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Influence of Maternal Obesity on Labor Induction: A Systematic Review and Meta-Analysis

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Abstract

Introduction: Studies have shown that women with obesity have longer labors. The purpose of this systematic review and meta-analysis is to examine existing evidence regarding labor induction in women with obesity, including processes and outcomes. The primary outcome was cesarean birth following labor induction. Secondary outcomes were the timing and dosage of prostaglandins, the success of mechanical cervical ripening methods, and synthetic oxytocin dose and timing.

Methods: Searches were performed in PubMed MEDLINE, Embase, CINAHL, EBSCO, the Cochrane Database of Systematic Reviews, the Database of Abstracts of Effects, Google Scholar, and ClinicalTrials.gov. Searches were limited to studies published in English after 1990. Ten studies published between 2009 and 2017 were included in this review. All were observational studies comparing processes and outcomes of induction of labor in relation to maternal body mass index. The primary outcome was cesarean birth following labor induction. We assessed heterogeneity using Cochran's Q test and tau-squared and I^2 statistics. We also calculated fixed-effect models to estimate pooled relative risks and weighted mean differences.

Results: Ten cohort studies met inclusion criteria; 8 studies had data available for a meta-analysis of the primary outcome. Cesarean birth was more common among women with obesity compared with women of normal weight following labor induction (Mantel-Haenszel fixed-effect

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CONFLICT OF INTEREST

The authors have no conflicts of interest to disclose.

odds ratio, 1.82; 95% CI, 1.55–2.12; $P < .001$). Maternal obesity was associated with a longer time to birth, higher doses of prostaglandins, less frequent success of cervical ripening methods, and higher dose of synthetic oxytocin, as well as a longer time to birth after oxytocin use.

Discussion: Women with obesity are more likely than women with a normal weight to end labor induction with cesarean birth. Additionally, women with obesity require longer labor inductions involving larger, more frequent applications of both cervical ripening methods and synthetic oxytocin.

Keywords

obesity; induction of labor; oxytocin; prostaglandins; cervical ripening; transcervical catheters; intrapartum

INTRODUCTION

The rising prevalence of obesity in the United States over the past 3 decades has resulted in a much higher incidence of women becoming pregnant with a high body mass index (BMI).¹ A recent report of US birth certificate data revealed that more than 50% of women who gave birth in 2014 had a prepregnancy BMI that was classified as either overweight (25.6%) or obese (24.8%).² The incidence of maternal obesity is even higher among some racial and ethnic minority groups, with nearly half of Hispanic women (45.75%) and more than half of non-Hispanic black women (56.9%) being obese during childbearing years.³

Use of labor induction for all women in the United States has also increased significantly, rising from 9.9% in 1990 to 23% in 2008, and remains at this level today.^{4,5} Notably, rates of labor induction are highest among women with obesity.⁶ Use of labor induction increases in a dose-dependent manner with each increase in a woman's BMI category.^{6,7} Higher maternal BMI is positively associated with complications such as gestational diabetes, hypertensive disorders of pregnancy, fetal macrosomia, and stillbirth in the third trimester.⁸ These complications in turn result in increased use of labor induction among women with obesity.⁸ In addition, women who are obese are less likely than women of normal weight to initiate spontaneous labor before 41 weeks' gestation, placing them at higher risk for post-term pregnancy and labor induction.^{6,9}

In all women, labor induction is associated with longer labor course, more dysfunctional labor patterns, increased use of interventions (epidural analgesia, invasive fetal monitoring, and instrumental or operative birth), and extended hospital stays.^{10–12} In addition, there is some evidence that induction of labor is more likely to be unsuccessful in women with an increased BMI.⁷ In a retrospective population cohort study of 80,887 women, women with a BMI of at least 40 kg/m² had a 29% risk of an unsuccessful labor induction compared with a 13% risk among women of normal weight (odds ratio [OR], 2.73; 95% CI, 2.53–2.96).⁷

