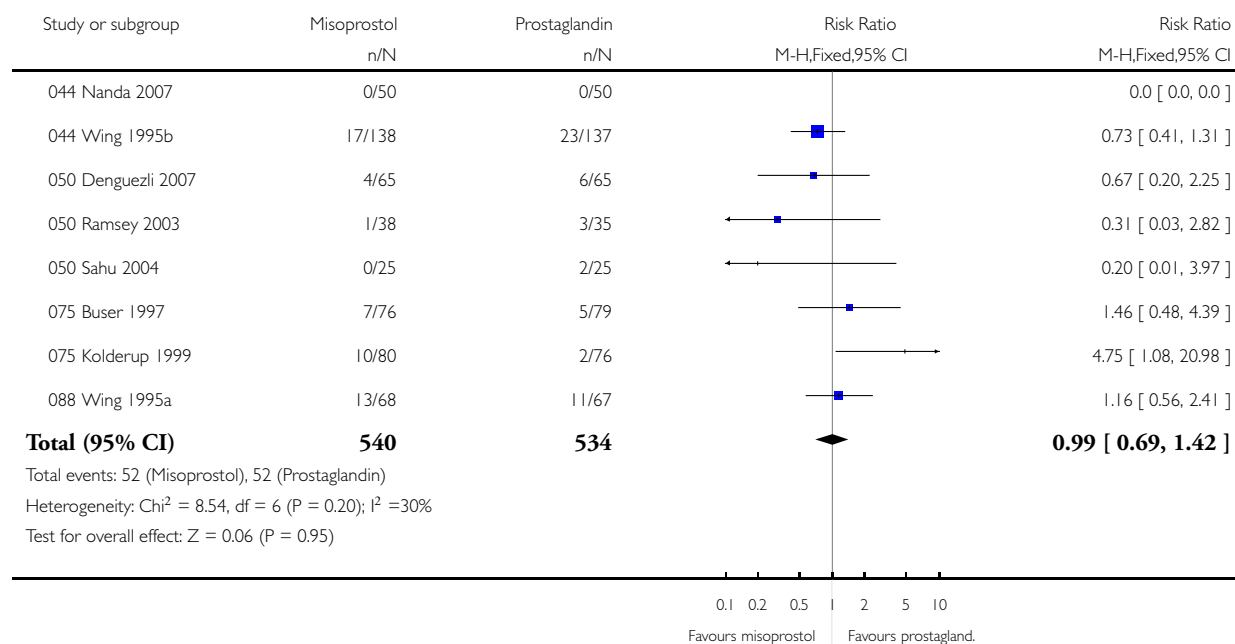


## Analysis 27.12. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 12 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 12 Neonatal intensive care unit admission



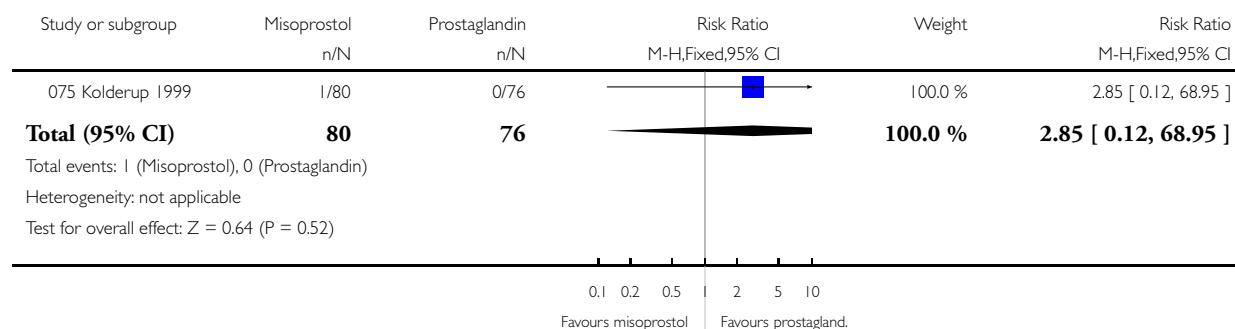


### Analysis 27.13. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 13 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 13 Perinatal death

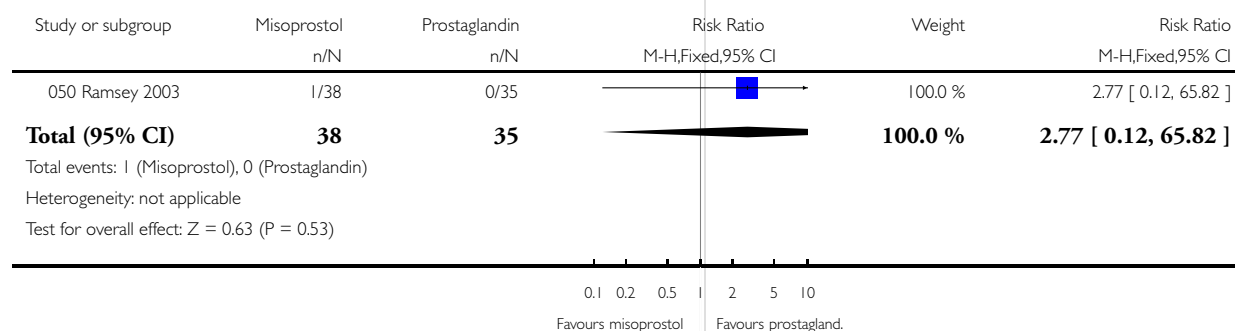


### Analysis 27.14. Comparison 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix, Outcome 14 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 27 Misoprostol versus intracervical prostaglandin: all women with intact membranes and unfavourable cervix

Outcome: 14 Postpartum haemorrhage

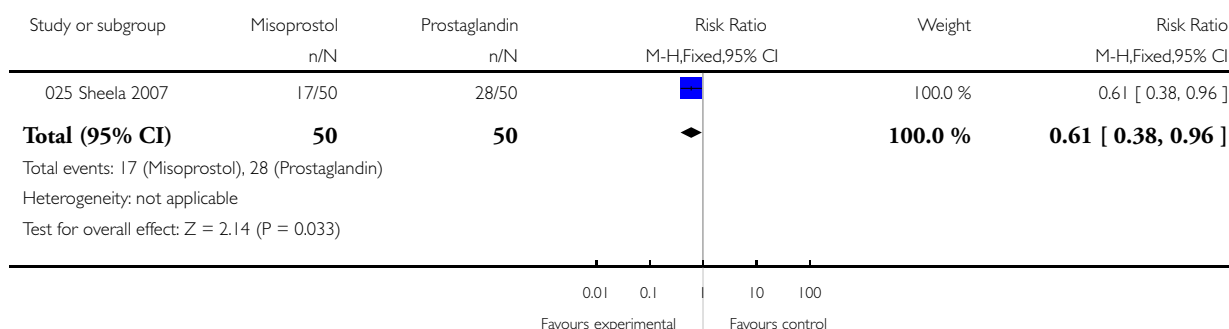


### Analysis 28.1. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 1 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 1 Vaginal delivery not achieved in 24 hours

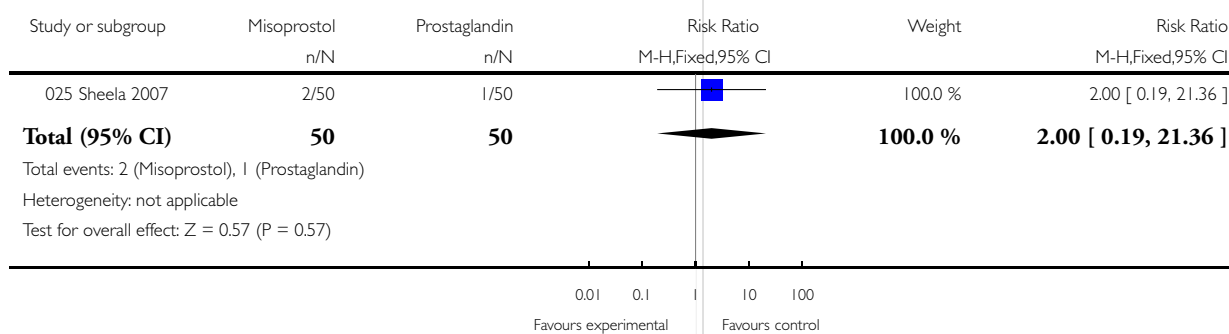


### Analysis 28.2. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

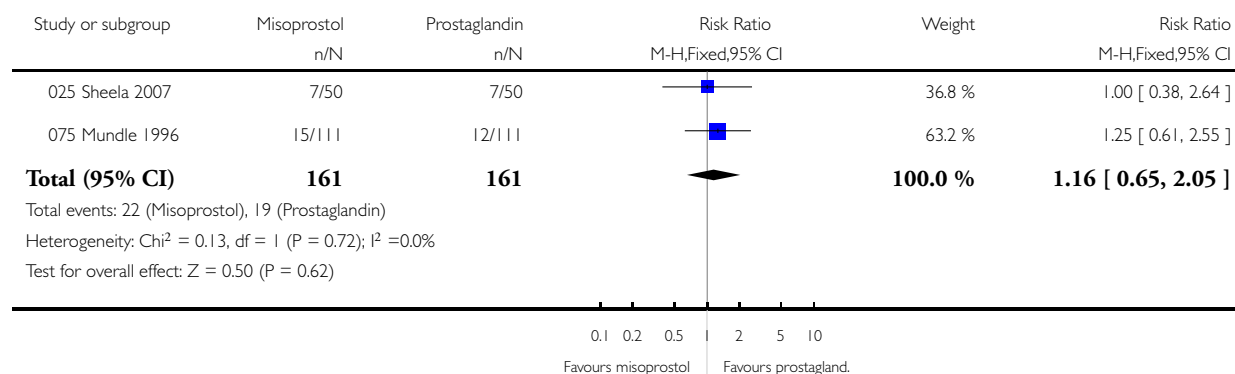


### Analysis 28.3. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 3 Caesarean section

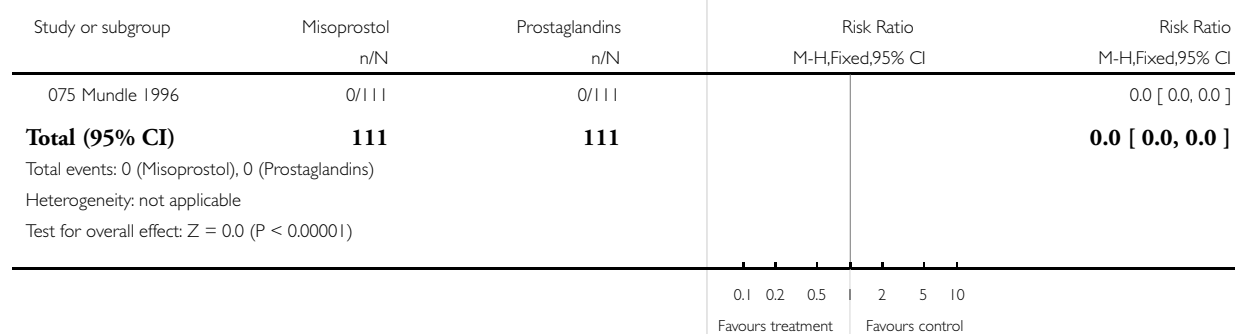


### Analysis 28.4. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 4 Serious neonatal morbidity/perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 4 Serious neonatal morbidity/perinatal death

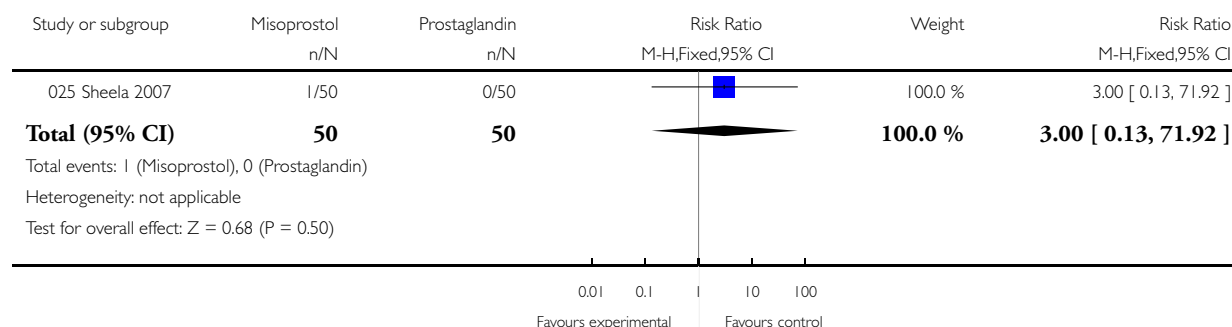


### Analysis 28.5. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 5 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 5 Neonatal intensive care unit admission

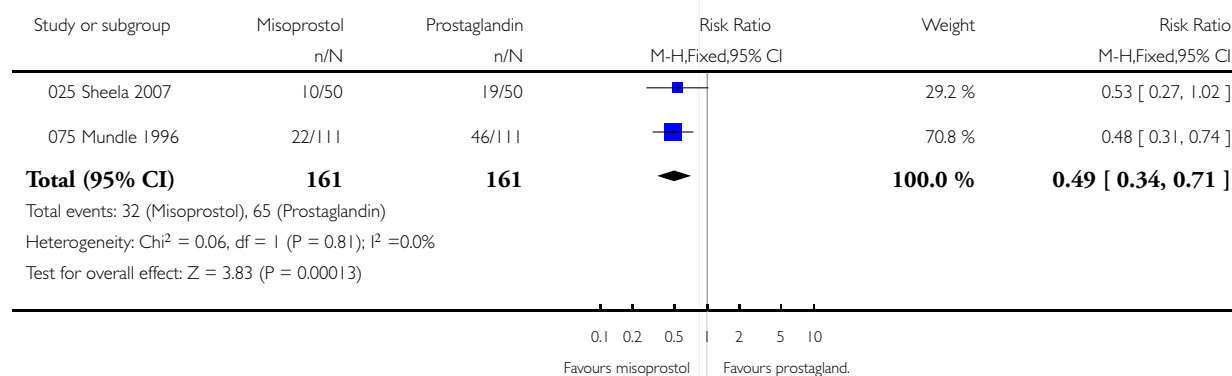


### Analysis 28.6. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 6 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 6 Oxytocin augmentation

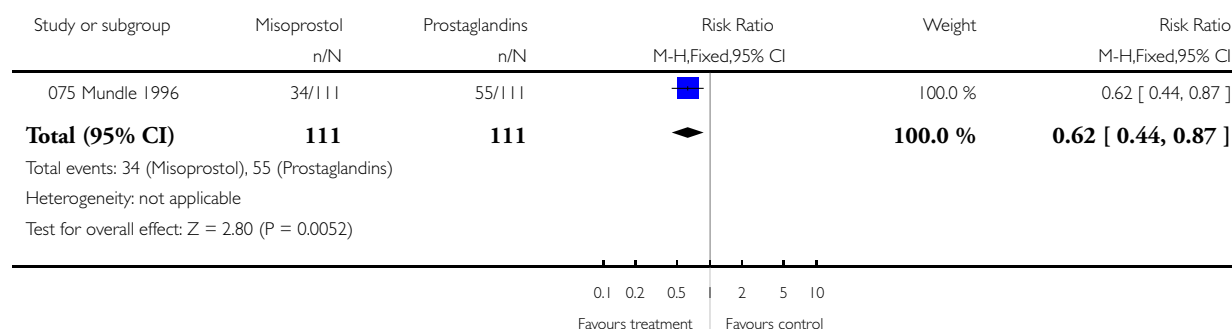


### Analysis 28.7. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 7 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 7 Epidural analgesia

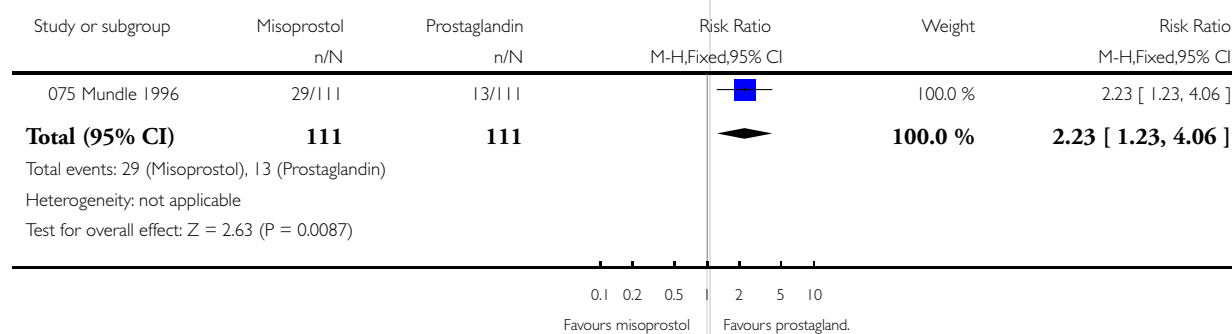


### Analysis 28.8. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 8 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 8 Instrumental vaginal delivery

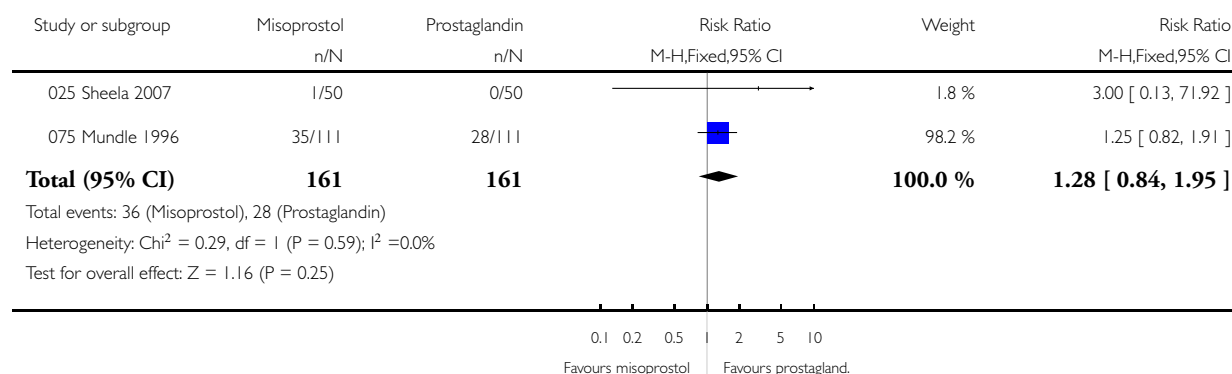


### Analysis 28.9. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 9 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 9 Meconium-stained liquor

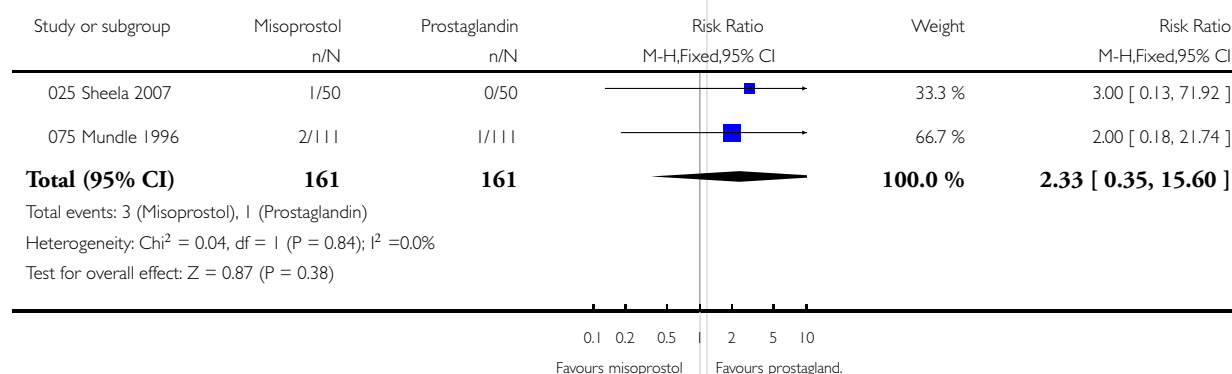


### Analysis 28.10. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 10 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 10 Apgar score < 7 at 5 minutes

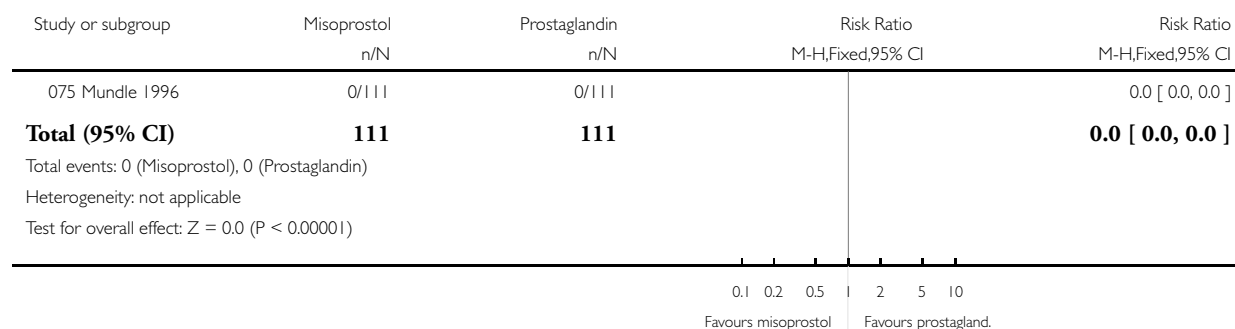


### Analysis 28.11. Comparison 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix, Outcome 11 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 28 Misoprostol versus intracervical prostaglandin: all women with intact membranes and variable or undefined cervix

Outcome: 11 Perinatal death

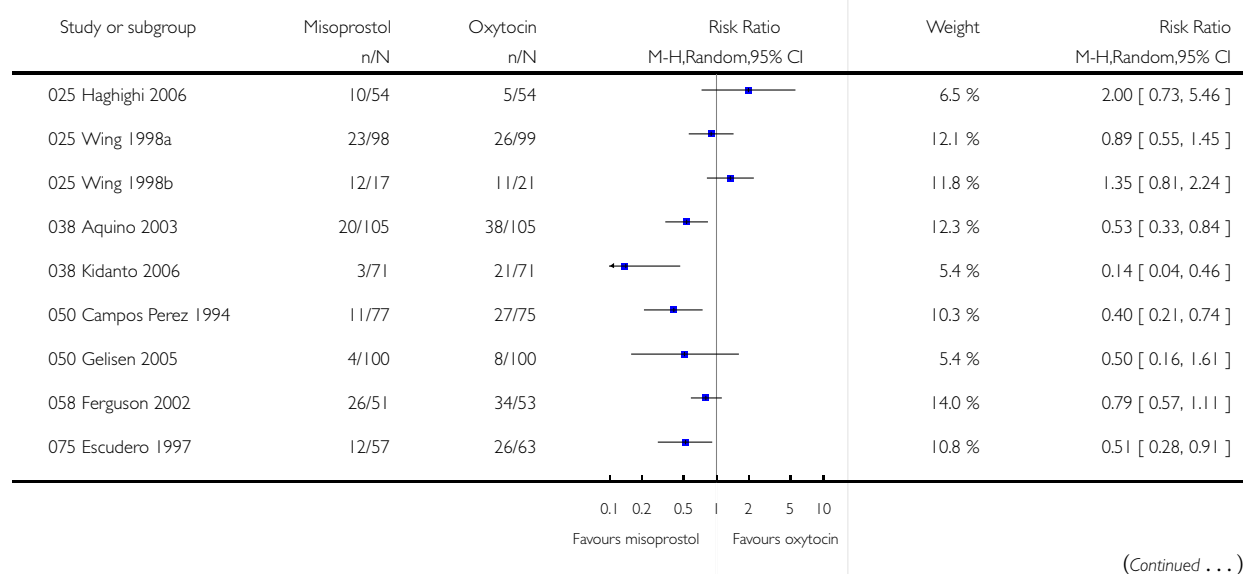


### Analysis 29.1. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 1 Vaginal delivery not achieved within 24 hours.

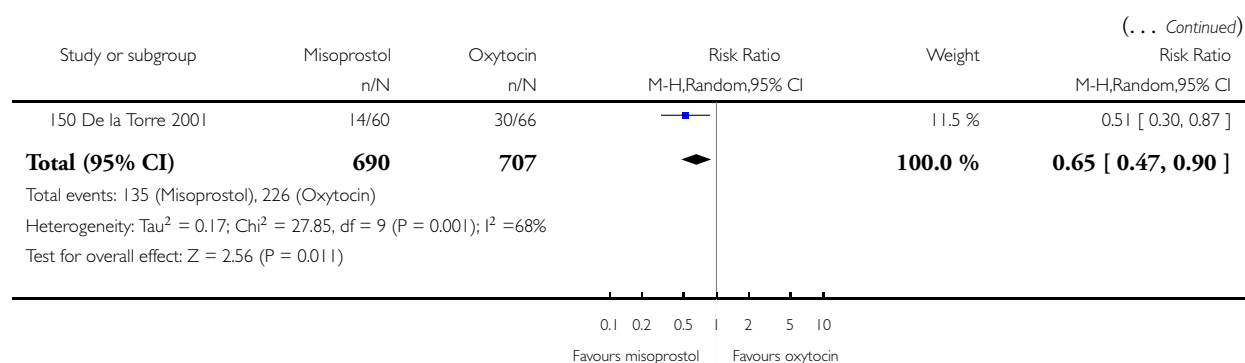
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 1 Vaginal delivery not achieved within 24 hours



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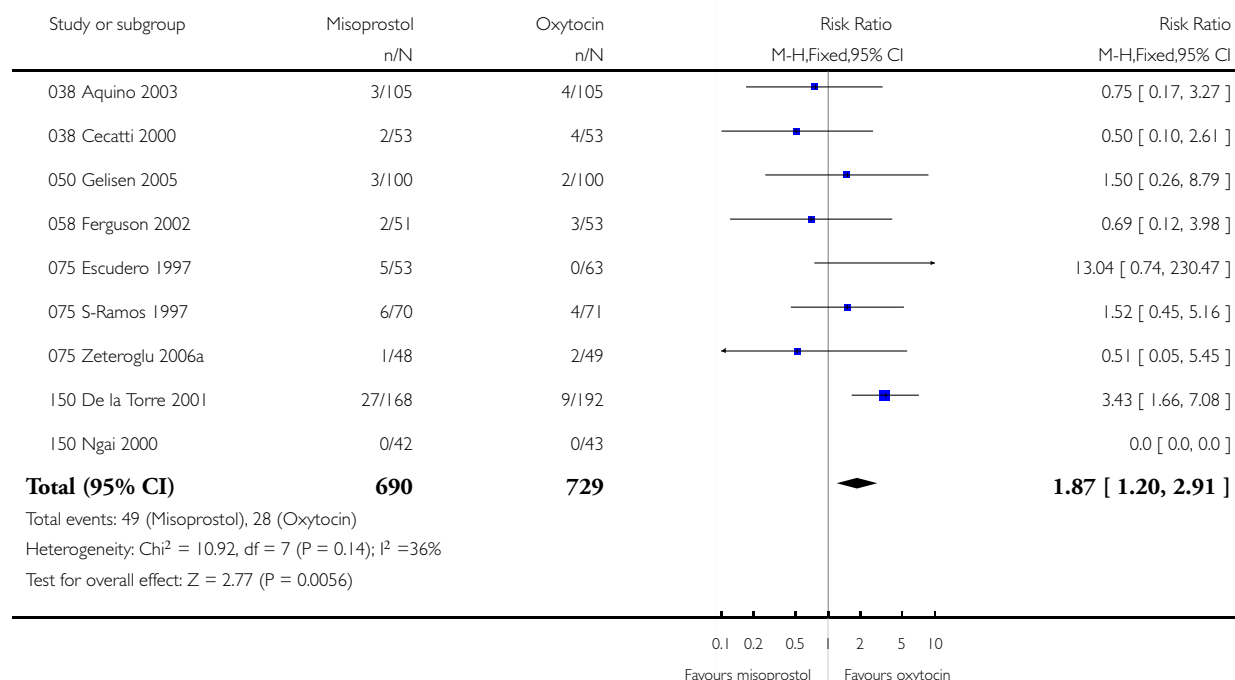


## Analysis 29.2. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 2 Uterine hyperstimulation with FHR changes



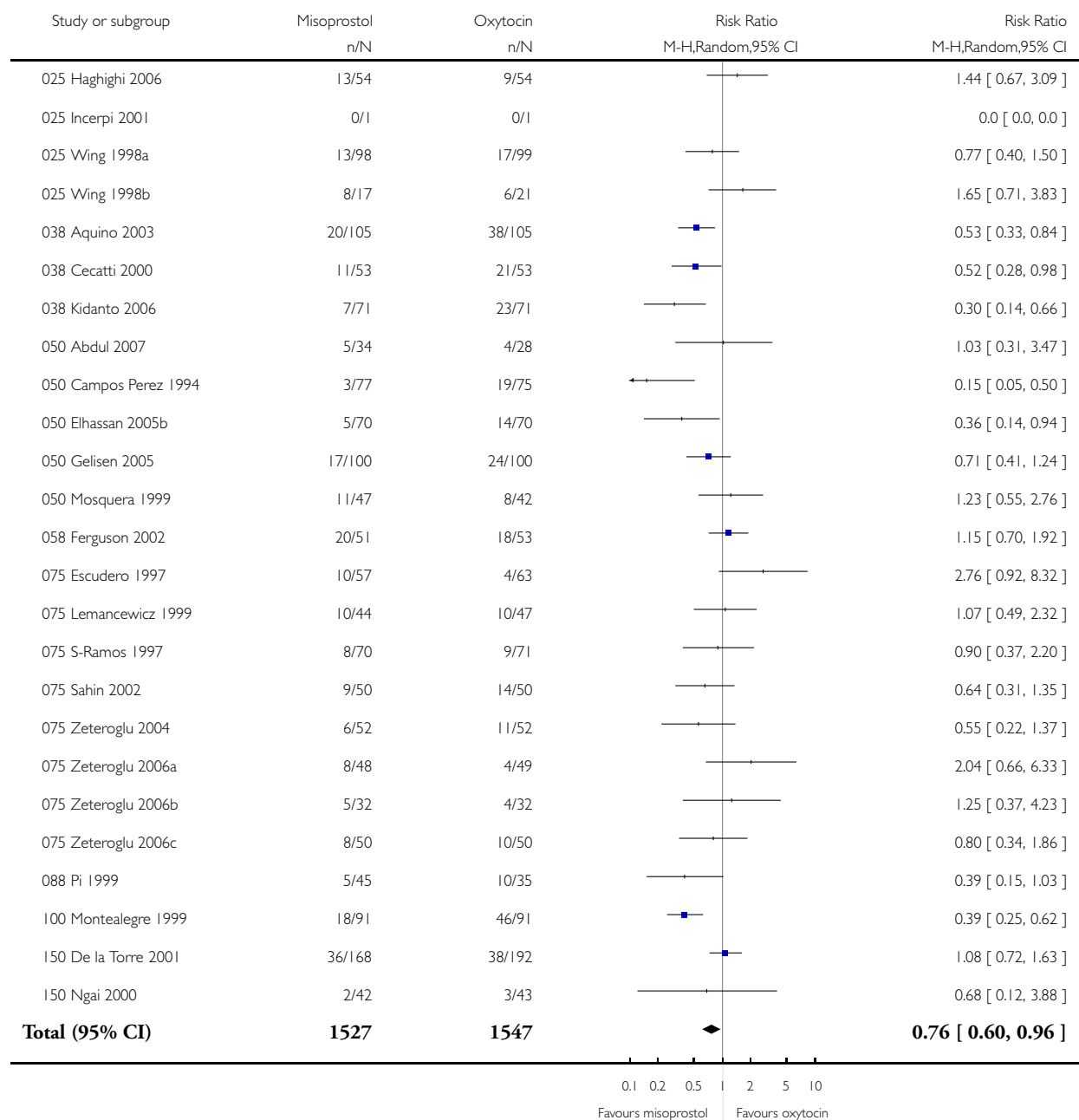


### Analysis 29.3. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 3 Caesarean section.

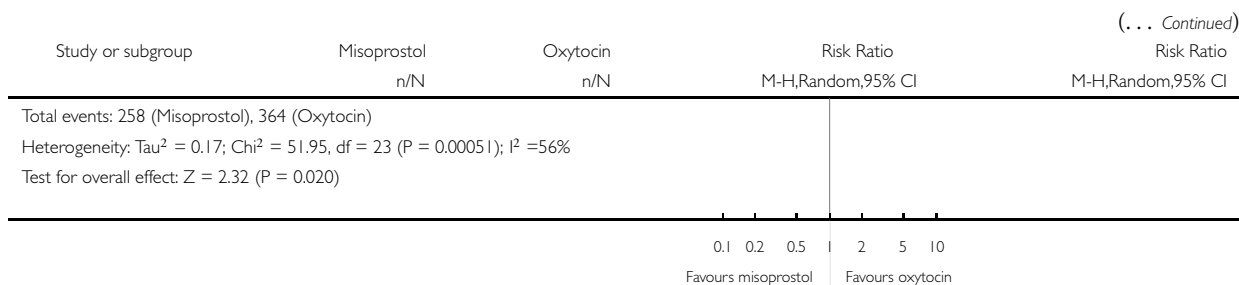
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 3 Caesarean section



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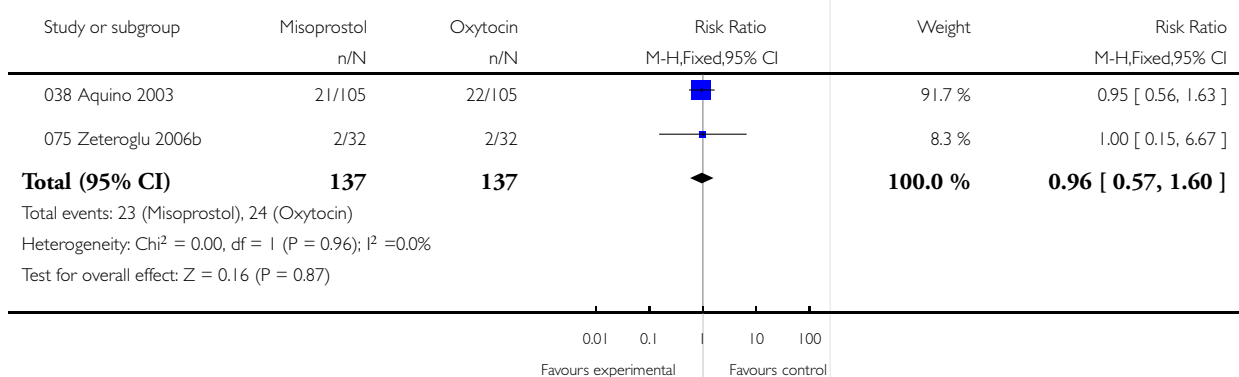


#### Analysis 29.4. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 4 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 4 Serious neonatal morbidity

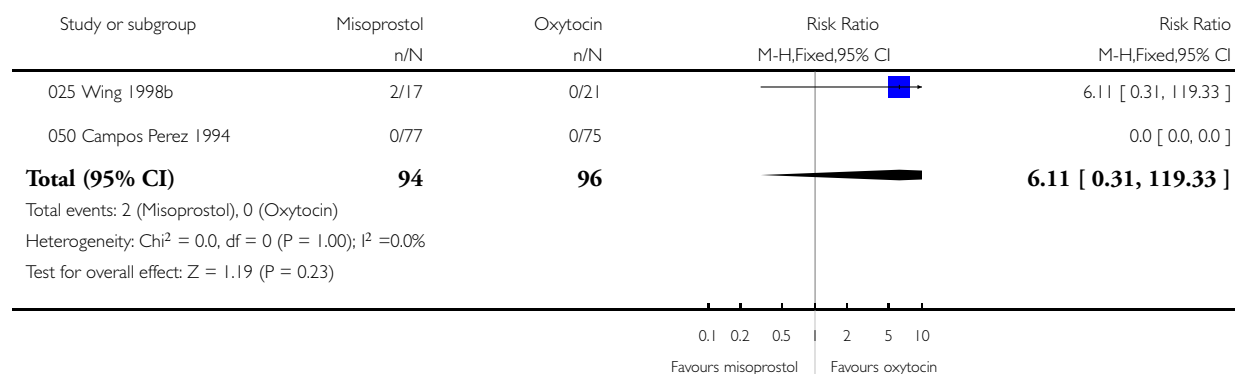


### Analysis 29.5. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 5 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 5 Serious maternal morbidity or death

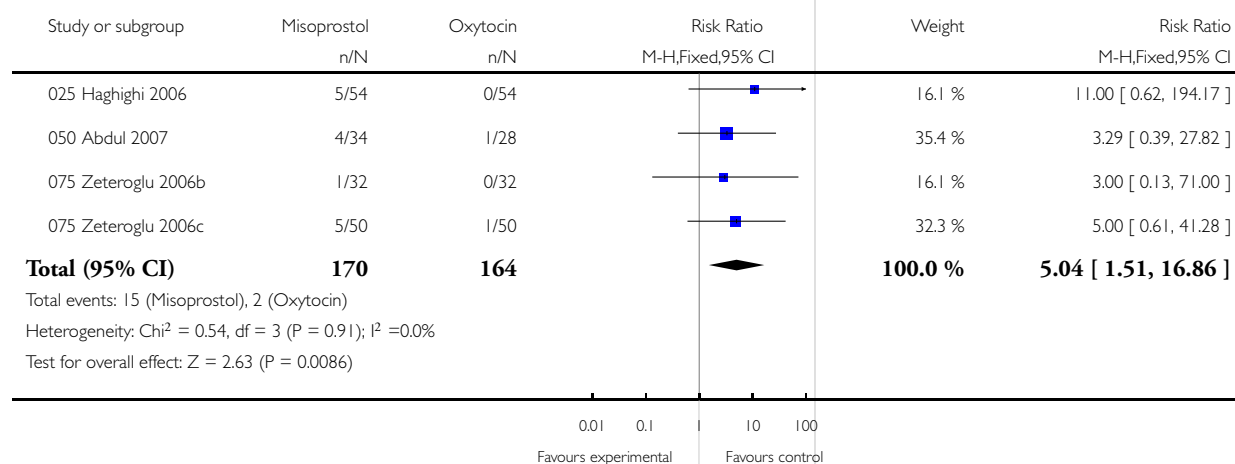


### Analysis 29.6. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 6 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 6 Maternal side effects (eg nausea, vomiting, diarrhoea)

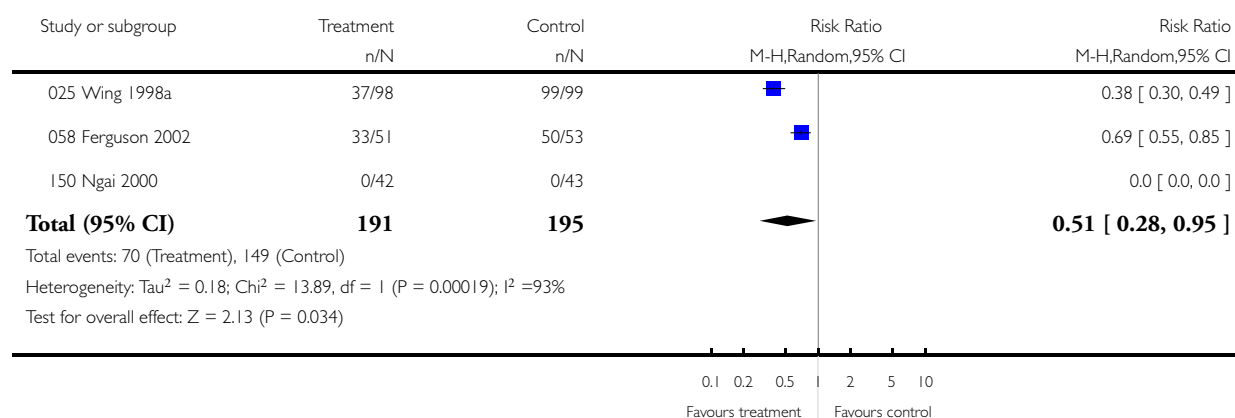


## Analysis 29.7. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 7 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 7 Oxytocin augmentation

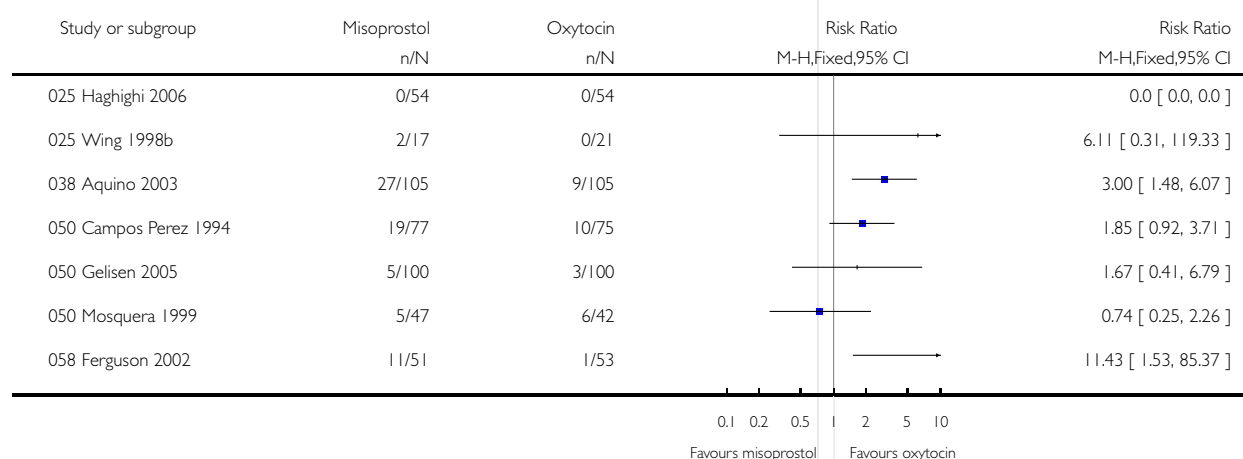


## Analysis 29.8. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 8 Uterine hyperstimulation without FHR changes.

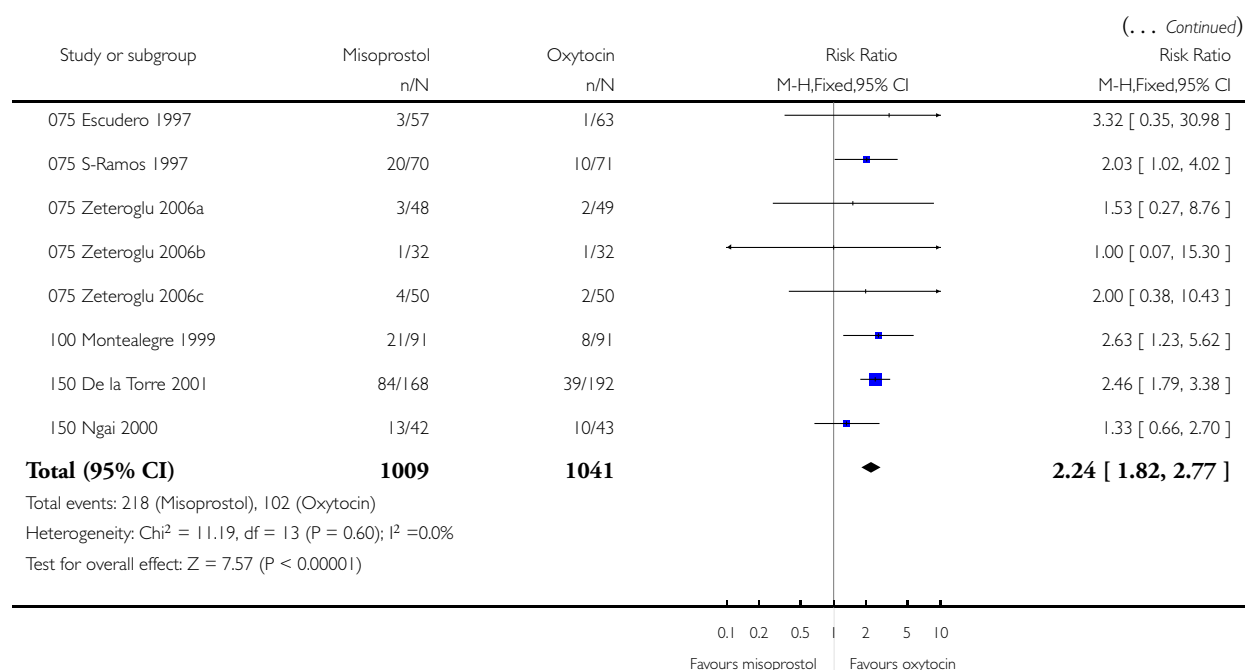
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 8 Uterine hyperstimulation without FHR changes



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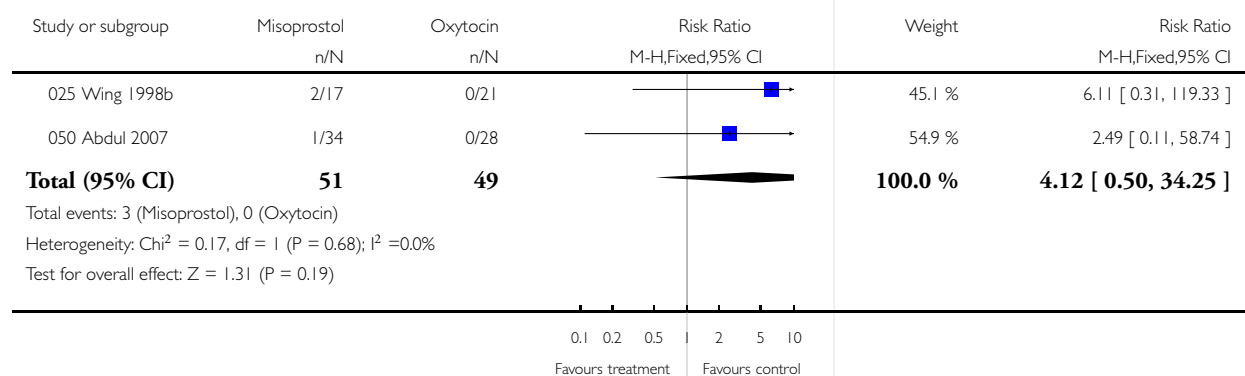


### Analysis 29.9. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 9 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 9 Uterine rupture

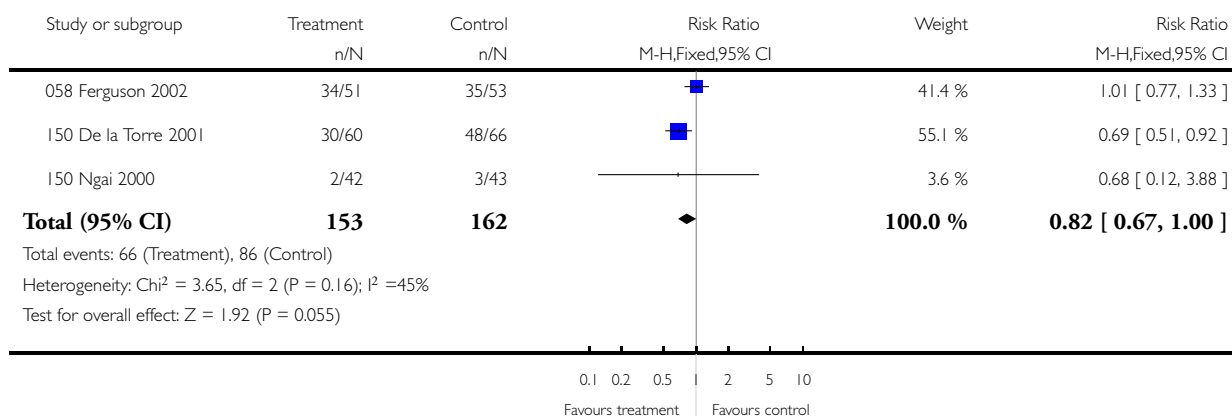


### Analysis 29.10. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 10 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 10 Epidural analgesia