With several recent meta-analyses and clinical trials reporting a decreased risk of cesarean birth among women who have labor induction compared with expectant management, use of this intervention may increase in the future.^{13,14} However, we lack evidence regarding the most effective methods of labor induction among women who are obese and the risk of

cesarean birth following labor induction in this population.¹⁵ The purpose of this systematic review and meta-analysis was to collect and examine existing evidence to estimate the influence of maternal obesity on labor induction processes and outcomes. The primary outcome was cesarean birth following labor induction. Secondary outcomes were prostaglandin dosage and timing, the success of mechanical cervical ripening methods, and synthetic oxytocin dosage and timing.

METHODS

We searched electronic databases including PubMed, MEDLINE, Embase, CINAHL, EBSCO, the Cochrane Database of Systematic Reviews, the Database of Abstracts of Effects, Google Scholar, and [ClinicalTrials.gov](https://clinicaltrials.gov). Primary Medical Subject Heading (MeSH) terms included *obese*, *obesity*, *labor induced*, *labor induction methods*, *cervical ripening*, *oxytocin*, *transcervical catheter* and *prostaglandins*. Also, *artificial rupture of membranes* and *mechanical cervical ripening* were searched. Searches took place in September and October of 2016, with a search of [ClinicalTrials.gov](https://clinicaltrials.gov) following in February 2017. All searches were updated in September 2017.

Searches included English-language studies published between 1990 and March 2017. We limited the search to investigations published after 1990 because of the rapid increase in obesity among women and substantive changes in labor induction practices over the past several decades. The exposure for this review was labor induction. Observational studies that provided information about labor induction outcomes and/or cervical ripening outcomes stratified by maternal BMI were included. We included induction of labor with prostaglandins, transcervical catheters, oxytocin, and combinations of these methods. We did not include women undergoing induction of labor who had artificial rupture of membranes alone. The use of other methods for labor induction, such as stripping membranes and laminaria, were also not included. The primary outcome was cesarean birth following a trial of labor after induction. We also report secondary outcomes by the induction agent, including time to birth, the dose and timing of prostaglandins, achievement of active-phase labor, success of the cervical ripening attempt, and total dose and timing of oxytocin required for induction.

The initial search of the literature by the primary author was repeated using the same criteria by a university research librarian. Results of these 2 independent searches were combined (Figure 1).¹⁶ EndNote (Clarivate Analytics, Philadelphia, PA) was used to remove duplicates. Titles and abstracts were screened using inclusion criteria; then full-text articles of selected studies were retrieved. Any discrepancies in decisions regarding study inclusion were discussed by the authors until agreement was reached. The Newcastle-Ottawa scale was used to rate the quality of observational studies included in this review.¹⁷

For the primary outcome of cesarean birth, the *meta* package in R version 4.8.1 (R Foundation, Vienna, Austria) was used to perform a Mantel-Haenszel method fixed-effect meta-analysis of the influence of maternal obesity on cesarean birth among women who were induced to calculate ORs. Fixed-effect analyses were selected over random-effects analyses because of the low number of studies available; statistical analyses are considered

descriptive only.¹⁸ Presence of heterogeneity between studies was assessed with Cochran's Q test; heterogeneity magnitude was assessed with tau-squared and I^2 statistics. One study author was contacted for additional data for the analysis.¹⁹

For secondary outcomes, time to birth, the number of prostaglandin doses required, achievement of active-phase labor, success of cervical ripening attempt, and total dose and timing of the induction agent required for induction of labor were calculated. If an OR was not provided in the reviewed studies, we calculated one from raw data using MedCalc for Windows, version 17.2 (MedCalc Software, Ostend, Belgium).