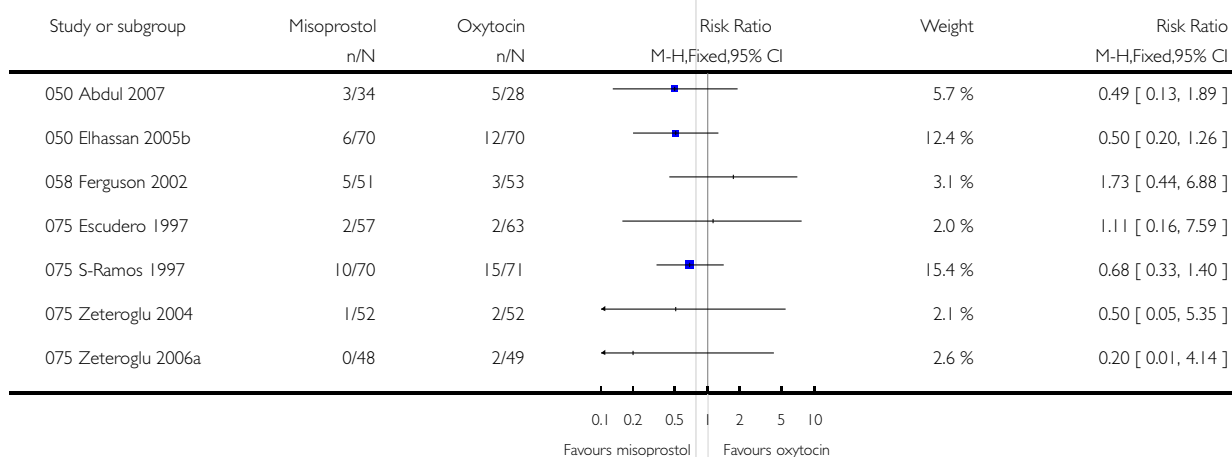


### Analysis 29.11. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 11 Instrumental vaginal delivery.

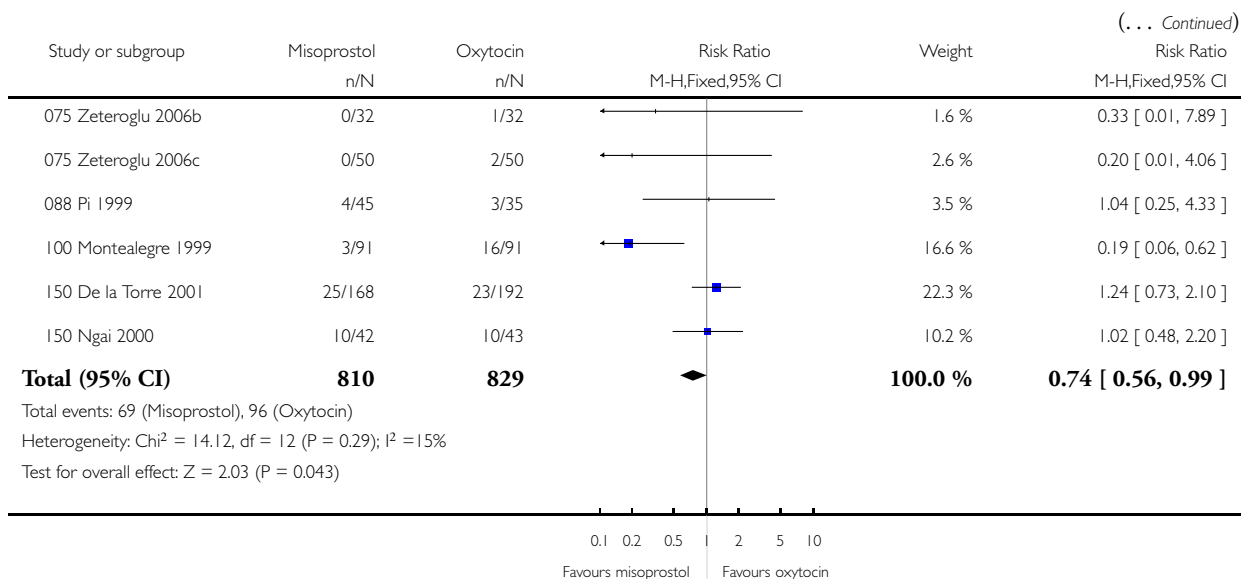
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 11 Instrumental vaginal delivery



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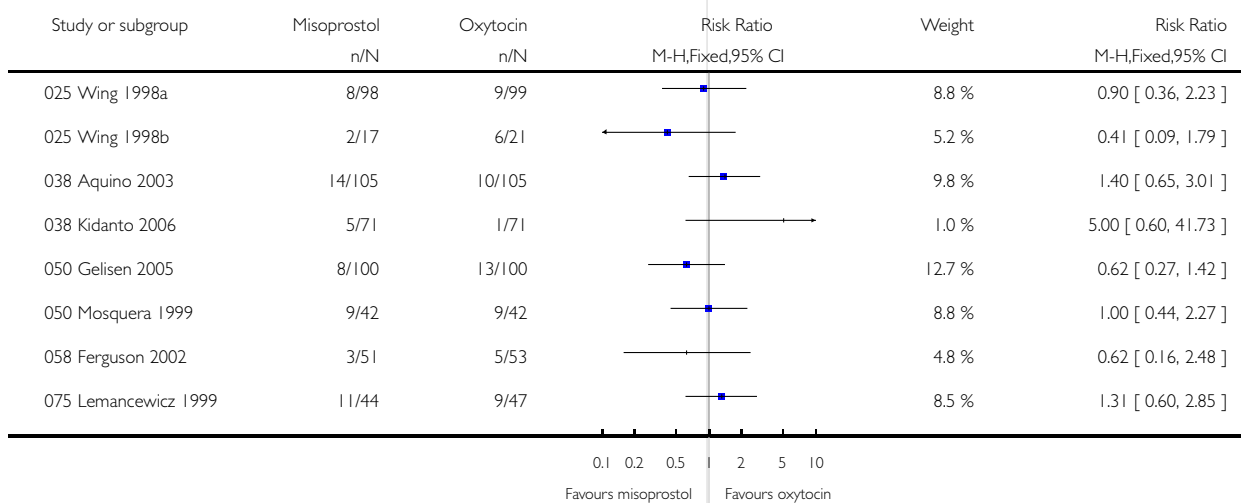


### Analysis 29.12. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 12 Meconium-stained liquor.

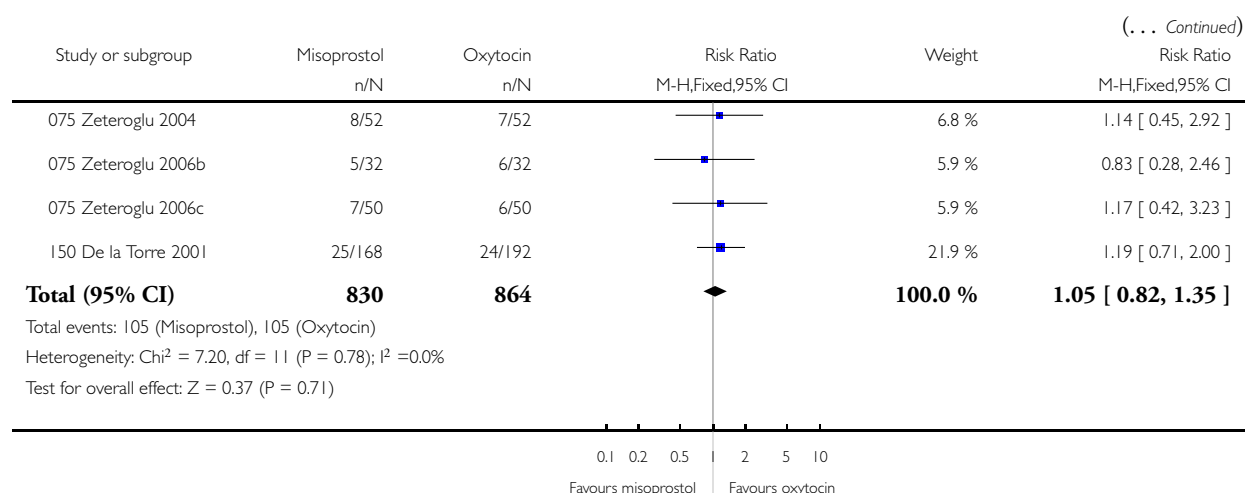
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 12 Meconium-stained liquor



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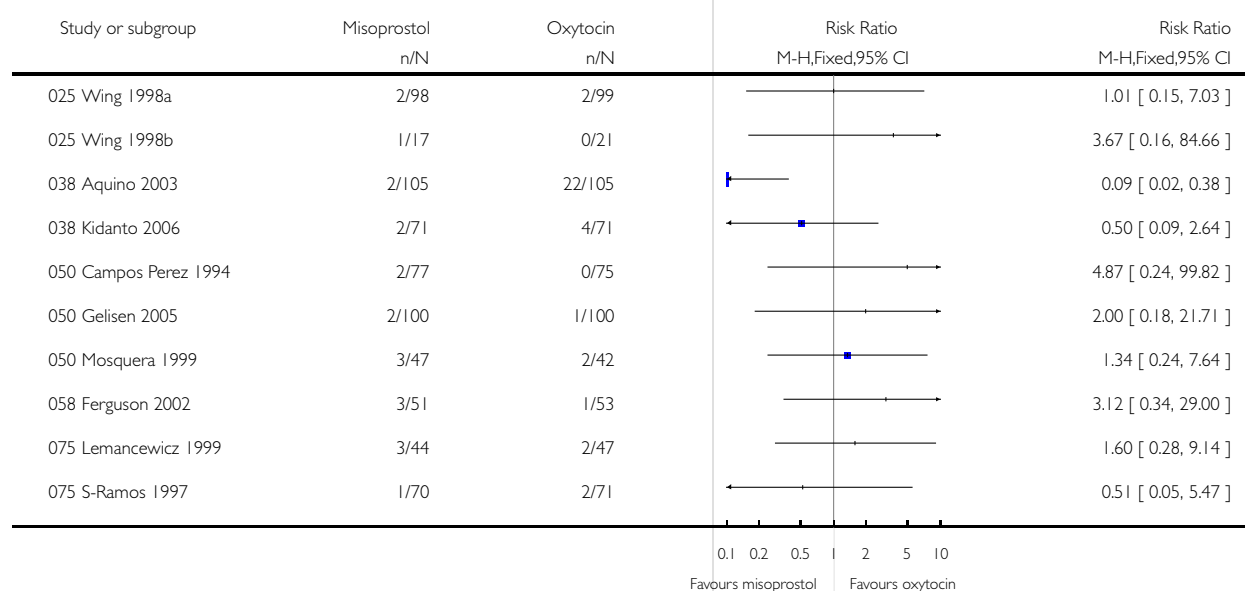


### Analysis 29.13. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 13 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

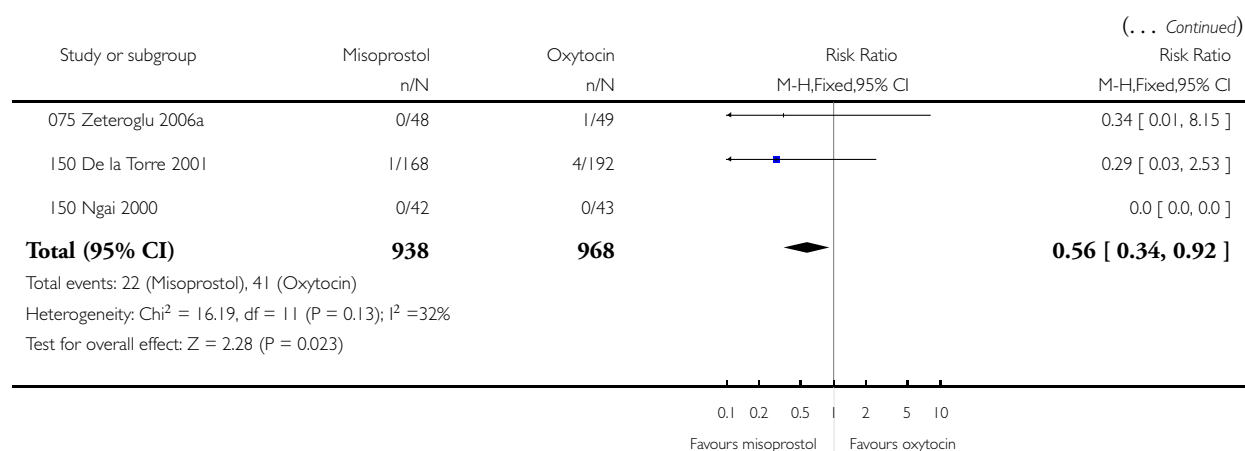
Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 13 Apgar score < 7 at 5 minutes



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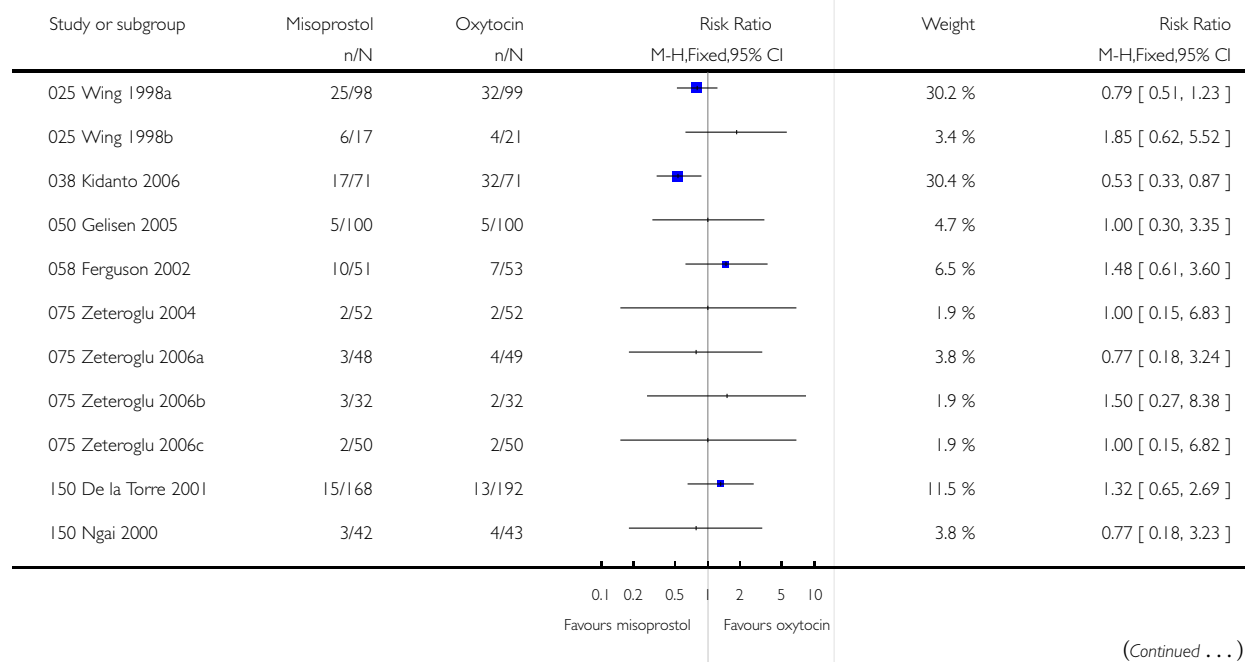


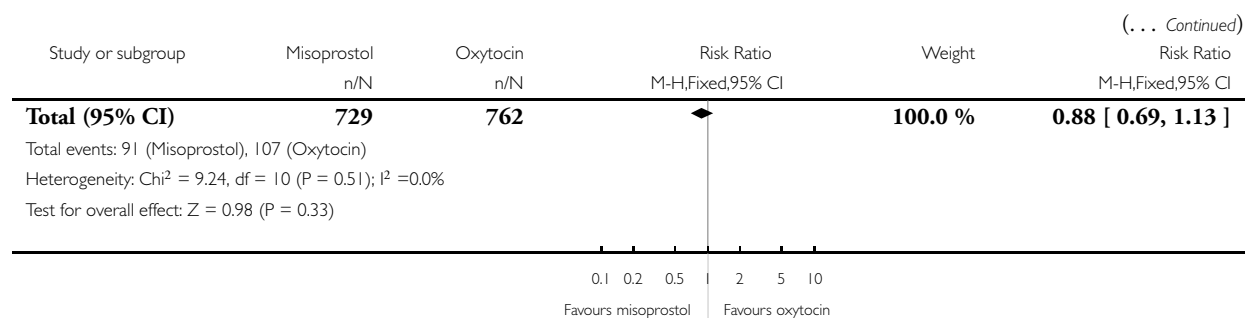
#### Analysis 29.14. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 14 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 14 Neonatal intensive care unit admission



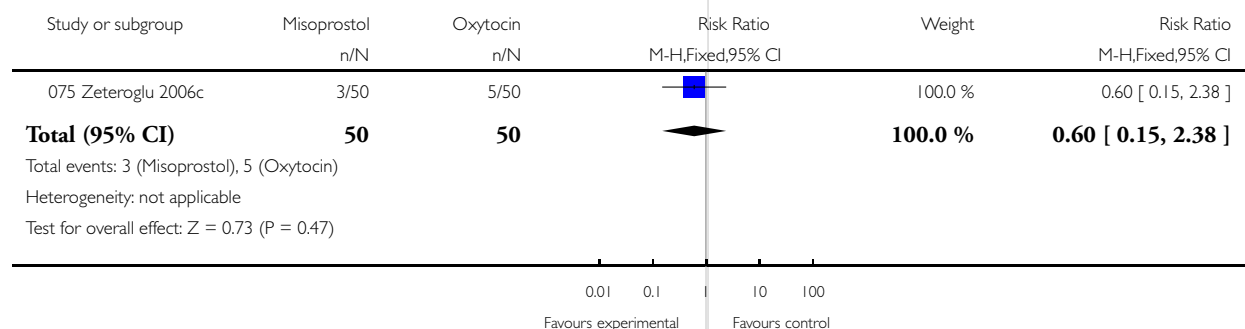


### Analysis 29.15. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 15 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 15 Serious maternal complications

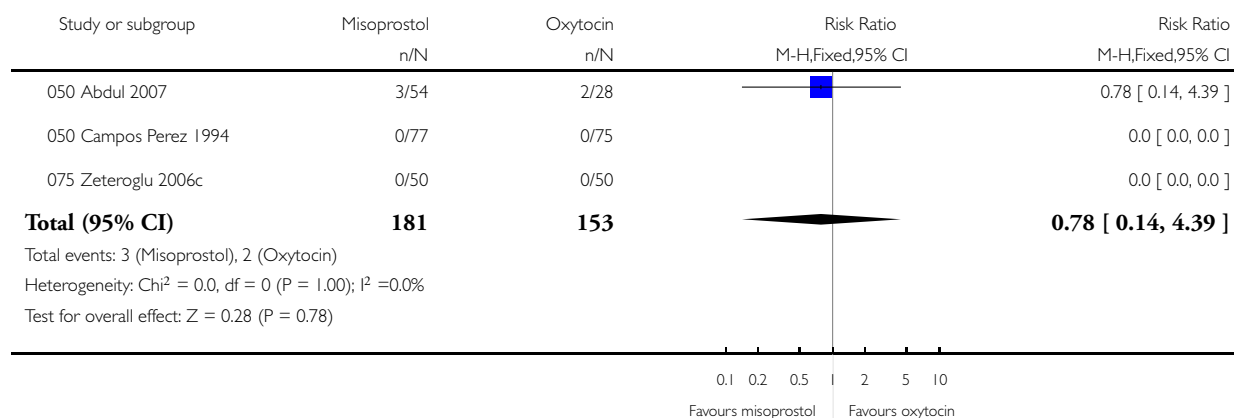


### Analysis 29.16. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 16 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 16 Perinatal death

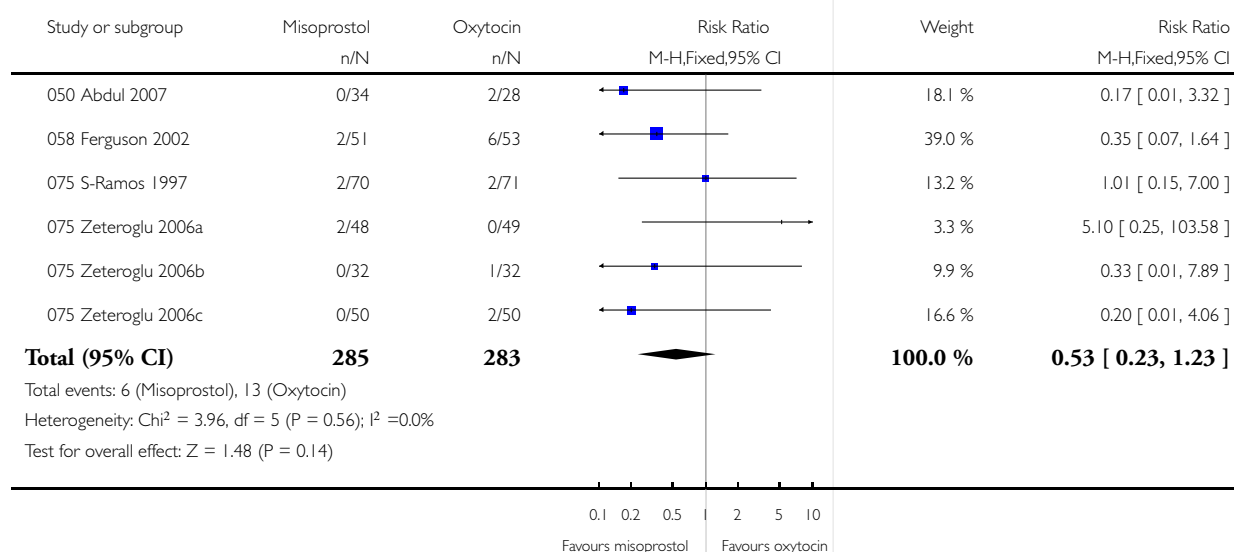


### Analysis 29.17. Comparison 29 Misoprostol versus oxytocin: all women, Outcome 17 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 29 Misoprostol versus oxytocin: all women

Outcome: 17 Postpartum haemorrhage

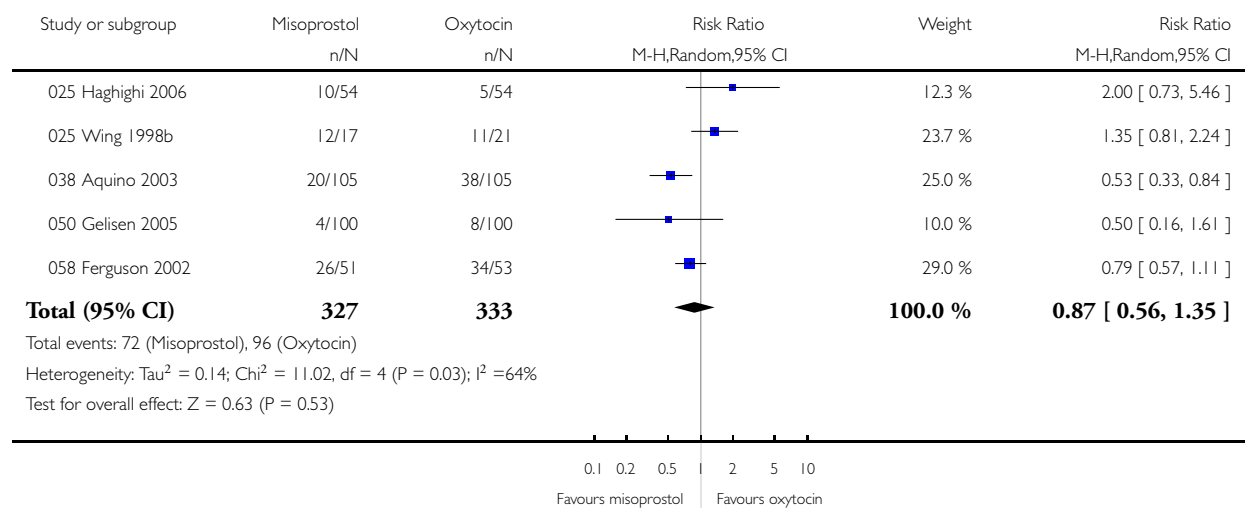


**Analysis 30.1. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

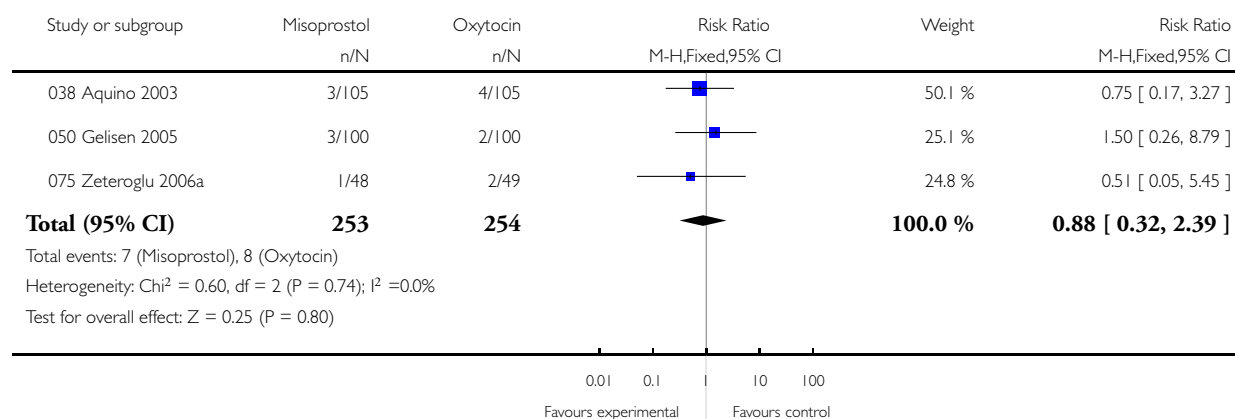


## Analysis 30.2. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

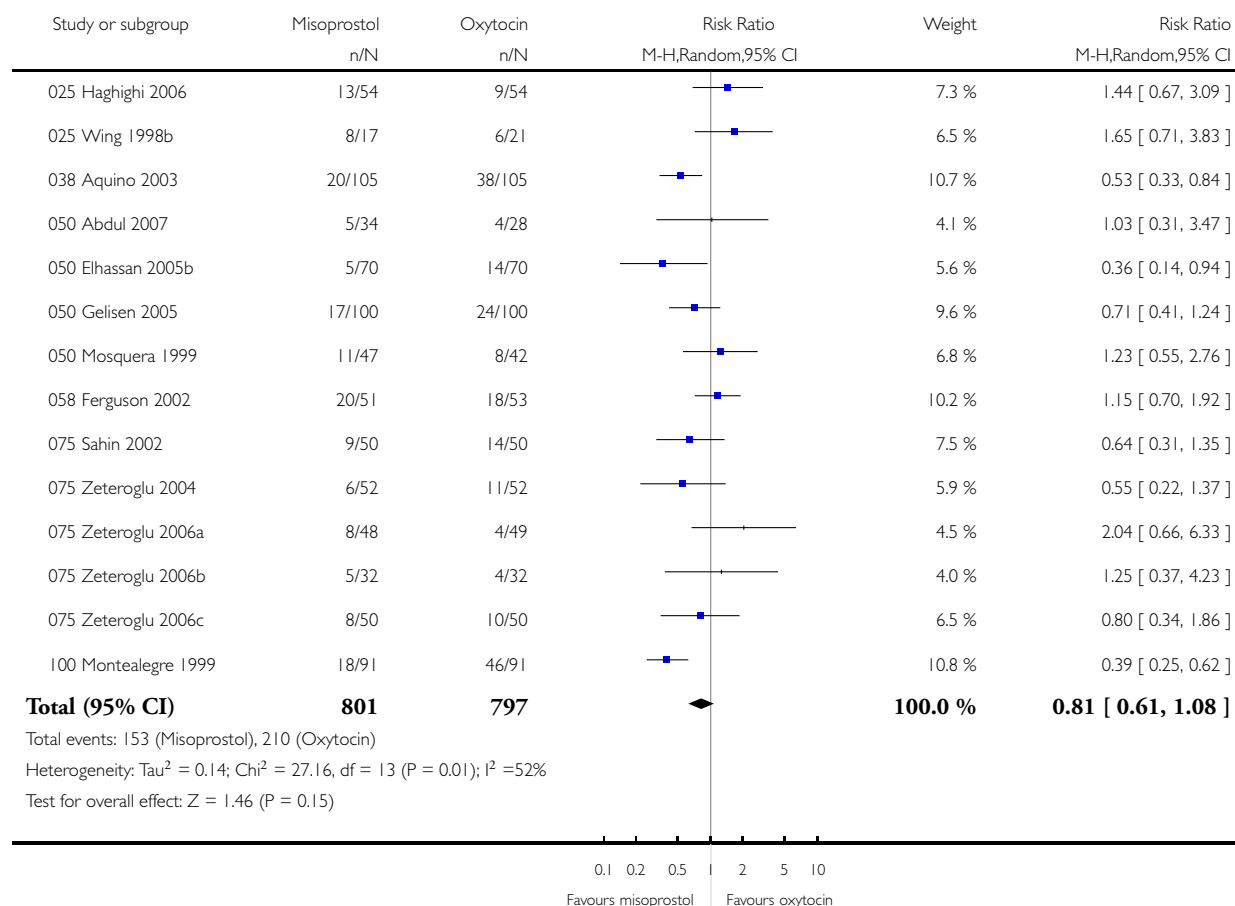


### Analysis 30.3. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 3 Caesarean section

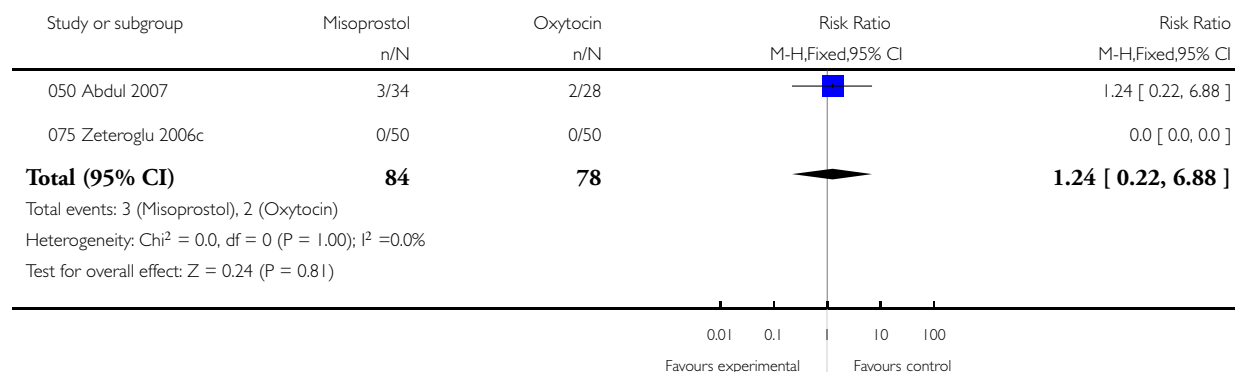


### Analysis 30.4. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 4 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 4 Perinatal death

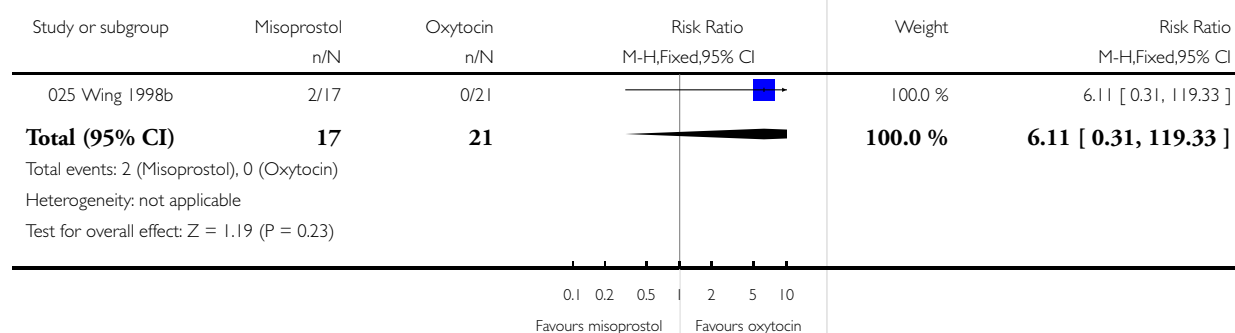


### Analysis 30.5. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 5 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 5 Serious maternal morbidity or death

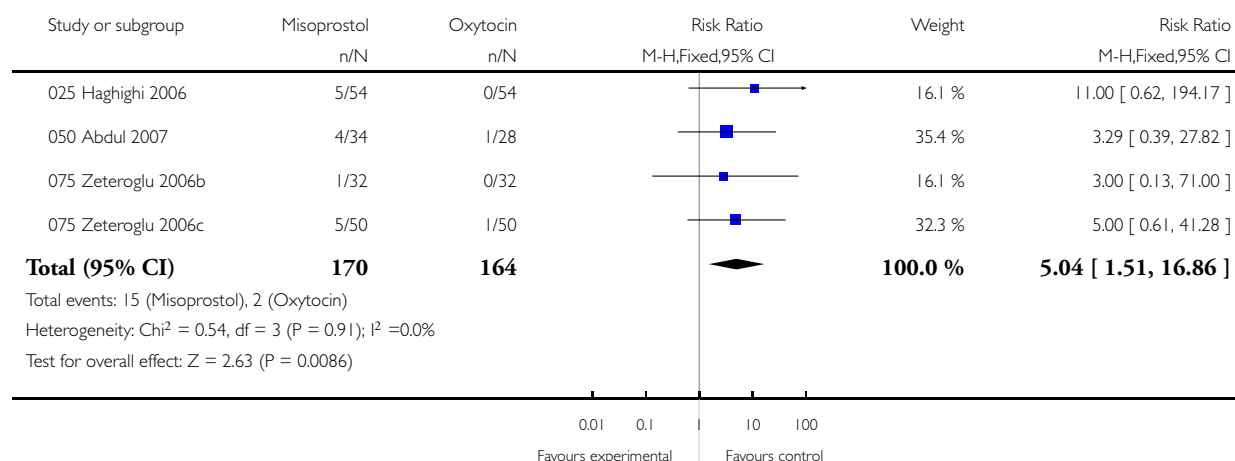


### Analysis 30.6. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 6 Maternal side effects.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 6 Maternal side effects

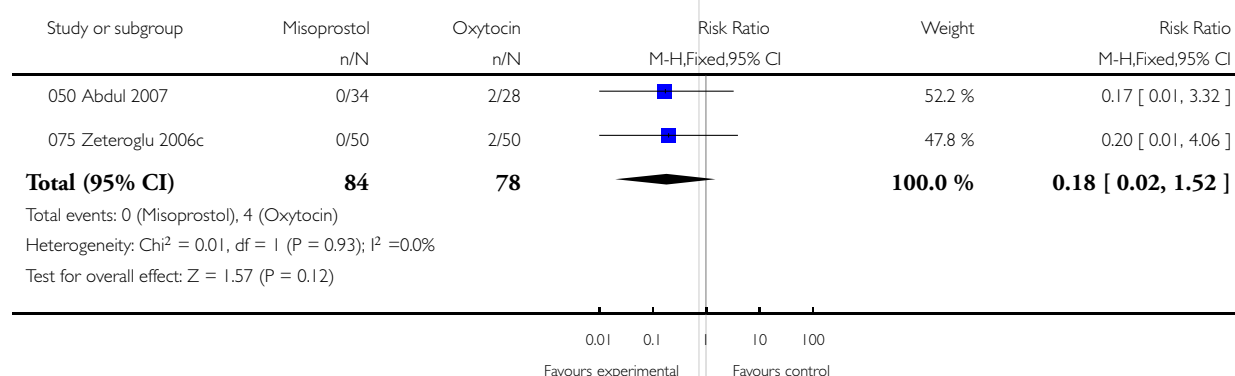


### Analysis 30.7. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 7 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 7 Postpartum haemorrhage



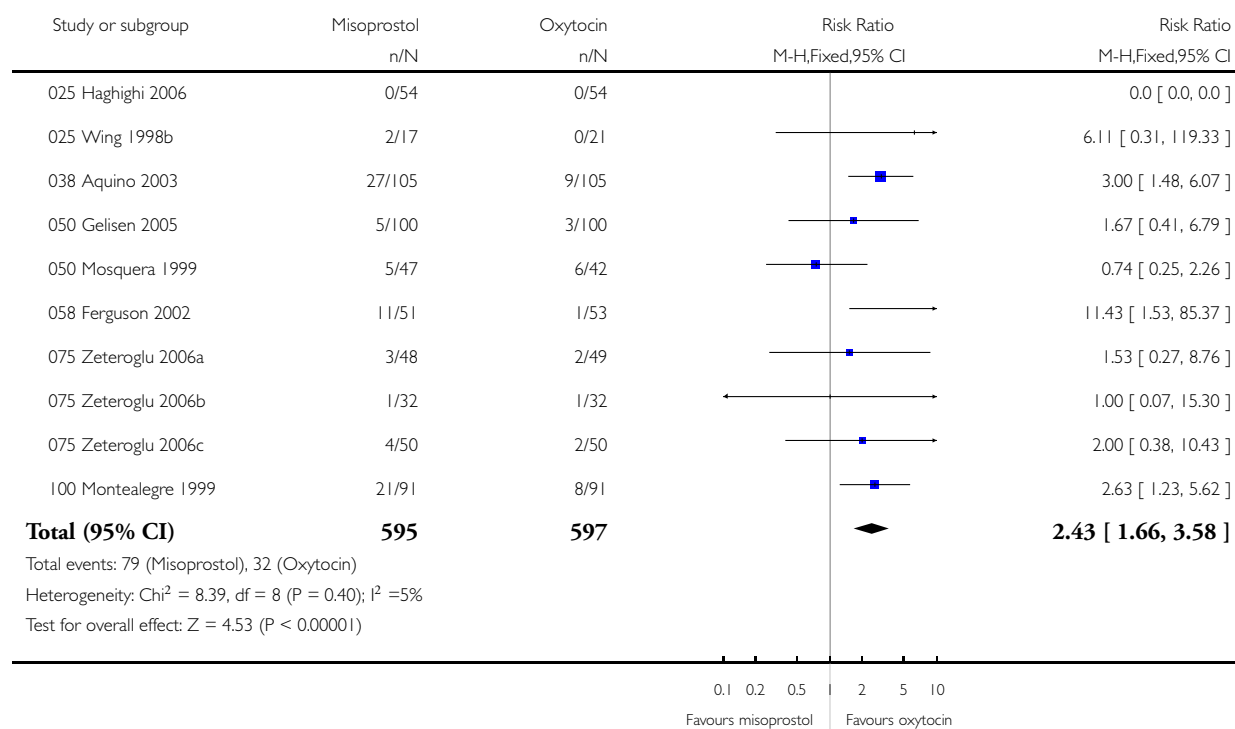


### Analysis 30.8. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 8 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 8 Uterine hyperstimulation without FHR changes

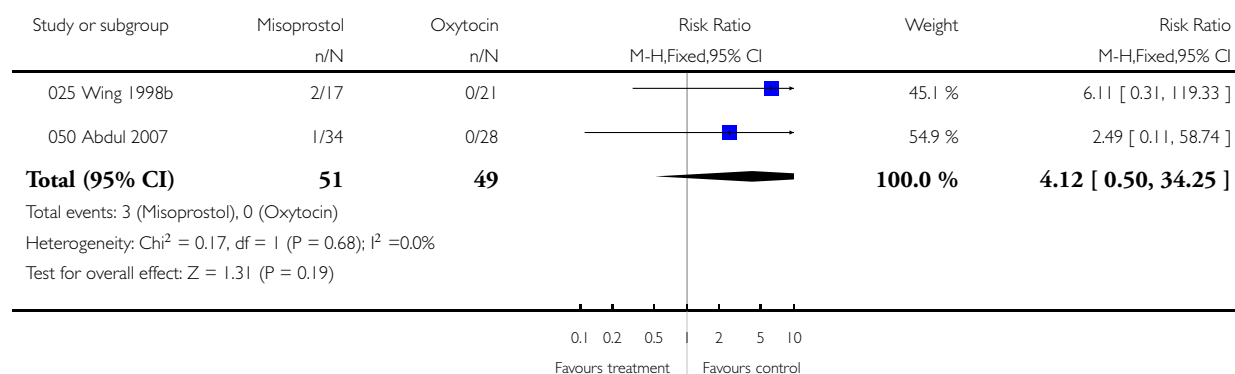


### Analysis 30.9. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 9 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 9 Uterine rupture

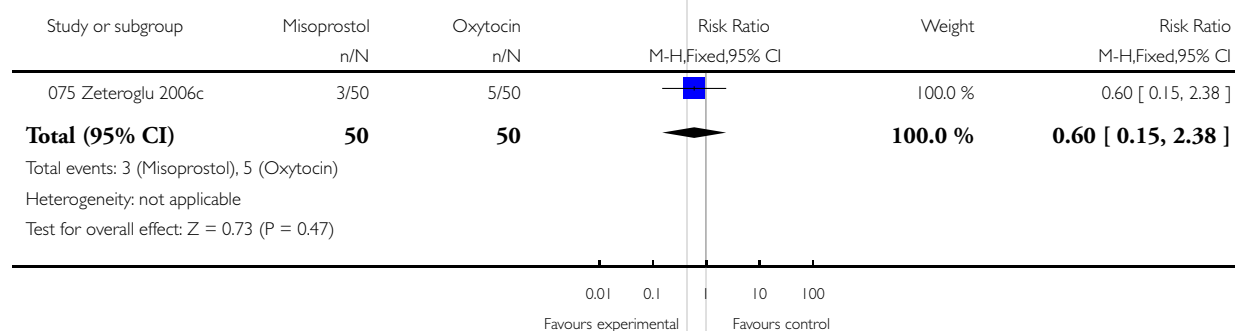


### Analysis 30.10. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 10 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 10 Serious maternal complications

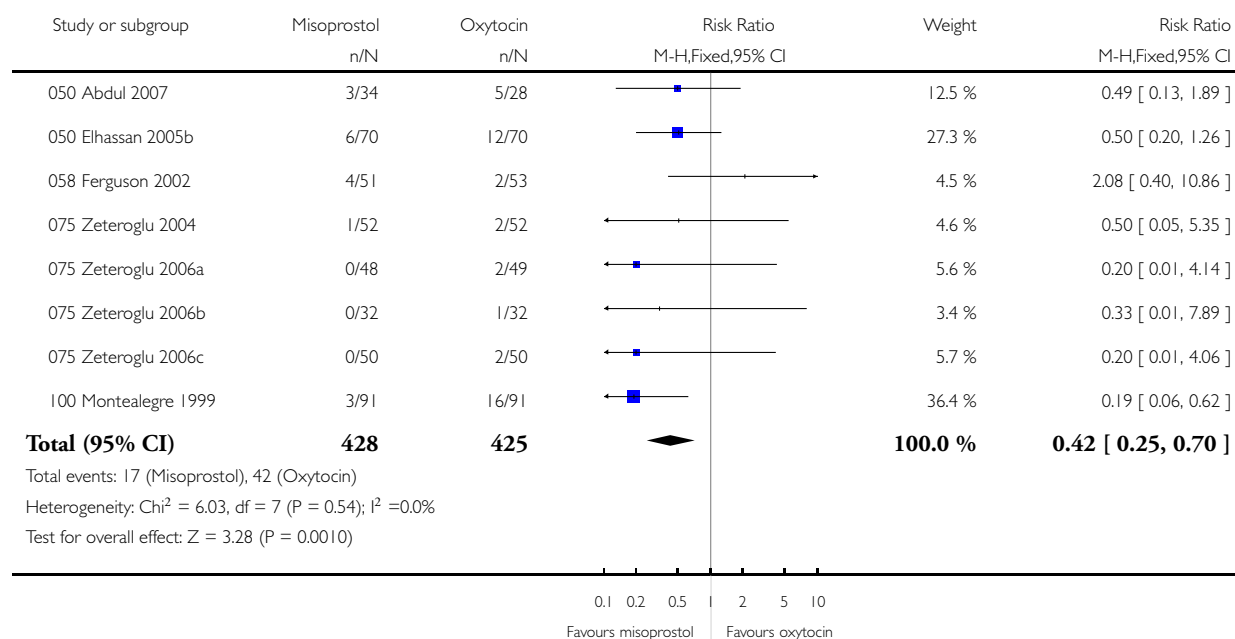


### Analysis 30.11. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 11 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 11 Instrumental vaginal delivery

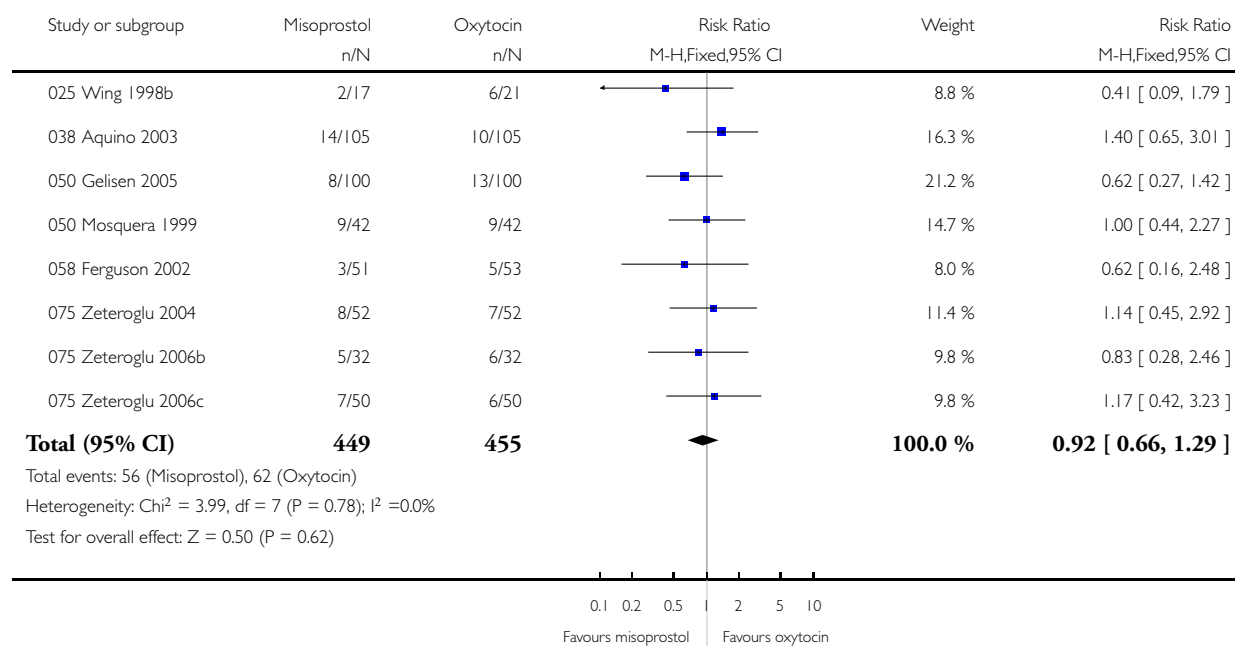


## Analysis 30.12. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 12 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 12 Meconium-stained liquor

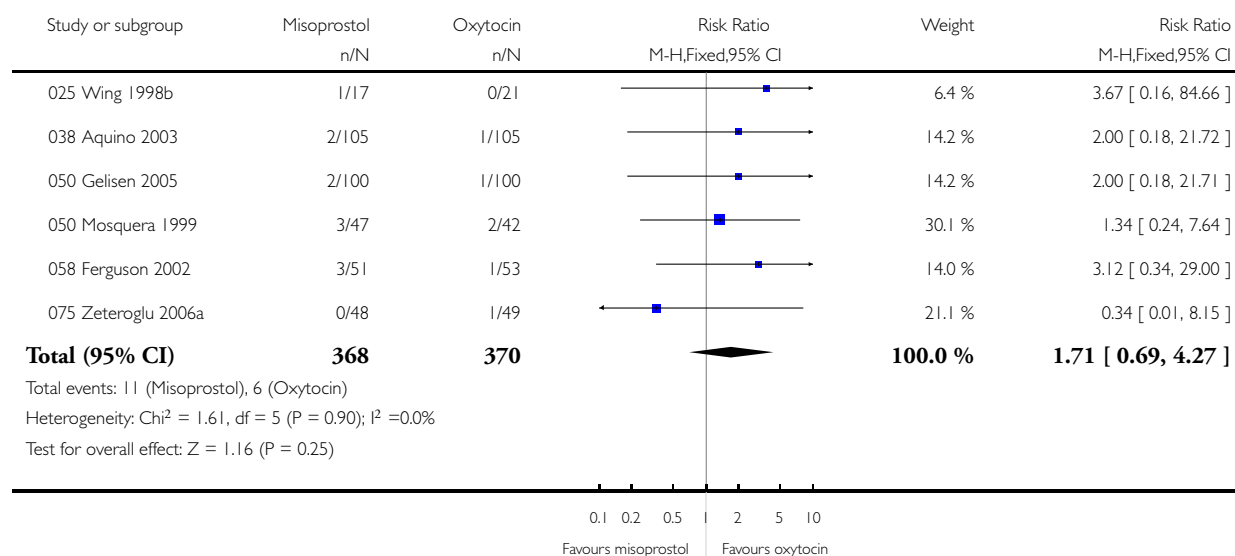


### Analysis 30.13. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 13 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 13 Apgar score < 7 at 5 minutes

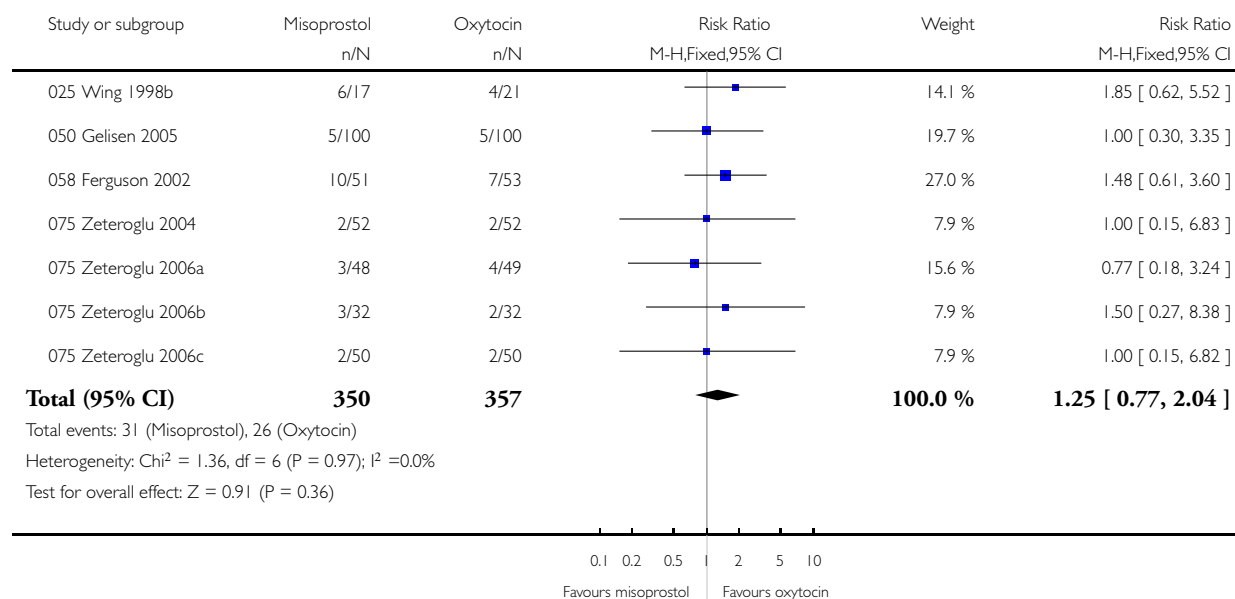


### Analysis 30.14. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 14 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 14 Neonatal intensive care unit admission

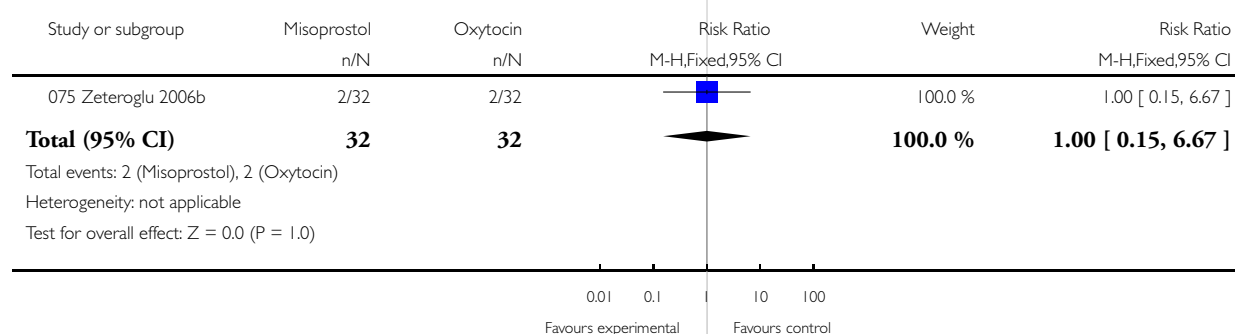


### Analysis 30.15. Comparison 30 Misoprostol versus oxytocin: all women with unfavourable cervix, Outcome 15 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 30 Misoprostol versus oxytocin: all women with unfavourable cervix

Outcome: 15 Serious neonatal morbidity

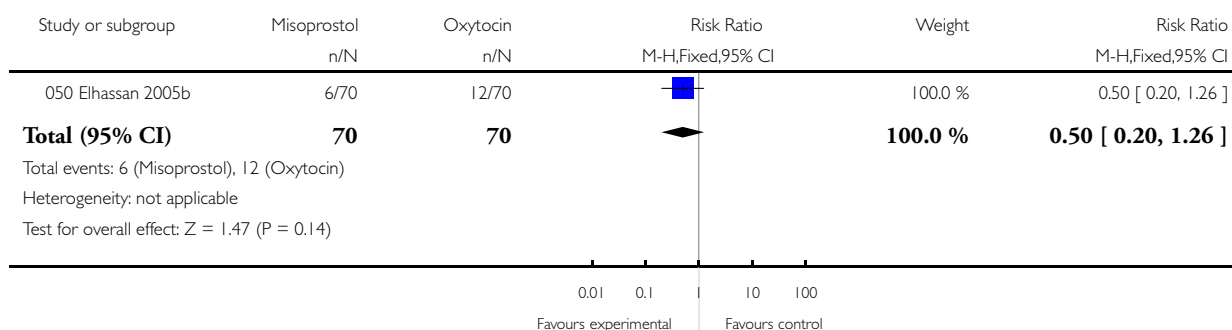


### Analysis 31.1. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 1 Instrumental delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 1 Instrumental delivery

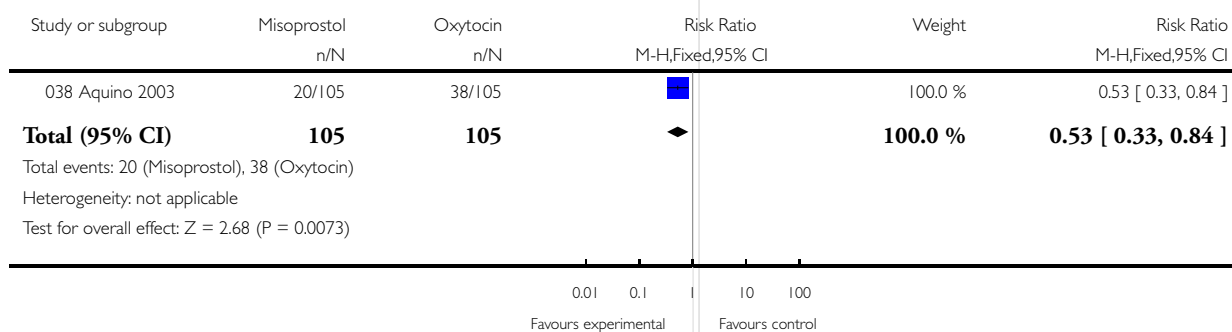


### Analysis 31.2. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 2 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 2 Vaginal delivery not achieved in 24 hours

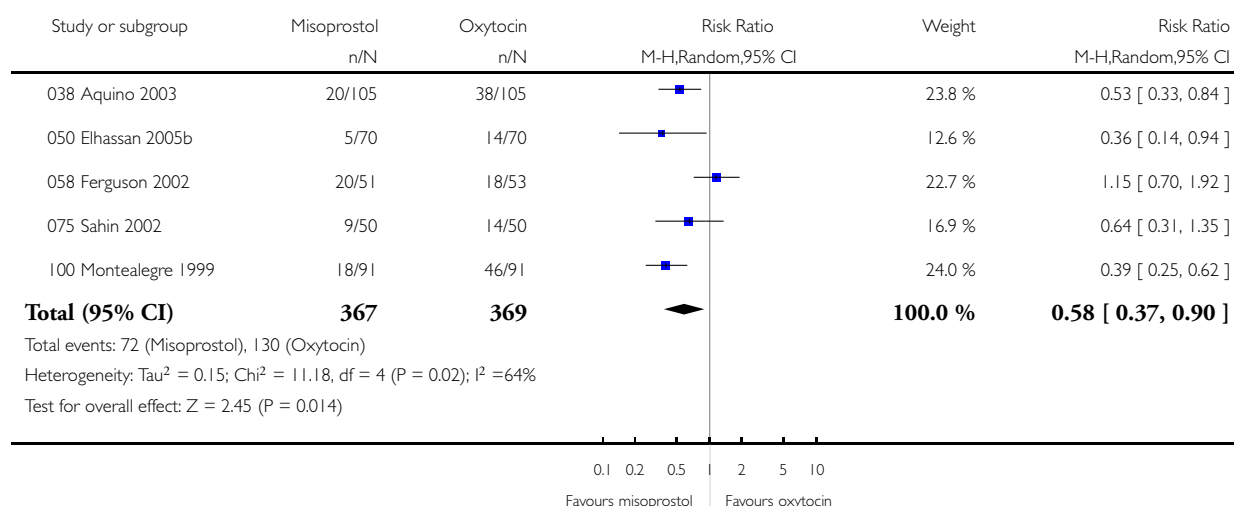


### Analysis 31.3. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 3 Caesarean section

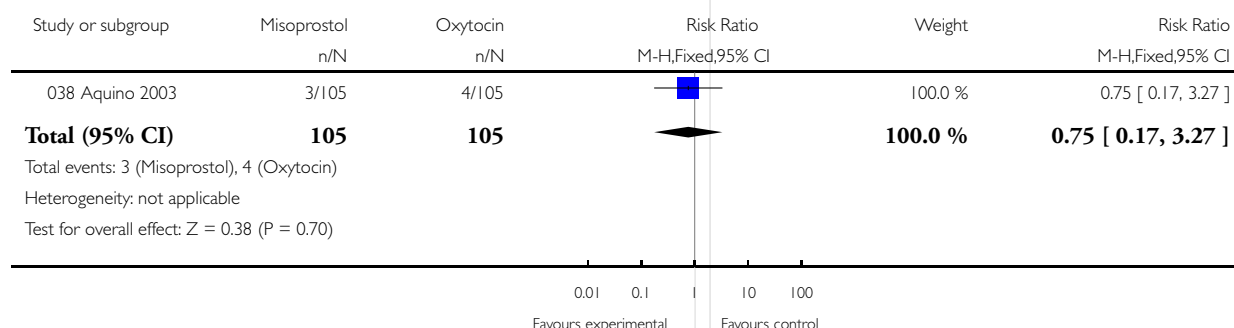


### Analysis 31.4. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 4 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 4 Uterine hyperstimulation with FHR changes



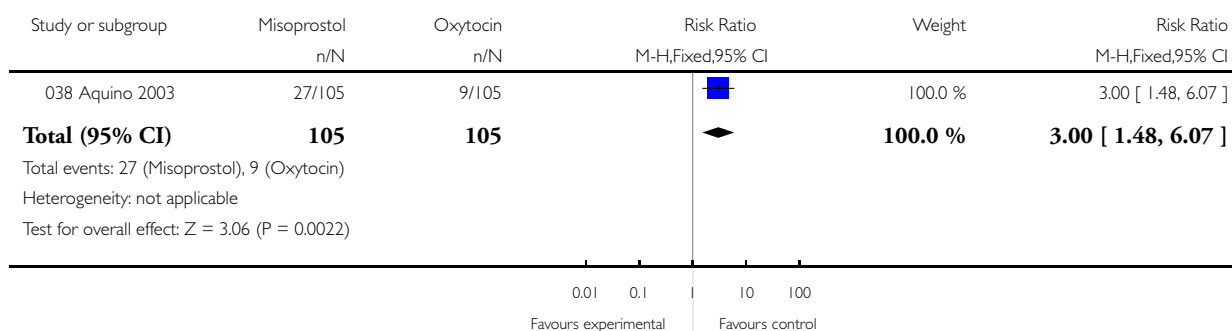


### Analysis 31.5. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 5 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 5 Uterine hyperstimulation without FHR changes

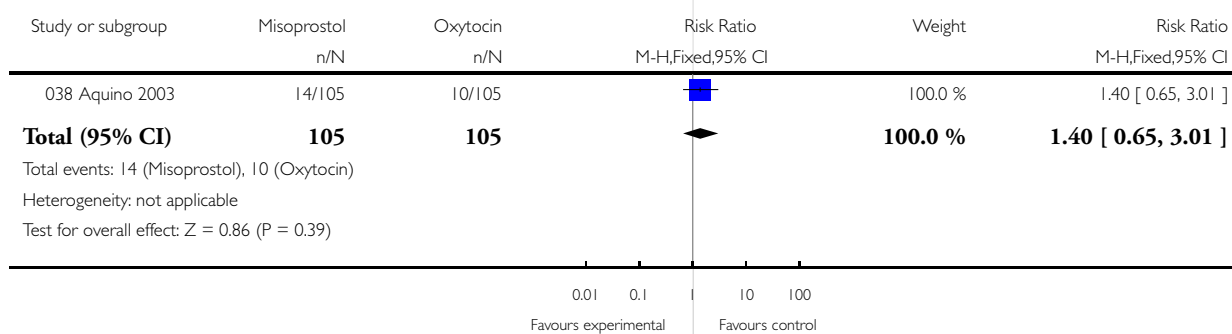


### Analysis 31.6. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 6 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 6 Meconium-stained liquor

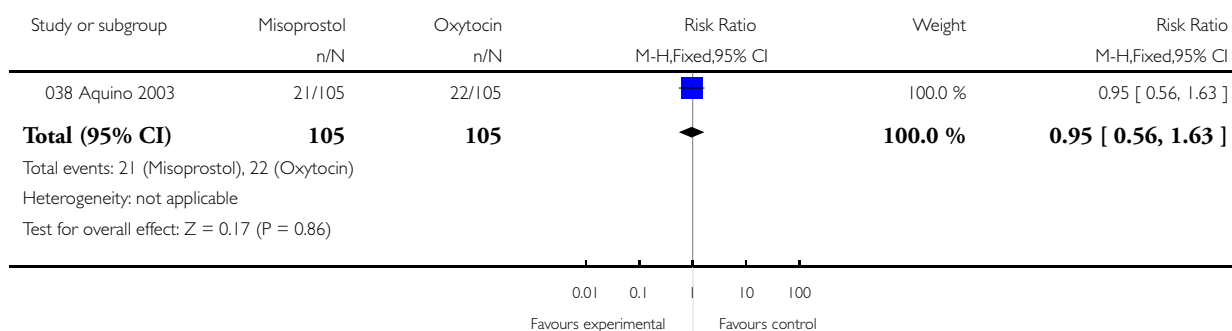


### Analysis 31.7. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 7 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 7 Serious neonatal morbidity

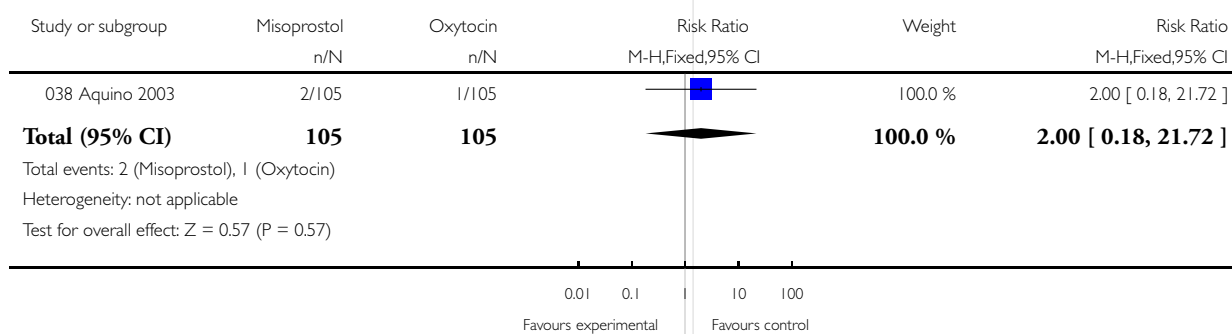


### Analysis 31.8. Comparison 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix, Outcome 8 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 31 Misoprostol versus oxytocin: all women with intact membranes and unfavourable cervix

Outcome: 8 Apgar score < 7 at 5 minutes

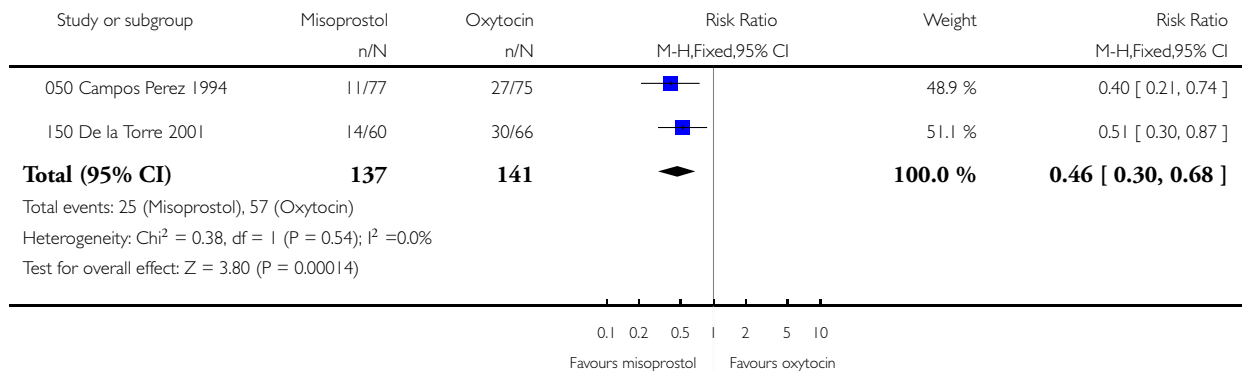


### Analysis 32.1. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

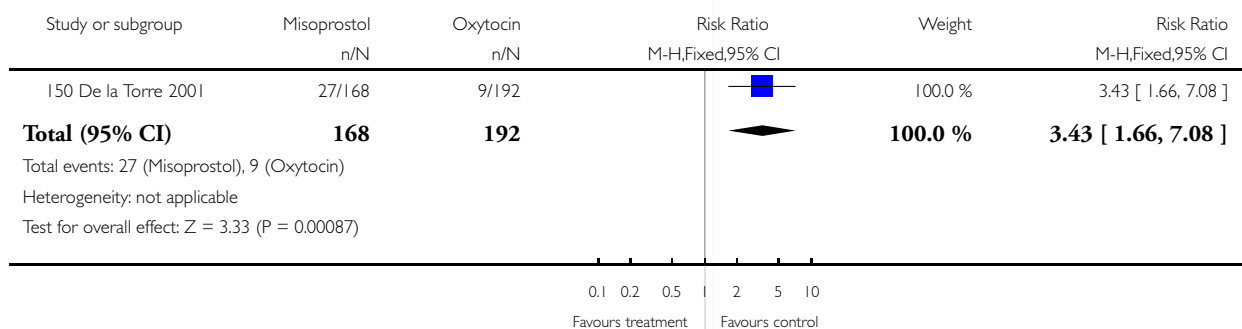


### Analysis 32.2. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

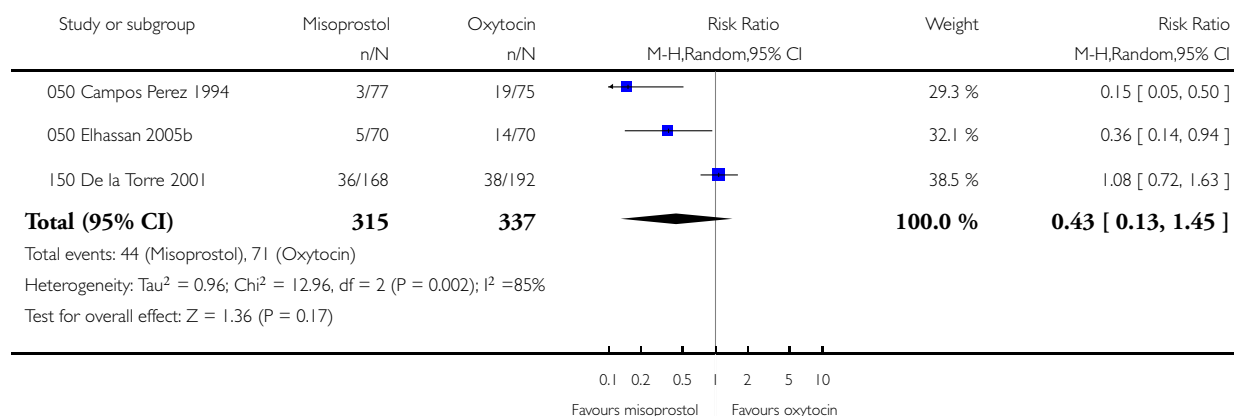


### Analysis 32.3. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 3 Caesarean section

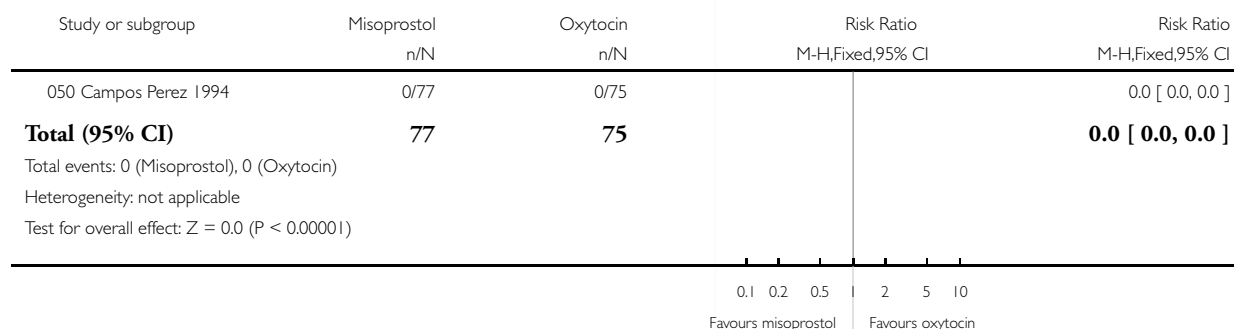


### Analysis 32.4. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 4 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 4 Serious maternal morbidity or death

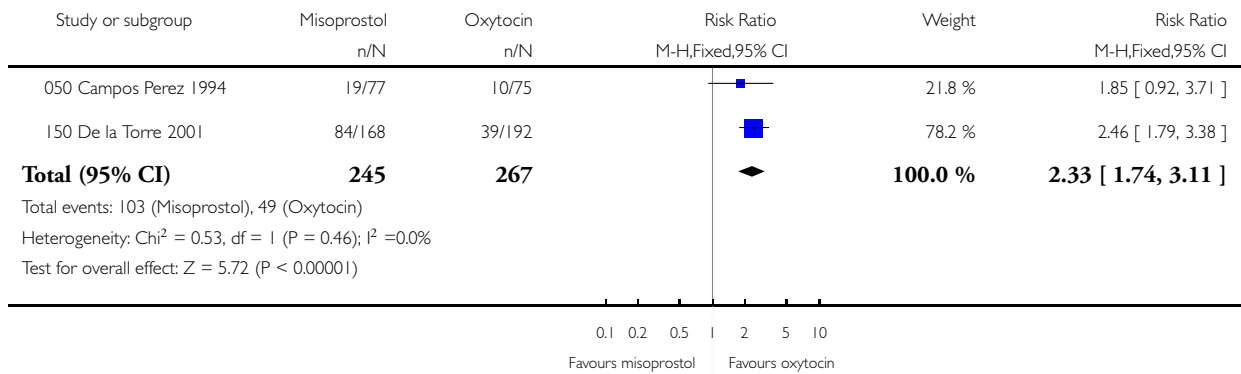


### Analysis 32.5. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 5 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 5 Uterine hyperstimulation without FHR changes

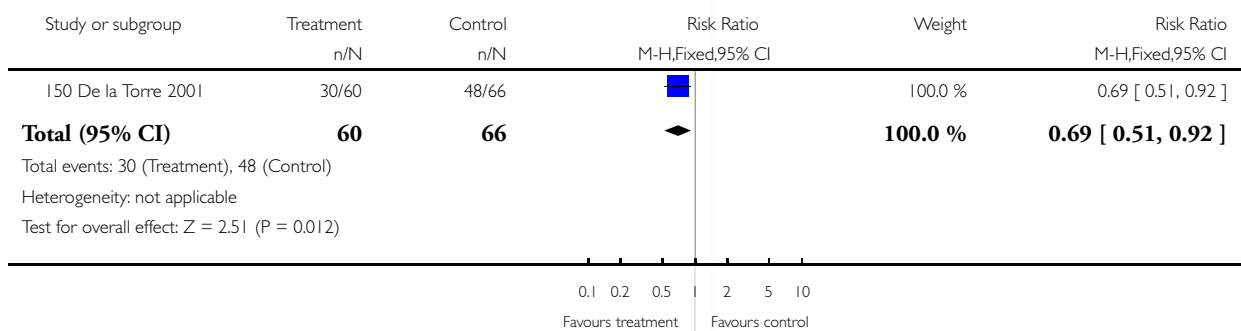


### Analysis 32.6. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 6 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 6 Epidural analgesia

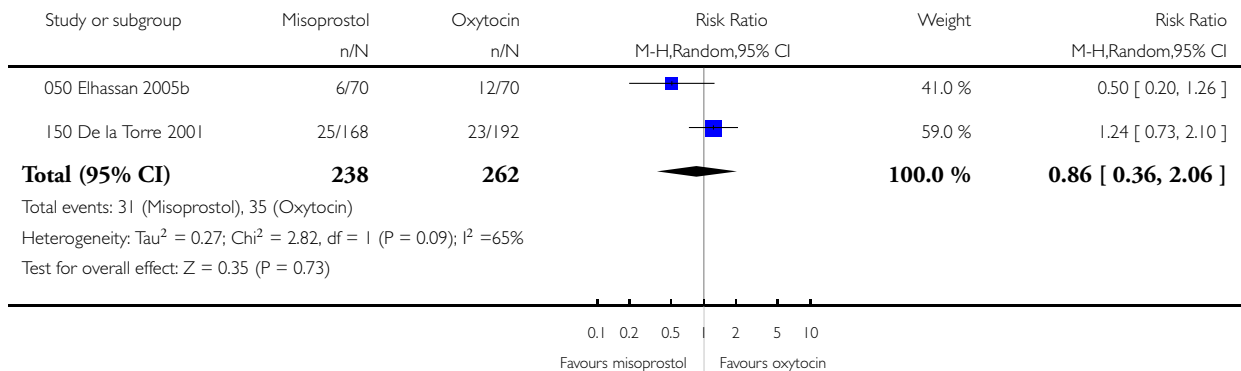


### Analysis 32.7. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 7 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 7 Instrumental vaginal delivery

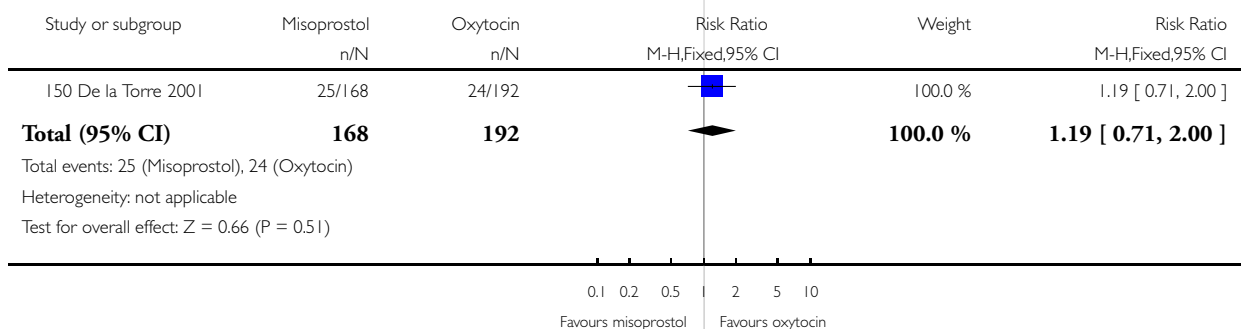


### Analysis 32.8. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 8 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 8 Meconium-stained liquor

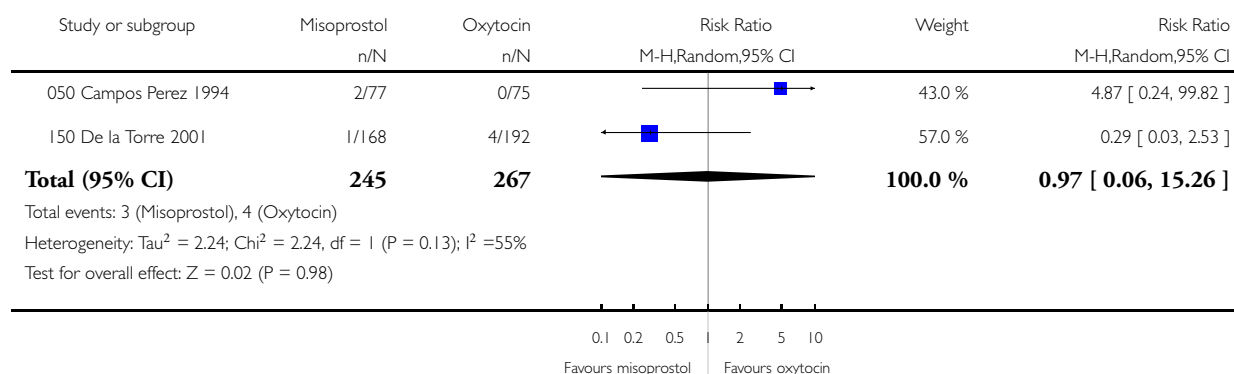


### Analysis 32.9. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 9 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 9 Apgar score < 7 at 5 minutes

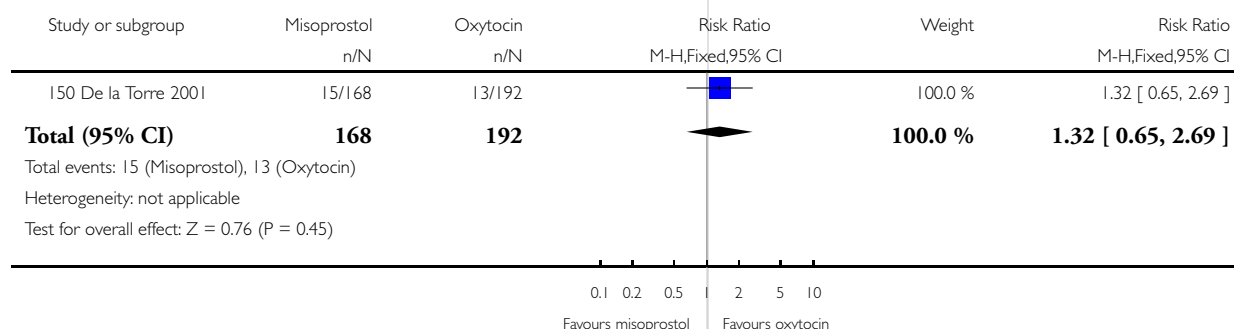


### Analysis 32.10. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 10 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 10 Neonatal intensive care unit admission

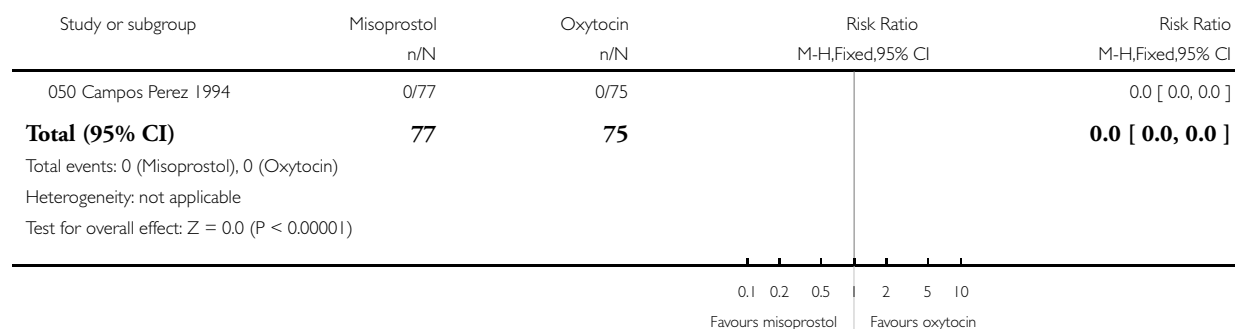


### Analysis 32.1.1. Comparison 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix, Outcome 1.1 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 32 Misoprostol versus oxytocin: all women with intact membranes and variable or undefined cervix

Outcome: 1.1 Perinatal death

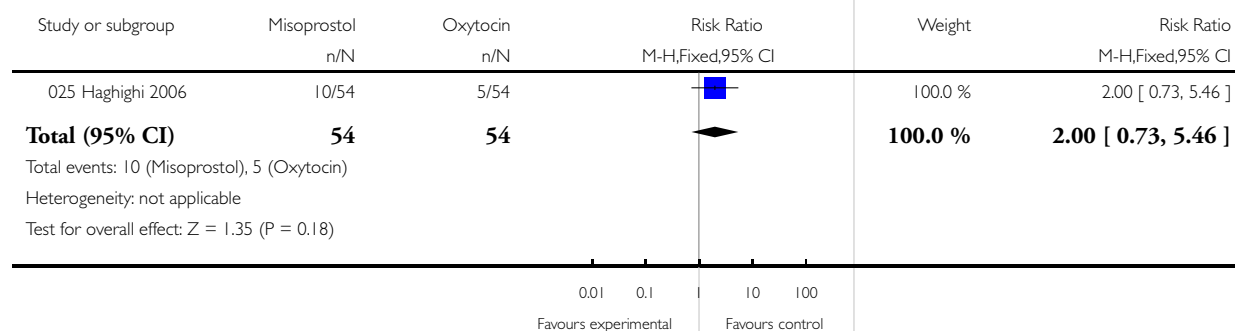


### Analysis 33.1. Comparison 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix, Outcome 1 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix

Outcome: 1 Vaginal delivery not achieved in 24 hours



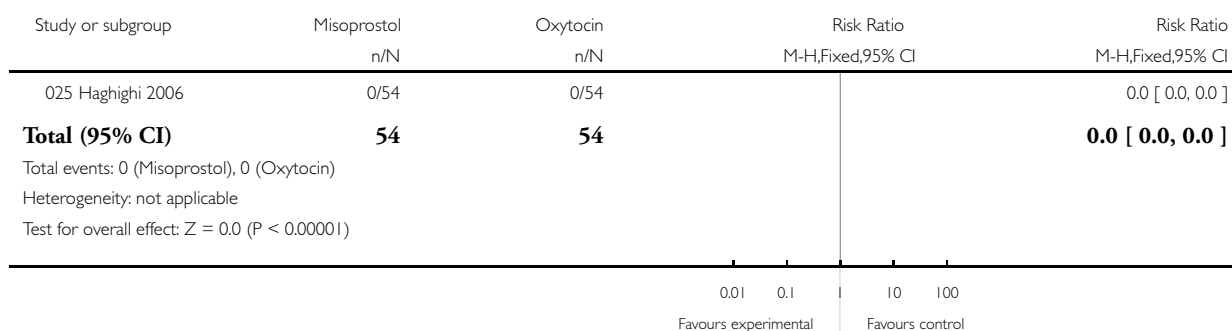


### Analysis 33.2. Comparison 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix, Outcome 2 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix

Outcome: 2 Uterine hyperstimulation without FHR changes

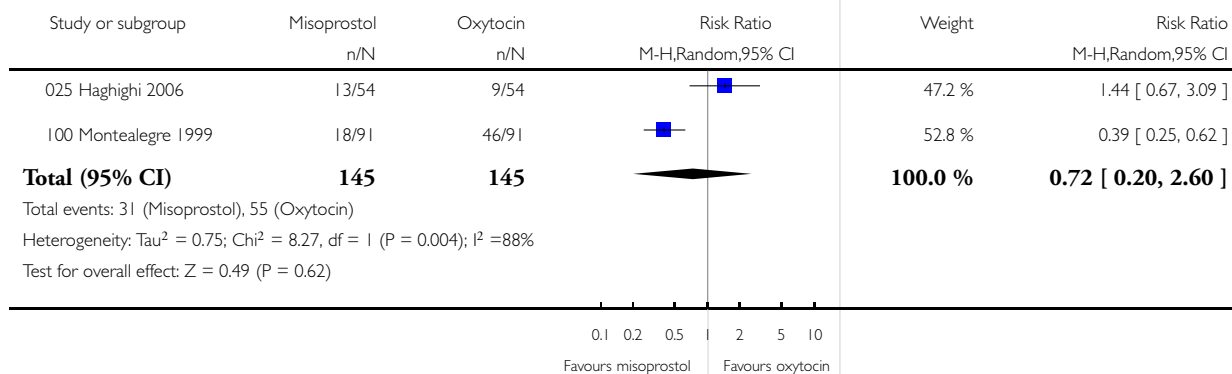


### Analysis 33.3. Comparison 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix

Outcome: 3 Caesarean section

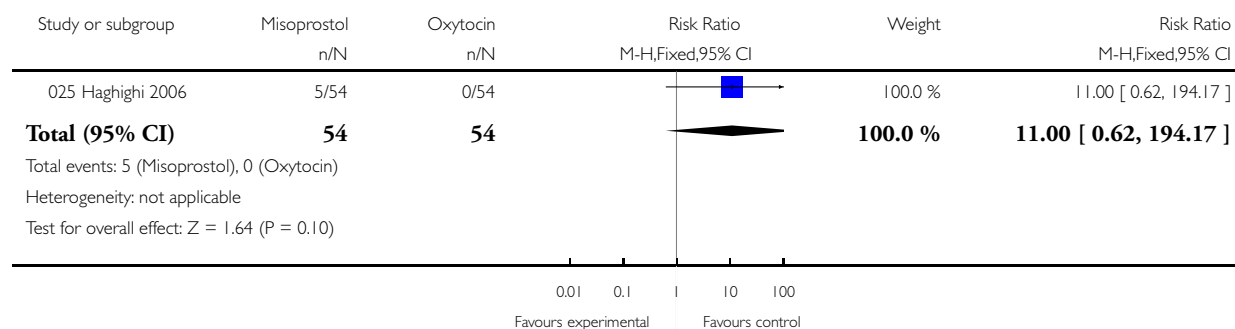


### Analysis 33.4. Comparison 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix, Outcome 4 Maternal side effects.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 33 Misoprostol versus oxytocin: all women with ruptured membranes and unfavourable cervix

Outcome: 4 Maternal side effects

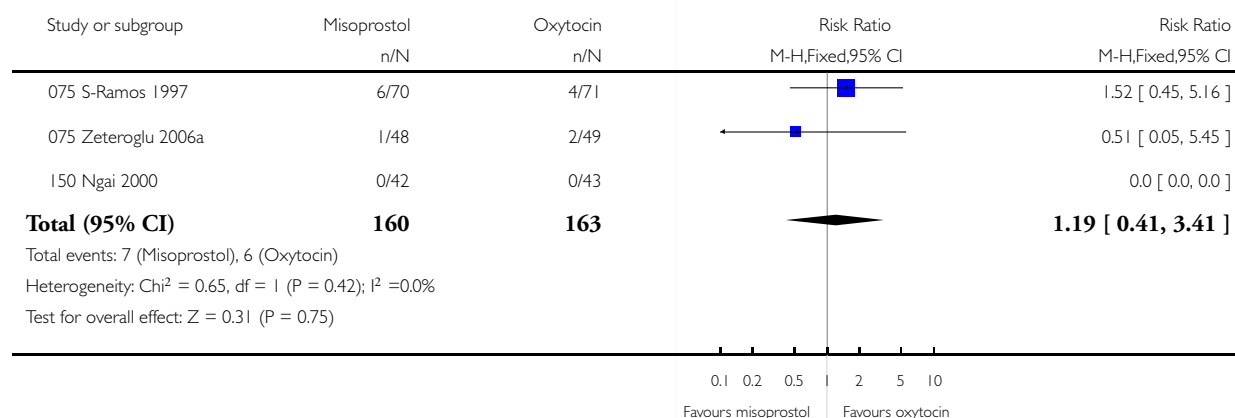


### Analysis 34.1. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 1 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 1 Uterine hyperstimulation with FHR changes

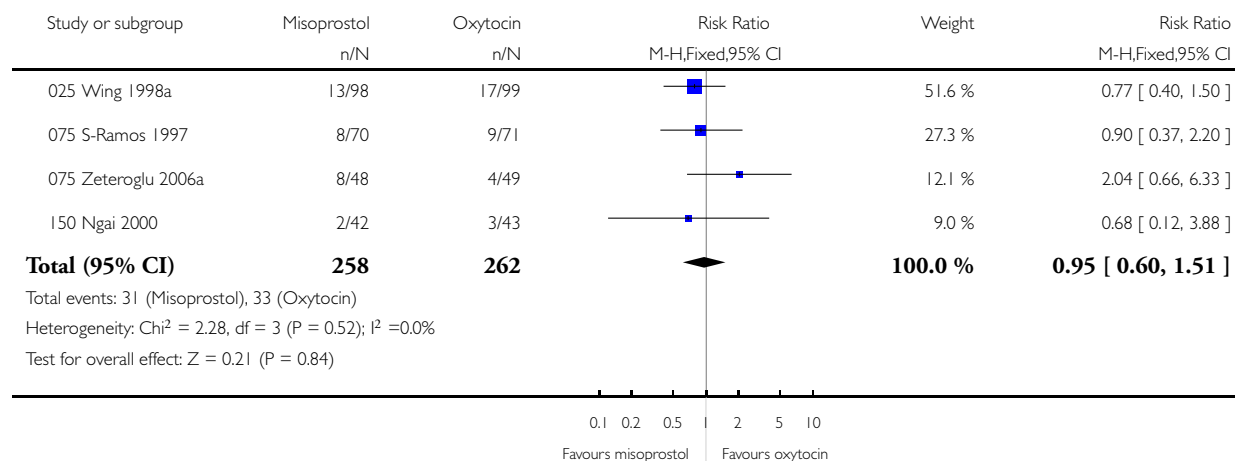


### Analysis 34.2. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 2 Caesarean section

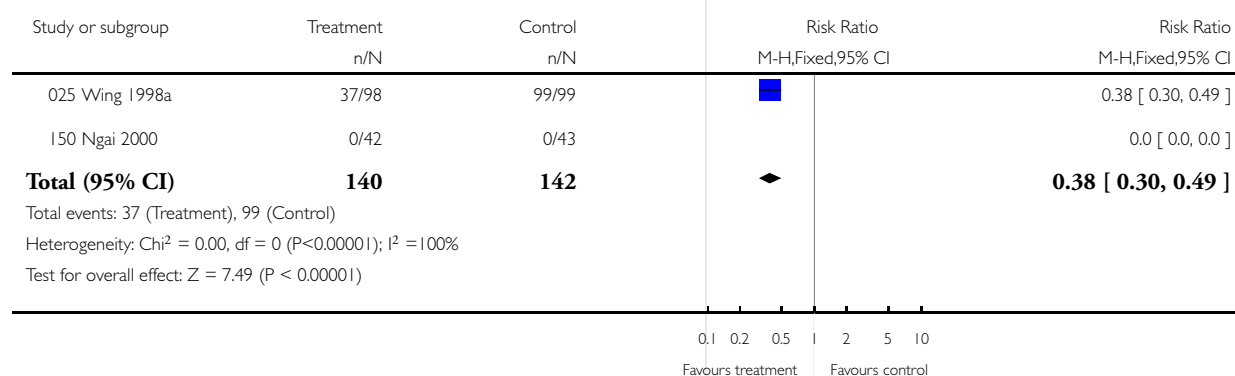


### Analysis 34.3. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 3 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 3 Oxytocin augmentation

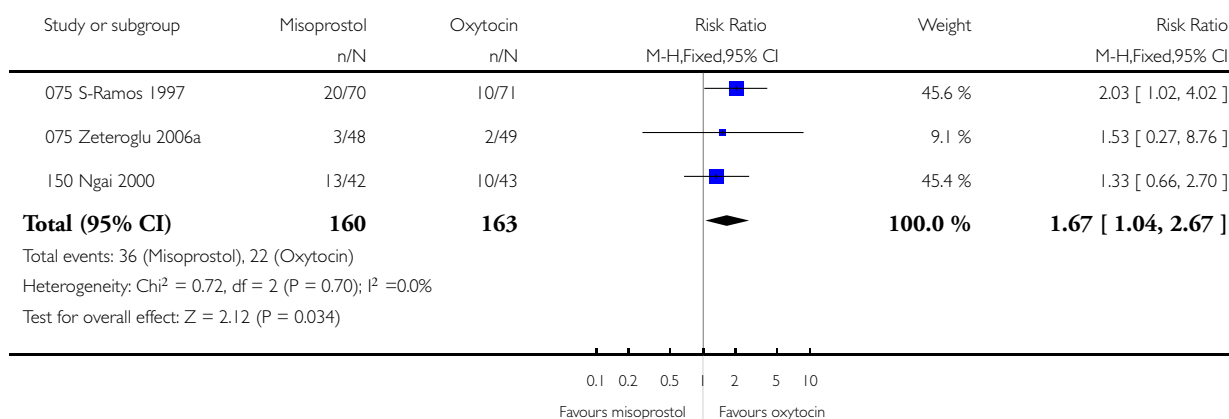


### Analysis 34.4. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 4 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 4 Uterine hyperstimulation without FHR changes