RESULTS

The initial searches returned a total of 505 articles (Figure 1).¹⁶ Once duplicates were removed, 274 unique articles remained. Screening of the titles and abstracts to identify those that met the inclusion criteria of English language articles, articles pertaining to induction of labor, and peer-reviewed published literature from 1990 to September 2017 resulted in the exclusion of 215 articles. Full-text copies of the remaining 59 articles were obtained and analyzed again for eligibility. An additional 49 studies were excluded for the following reasons: not focused on induction of labor ($n = 27$), BMI not stratified ($n = 12$), no details for induction of labor protocol ($n = 7$), included only nulliparous women ($n = 2$), used nonstandard BMI categorizations ($n = 1$). Ten studies remained for the analysis.^{19–28}

The final group for the systematic review was 10 studies that included a total of 7881 women. Eight of the 10 studies reporting outcomes of cesarean births were included in the meta-analysis ($n = 5450$ participants). Five of the included studies were conducted in the United States;^{21,22,24,26,28} the remaining studies were carried out in Israel,^{20,25} Ireland,²⁷ France,²³ and Canada¹⁹ (Table 1). The studies included in this review scored between 7 and 9 stars on the Newcastle-Ottawa scale, indicating high quality.¹⁷ Cochran's Q statistic for the studies included in the meta-analysis indicated minimal heterogeneity between studies ($Q [7] = 7.56$; $P > .05$; tau-squared = 0.00; $I^2 = 7.4\%$ [0%–70.0%]).

Body Mass Index Categories

The timing of BMI measurement differed between studies (Table 1), including prepregnancy,²² the first prenatal visit,^{23,25} prenatal enrollment in the first trimester,²⁷ perinatal enrollment before 25 weeks' gestation,¹⁹ and the time of labor admission.^{19–22,24,26} Beckwith et al²² were the only investigators who analyzed outcomes using both the prepregnancy BMI and the time of hospital admission BMI with the rationale that a difference in effect could be due to a difference in volume distribution at the time of hospital admission. In addition, obesity was defined differently in several of the included studies (Tables 2, 3, and 4). Most of the studies used standard BMI categories defined by the Institute of Medicine²⁹ or the World Health Organization,³⁰ but 3 studies^{20,22,26} included only obese and nonobese categories, using a BMI of 30 kg/m² to separate the categories. In one of the included studies, the normal and overweight BMI categories were described but used differently in the analysis.²⁷

Labor Induction Protocols

There were also variations among included studies in labor induction protocols. For example, successful cervical ripening was variously defined as a Bishop score from 3 to greater than 7 (Table 1).^{20,25} Dosage of prostaglandins, route of administration, and timing of induction agents also varied between studies included in this review (Table 1). Two different doses were reported by the investigators of studies involving dinoprostone (Cervidil: 3 mg or 10 mg vaginally), and both dose and route varied for studies involving misoprostol (Cytotec; 25–100 mcg administered orally or vaginally). Among studies reporting results of labor inductions involving transcervical catheters, both involved Foley catheters,^{20,22} with one comparing Cook catheters with Foley catheters.²⁰ We identified no studies that included a combination of prostaglandins and transcervical catheters.

Cesarean Birth

A fixed-effect meta-analysis of 8 studies with usable data indicated that cesarean birth following labor induction was more common among women with obesity compared with women of normal weight (OR, 1.82; 95% CI, 1.55–2.12; $P < .001$; Figure 2). Moreover, an increasing degree of maternal obesity was associated with a higher risk of cesarean birth following labor induction in a dose-dependent manner in both of the 2 studies that reported more than one category of maternal obesity (Table 2).^{21,24} As in the other studies, Beckwith et al²² found that the risk of a cesarean birth following labor induction was significantly higher among women with obesity compared with women of normal weight (OR, 1.5; 95% CI, 1.03–2.20). However, these investigators did not find a significant association between labor induction and cesarean birth among the subgroup of women who used transcervical catheters (OR, 1.14; 95% CI, 0.59–2.19).²²

Secondary Outcomes

In addition to the primary outcome of cesarean birth following labor induction, associations between maternal obesity and secondary outcomes of timing, dosage, success of cervical ripening methods, and synthetic oxytocin dosage and timing were examined (Tables 3 and 4) in those studies that included the secondary outcomes of interest. For these secondary outcomes, we analyzed the results by method of labor induction (eg, prostaglandins, transcervical catheters, oxytocin).