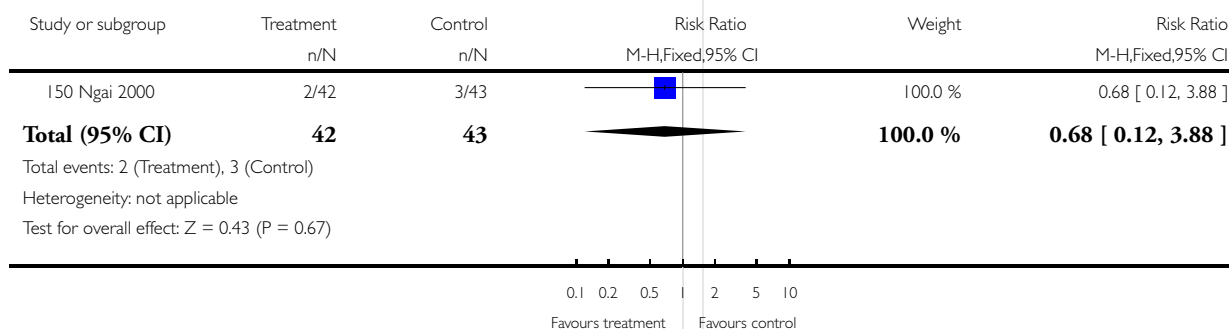


### Analysis 34.5. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 5 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 5 Epidural analgesia

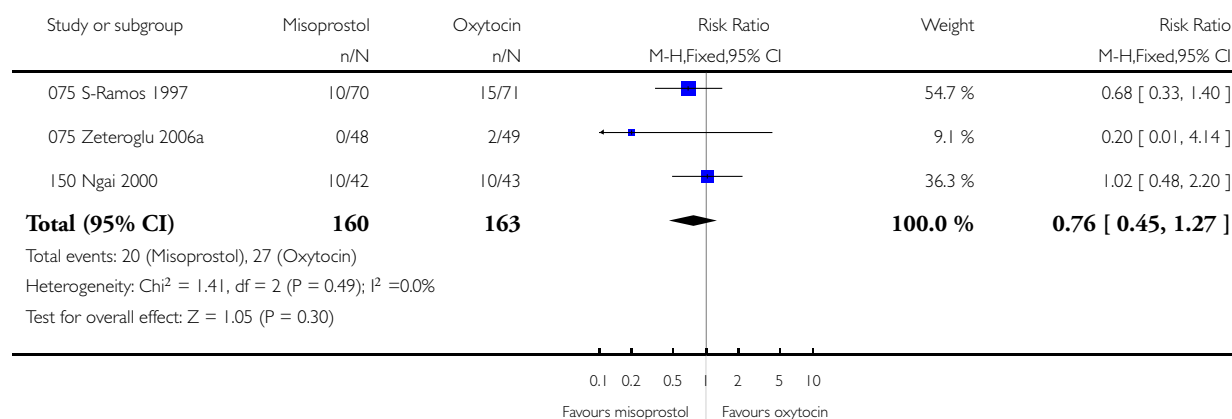


### Analysis 34.6. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 6 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 6 Instrumental vaginal delivery

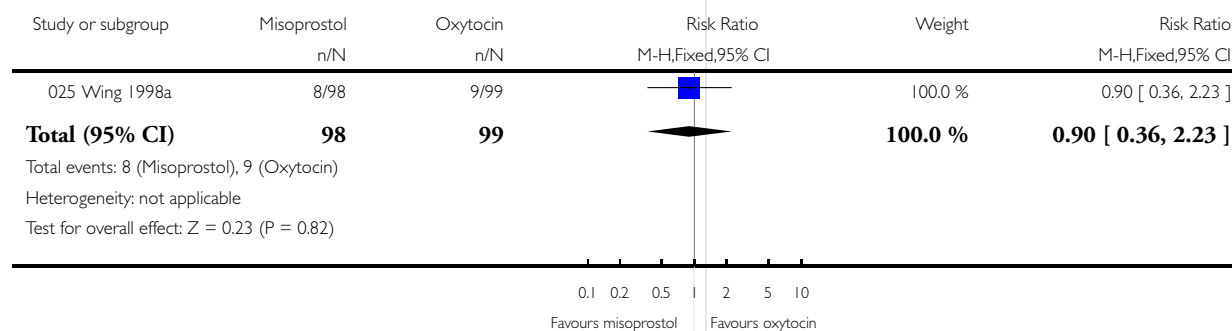


### Analysis 34.7. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 7 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 7 Meconium-stained liquor

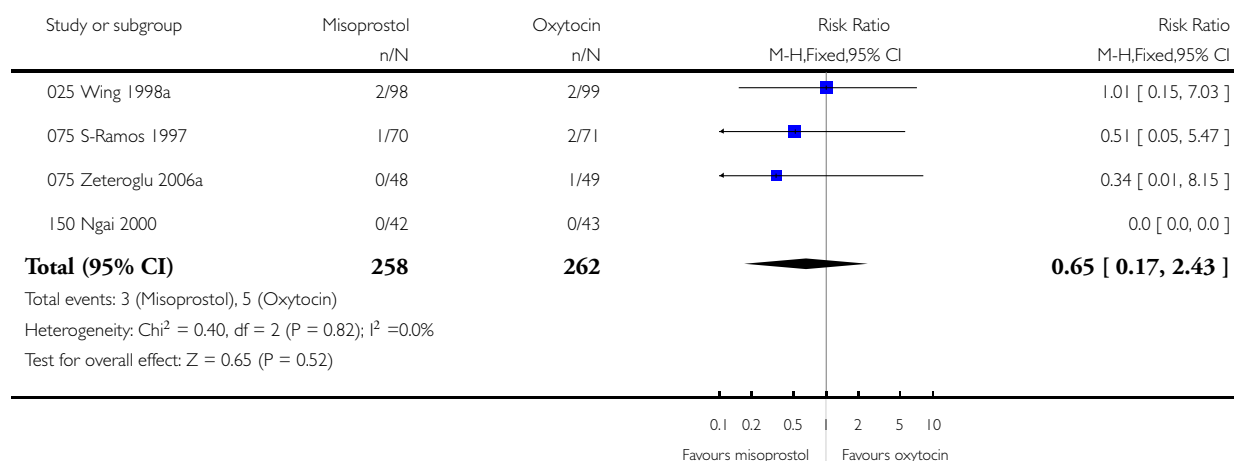


### Analysis 34.8. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 8 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 8 Apgar score < 7 at 5 minutes

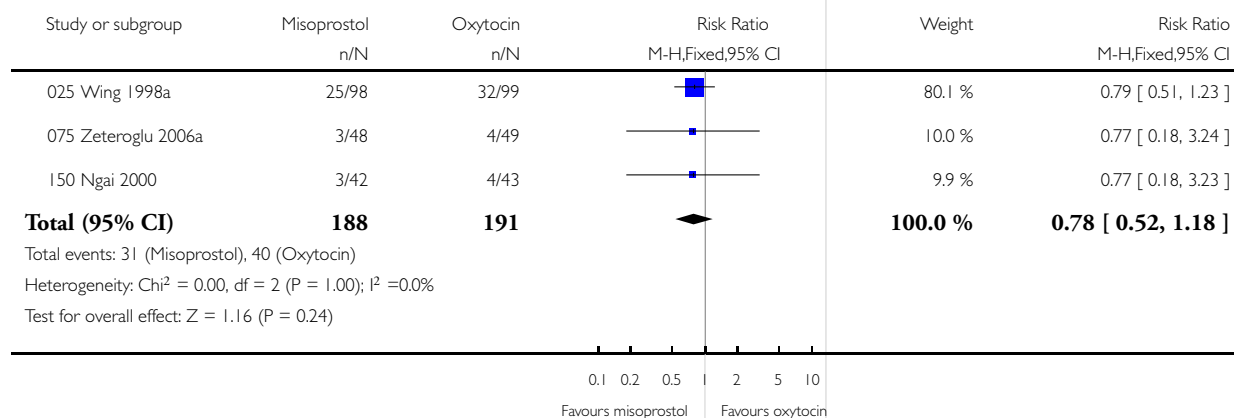


### Analysis 34.9. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 9 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 9 Neonatal intensive care unit admission

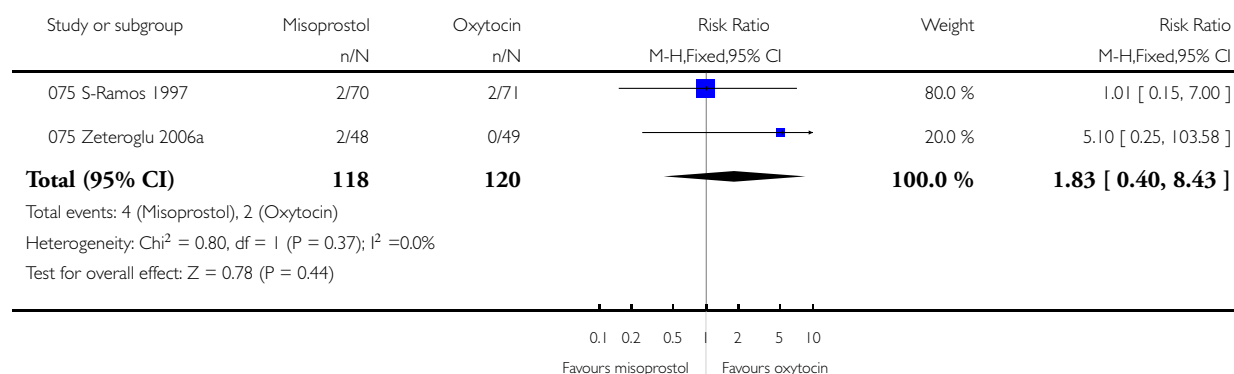


### Analysis 34.10. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 10 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 10 Postpartum haemorrhage

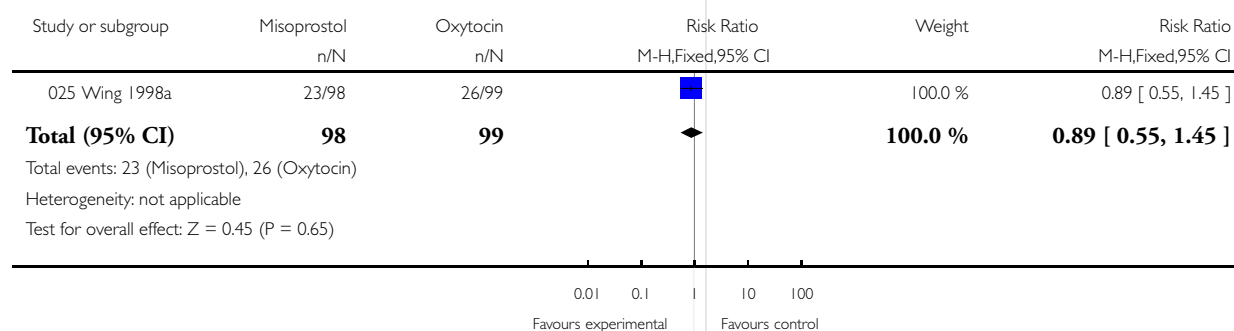


### Analysis 34.11. Comparison 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix, Outcome 11 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 34 Misoprostol versus oxytocin: all women with ruptured membranes and variable or undefined cervix

Outcome: 11 Vaginal delivery not achieved within 24 hours

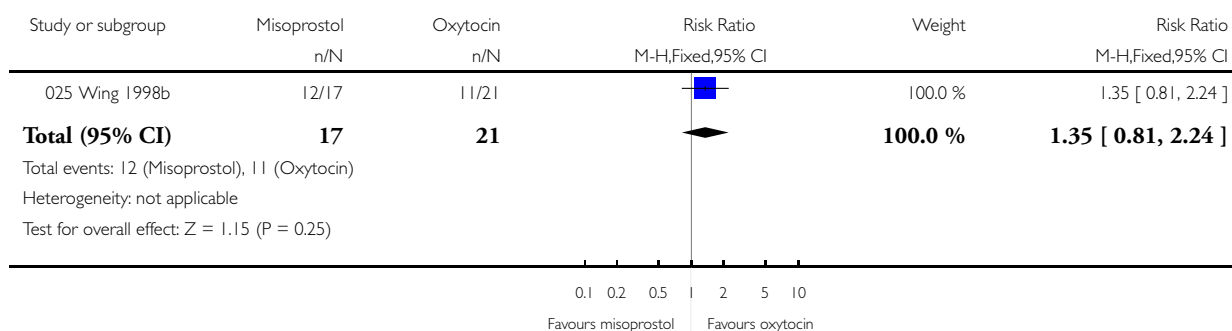


### Analysis 35.1. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 1 Vaginal delivery not achieved within 24 hours

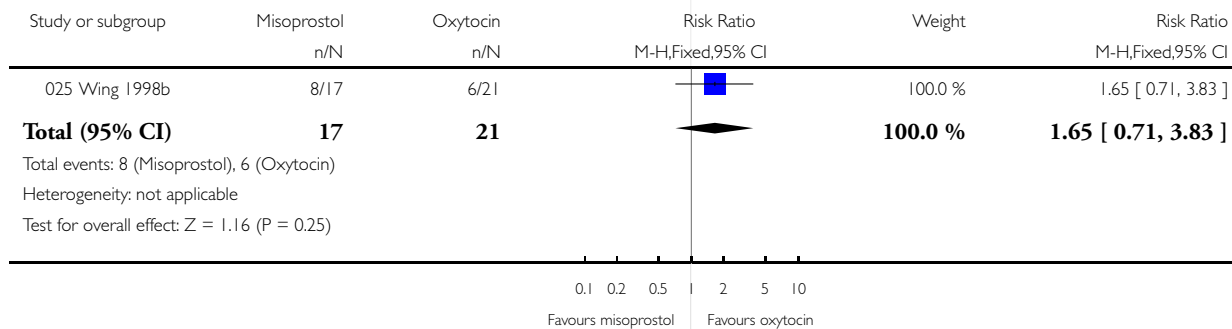


### Analysis 35.2. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 2 Caesarean section



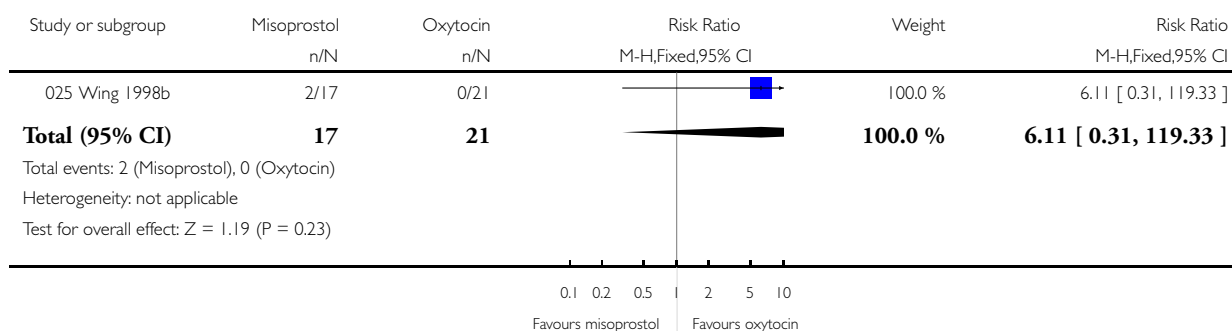


### Analysis 35.3. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 3 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 3 Serious maternal morbidity or death

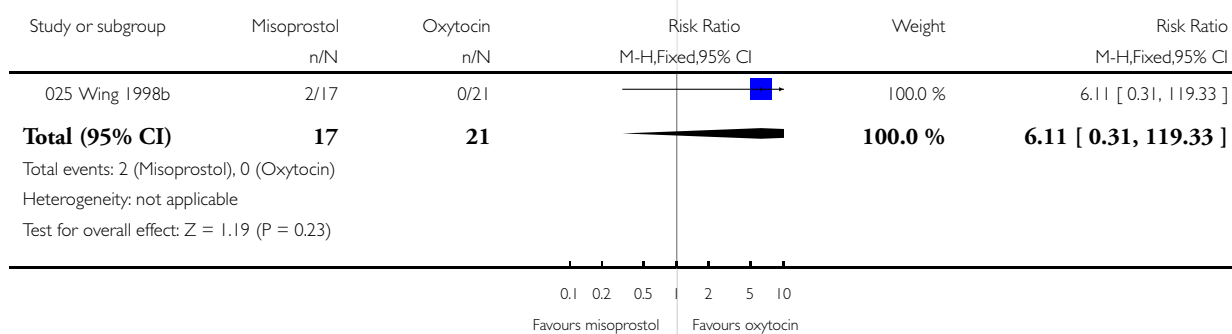


### Analysis 35.4. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 4 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 4 Uterine hyperstimulation without FHR changes

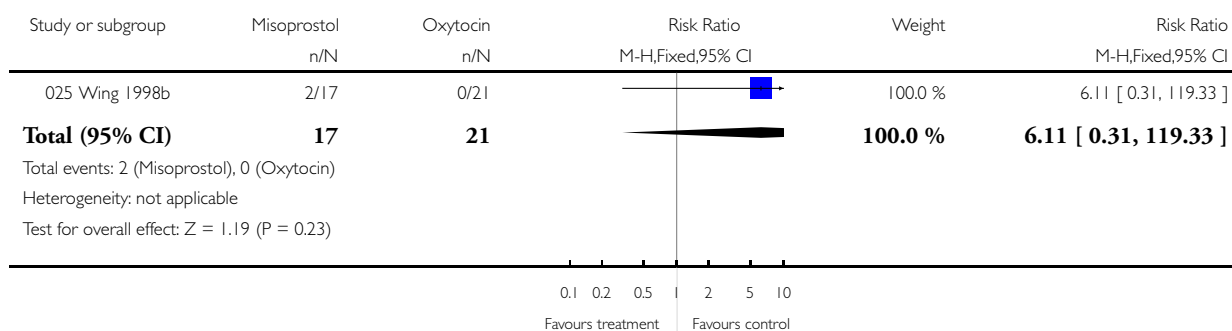


### Analysis 35.5. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 5 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 5 Uterine rupture

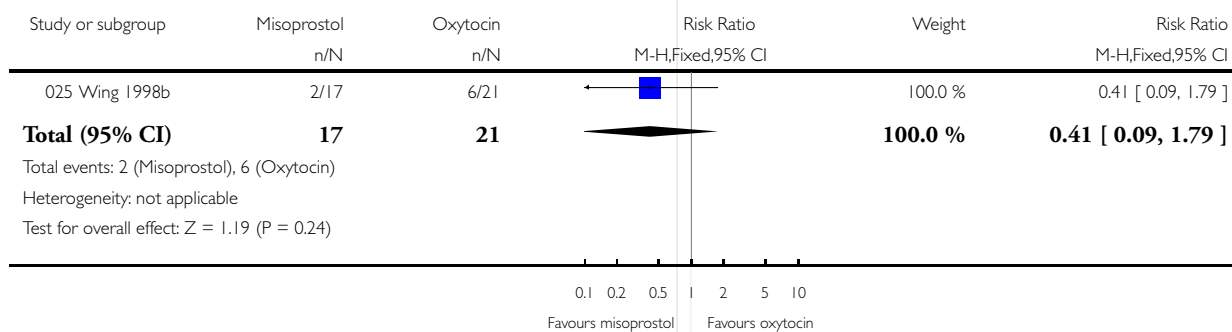


### Analysis 35.6. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 6 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 6 Meconium-stained liquor

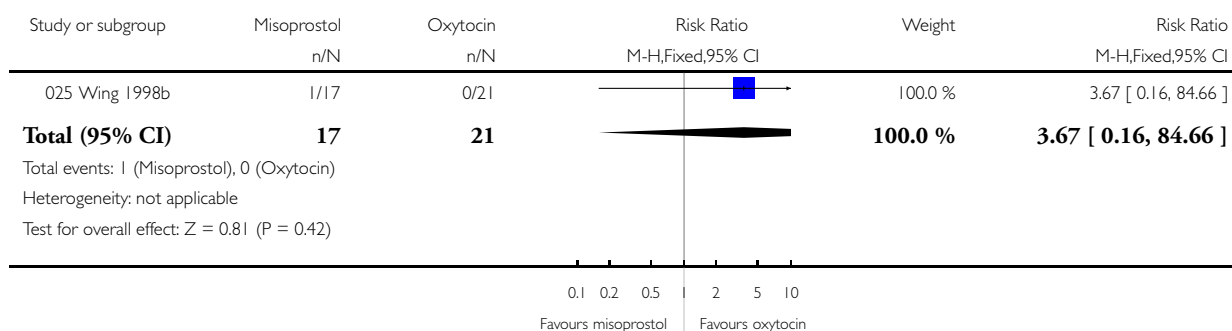


**Analysis 35.7. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 7 Apgar score < 7 at 5 minutes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 7 Apgar score < 7 at 5 minutes

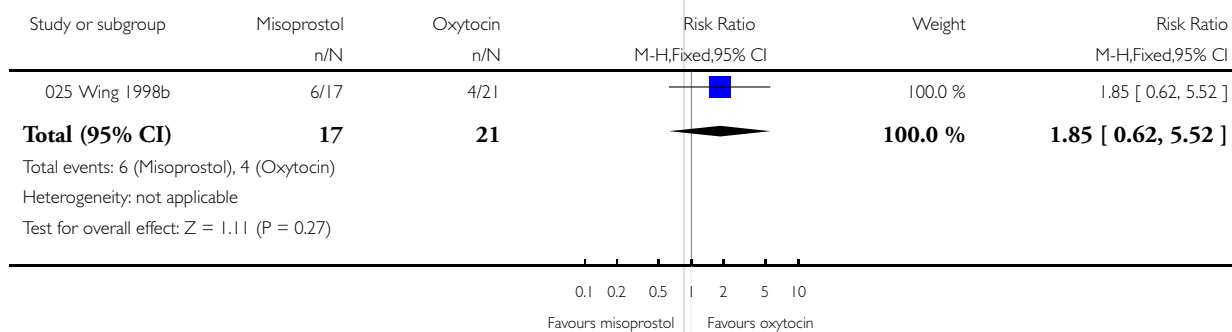


**Analysis 35.8. Comparison 35 Misoprostol versus oxytocin: all women with previous caesarean section, Outcome 8 Neonatal intensive care unit admission.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 35 Misoprostol versus oxytocin: all women with previous caesarean section

Outcome: 8 Neonatal intensive care unit admission

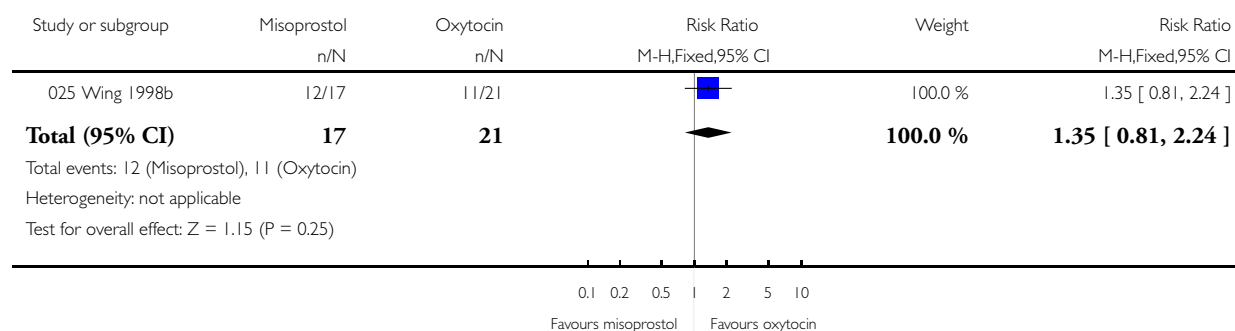


### Analysis 36.1. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

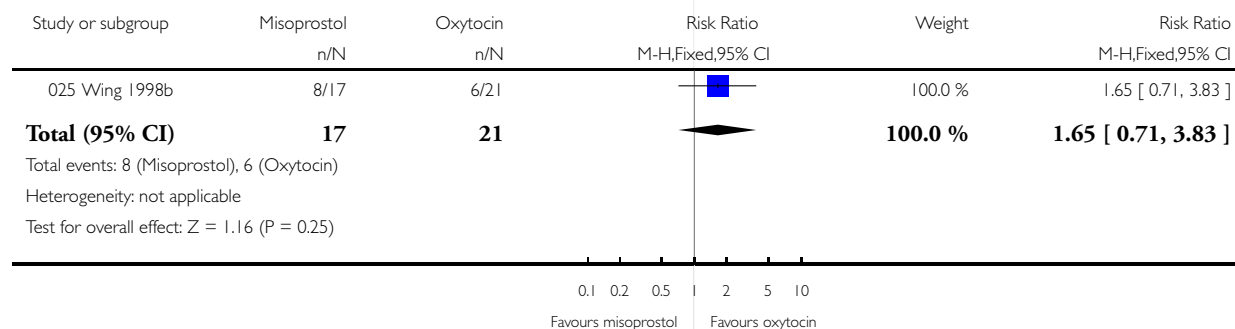


### Analysis 36.2. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 2 Caesarean section

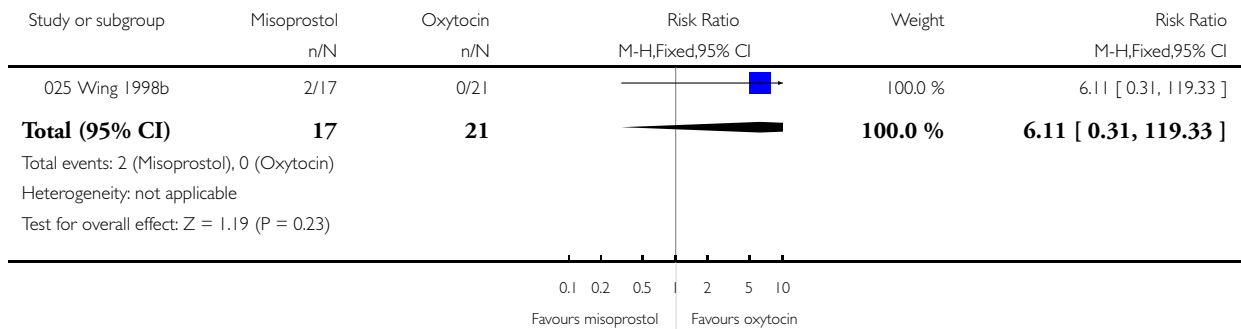


### Analysis 36.3. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 3 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 3 Serious maternal morbidity or death

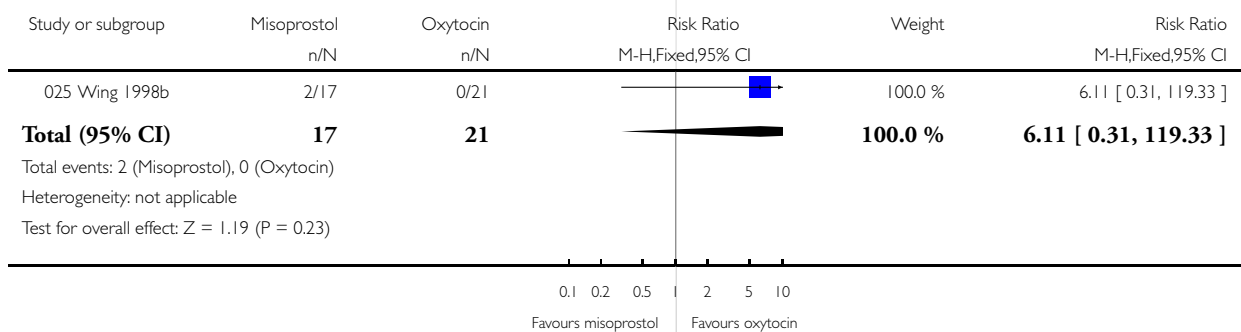


### Analysis 36.4. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 4 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 4 Uterine hyperstimulation without FHR changes

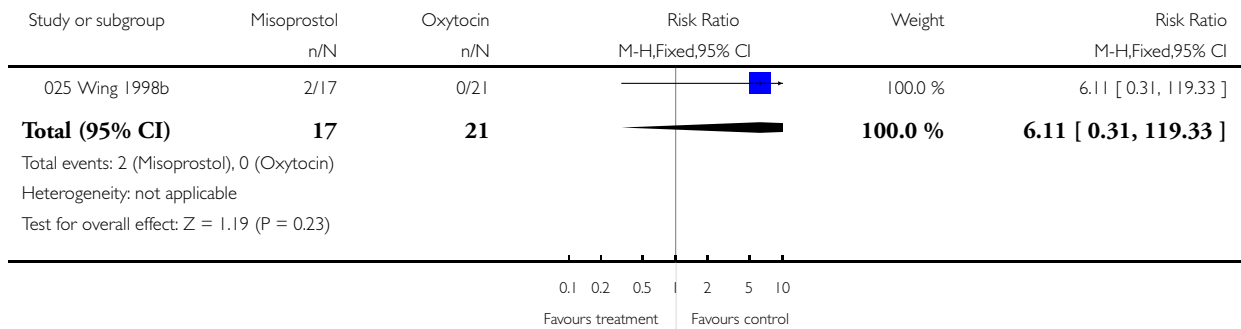


### Analysis 36.5. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 5 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 5 Uterine rupture

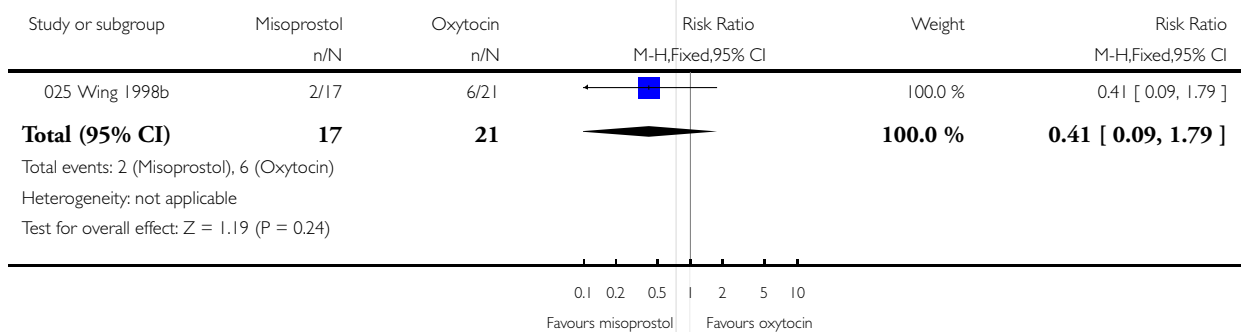


### Analysis 36.6. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 6 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 6 Meconium-stained liquor

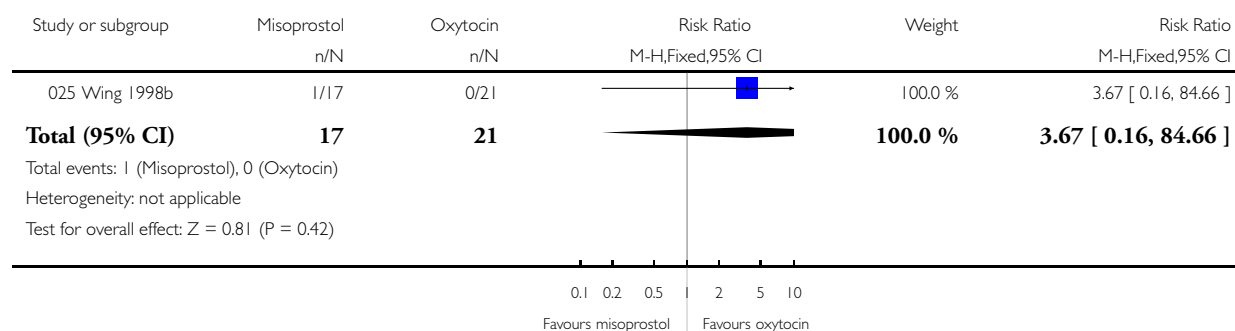


### Analysis 36.7. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 7 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 7 Apgar score < 7 at 5 minutes

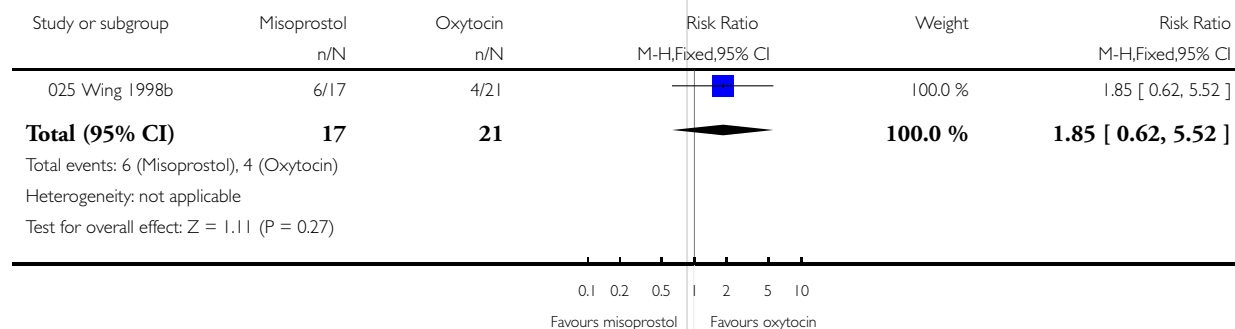


### Analysis 36.8. Comparison 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix, Outcome 8 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 36 Misoprostol versus oxytocin: all women with previous caesarean section and unfavourable cervix

Outcome: 8 Neonatal intensive care unit admission

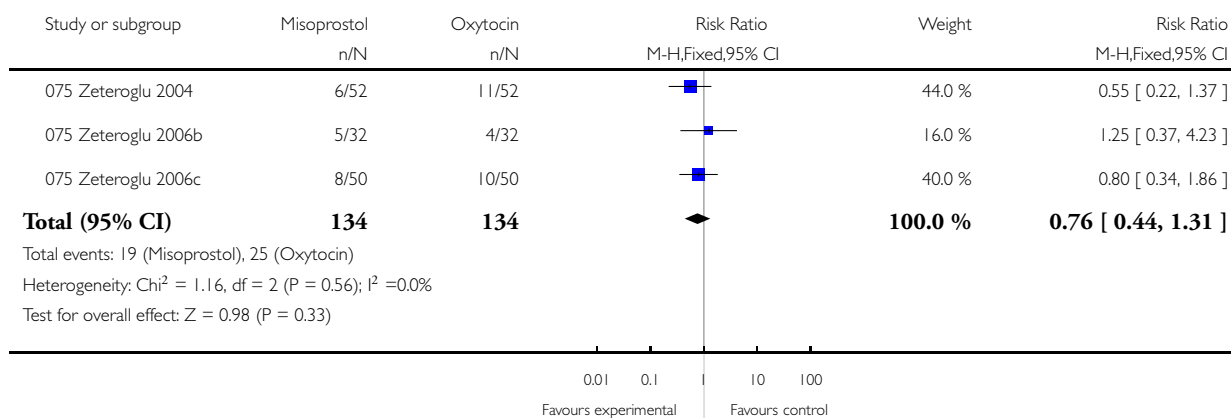


### Analysis 37.1. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 1 Caesarean section

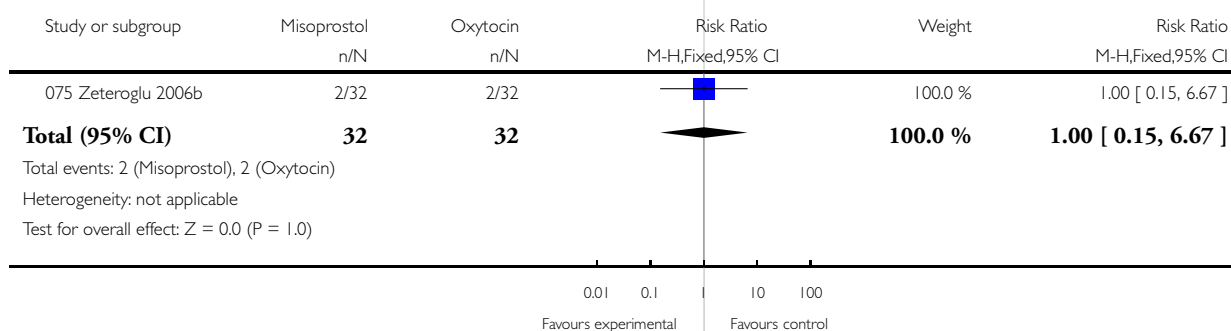


### Analysis 37.2. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 2 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 2 Serious neonatal morbidity



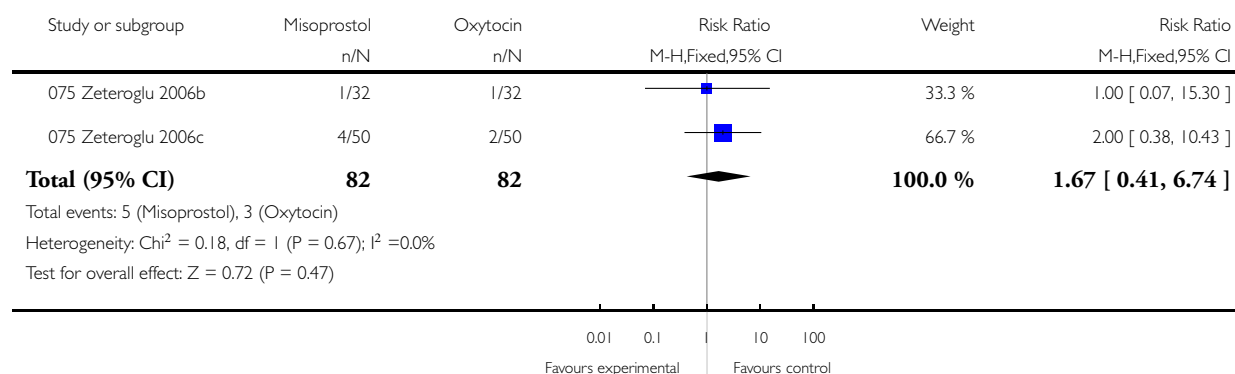


### Analysis 37.3. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 3 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 3 Uterine hyperstimulation without FHR changes

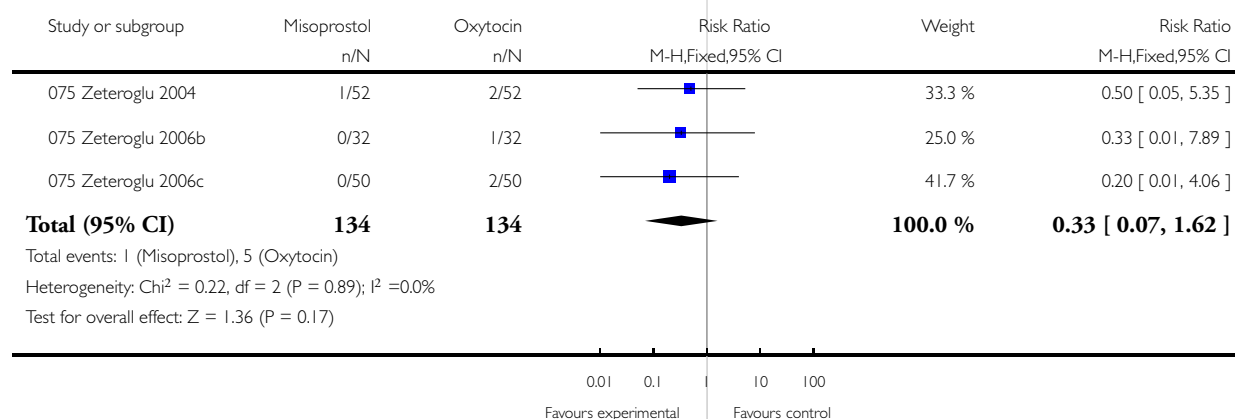


### Analysis 37.4. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 4 Instrumental delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 4 Instrumental delivery

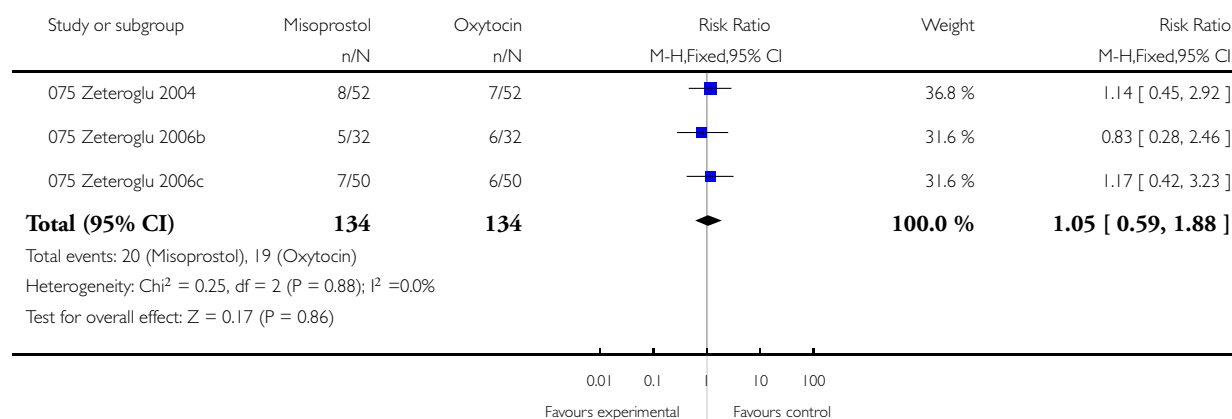


### Analysis 37.5. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 5 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 5 Meconium-stained liquor

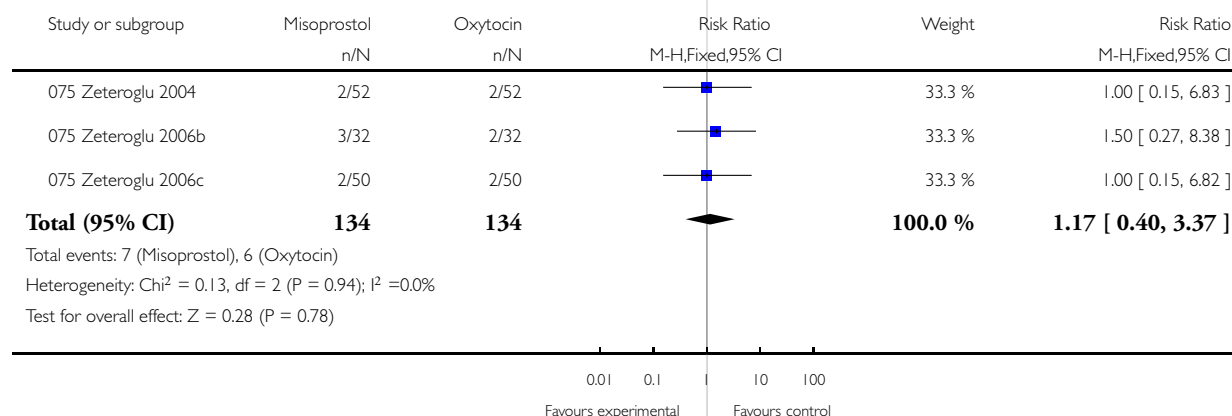


### Analysis 37.6. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 6 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 6 Neonatal intensive care unit admission

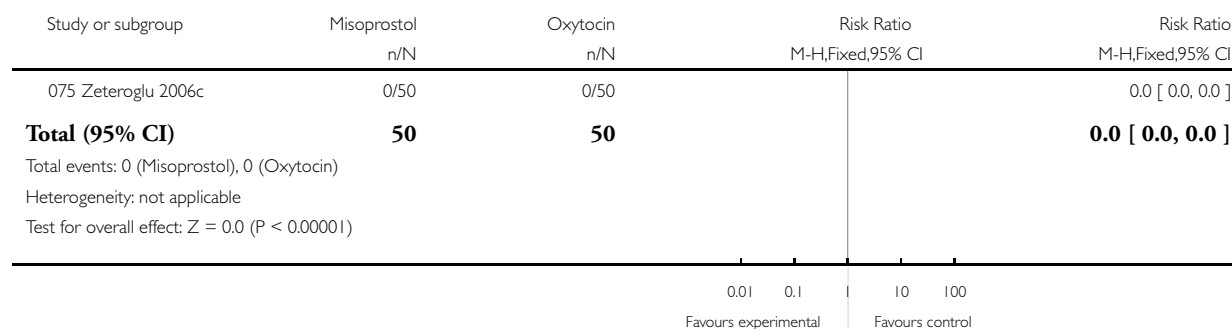


### Analysis 37.7. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 7 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 7 Perinatal death

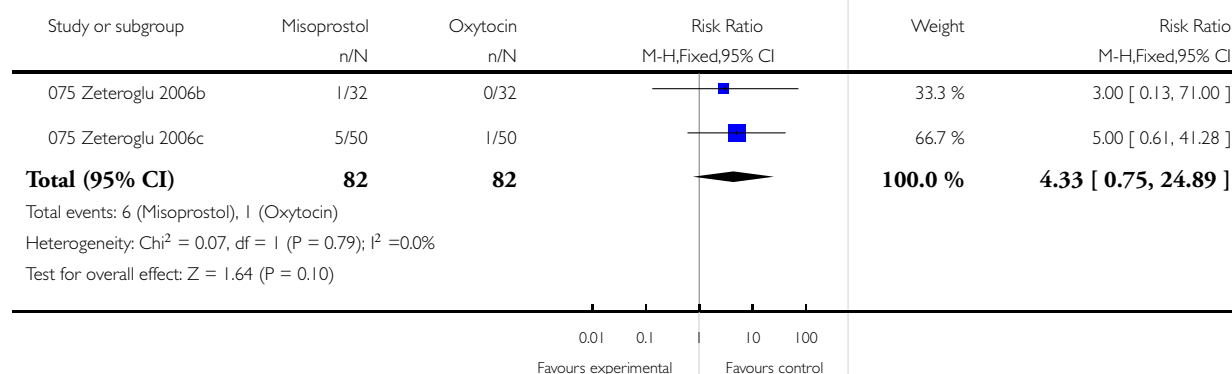


### Analysis 37.8. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 8 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 8 Maternal side effects (eg nausea, vomiting, diarrhoea)

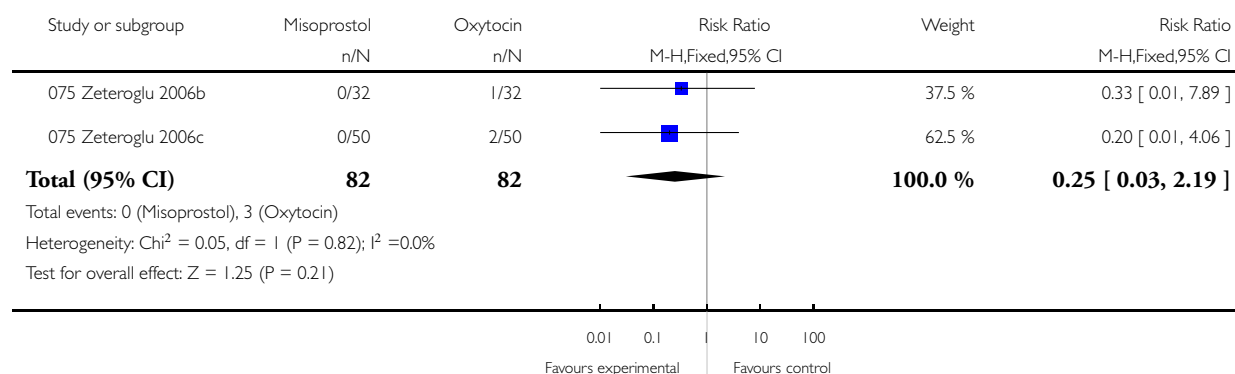


### Analysis 37.9. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 9 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 9 Postpartum haemorrhage

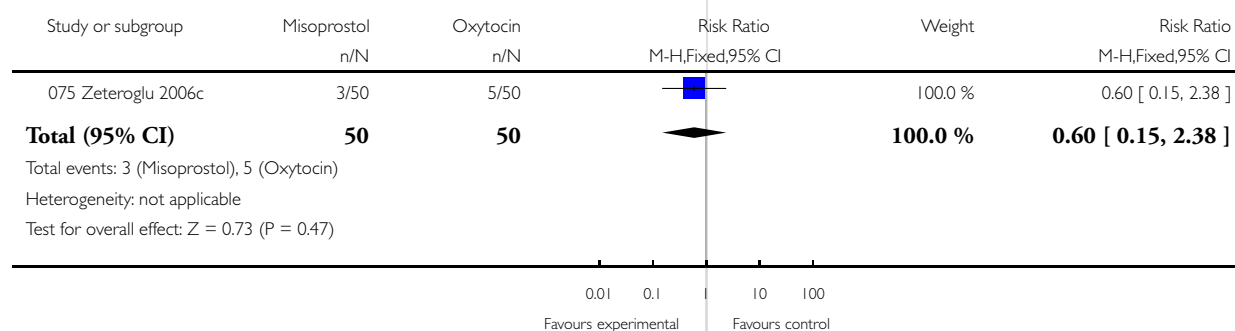


### Analysis 37.10. Comparison 37 Misoprostol versus oxytocin: all multiparae, Outcome 10 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 37 Misoprostol versus oxytocin: all multiparae

Outcome: 10 Serious maternal complications

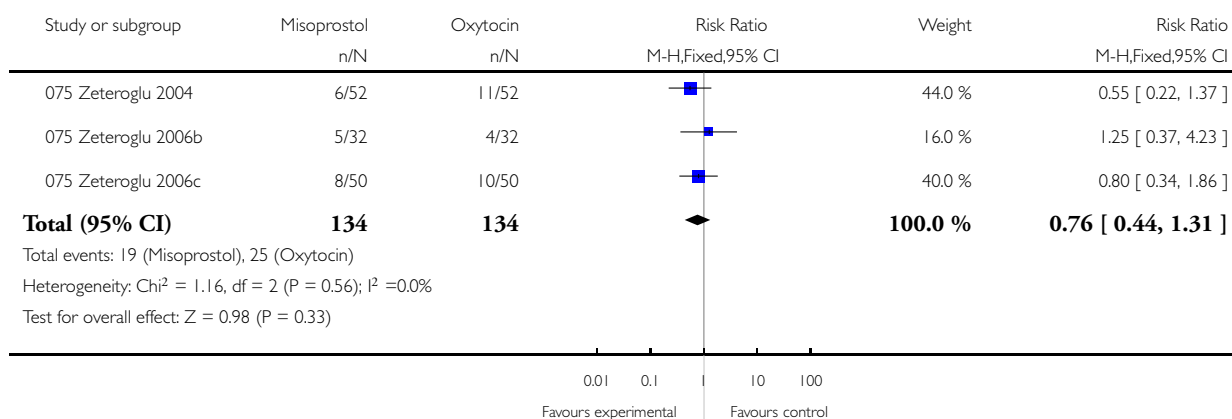


### Analysis 38.1. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 1 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 1 Caesarean section

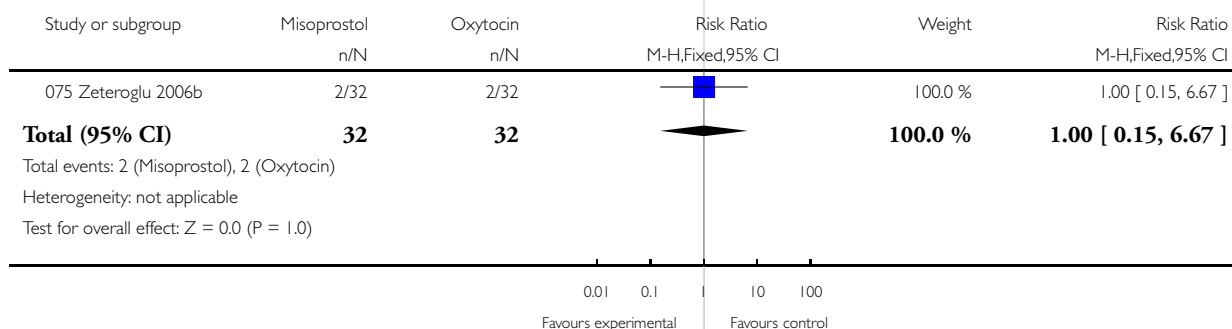


### Analysis 38.2. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 2 Serious neonatal morbidity.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 2 Serious neonatal morbidity

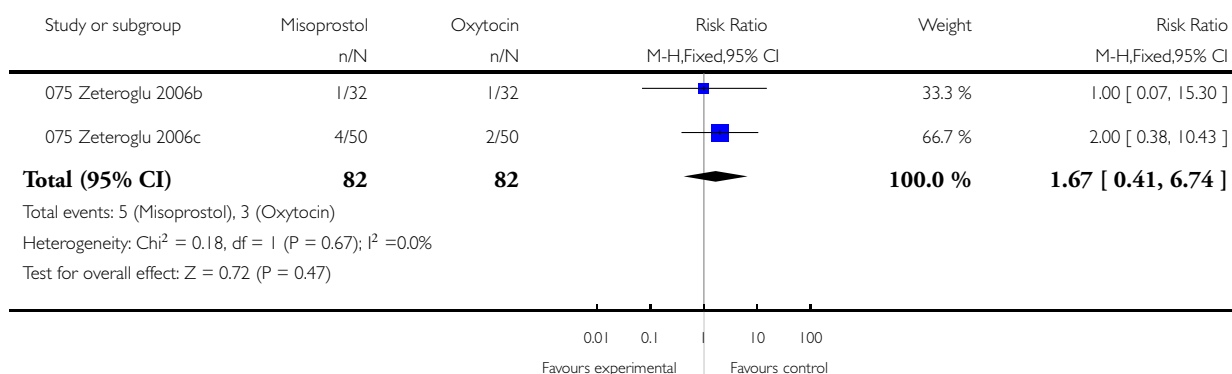


### Analysis 38.3. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 3 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 3 Uterine hyperstimulation without FHR changes

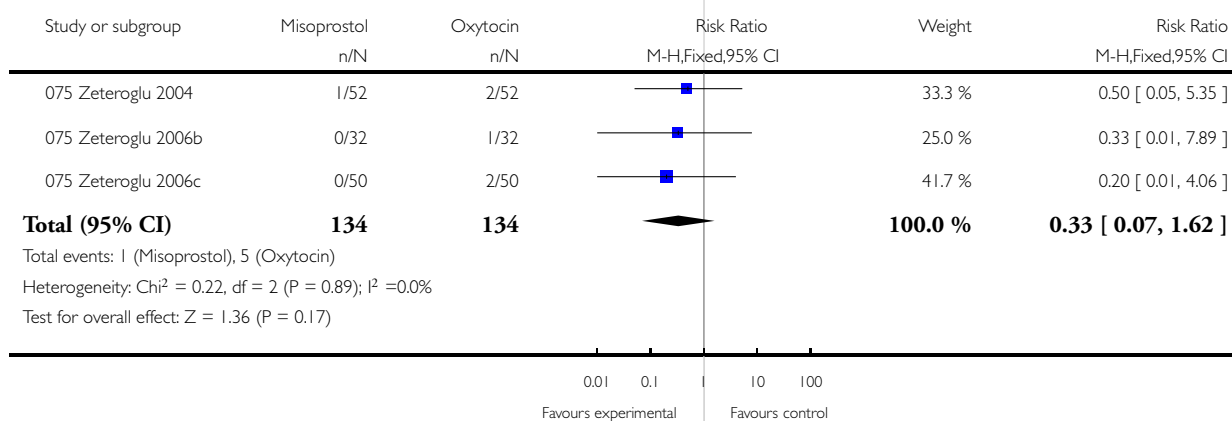


### Analysis 38.4. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 4 Instrumental delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 4 Instrumental delivery

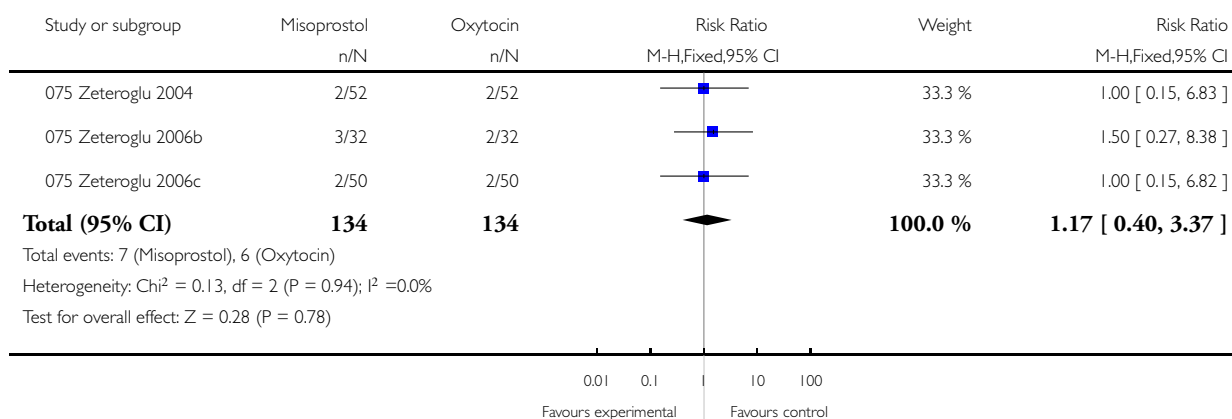


### Analysis 38.5. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 5 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 5 Neonatal intensive care unit admission

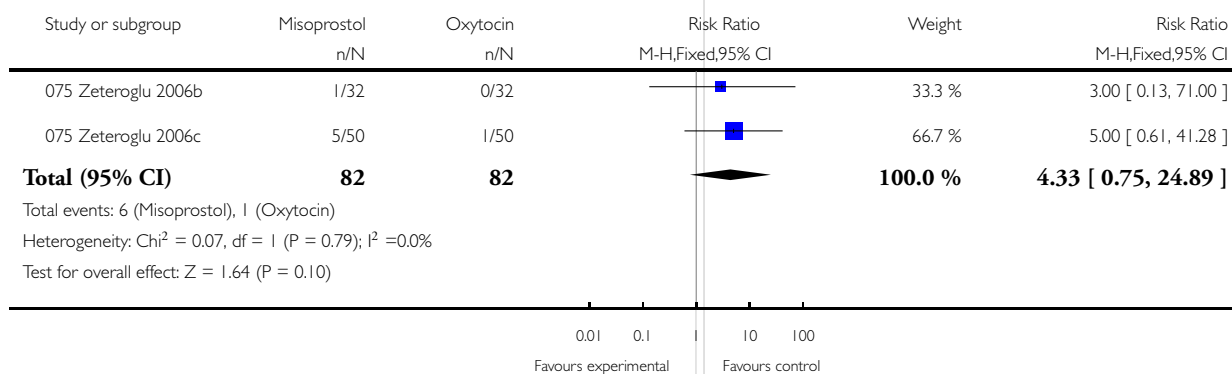


### Analysis 38.6. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 6 Maternal side effects.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 6 Maternal side effects

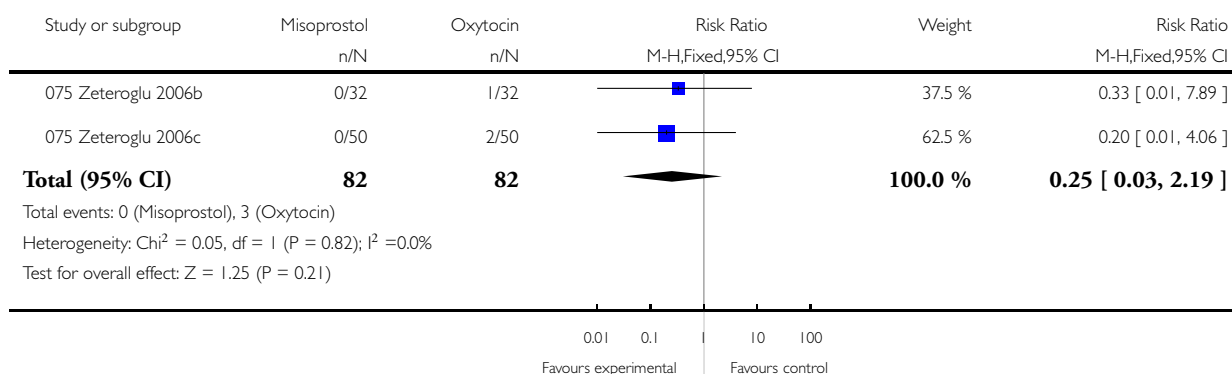


### Analysis 38.7. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 7 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 7 Postpartum haemorrhage

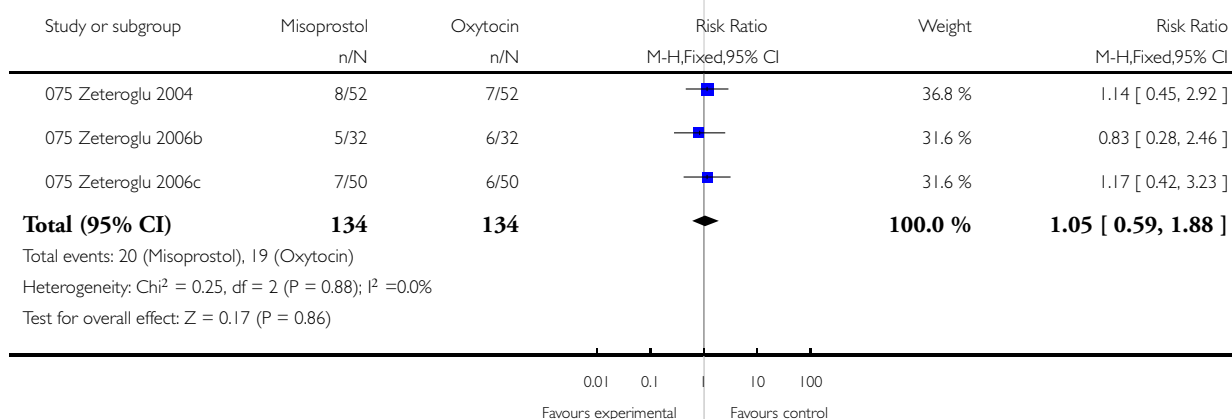


### Analysis 38.8. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 8 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 8 Meconium-stained liquor



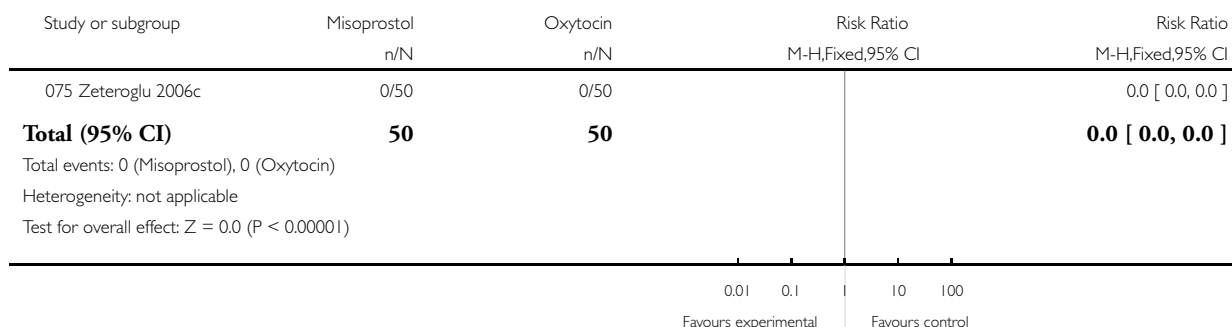


### Analysis 38.9. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 9 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 9 Perinatal death

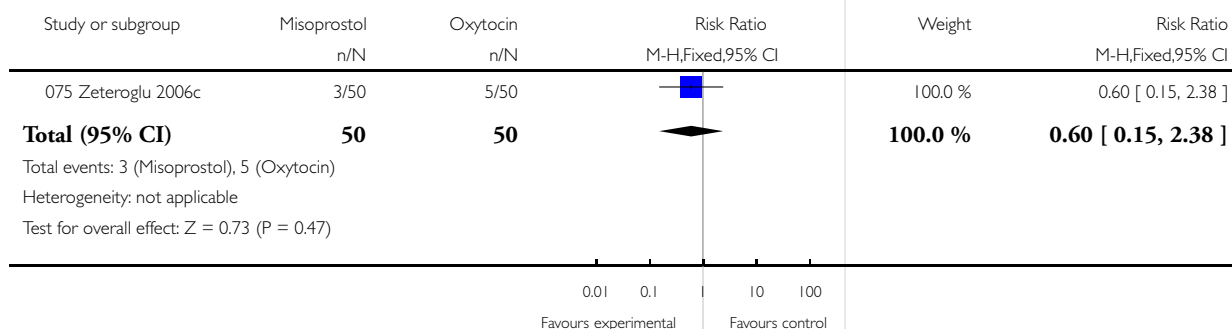


### Analysis 38.10. Comparison 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix, Outcome 10 Serious maternal complications.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 38 Misoprostol versus oxytocin: all multiparae with unfavourable cervix

Outcome: 10 Serious maternal complications

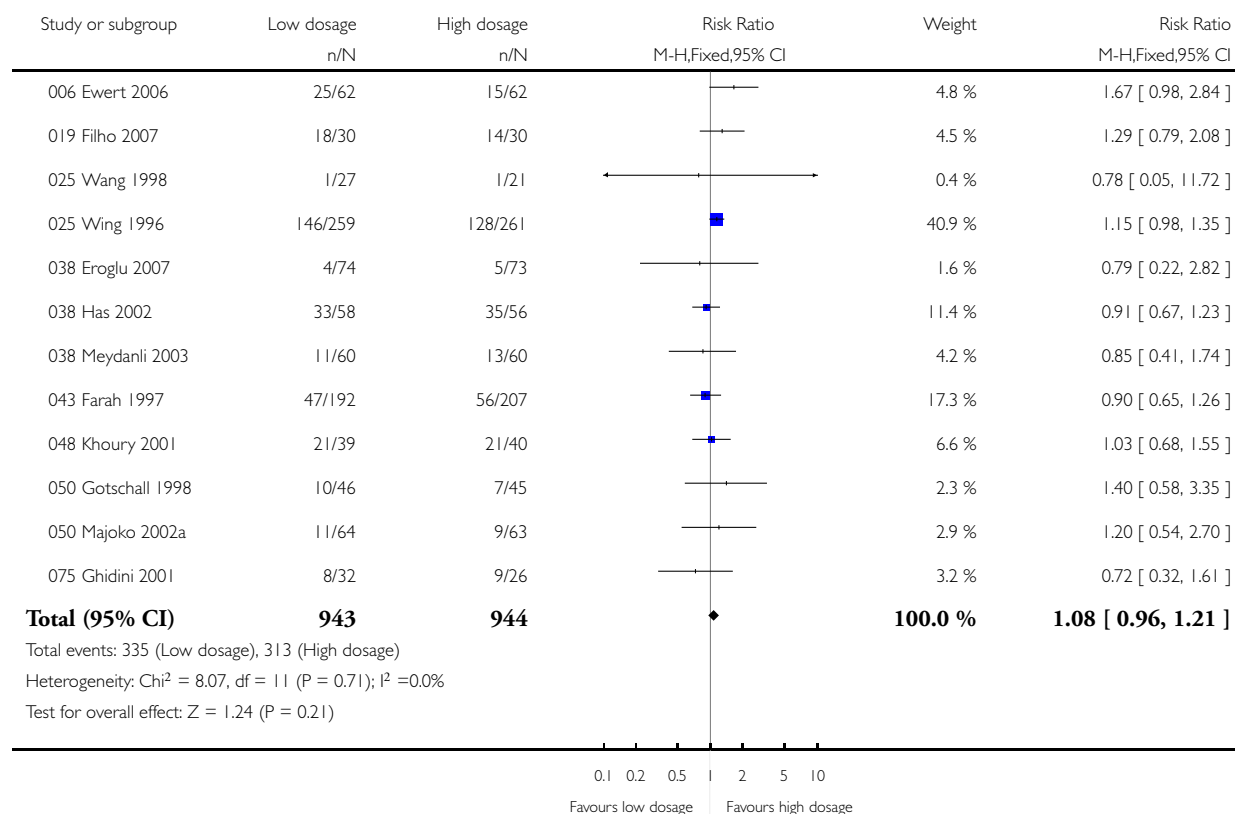


### Analysis 39.1. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 1 Vaginal delivery not achieved within 24 hours

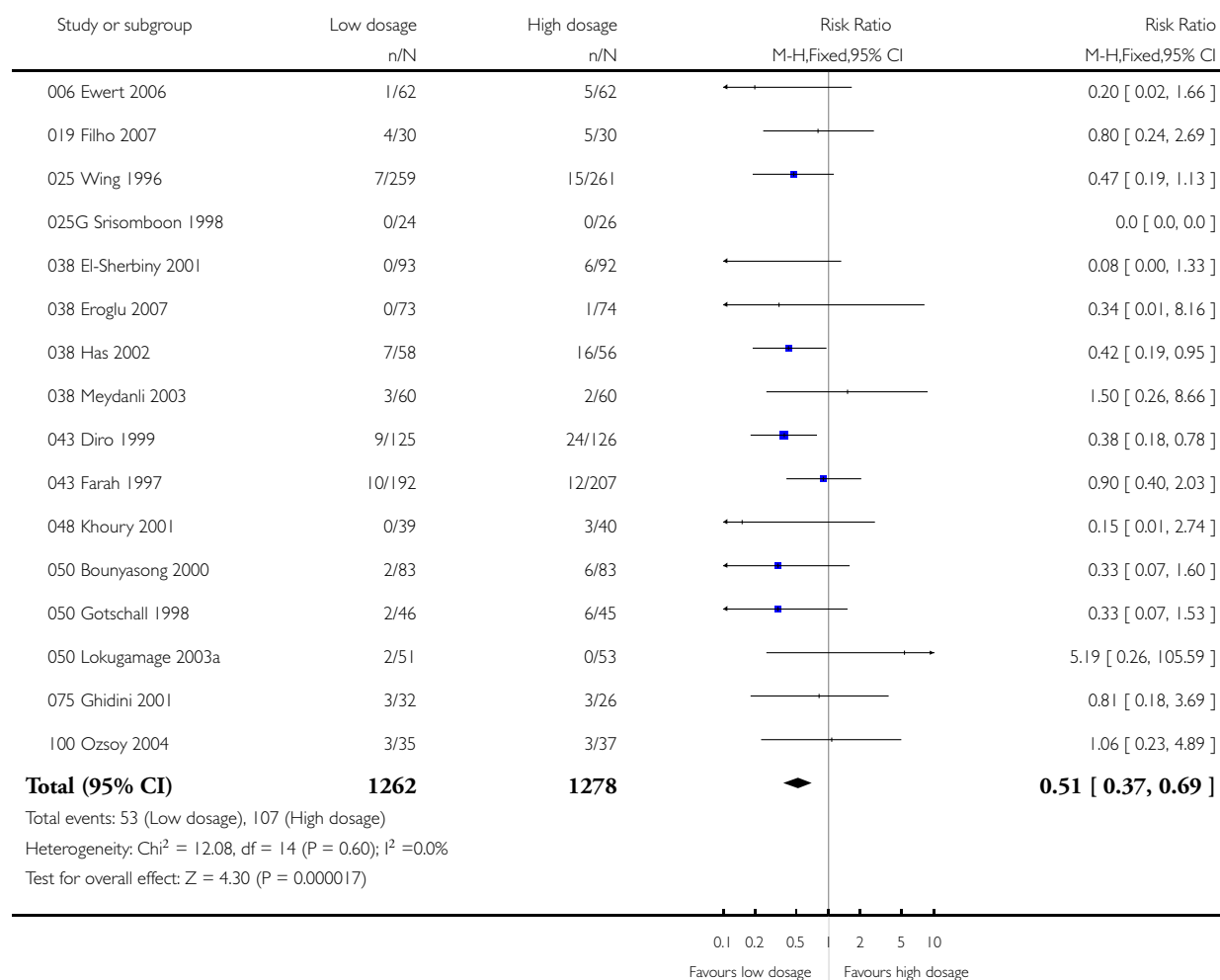


### Analysis 39.2. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 2 Uterine hyperstimulation with FHR changes

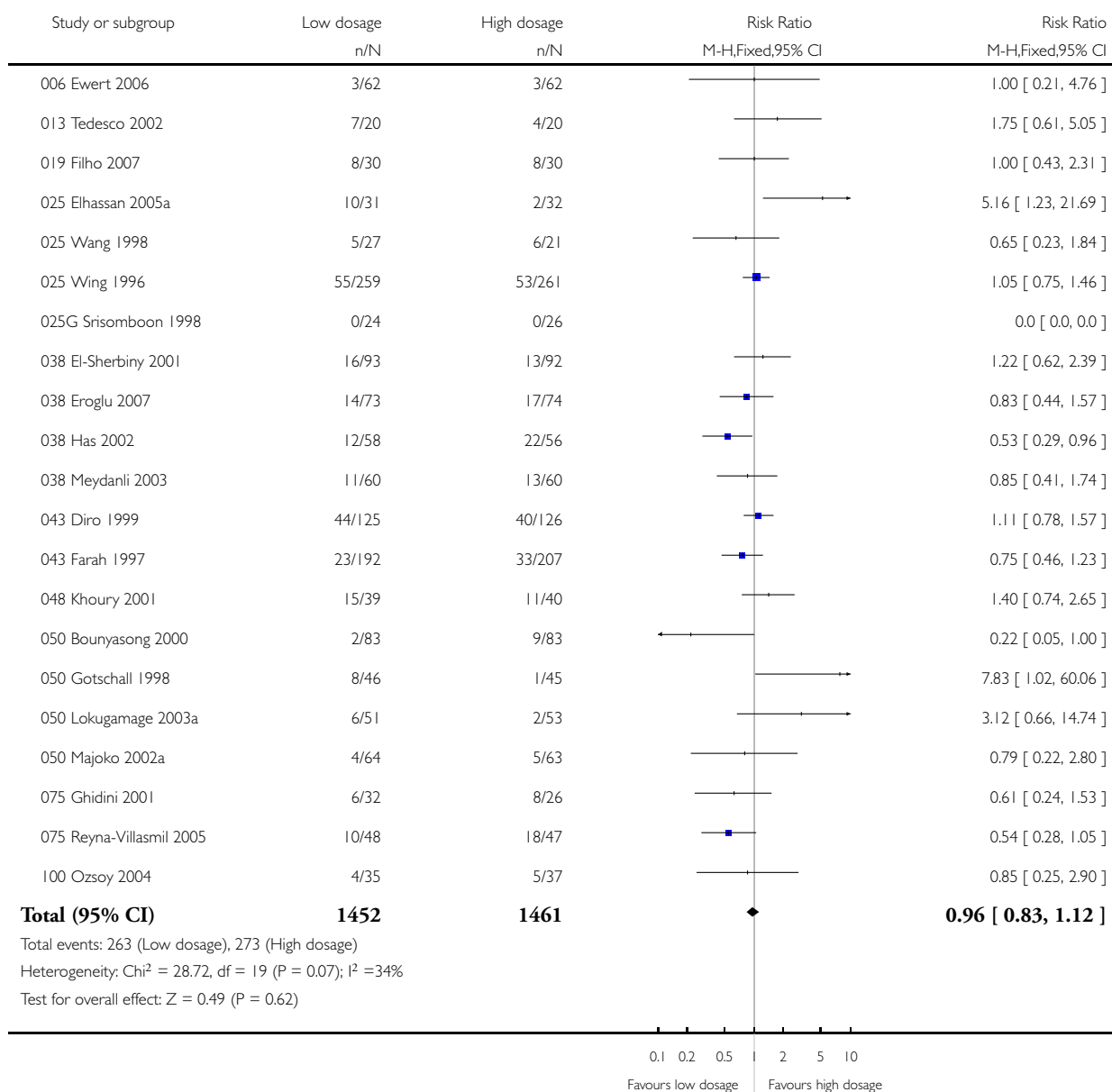


### Analysis 39.3. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 3 Caesarean section



### Analysis 39.4. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 4 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 4 Serious maternal morbidity or death

Study or subgroup	Lower dose n/N	Higher dose n/N	Risk Ratio M-H,Fixed,95% CI	Risk Ratio M-H,Fixed,95% CI
025 Stitely 2000	0/1	0/1		0.0 [ 0.0, 0.0 ]
025 Wing 1996	0/1	0/1		0.0 [ 0.0, 0.0 ]
025G Srisomboon 1998	0/1	0/1		0.0 [ 0.0, 0.0 ]
200 Lee 1997	0/1	0/1		0.0 [ 0.0, 0.0 ]
<b>Total (95% CI)</b>	<b>4</b>	<b>4</b>		<b>0.0 [ 0.0, 0.0 ]</b>
Total events: 0 (Lower dose), 0 (Higher dose)				
Heterogeneity: Chi <sup>2</sup> = 0.0, df = 0 (P<0.00001); I <sup>2</sup> = 0.0%				
Test for overall effect: Z = 0.0 (P = 0.00001)				



0.1 0.2 0.5 2 5 10  
Favours treatment Favours control

### Analysis 39.5. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 5 Cervix unfavourable/unchanged after 12-24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 5 Cervix unfavourable/unchanged after 12-24 hours

Study or subgroup	Lower dose n/N	Higher dose n/N	Risk Ratio M-H,Fixed,95% CI	Weight	Risk Ratio M-H,Fixed,95% CI
048 Khoury 2001	19/34	21/35		100.0 %	0.93 [ 0.62, 1.39 ]
<b>Total (95% CI)</b>	<b>34</b>	<b>35</b>		<b>100.0 %</b>	<b>0.93 [ 0.62, 1.39 ]</b>
Total events: 19 (Lower dose), 21 (Higher dose)					
Heterogeneity: not applicable					
Test for overall effect: Z = 0.35 (P = 0.73)					

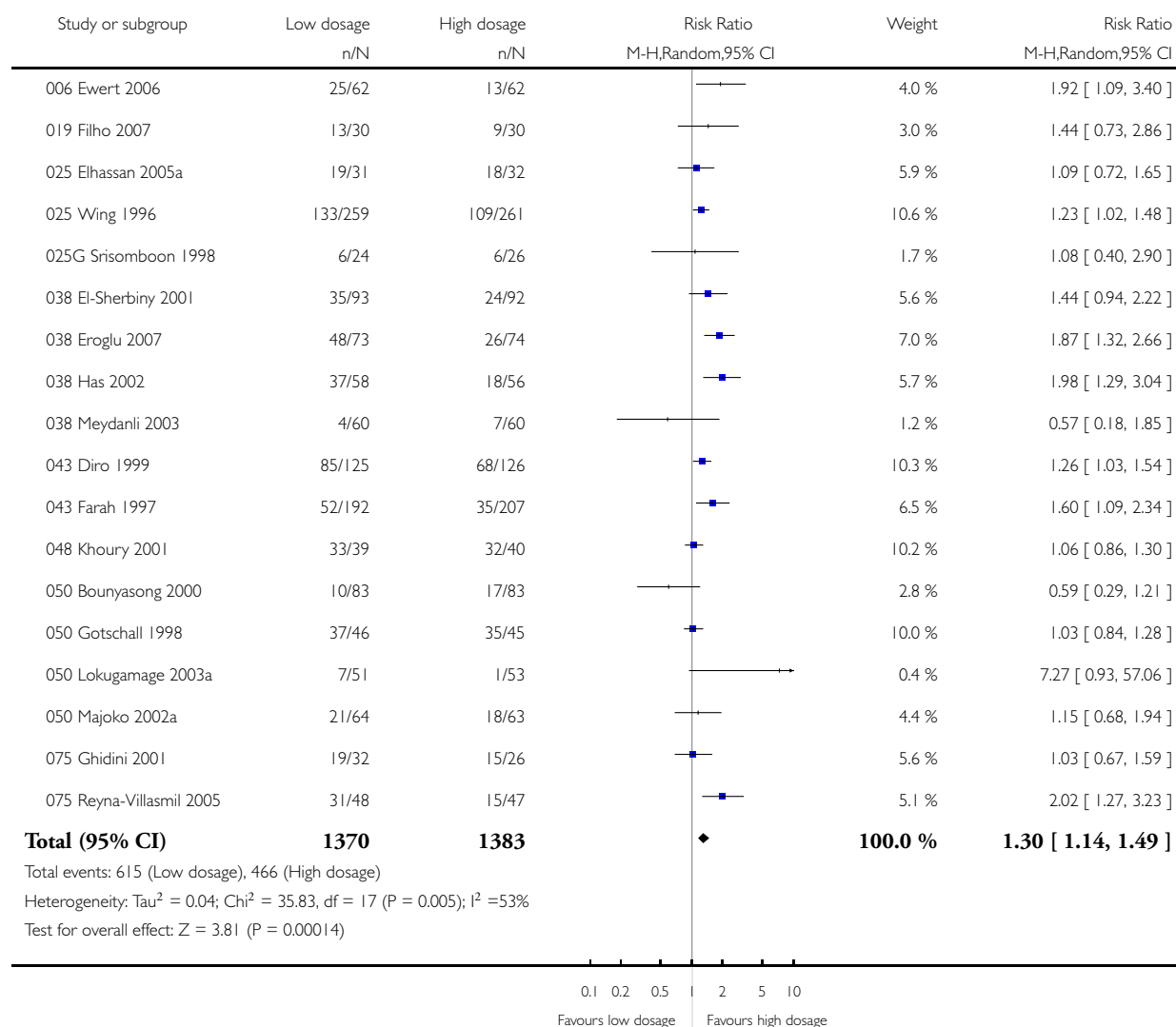
0.1 0.2 0.5 2 5 10  
Favours treatment Favours control

### Analysis 39.6. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 6 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 6 Oxytocin augmentation

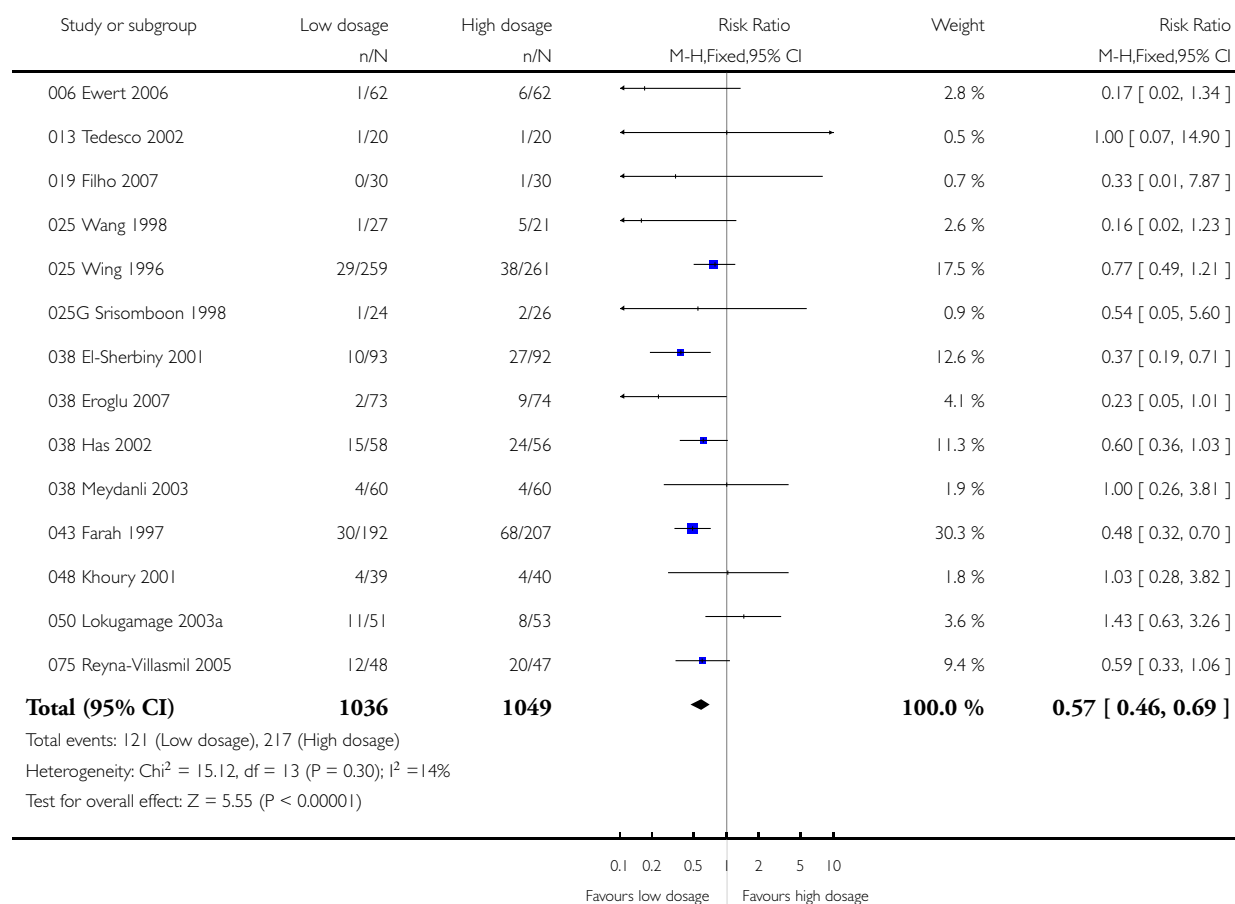


### Analysis 39.7. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 7 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 7 Uterine hyperstimulation without FHR changes

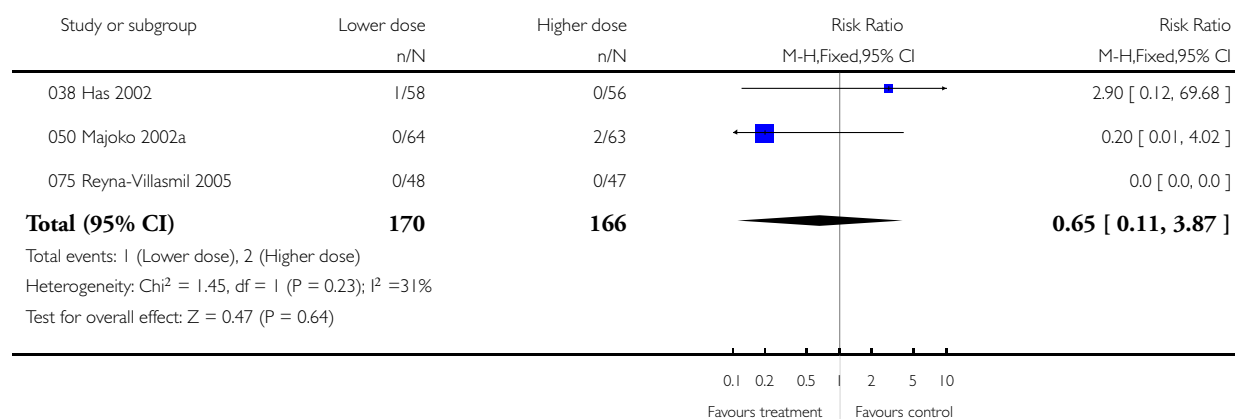


### Analysis 39.8. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 8 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 8 Uterine rupture

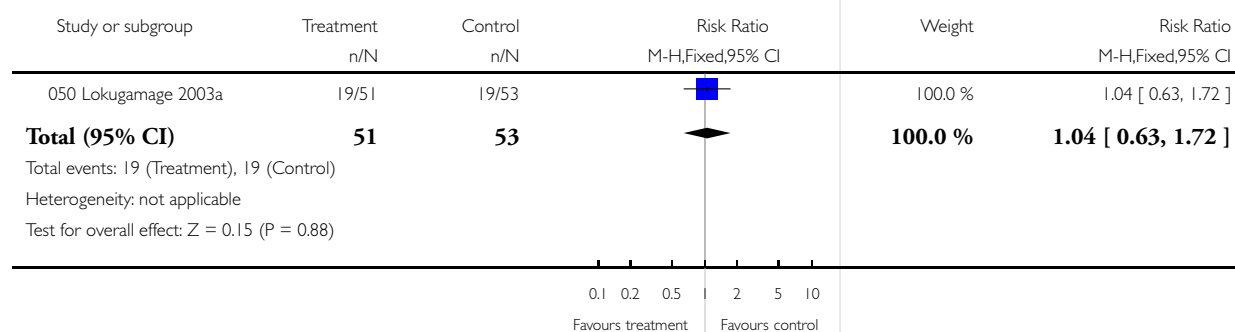


### Analysis 39.9. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 9 Epidural.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 9 Epidural



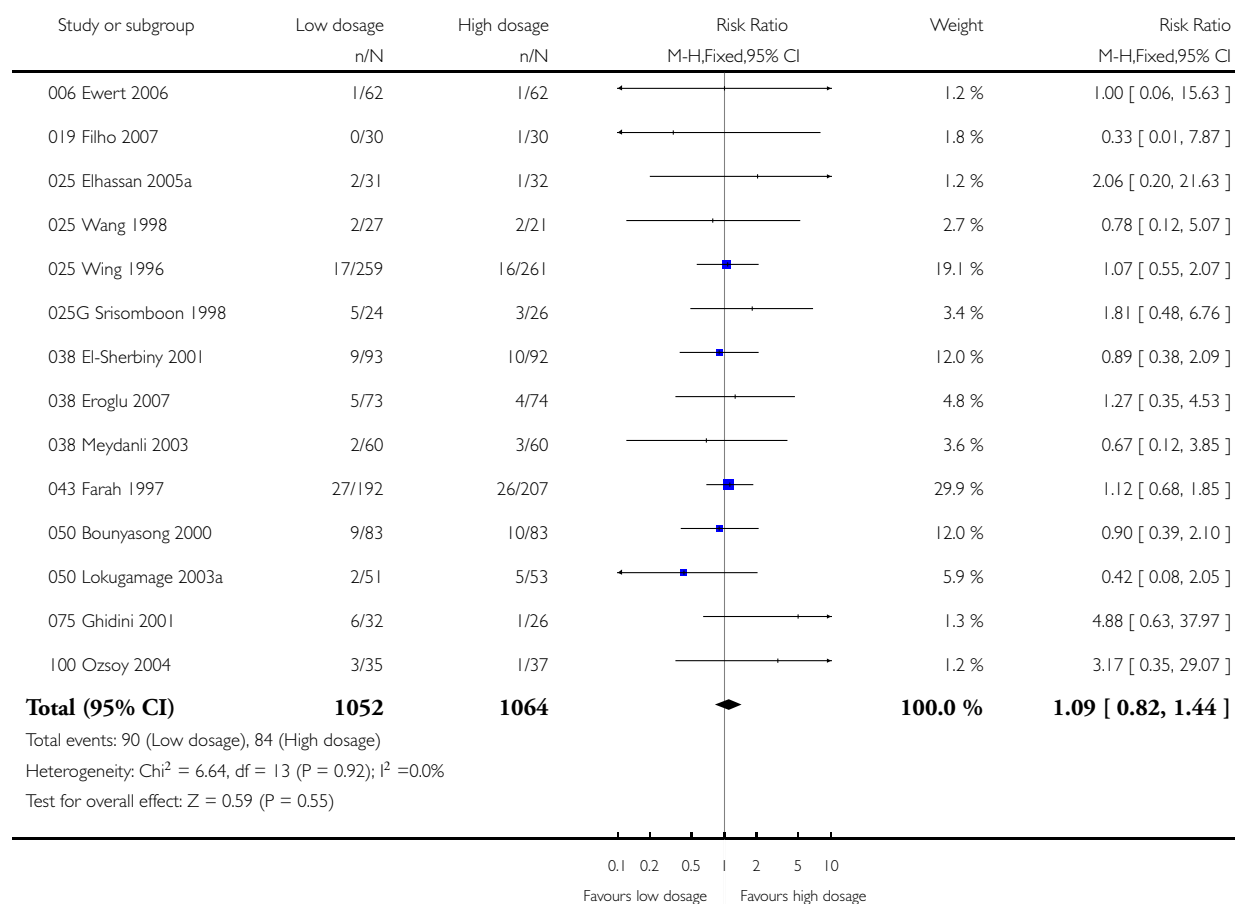


### Analysis 39.10. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 10 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 10 Instrumental vaginal delivery

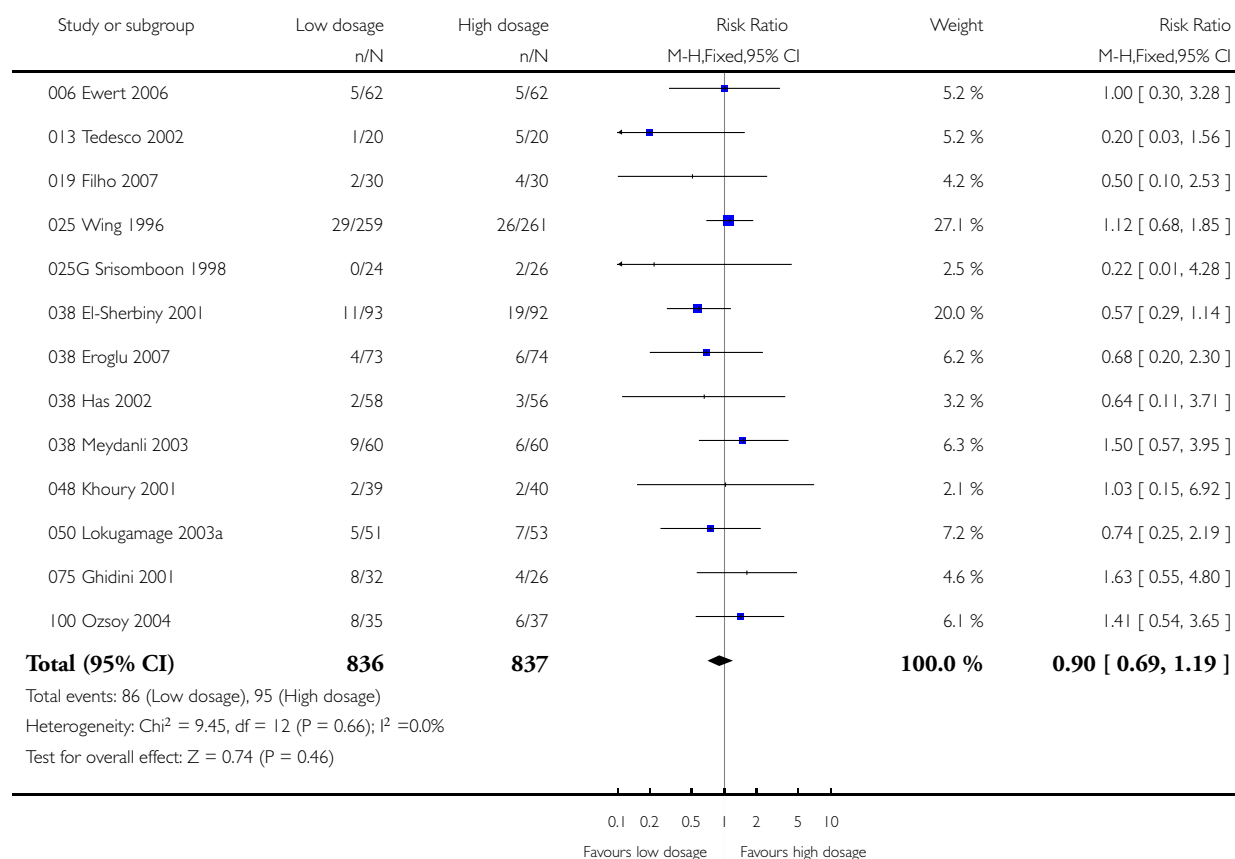


### Analysis 39.11. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 11 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 11 Meconium-stained liquor

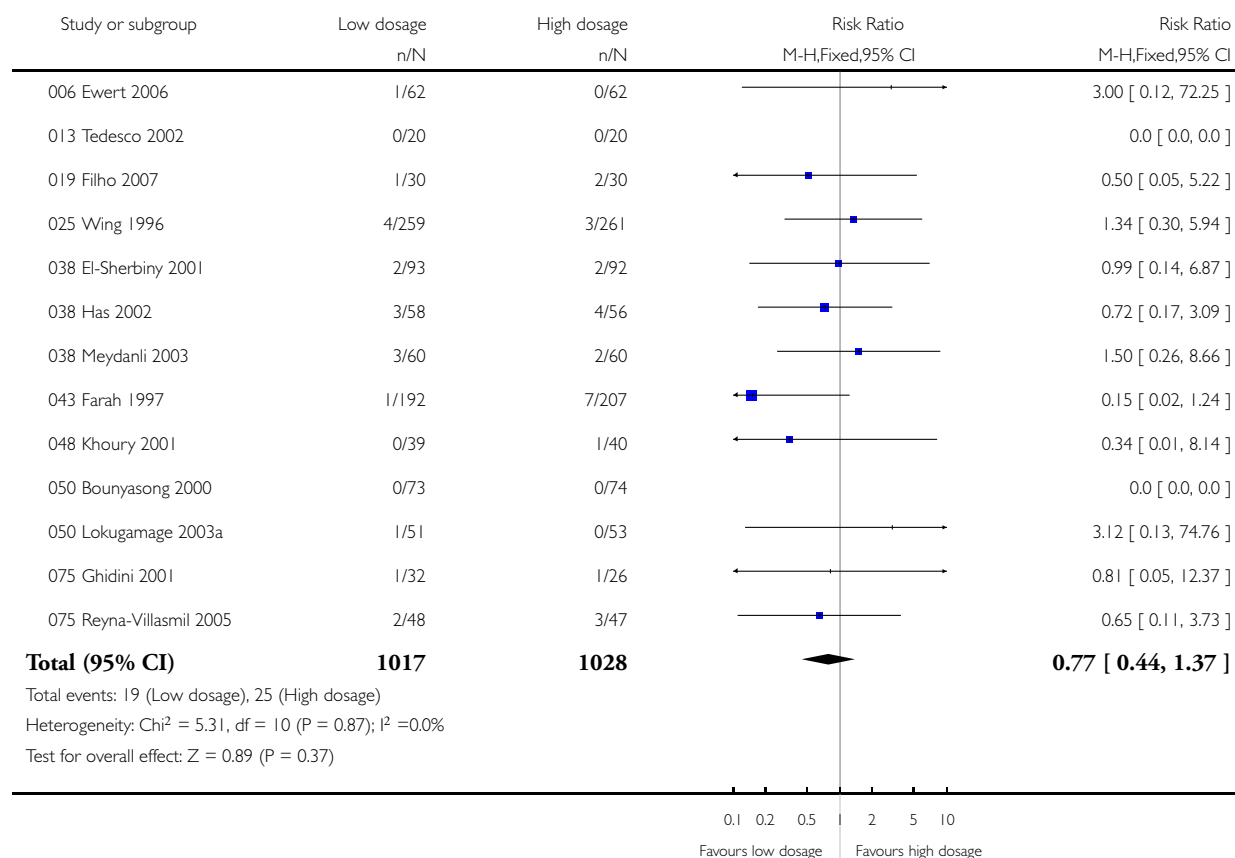


### Analysis 39.12. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 12 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 12 Apgar score < 7 at 5 minutes

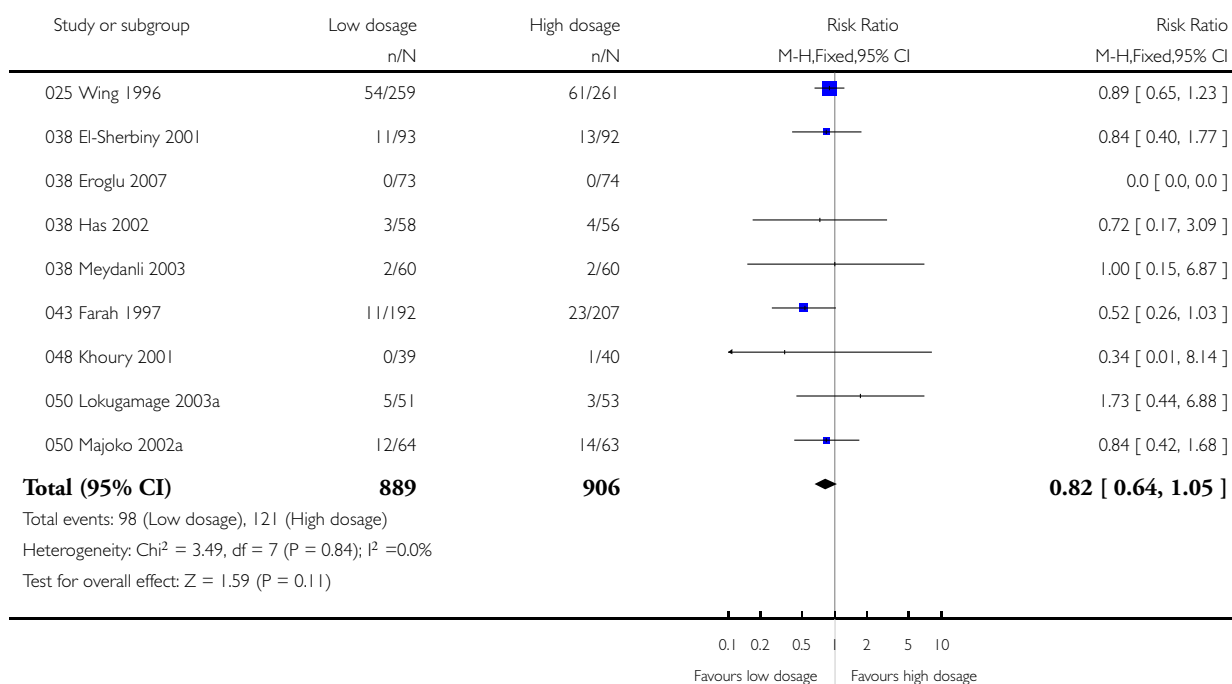


### Analysis 39.13. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 13 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 13 Neonatal intensive care unit admission

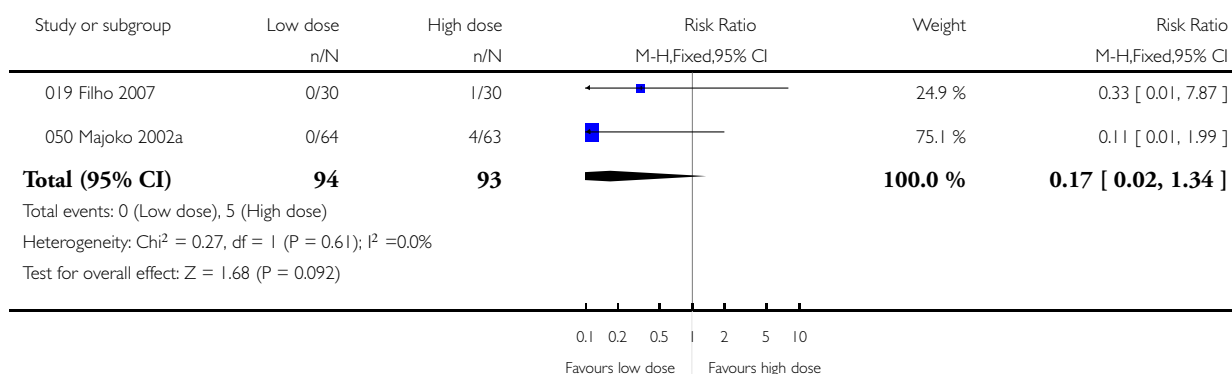


### Analysis 39.14. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 14 Perinatal death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 14 Perinatal death

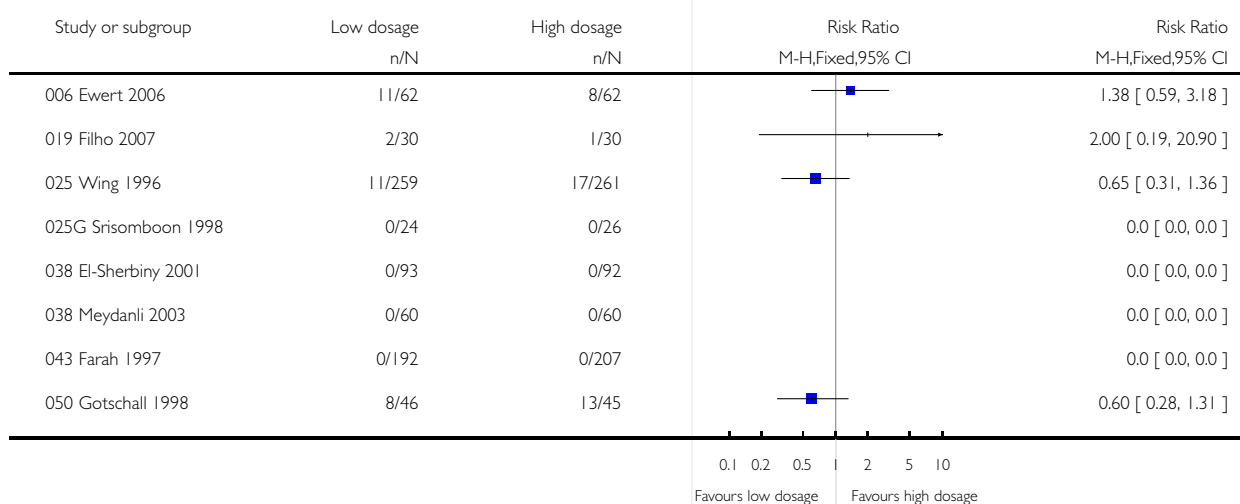


### Analysis 39.15. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 15 Maternal side effects (eg nausea, vomiting, diarrhoea).

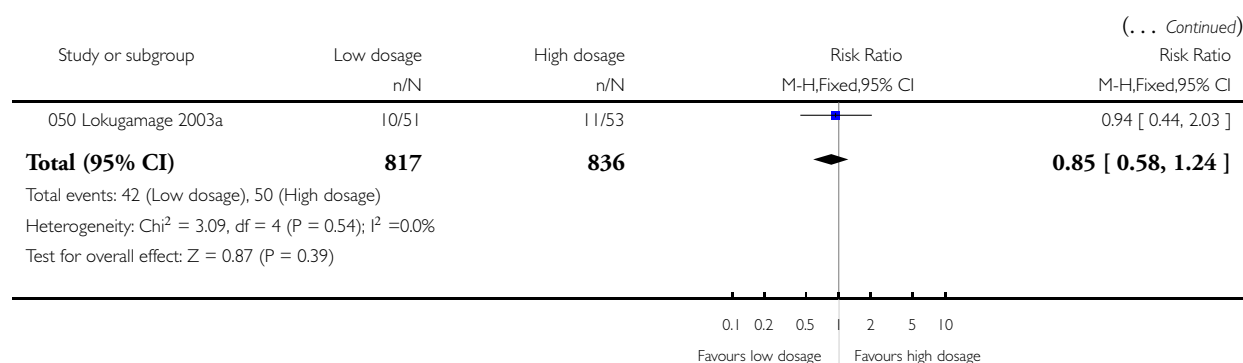
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 15 Maternal side effects (eg nausea, vomiting, diarrhoea)



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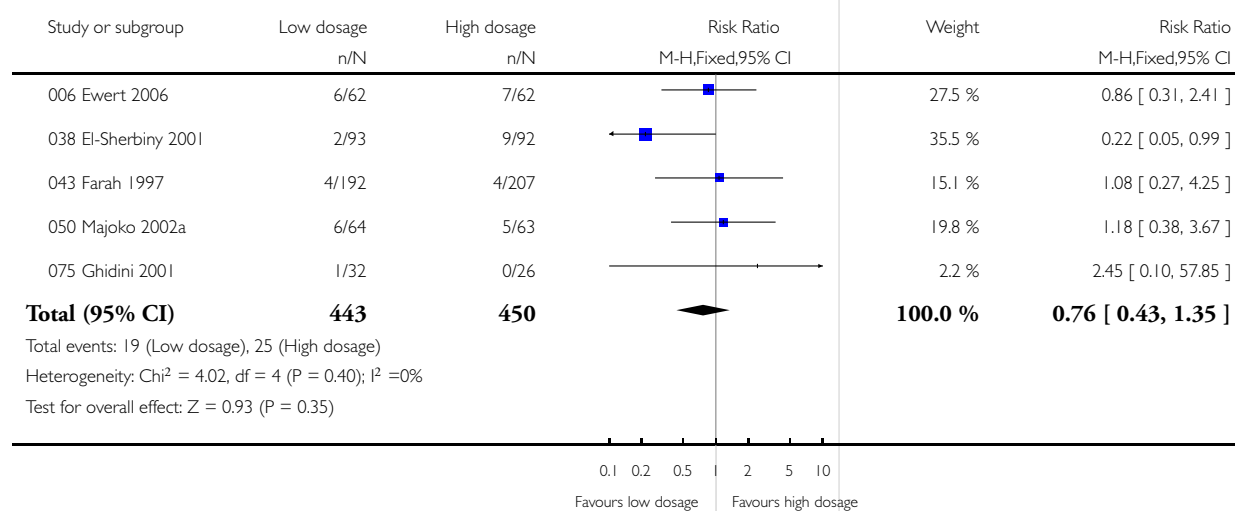


### Analysis 39.16. Comparison 39 Misoprostol lower versus higher dose: all women, Outcome 16 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 39 Misoprostol lower versus higher dose: all women

Outcome: 16 Postpartum haemorrhage

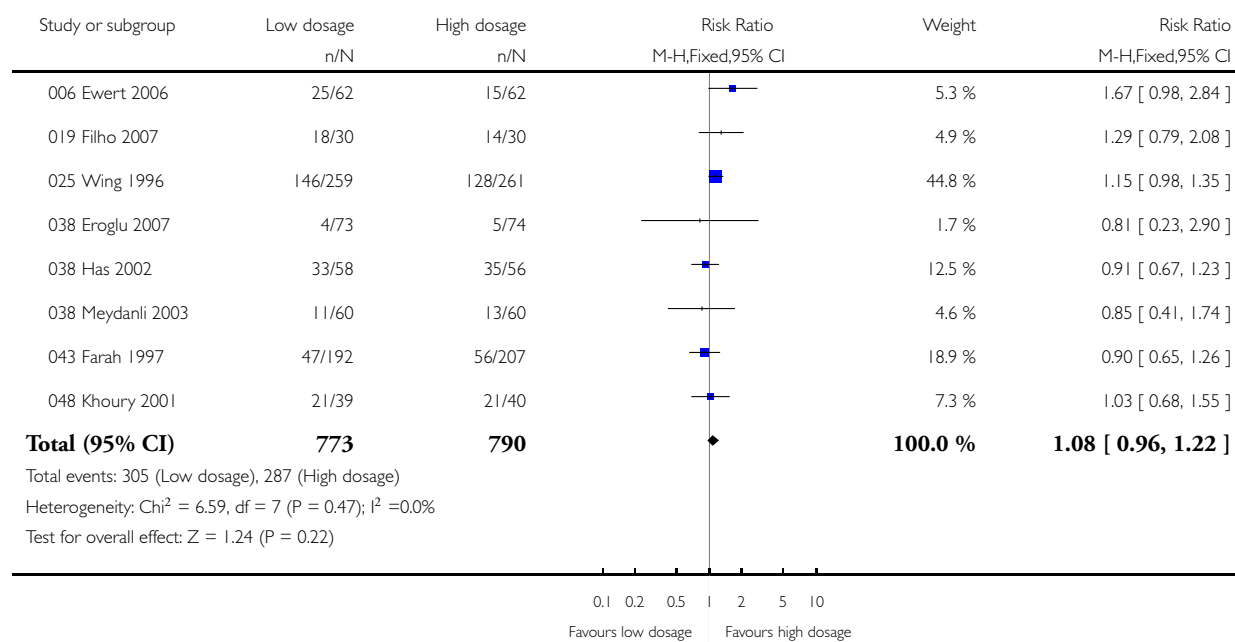


# **Analysis 40.1. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

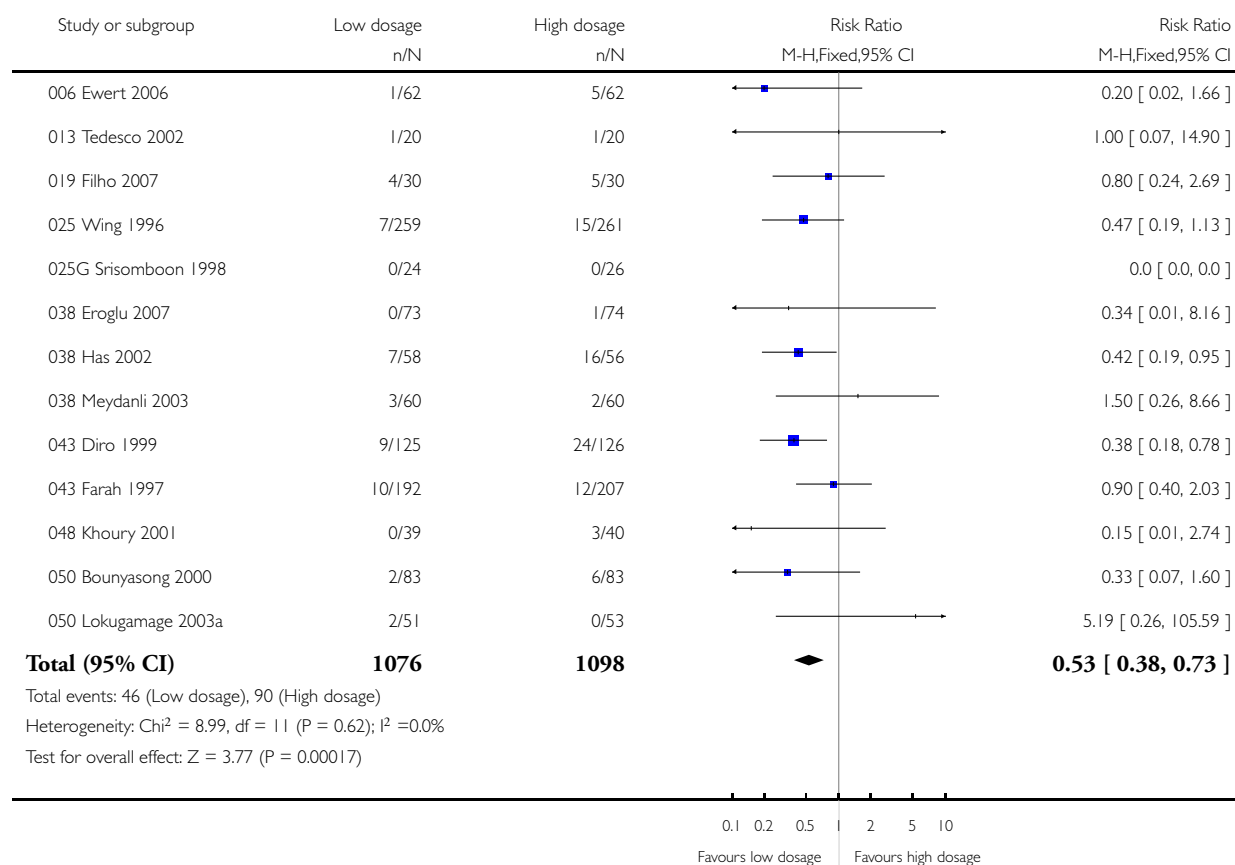


## Analysis 40.2. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes



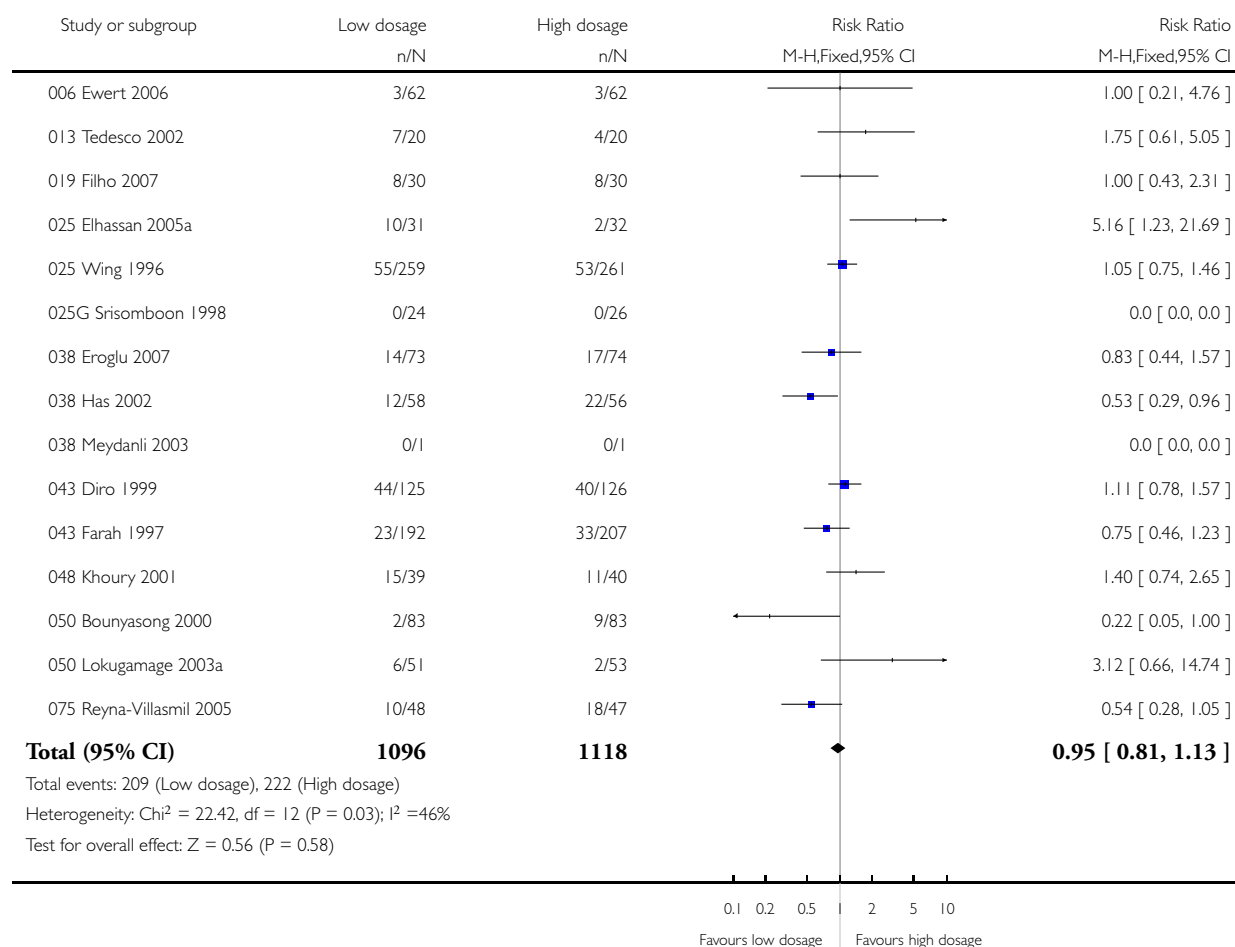


### Analysis 40.3. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 3 Caesarean section

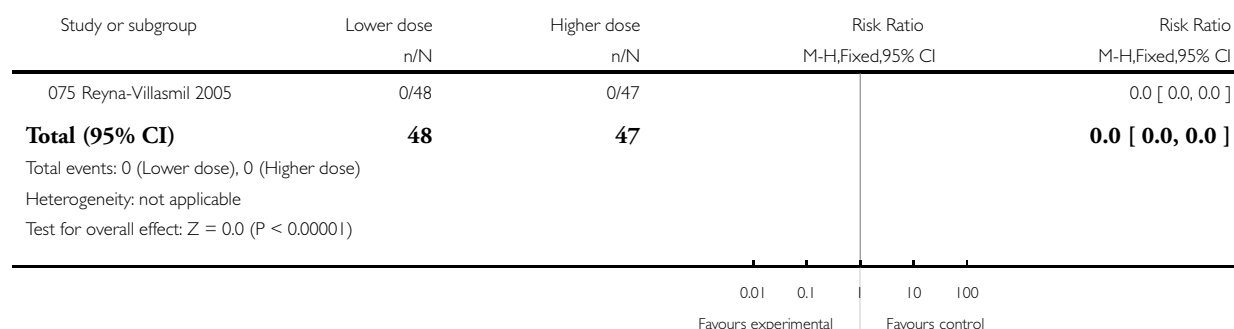


#### Analysis 40.4. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 4 Uterine rupture.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 4 Uterine rupture

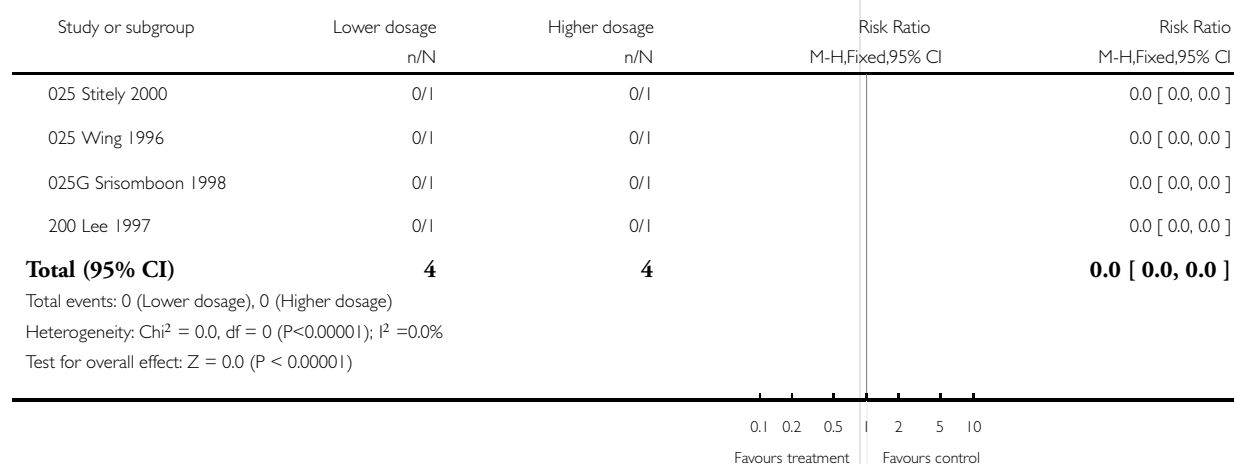


#### Analysis 40.5. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 5 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 5 Serious maternal morbidity or death

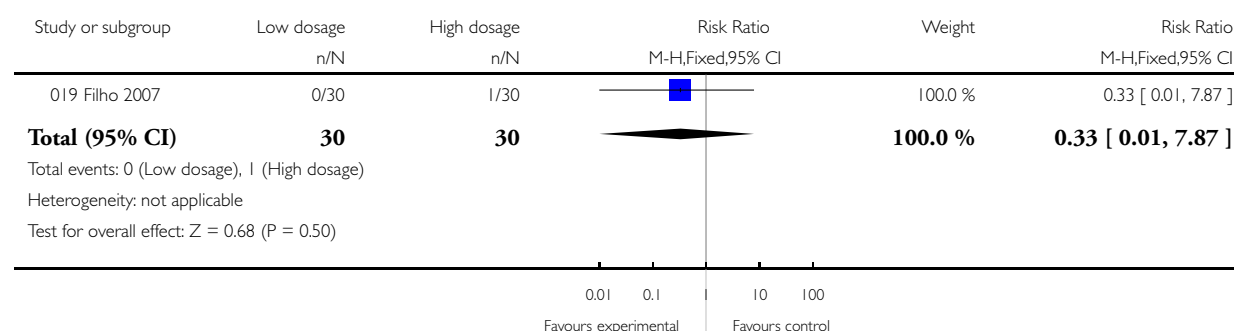


# **Analysis 40.6. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 6 Perinatal death.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 6 Perinatal death

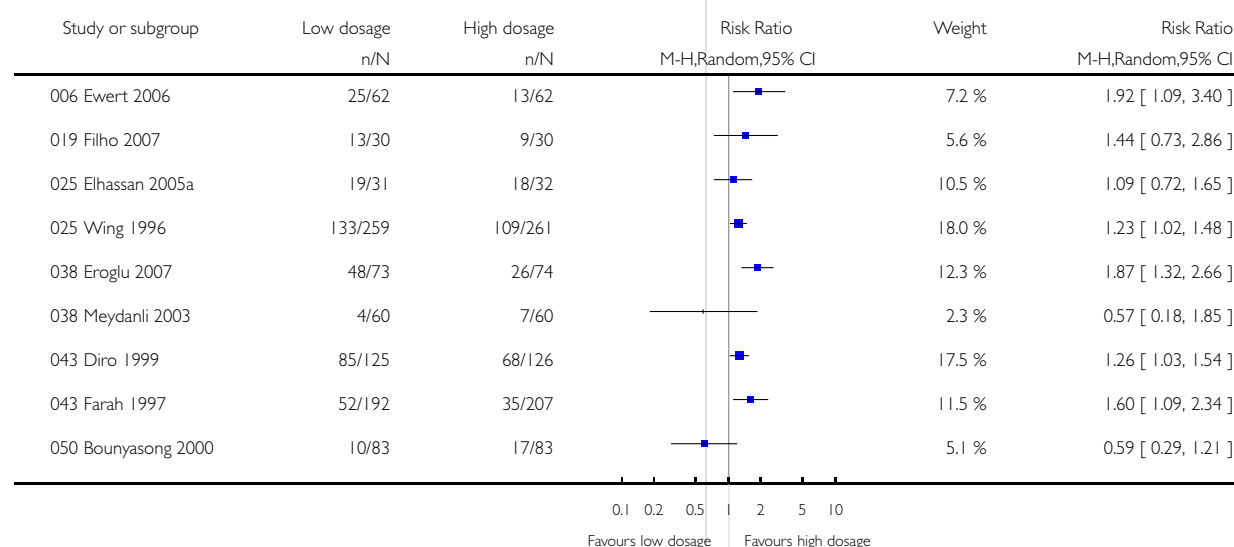


# **Analysis 40.7. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 7 Oxytocin augmentation.**

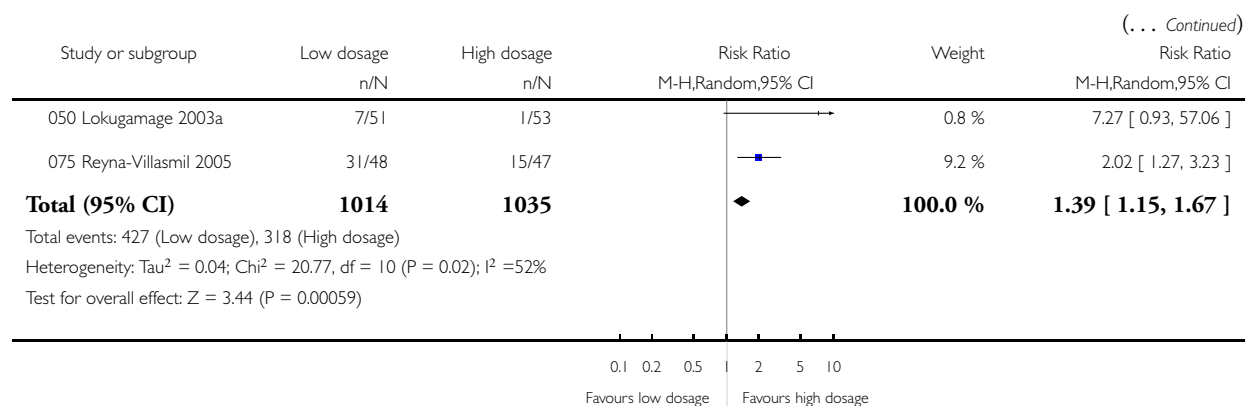
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 7 Oxytocin augmentation



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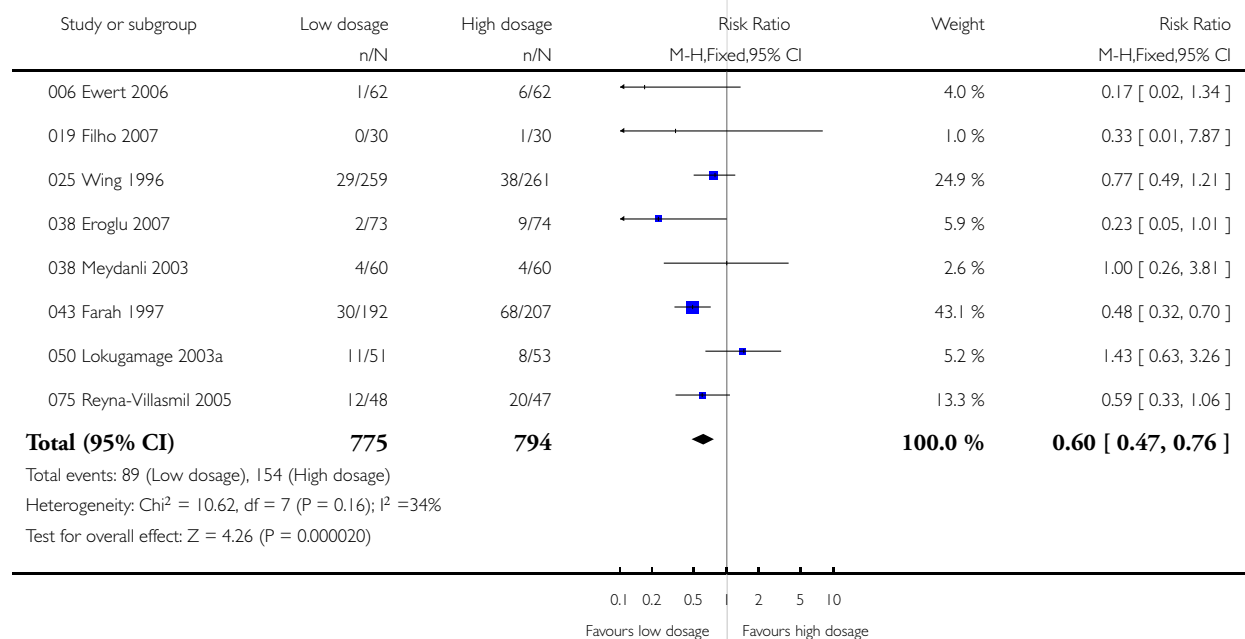


**Analysis 40.8. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 8 Uterine hyperstimulation without FHR changes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 8 Uterine hyperstimulation without FHR changes

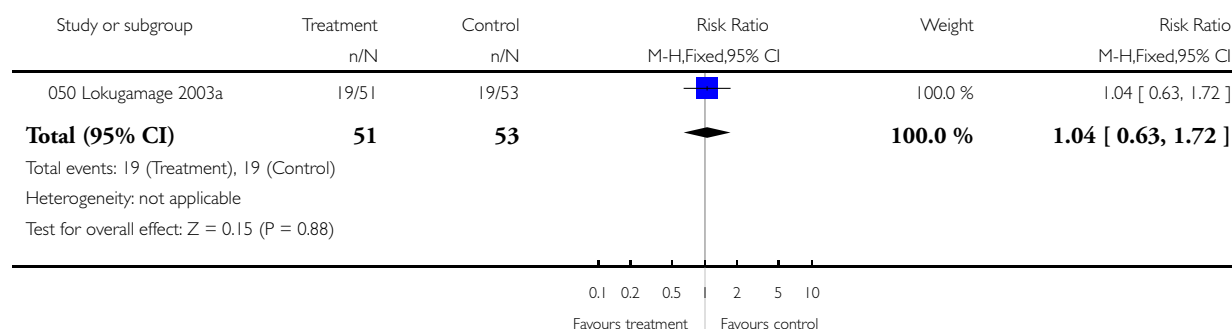


### Analysis 40.9. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 9 Epidural.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 9 Epidural

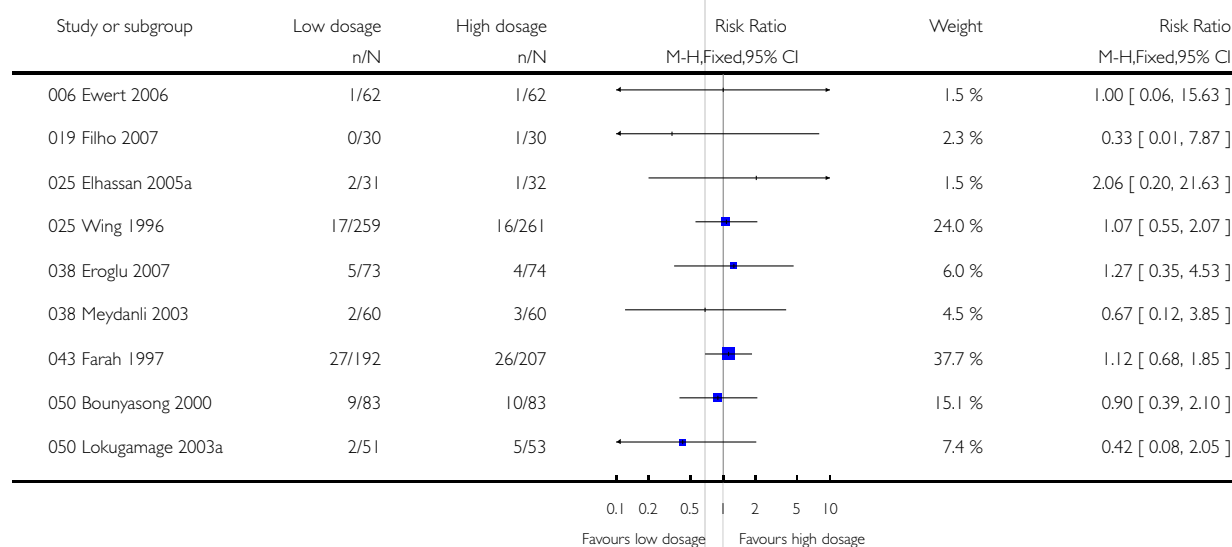


### Analysis 40.10. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 10 Instrumental vaginal delivery.

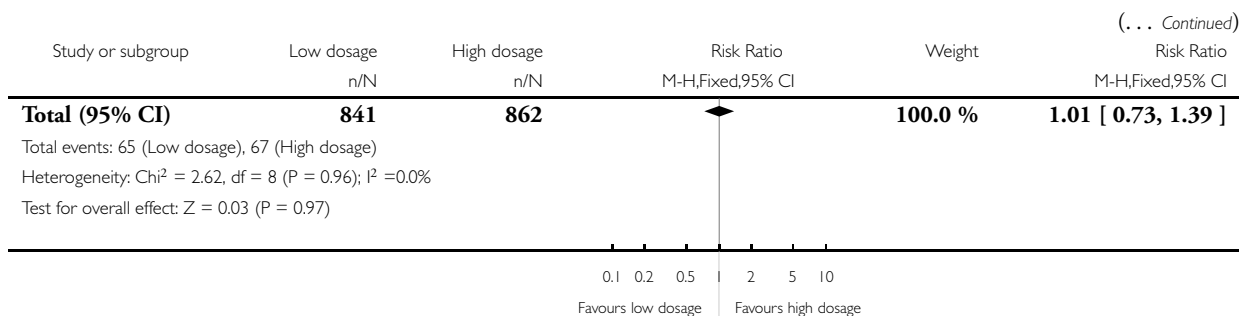
Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 10 Instrumental vaginal delivery



(Continued ...)

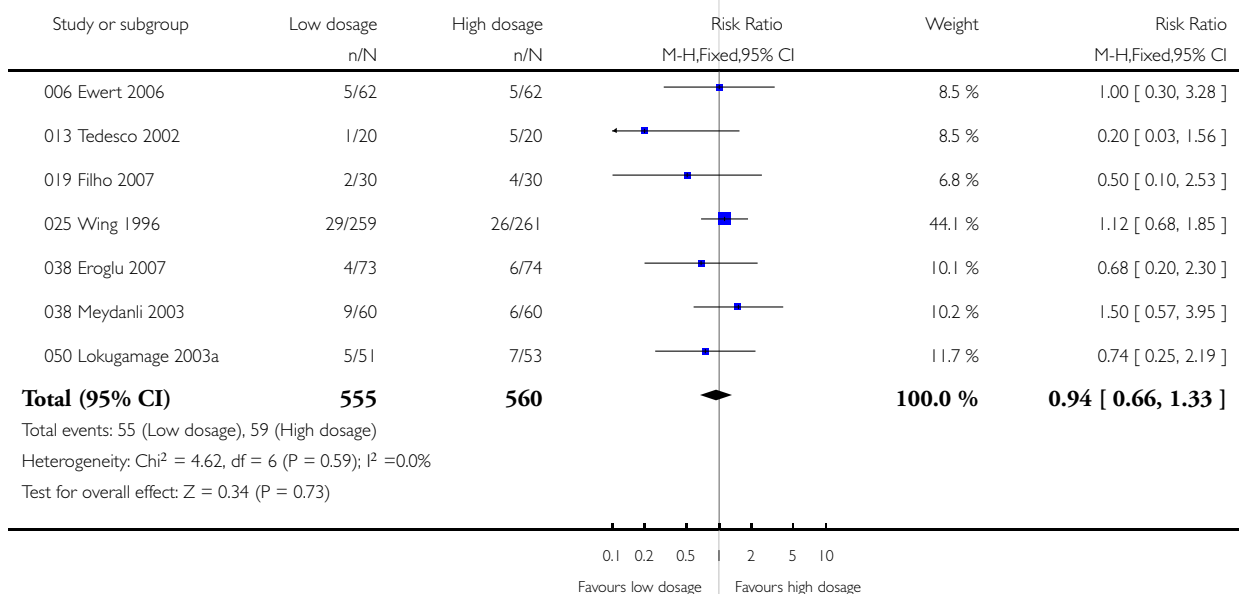


**Analysis 40.11. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 11 Meconium-stained liquor.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 11 Meconium-stained liquor

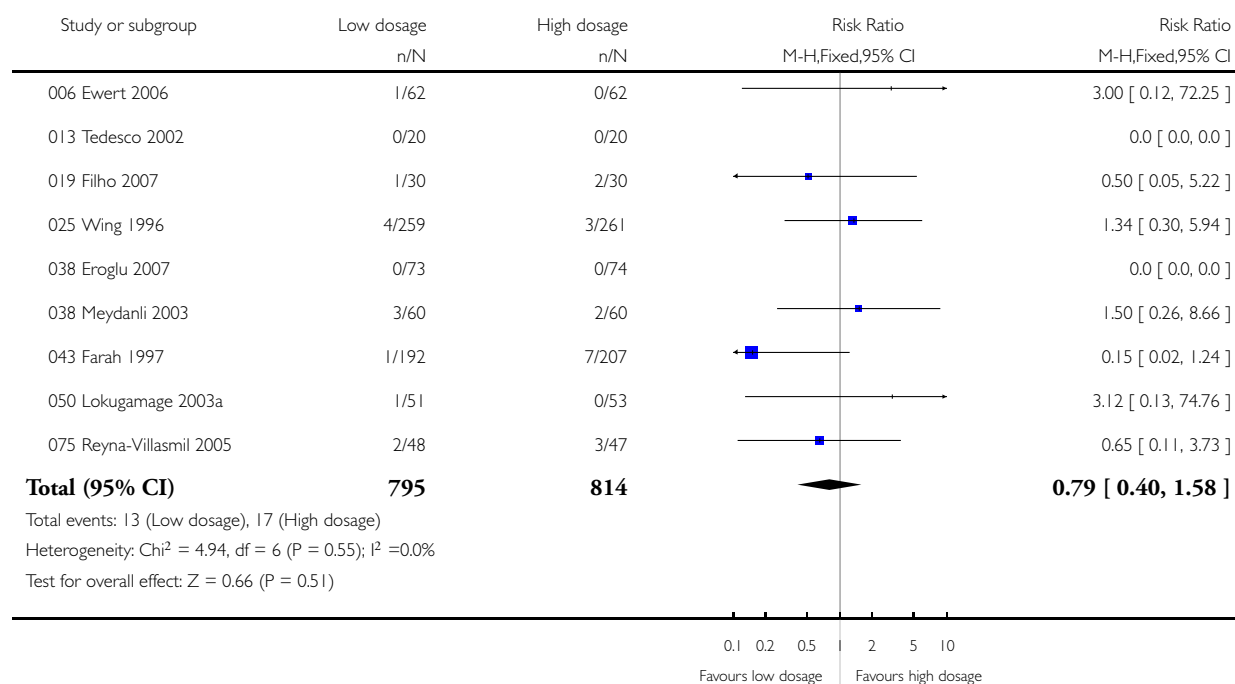


**Analysis 40.12. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 12 Apgar score < 7 at 5 minutes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 12 Apgar score < 7 at 5 minutes

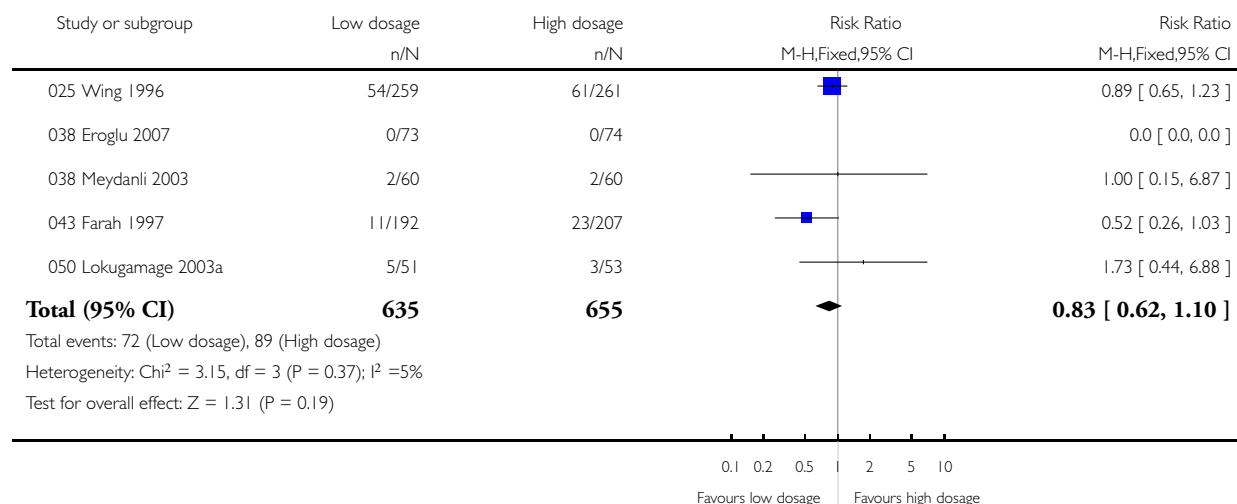


**Analysis 40.13. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 13 Neonatal intensive care unit admission.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 13 Neonatal intensive care unit admission



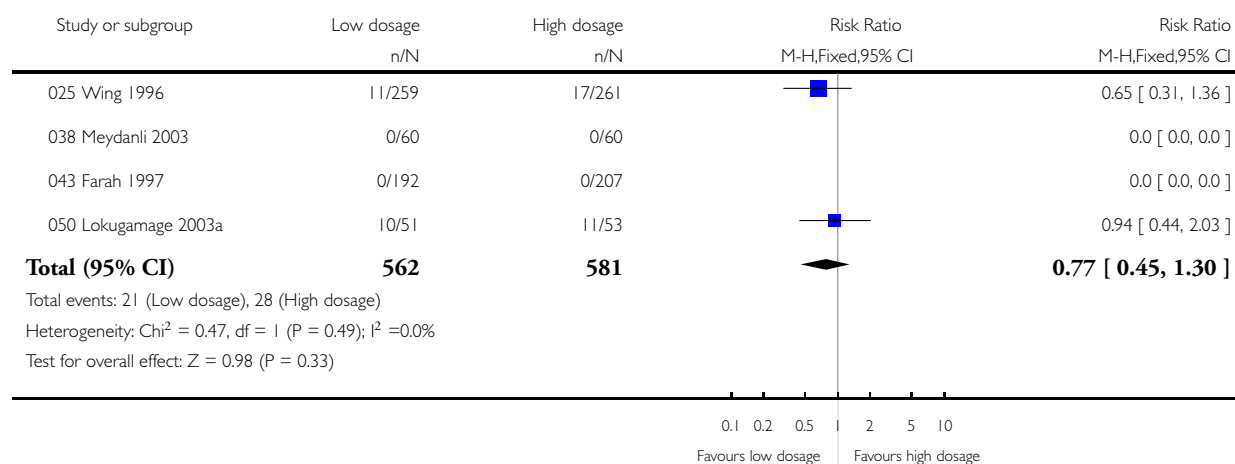


#### Analysis 40.14. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 14 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 14 Maternal side effects (eg nausea, vomiting, diarrhoea)

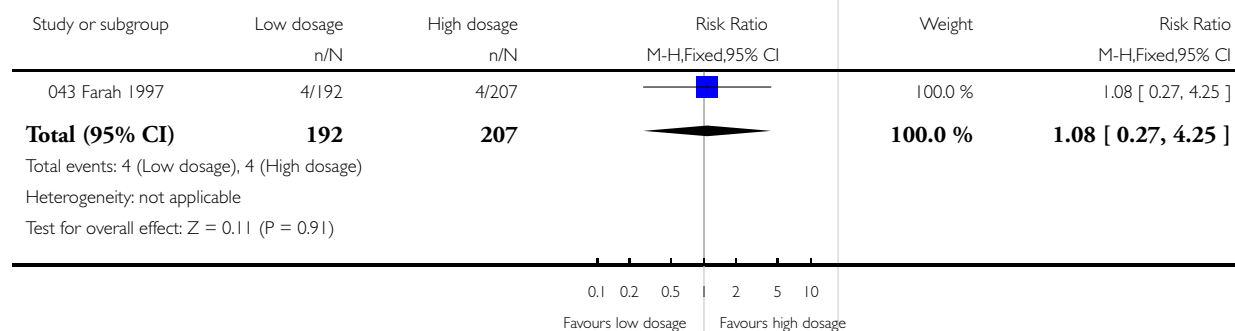


#### Analysis 40.15. Comparison 40 Misoprostol lower versus higher dose: all women with unfavourable cervix, Outcome 15 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 40 Misoprostol lower versus higher dose: all women with unfavourable cervix

Outcome: 15 Postpartum haemorrhage

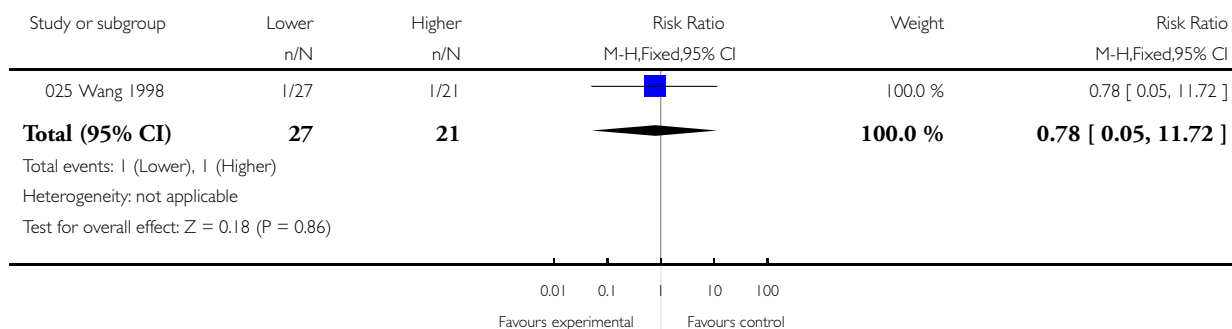


### Analysis 41.1. Comparison 41 Misoprostol lower versus higher dose: all women with intact membranes, Outcome 1 Vaginal delivery not achieved in 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 41 Misoprostol lower versus higher dose: all women with intact membranes

Outcome: 1 Vaginal delivery not achieved in 24 hours

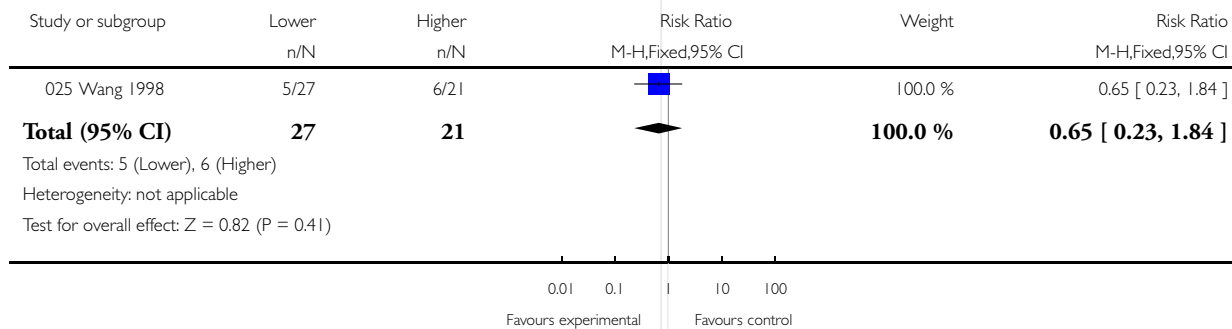


### Analysis 41.2. Comparison 41 Misoprostol lower versus higher dose: all women with intact membranes, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 41 Misoprostol lower versus higher dose: all women with intact membranes

Outcome: 2 Caesarean section

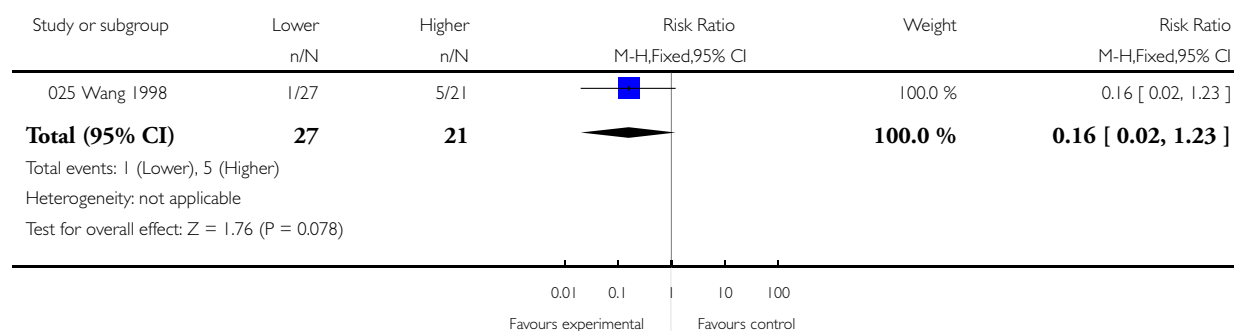


### Analysis 41.3. Comparison 41 Misoprostol lower versus higher dose: all women with intact membranes, Outcome 3 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 41 Misoprostol lower versus higher dose: all women with intact membranes

Outcome: 3 Uterine hyperstimulation without FHR changes

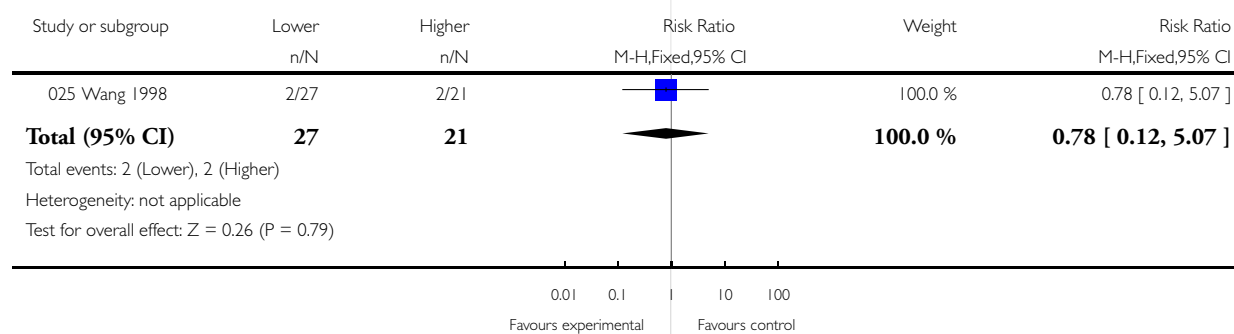


### Analysis 41.4. Comparison 41 Misoprostol lower versus higher dose: all women with intact membranes, Outcome 4 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 41 Misoprostol lower versus higher dose: all women with intact membranes

Outcome: 4 Meconium-stained liquor

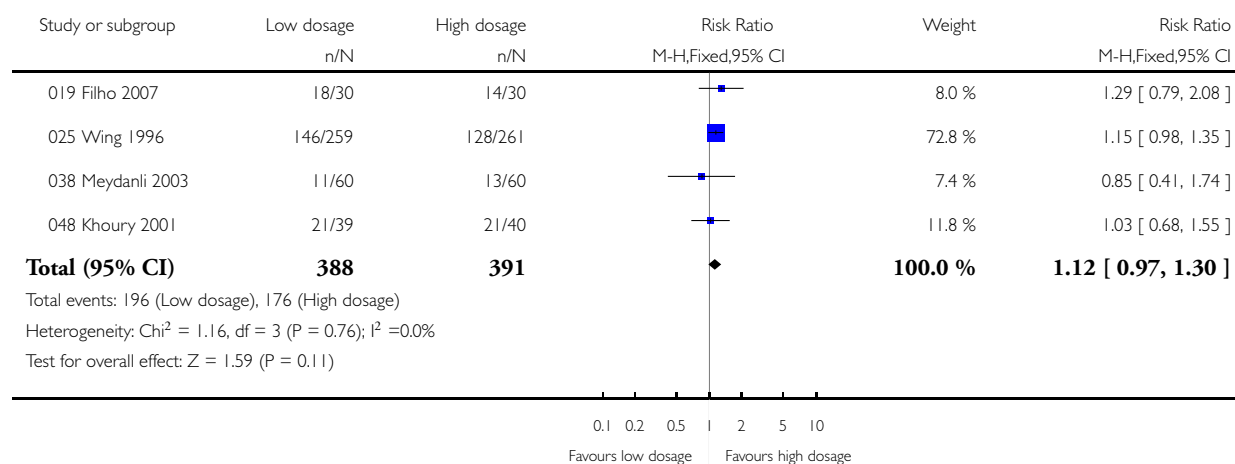


**Analysis 42.1. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours

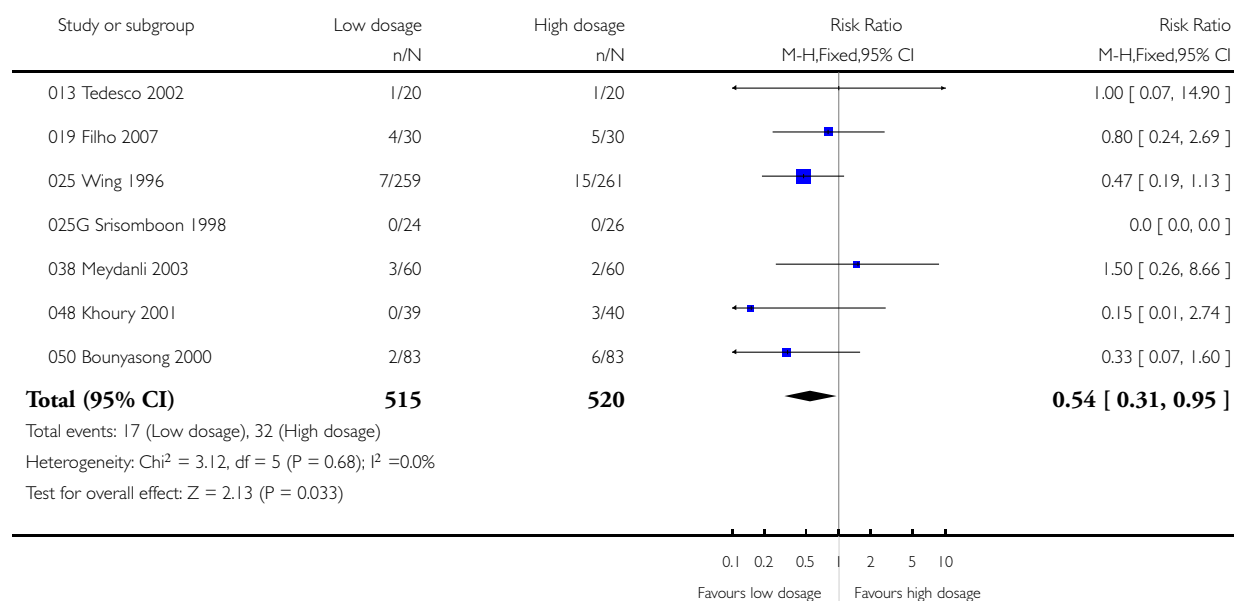


## Analysis 42.2. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

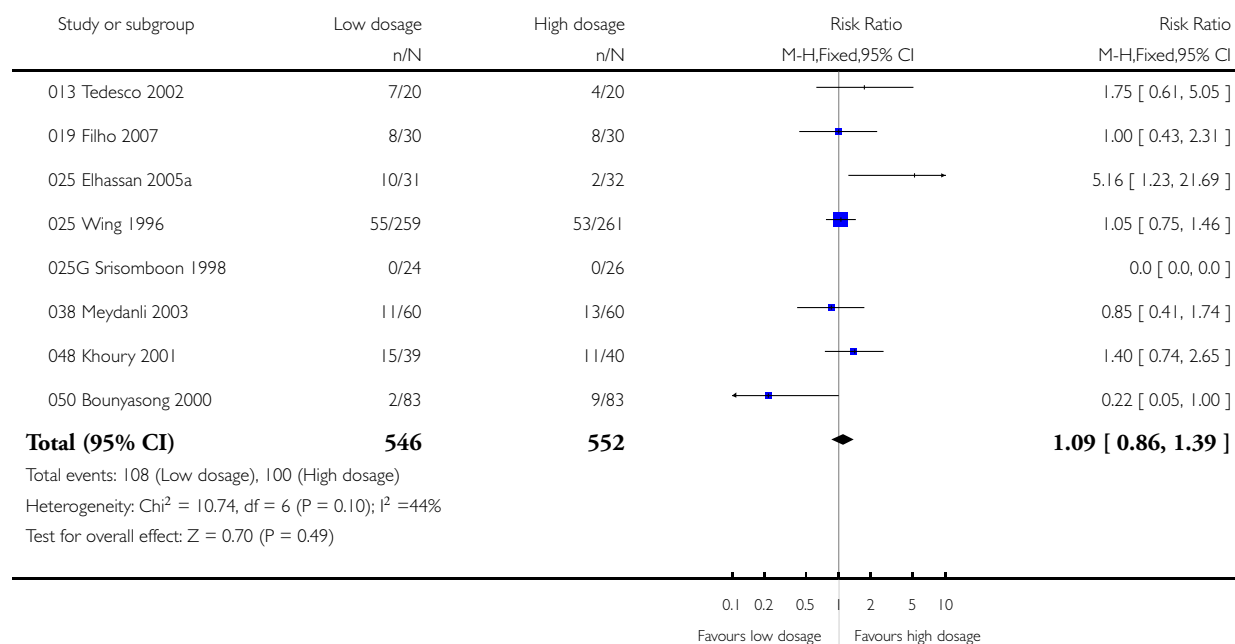


### Analysis 42.3. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 3 Caesarean section

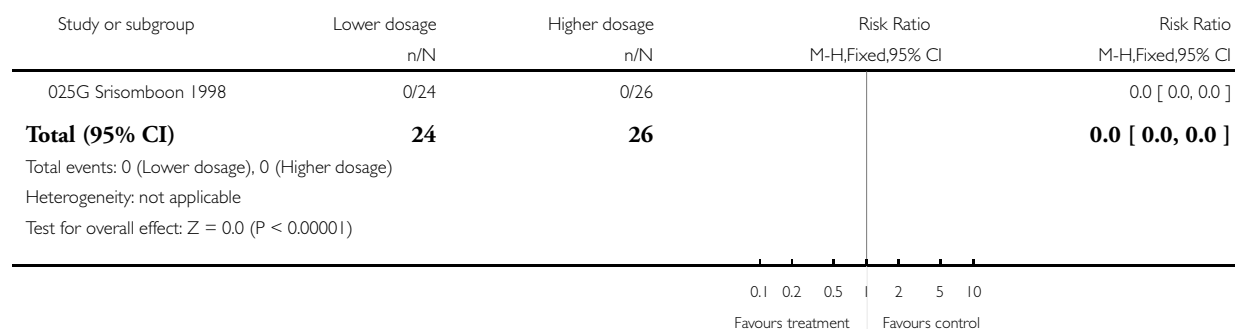


#### Analysis 42.4. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 4 Serious maternal morbidity or death.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 4 Serious maternal morbidity or death

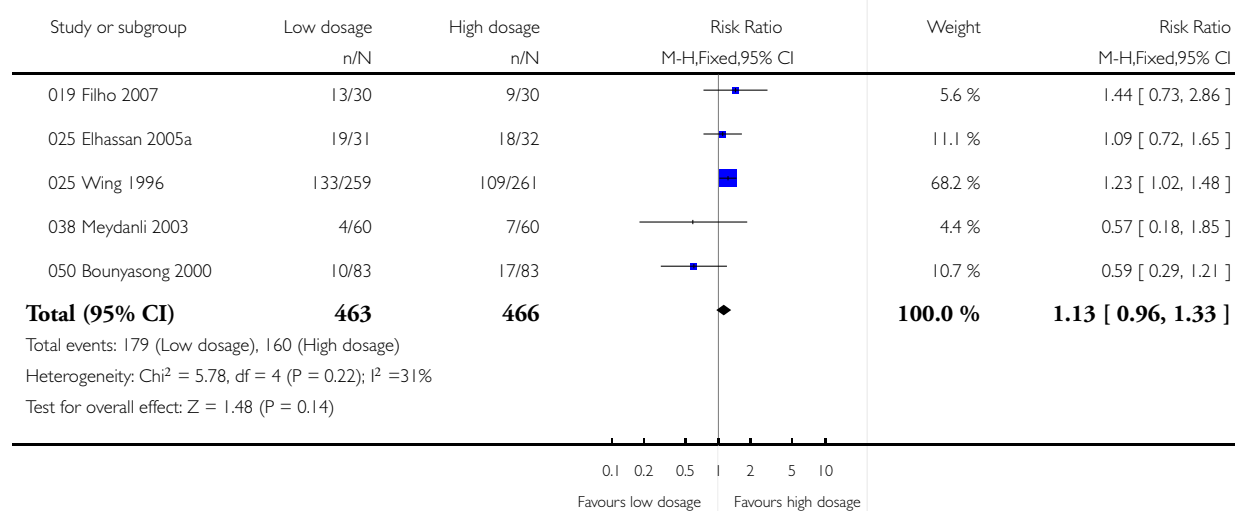


#### Analysis 42.5. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 5 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 5 Oxytocin augmentation

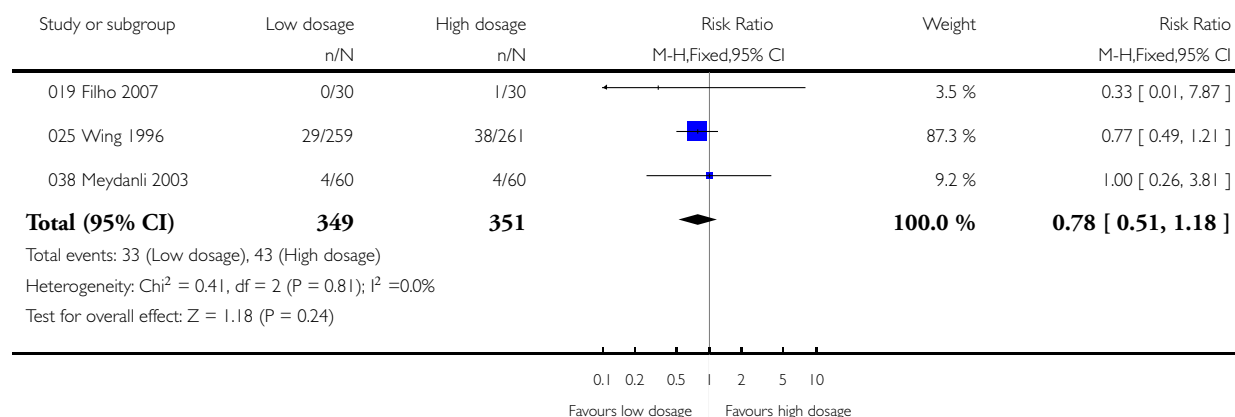


# **Analysis 42.6. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 6 Uterine hyperstimulation without FHR changes.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 6 Uterine hyperstimulation without FHR changes

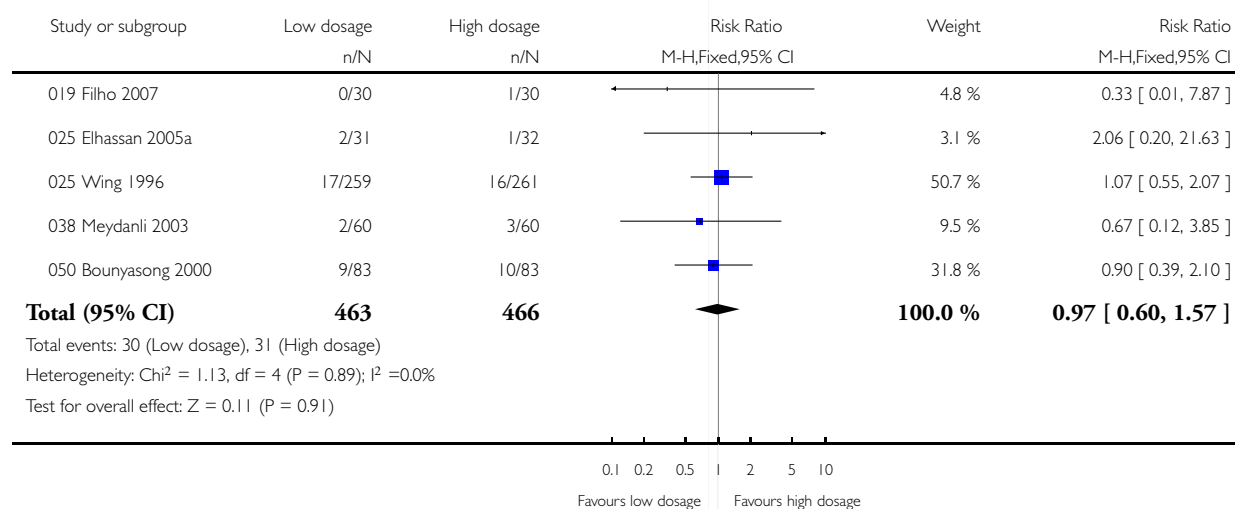


# **Analysis 42.7. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 7 Instrumental vaginal delivery.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 7 Instrumental vaginal delivery



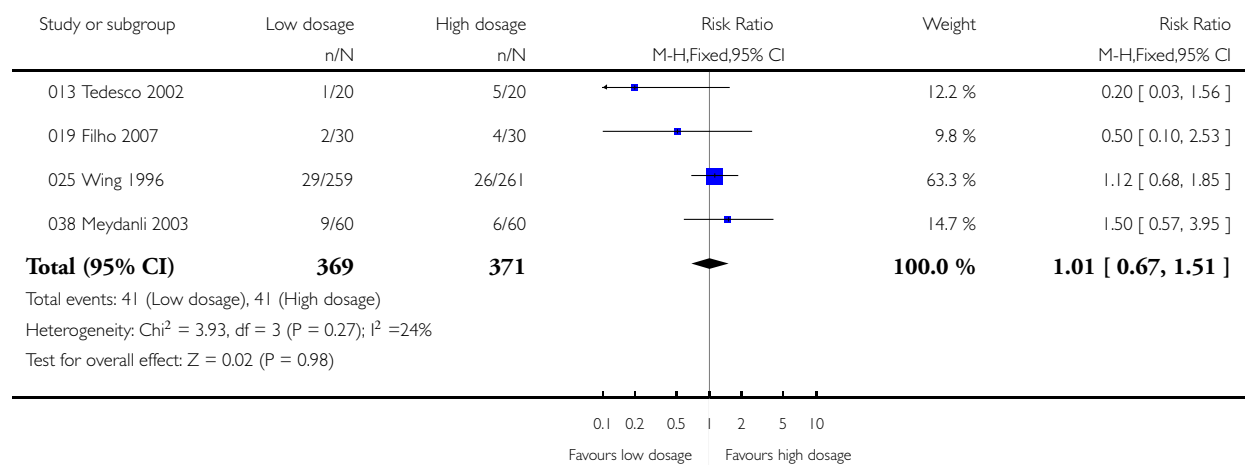


**Analysis 42.8. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 8 Meconium-stained liquor.**

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 8 Meconium-stained liquor

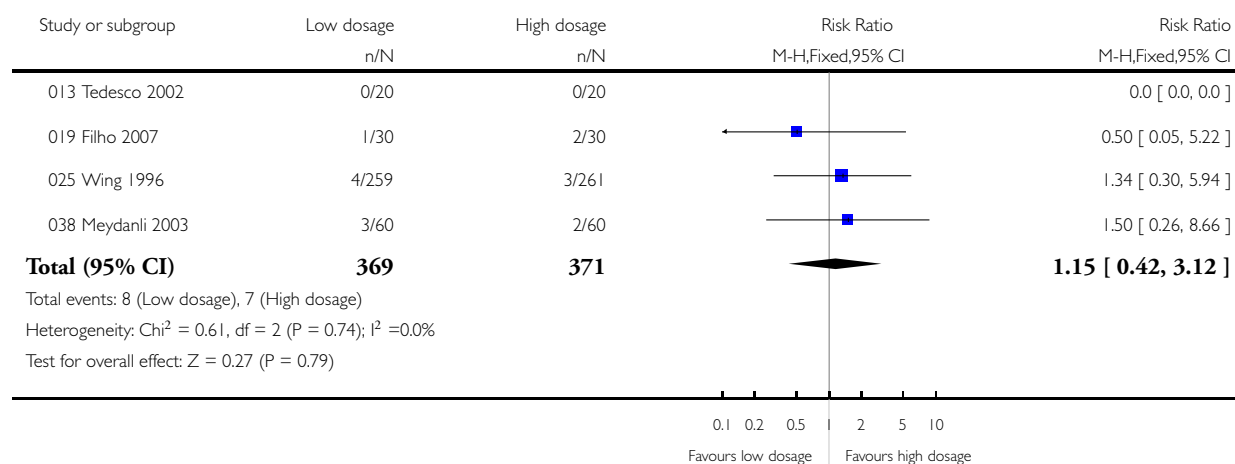


### Analysis 42.9. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 9 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 9 Apgar score < 7 at 5 minutes

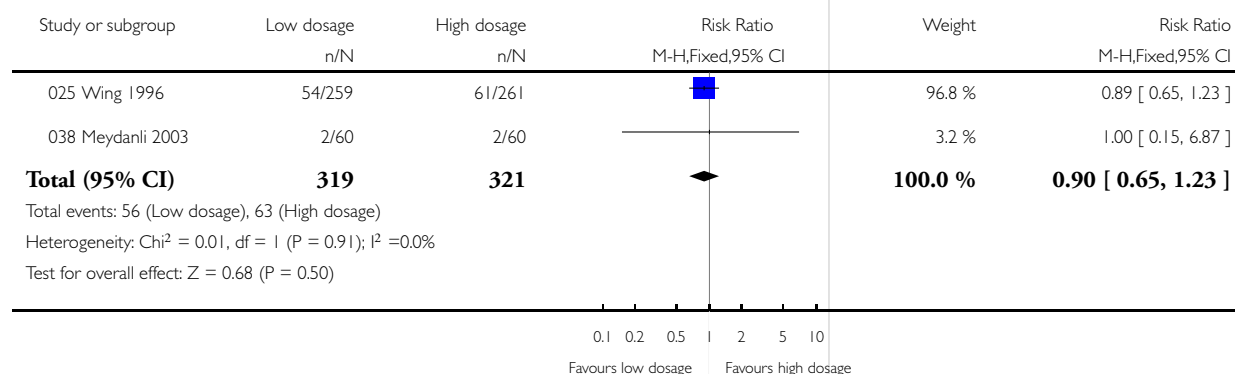


### Analysis 42.10. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 10 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 10 Neonatal intensive care unit admission

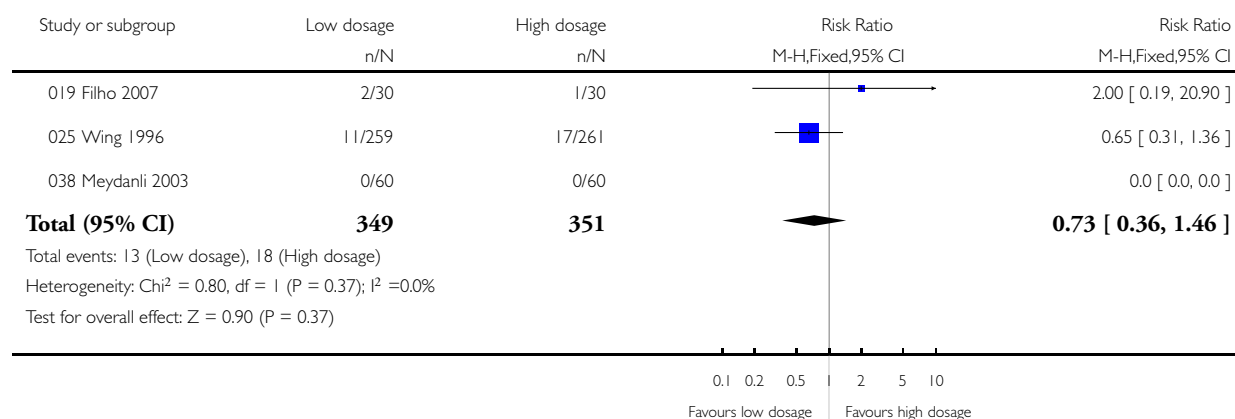


### Analysis 42.11. Comparison 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix, Outcome 11 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 42 Misoprostol lower versus higher dose: all women with intact membranes and unfavourable cervix

Outcome: 11 Maternal side effects (eg nausea, vomiting, diarrhoea)

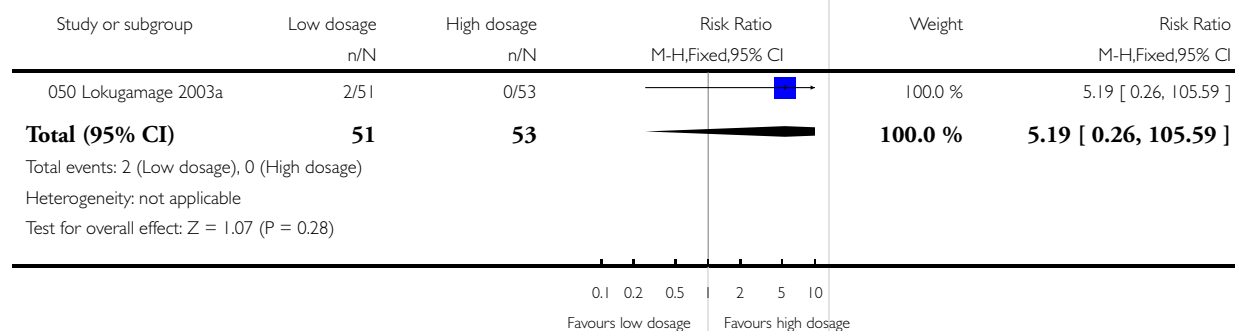


### Analysis 43.1. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 1 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 1 Uterine hyperstimulation with FHR changes

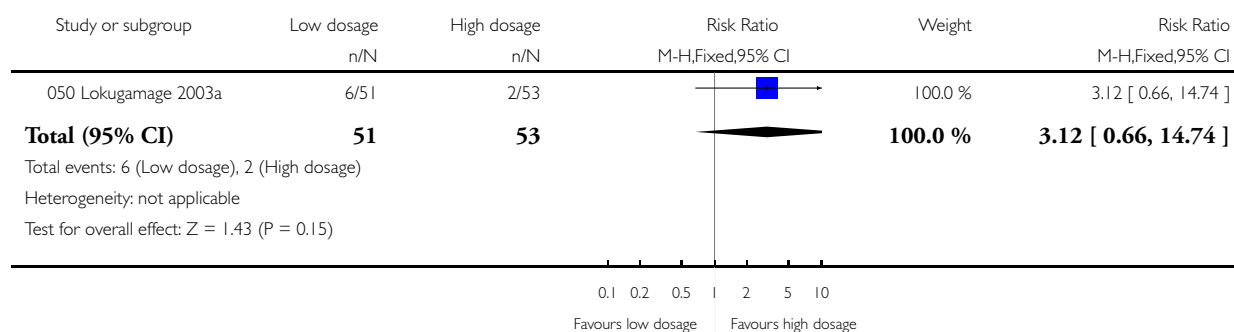


### Analysis 43.2. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 2 Caesarean section

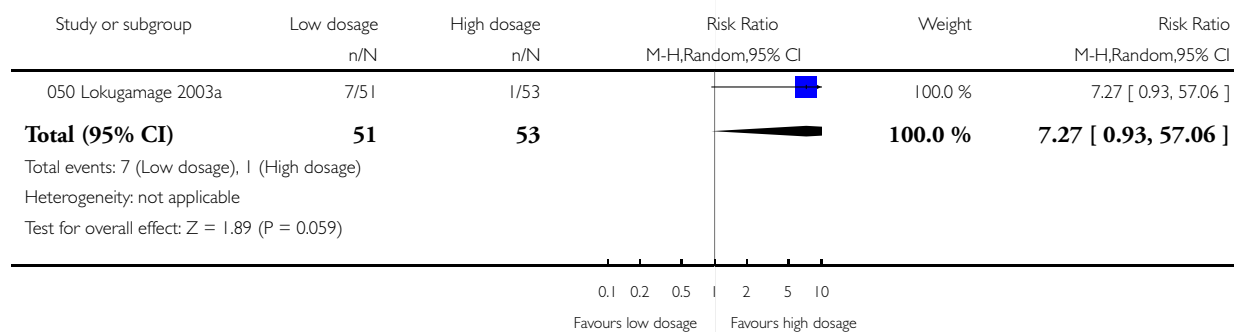


### Analysis 43.3. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 3 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 3 Oxytocin augmentation

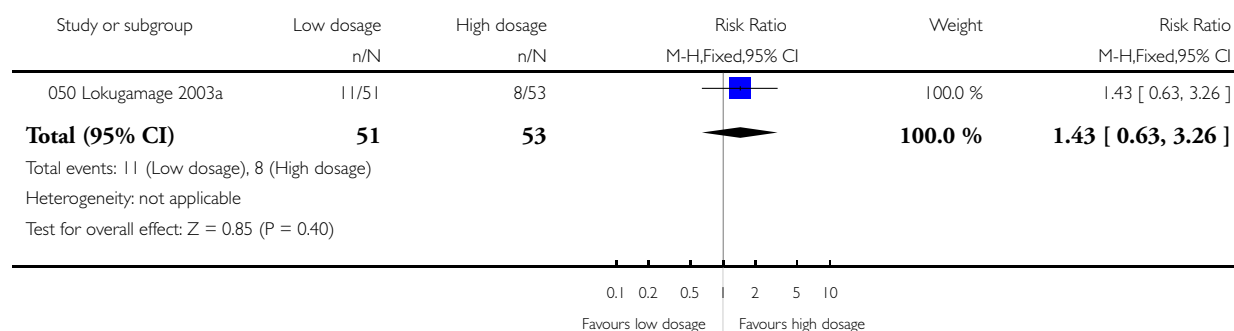


### Analysis 43.4. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 4 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 4 Uterine hyperstimulation without FHR changes

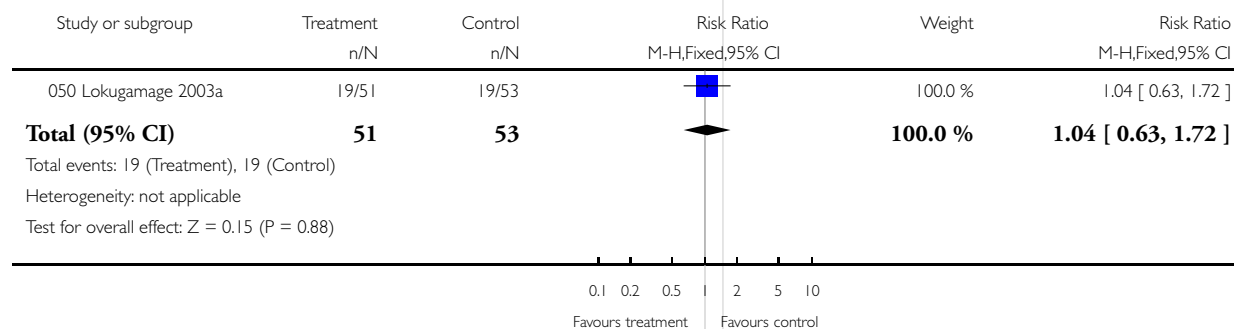


### Analysis 43.5. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 5 Epidural.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 5 Epidural

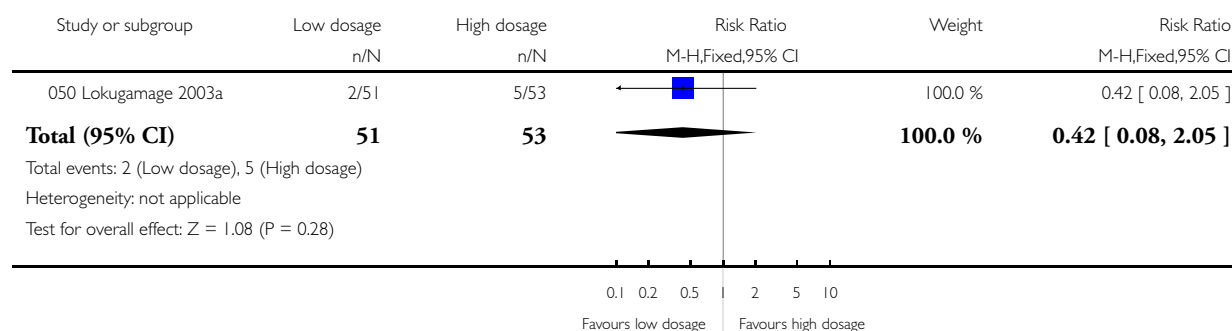


### Analysis 43.6. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 6 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 6 Instrumental vaginal delivery

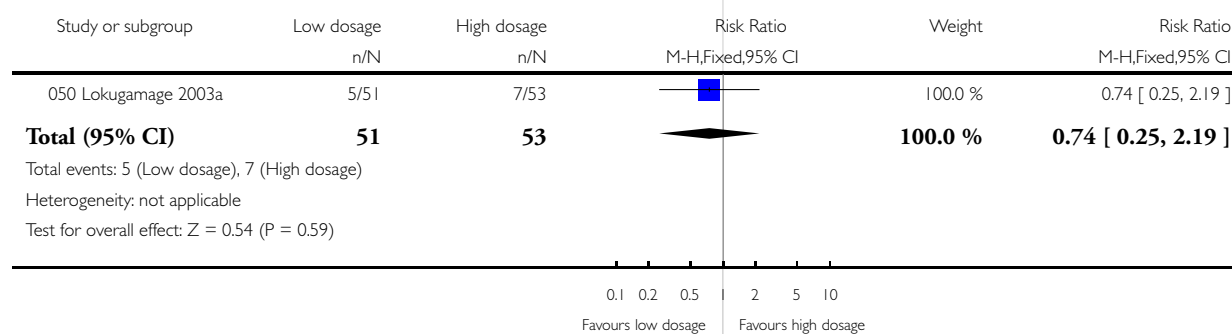


### Analysis 43.7. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 7 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 7 Meconium-stained liquor

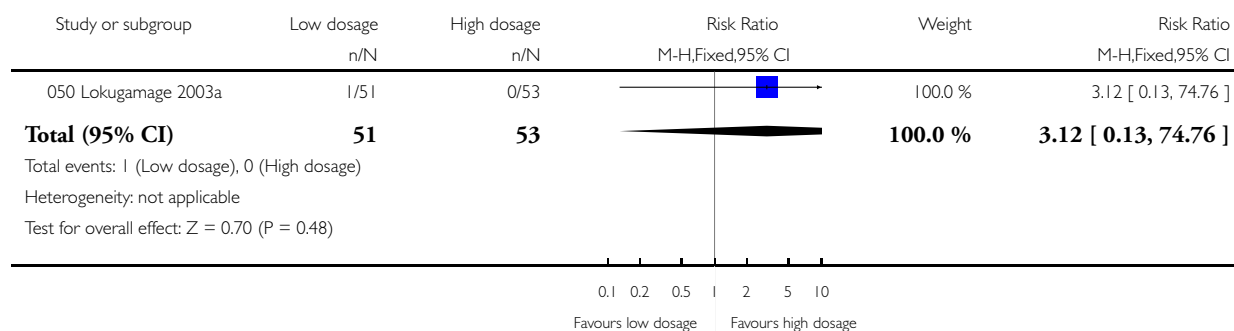


### Analysis 43.8. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 8 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 8 Apgar score < 7 at 5 minutes

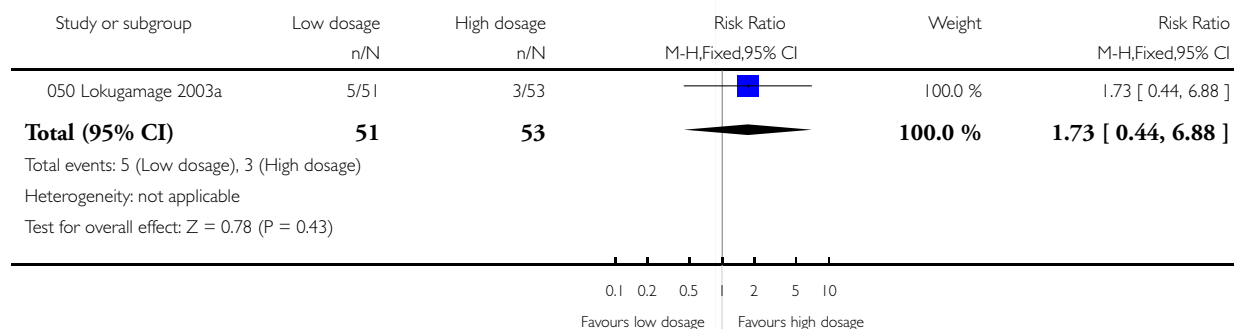


### Analysis 43.9. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 9 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 9 Neonatal intensive care unit admission

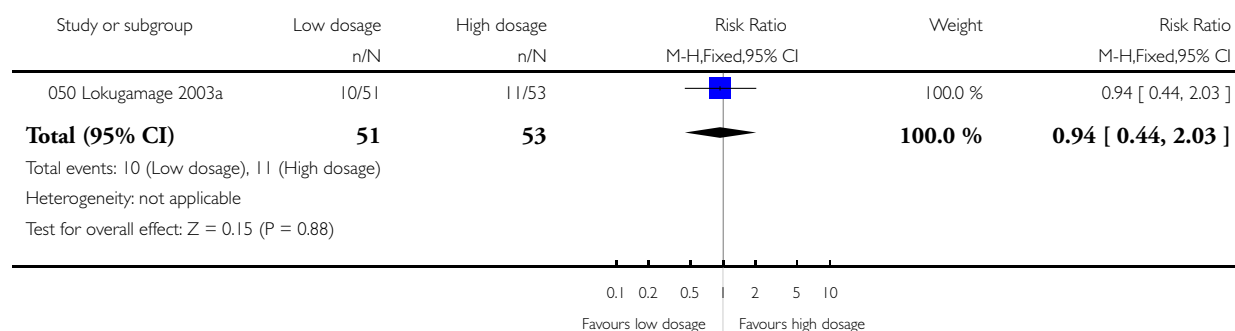


### Analysis 43.10. Comparison 43 Misoprostol lower versus higher dose: all multiparae, Outcome 10 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 43 Misoprostol lower versus higher dose: all multiparae

Outcome: 10 Maternal side effects (eg nausea, vomiting, diarrhoea)

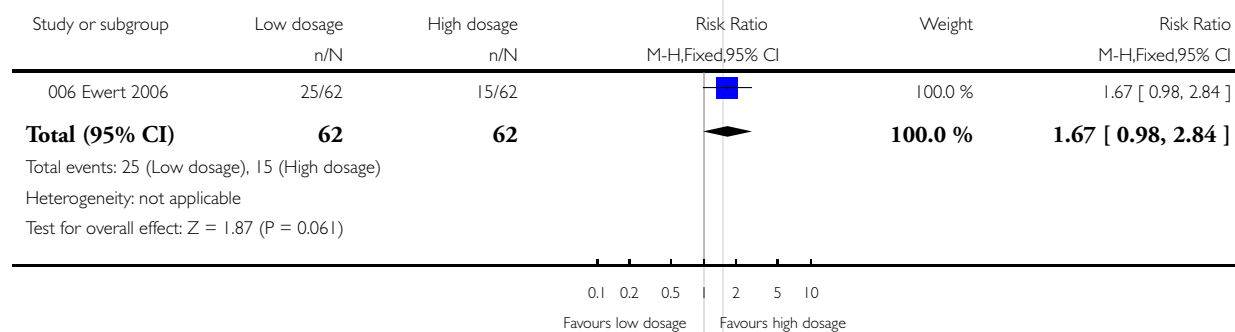


### Analysis 44.1. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 1 Vaginal delivery not achieved within 24 hours.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 1 Vaginal delivery not achieved within 24 hours



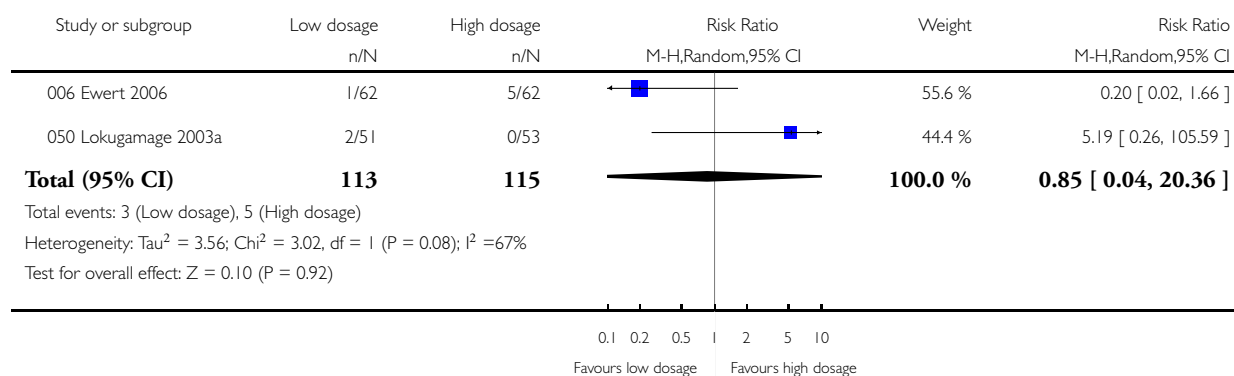


## Analysis 44.2. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 2 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 2 Uterine hyperstimulation with FHR changes

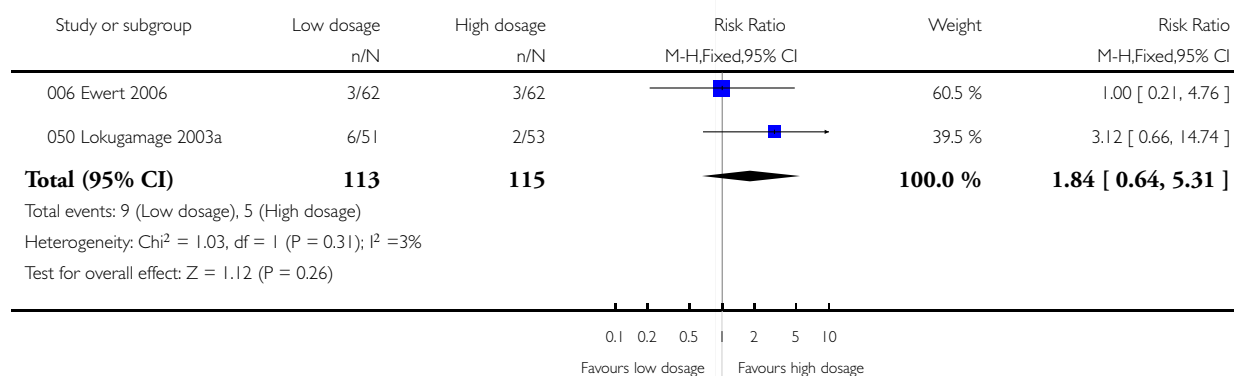


## Analysis 44.3. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 3 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 3 Caesarean section

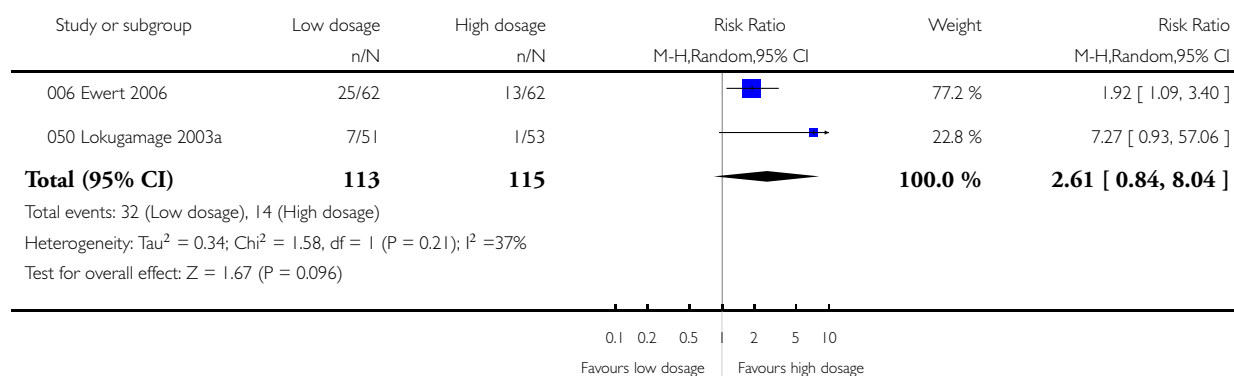


#### Analysis 44.4. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 4 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 4 Oxytocin augmentation

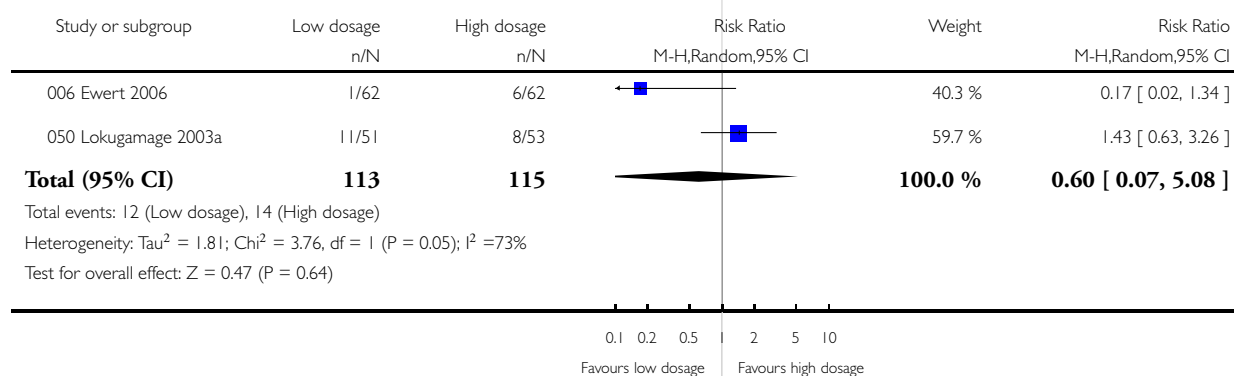


#### Analysis 44.5. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 5 Uterine hyperstimulation without FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 5 Uterine hyperstimulation without FHR changes

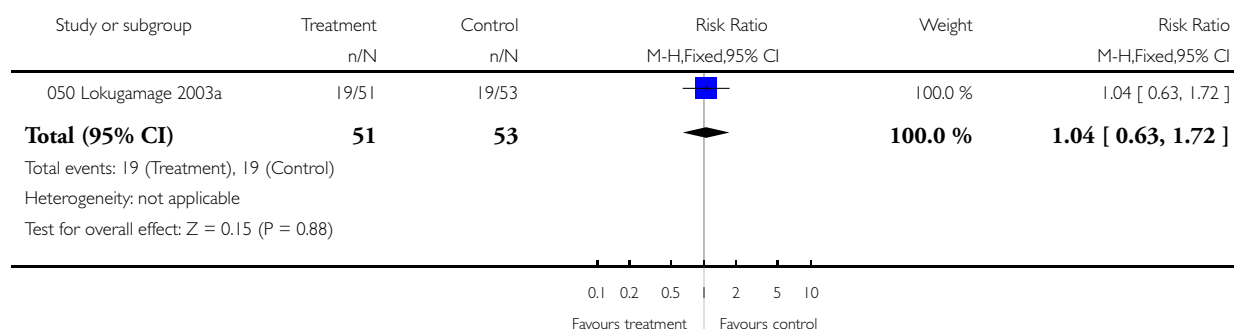


#### Analysis 44.6. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 6 Epidural.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 6 Epidural

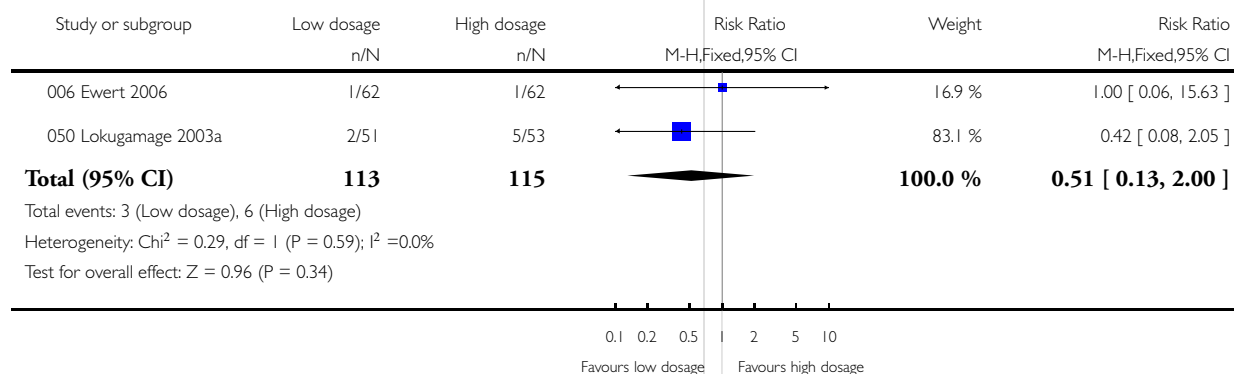


#### Analysis 44.7. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 7 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 7 Instrumental vaginal delivery

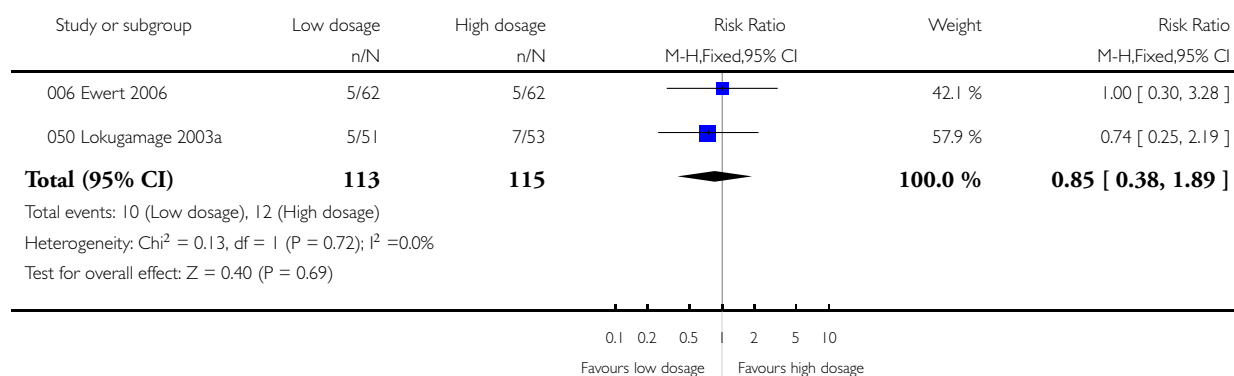


#### Analysis 44.8. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 8 Meconium-stained liquor.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 8 Meconium-stained liquor

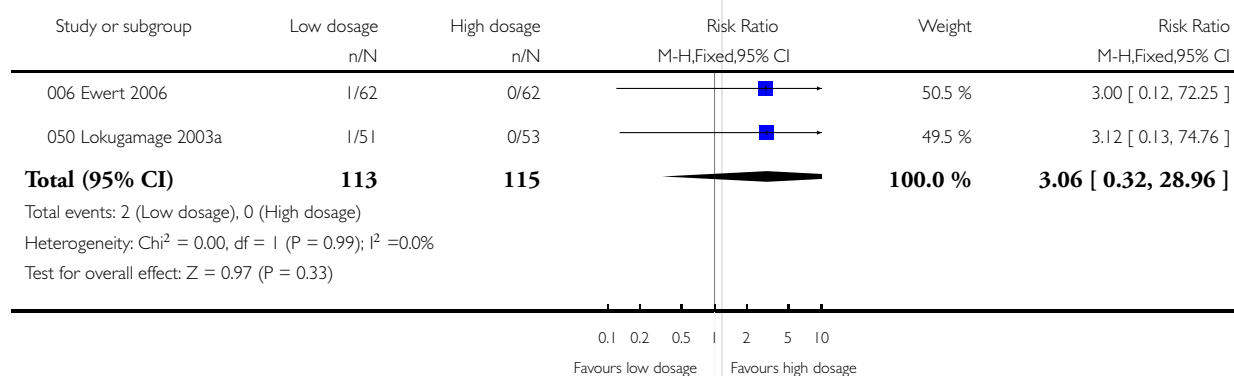


#### Analysis 44.9. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 9 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 9 Apgar score < 7 at 5 minutes

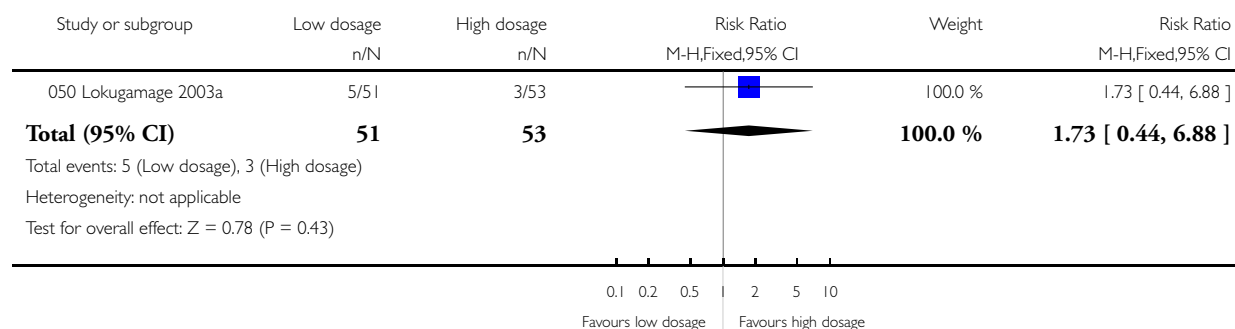


### Analysis 44.10. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 10 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 10 Neonatal intensive care unit admission

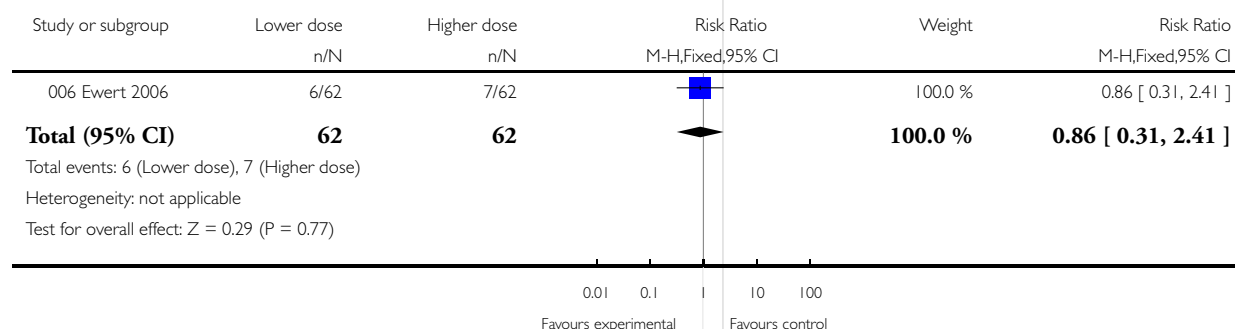


### Analysis 44.11. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 11 Postpartum haemorrhage.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 11 Postpartum haemorrhage

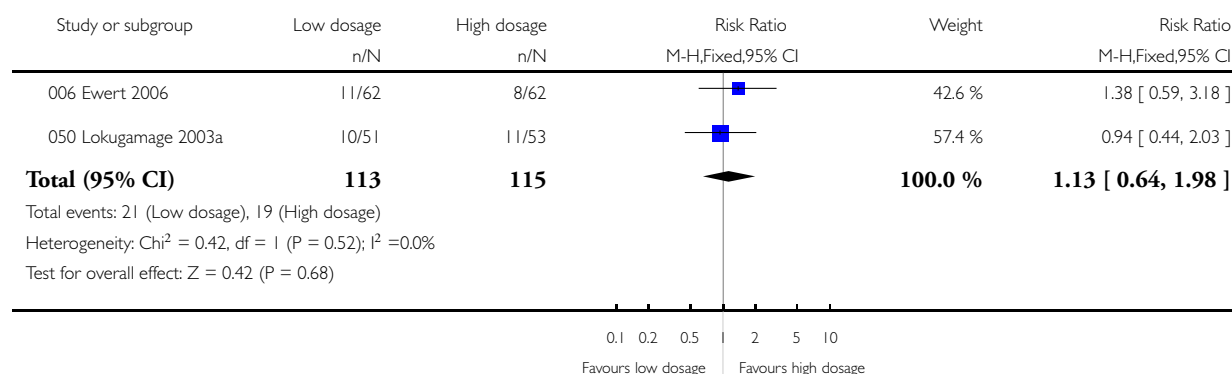


### Analysis 44.12. Comparison 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix, Outcome 12 Maternal side effects (eg nausea, vomiting, diarrhoea).

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 44 Misoprostol lower versus higher dose: all multiparae with unfavourable cervix

Outcome: 12 Maternal side effects (eg nausea, vomiting, diarrhoea)

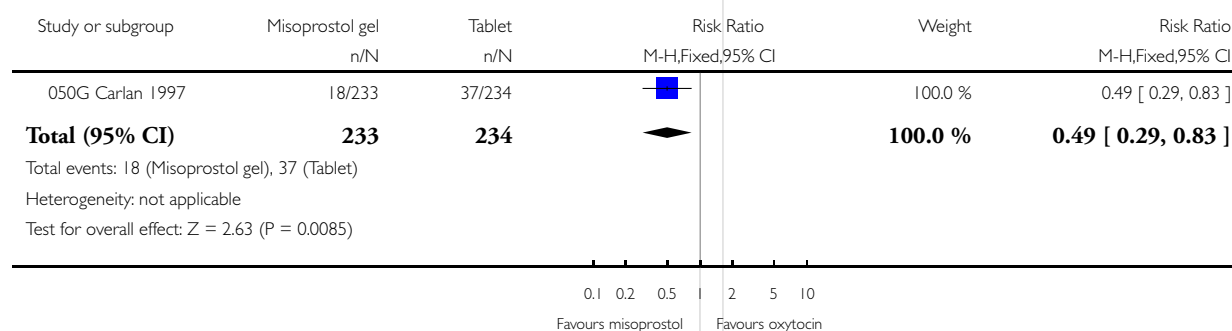


### Analysis 45.1. Comparison 45 Misoprostol gel versus tablet: all women, Outcome 1 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 45 Misoprostol gel versus tablet: all women

Outcome: 1 Uterine hyperstimulation with FHR changes

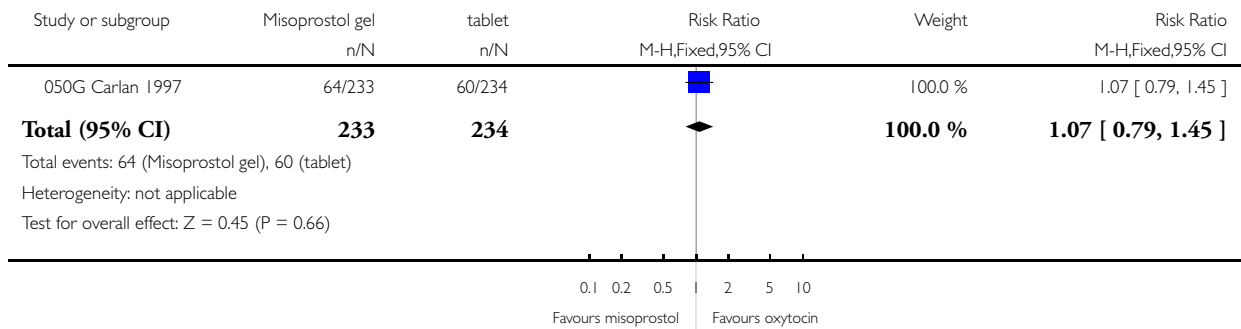


### Analysis 45.2. Comparison 45 Misoprostol gel versus tablet: all women, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 45 Misoprostol gel versus tablet: all women

Outcome: 2 Caesarean section

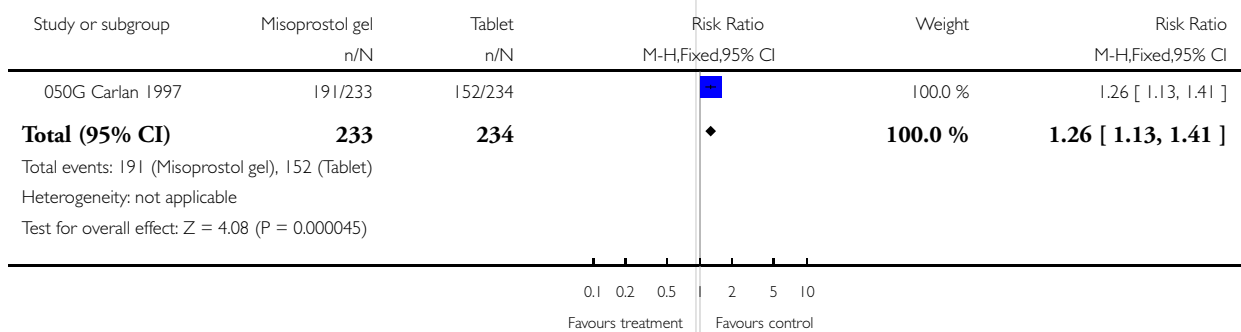


### Analysis 45.3. Comparison 45 Misoprostol gel versus tablet: all women, Outcome 3 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 45 Misoprostol gel versus tablet: all women

Outcome: 3 Oxytocin augmentation

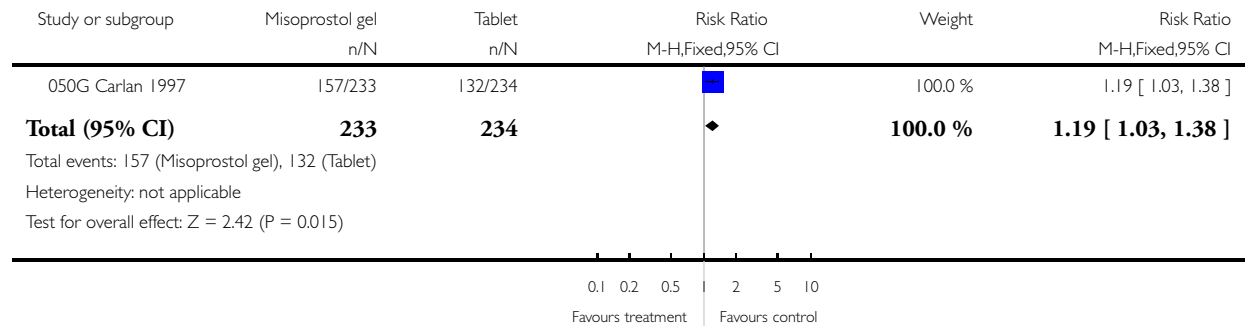


#### Analysis 45.4. Comparison 45 Misoprostol gel versus tablet: all women, Outcome 4 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 45 Misoprostol gel versus tablet: all women

Outcome: 4 Epidural analgesia

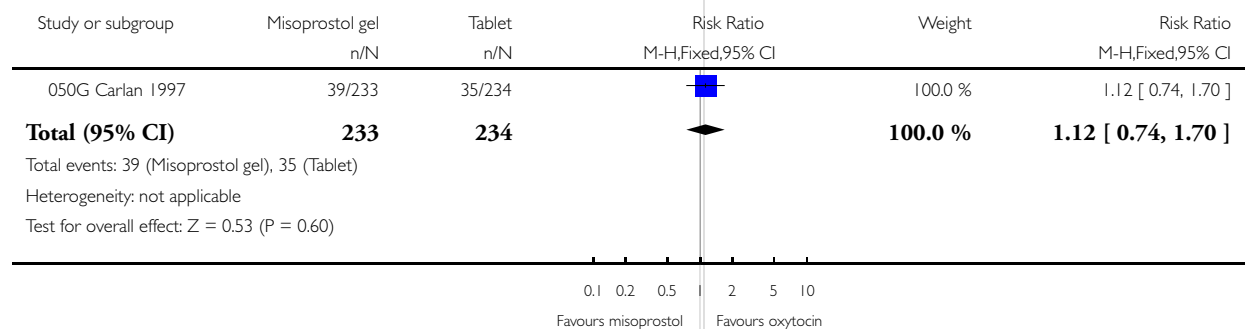


#### Analysis 45.5. Comparison 45 Misoprostol gel versus tablet: all women, Outcome 5 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 45 Misoprostol gel versus tablet: all women

Outcome: 5 Instrumental vaginal delivery



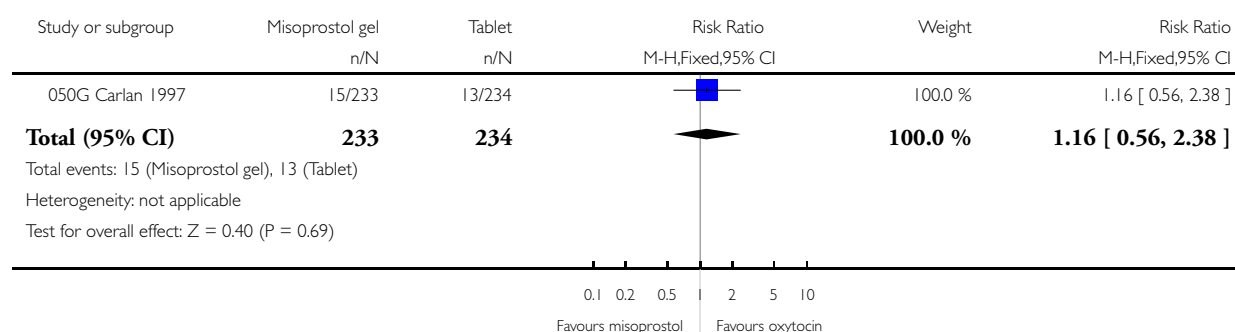


### Analysis 45.6. Comparison 45 Misoprostol gel versus tablet: all women, Outcome 6 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 45 Misoprostol gel versus tablet: all women

Outcome: 6 Apgar score < 7 at 5 minutes

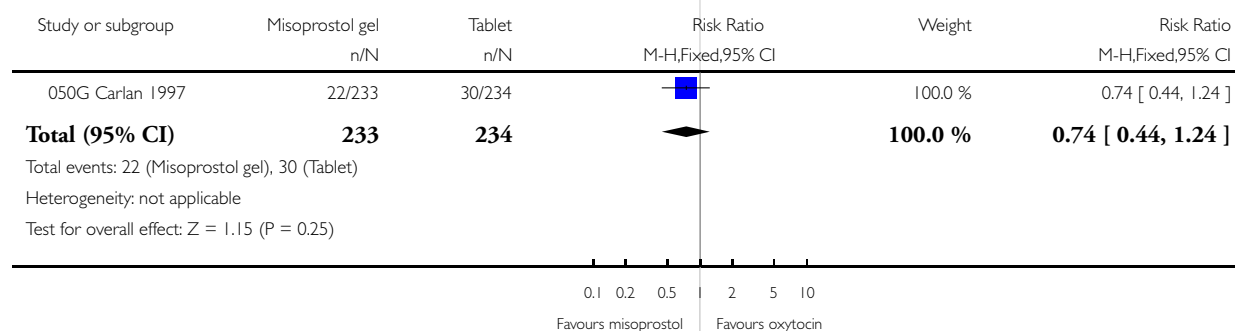


### Analysis 45.7. Comparison 45 Misoprostol gel versus tablet: all women, Outcome 7 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 45 Misoprostol gel versus tablet: all women

Outcome: 7 Neonatal intensive care unit admission

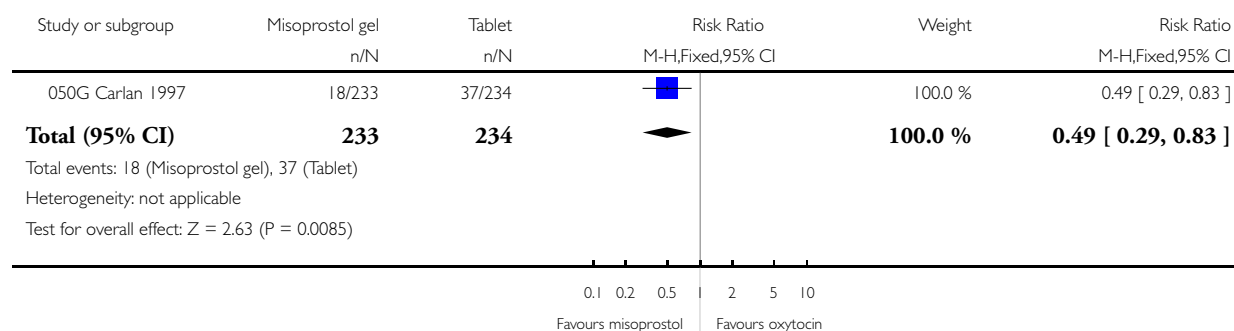


### Analysis 46.1. Comparison 46 Misoprostol gel versus tablet: all women with unfavourable cervix, Outcome 1 Uterine hyperstimulation with FHR changes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 46 Misoprostol gel versus tablet: all women with unfavourable cervix

Outcome: 1 Uterine hyperstimulation with FHR changes

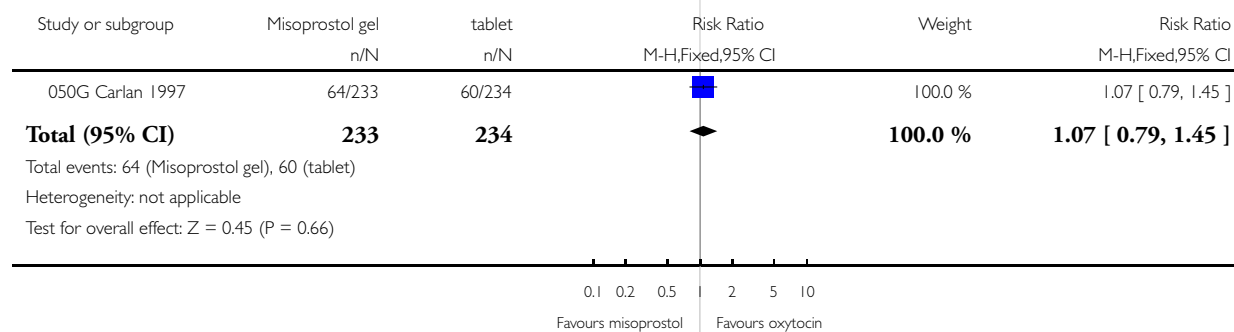


### Analysis 46.2. Comparison 46 Misoprostol gel versus tablet: all women with unfavourable cervix, Outcome 2 Caesarean section.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 46 Misoprostol gel versus tablet: all women with unfavourable cervix

Outcome: 2 Caesarean section

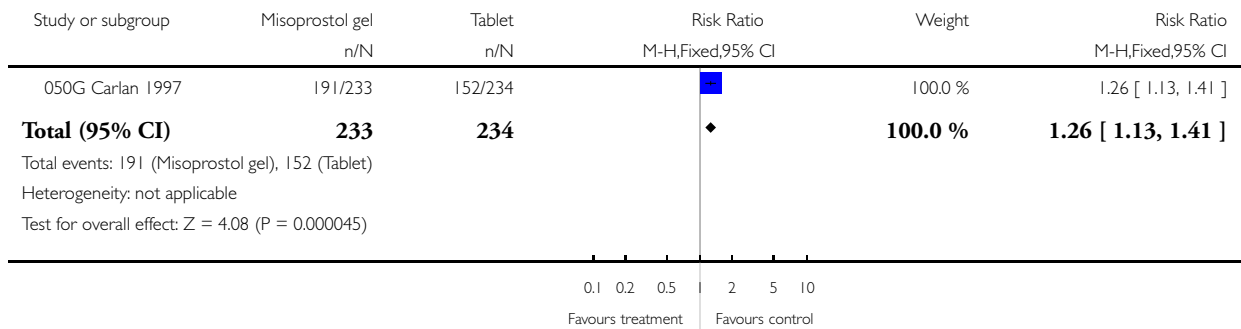


### Analysis 46.3. Comparison 46 Misoprostol gel versus tablet: all women with unfavourable cervix, Outcome 3 Oxytocin augmentation.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 46 Misoprostol gel versus tablet: all women with unfavourable cervix

Outcome: 3 Oxytocin augmentation

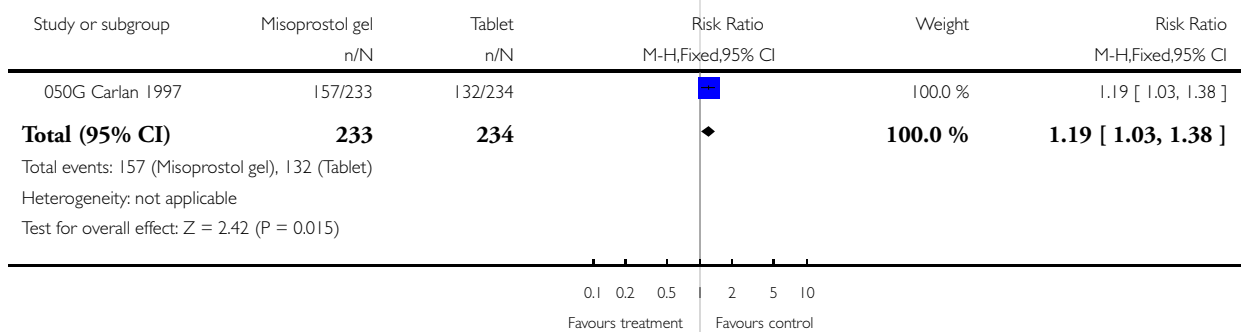


### Analysis 46.4. Comparison 46 Misoprostol gel versus tablet: all women with unfavourable cervix, Outcome 4 Epidural analgesia.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 46 Misoprostol gel versus tablet: all women with unfavourable cervix

Outcome: 4 Epidural analgesia

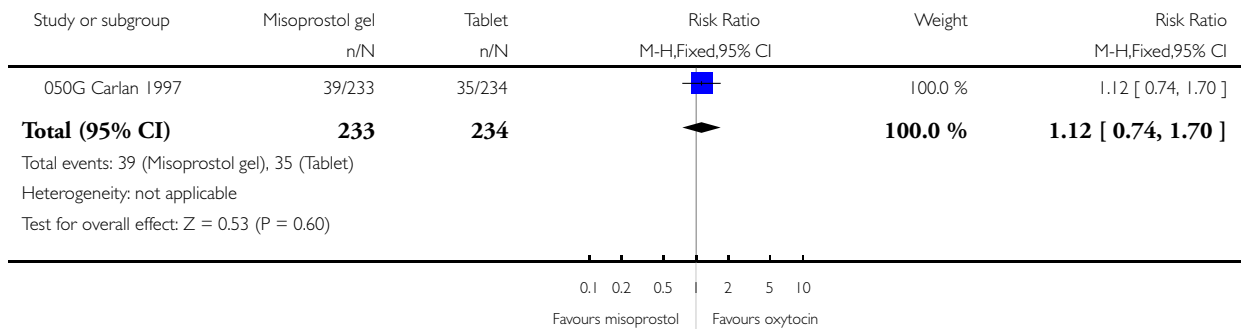


### Analysis 46.5. Comparison 46 Misoprostol gel versus tablet: all women with unfavourable cervix, Outcome 5 Instrumental vaginal delivery.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 46 Misoprostol gel versus tablet: all women with unfavourable cervix

Outcome: 5 Instrumental vaginal delivery

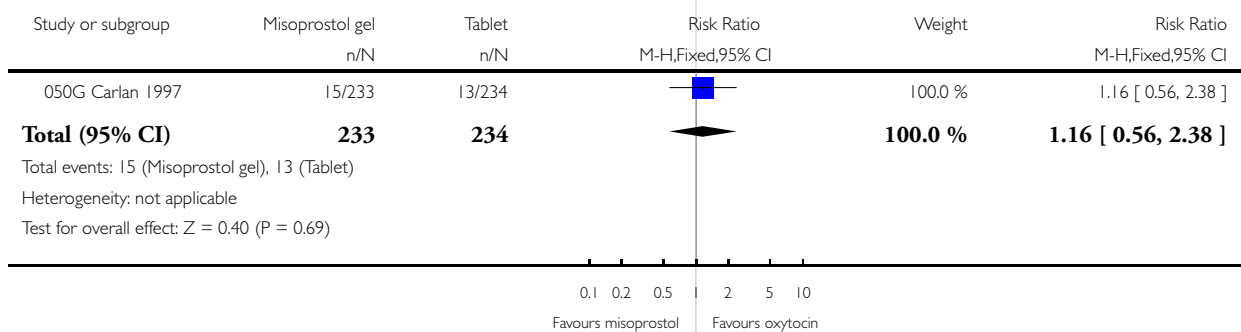


### Analysis 46.6. Comparison 46 Misoprostol gel versus tablet: all women with unfavourable cervix, Outcome 6 Apgar score < 7 at 5 minutes.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 46 Misoprostol gel versus tablet: all women with unfavourable cervix

Outcome: 6 Apgar score < 7 at 5 minutes

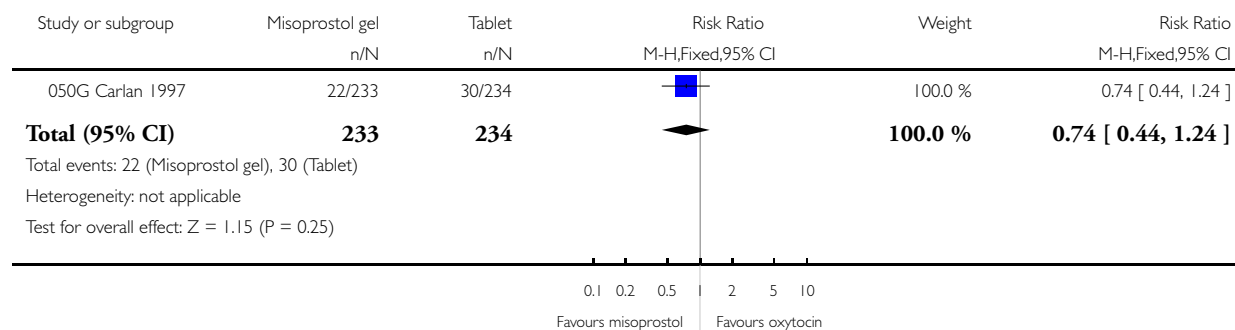


## Analysis 46.7. Comparison 46 Misoprostol gel versus tablet: all women with unfavourable cervix, Outcome 7 Neonatal intensive care unit admission.

Review: Vaginal misoprostol for cervical ripening and induction of labour

Comparison: 46 Misoprostol gel versus tablet: all women with unfavourable cervix

Outcome: 7 Neonatal intensive care unit admission



## APPENDICES

### Appendix I. Methods used to assess trials included in previous versions of this review

A strategy has been developed to deal with the large volume and complexity of trial data relating to labour induction. Many methods have been studied, in many different categories of women undergoing labour induction. Most trials are intervention-driven, comparing two or more methods in various categories of women. Clinicians and parents need the data arranged by category of woman, to be able to choose which method is best for a particular clinical scenario. To extract these data from several hundred trial reports in a single step would be very difficult. We have therefore developed a two-stage method of data extraction. The initial data extraction is done in a series of primary reviews arranged by methods of induction of labour, following a standardised methodology. The data will then be extracted from the primary reviews into a series of secondary reviews, arranged by category of woman.

To avoid duplication of data in the primary reviews, the labour induction methods have been listed in a specific order, from one to 25. Each primary review includes comparisons between one of the methods (from two to 25) with only those methods above it on the list. Thus, the review of intravenous oxytocin (4) will include only comparisons with intracervical prostaglandins (3), vaginal prostaglandins (2) or placebo (1). Methods identified in the future will be added to the end of the list. The current list is as follows:

- (1) placebo/no treatment;
- (2) vaginal prostaglandins;
- (3) intracervical prostaglandins;
- (4) intravenous oxytocin;
- (5) amniotomy;
- (6) intravenous oxytocin with amniotomy;
- (7) vaginal misoprostol;
- (8) oral misoprostol;
- (9) mechanical methods including extra-amniotic Foley catheter;
- (10) membrane sweeping;
- (11) extra-amniotic prostaglandins;
- (12) intravenous prostaglandins;
- (13) oral prostaglandins;

- (14) mifepristone;
- (15) estrogens;
- (16) corticosteroids;
- (17) relaxin;
- (18) hyaluronidase;
- (19) castor oil, bath, and/or enema;
- (20) acupuncture;
- (21) breast stimulation;
- (22) sexual intercourse;
- (23) homoeopathic methods;
- (24) nitric oxide;
- (25) buccal or sublingual misoprostol.

The primary reviews are analysed by the following subgroups:

- (1) previous caesarean section or not;
- (2) nulliparity or multiparity;
- (3) membranes intact or ruptured;
- (4) cervix favourable, unfavourable or undefined.

The secondary reviews will include all methods of labour induction for each of the categories of women for which subgroup analysis has been done in the primary reviews, and will include only five primary outcome measures. There will thus be six secondary reviews of methods of labour induction in the following groups of women:

- (1) nulliparous, intact membranes (unfavourable cervix, favourable cervix, cervix not defined);
- (2) nulliparous, ruptured membranes (unfavourable cervix, favourable cervix, cervix not defined);
- (3) multiparous, intact membranes (unfavourable cervix, favourable cervix, cervix not defined);
- (4) multiparous, ruptured membranes (unfavourable cervix, favourable cervix, cervix not defined);
- (5) previous caesarean section, intact membranes (unfavourable cervix, favourable cervix, cervix not defined);
- (6) previous caesarean section, ruptured membranes (unfavourable cervix, favourable cervix, cervix not defined).

Each time a primary review is updated with new data, those secondary reviews which include data which have changed, will also be updated.

The trials included in the primary reviews were extracted from an initial set of trials covering all interventions used in induction of labour (see above for details of search strategy). The data extraction process was conducted centrally. This was co-ordinated from the Clinical Effectiveness Support Unit (CESU) at the Royal College of Obstetricians and Gynaecologists, UK, in co-operation with The Pregnancy and Childbirth Group of The Cochrane Collaboration in 2000. This process has allowed the data extraction process to be standardised across all the reviews.

The trials are initially reviewed on eligibility criteria, using a standardised form and the basic selection criteria specified above. Following this, data are extracted to a standardised data extraction form which was piloted for consistency and completeness. The pilot process involved the researchers at the CESU and previous reviewers in the area of induction of labour.

Information is extracted regarding the methodological quality of trials on a number of levels. This process is completed without consideration of trial results. Assessment of selection bias examines the process involved in the generation of the random sequence and the method of allocation concealment separately. These are then judged as adequate or inadequate using the criteria described in Table 01 for the purpose of the reviews.

Performance bias is examined with regards to whom was blinded in the trials i.e. patient, caregiver, outcome assessor or analyst. In many trials the caregiver, assessor and analyst were the same party. Details of the feasibility and appropriateness of blinding at all levels is sought.

Individual outcome data are included in the analysis if they meet the pre stated criteria in 'Types of outcome measures'. Included trial data are processed as described in the Cochrane Collaboration Handbook (Clarke 2000). Data extracted from the trials are analysed on an intention to treat basis (when this was not done in the original report, re-analysis is performed if possible). Where data are missing, clarification is sought from the original authors. If the attrition was such that it might significantly affect the results, these data are excluded from the analysis. This decision rests with the reviewers of primary reviews and is clearly documented. Once missing data become available, they will be included in the analyses.

Data are extracted from all eligible trials to examine how issues of quality influence effect size in a sensitivity analysis. In trials where reporting is poor, methodological issues are reported as unclear or clarification sought.

Due to the large number of trials, double data extraction was not feasible and agreement between the three data extractors was therefore assessed on a random sample of trials.

Once the data had been extracted, they were distributed to individual reviewers for entry onto the Review Manager computer software (RevMan 1999), checked for accuracy, and analysed as above using the RevMan software. For dichotomous data, relative risks and 95% confidence intervals are calculated, and in the absence of heterogeneity, results are pooled using a fixed effects model. The predefined criteria for sensitivity analysis include all aspects of quality assessment as mentioned above, including aspects of selection, performance and attrition bias. Primary analysis is limited to the prespecified outcomes and sub-group analyses. In the event of differences in unspecified outcomes or sub-groups being found, these are analysed post hoc, but clearly identified as such to avoid drawing unjustified conclusions.

## FEEDBACK

### Mwanza, July 2002

#### Summary

From my experience of induction of labour I agree that the risk of failure is far less with misoprostol than with prostaglandin E2. If women are carefully selected and started with the lower dose (we used 50 micrograms in one hospital in Zambia) the complications of hyperstimulation and occasional excessive vomiting would be significantly reduced. The low cost of misoprostol and its effectiveness support its use in low-middle income countries.  
[Summary of comment from Moses Mabimba Mwanza, July 2002]

#### Reply

We agree with Dr Mwanza that if misoprostol is used for labour induction, the dosage should be kept to a minimum. Our findings suggest that the vaginal dosage should not exceed 25 mcg 4-hourly.  
[Reply from Justus Hofmeyr, August 2002]

#### Contributors

Moses Mabimba Mwanza

## WHAT'S NEW

Last assessed as up-to-date: 28 May 2009.

Date	Event	Description
10 September 2009	New citation required but conclusions have not changed	New author updated review with an additional 51 studies, which have provided more precise and robust conclusions.
10 September 2009	New search has been performed	We included 51 additional studies from an updated search in November 2008. We updated the search in April 2010 and added the results to Studies awaiting classification for consideration in the next update.

## HISTORY

Protocol first published: Issue 1, 1998

Review first published: Issue 1, 1998

Date	Event	Description
1 October 2002	New citation required and conclusions have changed	Substantive amendment

## CONTRIBUTIONS OF AUTHORS

GJ Hofmeyr prepared the original version, and is responsible for maintaining the review. AM Gulmezoglu quality-checked and revised the review.

C Pileggi critically appraised the studies and performed data extraction for the 2010 update of this review and contributed to the manuscript.

## DECLARATIONS OF INTEREST

None known.

## SOURCES OF SUPPORT

### Internal sources

- University of the Witwatersrand, South Africa.
- University of Fort Hare, Eastern Cape, South Africa.

### External sources

- South African Medical Research Council, South Africa.
- UNDP/UNFPA/WHO/World Bank (HRP), Switzerland.

## INDEX TERMS

### Medical Subject Headings (MeSH)

\*Cervical Ripening; \*Labor, Induced; Administration, Intravaginal; Misoprostol [\*administration & dosage]; Oxytocics [\*administration & dosage]; Pregnancy Trimester, Third; Randomized Controlled Trials as Topic



### **MeSH check words**

Female; Humans; Pregnancy