Prostaglandins

Prostaglandin use was evaluated for failure to achieve active labor and for the type of prostaglandin and dose. Investigators of 7 studies^{21–25,27,31} reported outcomes of labor inductions involving vaginal and oral prostaglandins stratified by maternal BMI. In all of these studies, women with obesity were significantly less likely than women of normal weight to successfully complete cervical ripening and/or achieve active-phase labor (Table 3). Similar to the findings for cesarean birth following labor induction, in studies including subcategories of women with different degrees of obesity, higher BMI was associated in a dose-dependent manner with a higher likelihood that cervical ripening was not successful.²⁵ Likewise, using normal weight women as a reference group, maternal BMI at the time of hospital admission was associated with higher odds that cervical ripening would be

unsuccessful (OR, 3.23; 95% CI, 1.37–4.0) compared with women with prepregnancy obesity (OR, 1.56; 95% CI, 1.18–2.05).²²

Three of the 6 studies reporting use of prostaglandins (Table 3) for labor induction provided information on the mean number of doses required for cervical ripening in women of different BMI ranges.^{23–25} Among women using dinoprostone, there were no differences in the number of doses required for cervical ripening related to maternal BMI.^{23,27} In contrast, in the study by Lassiter et al, there was an increase in the mean number of vaginal misoprostol doses required in women who had higher BMIs compared with women with lower BMIs (1.59 doses for women with a BMI <30 kg/m²; 2.05 and 2.32 doses required for women with BMI 30–40 kg/m² and >40 kg/m², respectively; *P* = .003).²⁴

Transcervical Catheters

Investigators of 2 studies^{20,22} included in this review examined the use of transcervical catheters among women with different BMI categories (Tables 3 and 4). Anabusi et al²⁰ found that the median time to birth following labor induction involving a transcervical catheter was similar for women with obesity and women of normal weight (16 hours for normal-weight and overweight women versus 16 hours and 57 minutes for women with obesity; *P* = .092), and women with obesity were not significantly more likely to have unsuccessful cervical ripening than were women of normal weight following transcervical catheter use (OR, 2.72; 95% CI, 0.48–15.24). Similarly, Beckwith and colleagues²² did not find significant differences in the abilities of women with obesity versus women of normal weight to achieve active-phase labor following transcervical catheter cervical ripening (OR, 1.14; 95% CI, 0.59–2.19).

Oxytocin

Use of synthetic oxytocin (Pitocin) for labor induction was the focus of 5 studies^{19,21,24,26,27} (Tables 3 and 4). Investigators of all 5 studies found that women with obesity who used oxytocin for labor induction either had a longer labor duration or required higher doses of oxytocin compared with women of normal weight. As with other outcomes in this review, the odds of requiring oxytocin for induction of labor were higher in women who were overweight or obese and with each increase in BMI category.

DISCUSSION

This systematic review and meta-analysis examined the influence of maternal obesity on labor induction processes and outcomes. In this meta-analysis, women with obesity were nearly 2 times more likely than women of normal weight to end labor with cesarean birth following labor induction.

We found evidence in this review that labor induction appears to take more time as maternal BMI increases, and it requires both increased number of doses and higher doses of induction agents. More women required synthetic oxytocin for induction of labor with each increase in BMI classification, which likely added time to their labor progress compared with women who went into labor spontaneously following cervical ripening. It is known that labor progress is altered in women with obesity, who are less likely to go into labor spontaneously

at term and who have longer durations of spontaneous labor compared with women of normal weight.¹⁰ These differences in labor initiation and labor progress may be caused by myriad endocrine and inflammatory alterations present in women with obesity.¹⁵ Taken together, the known lower likelihoods of initiating spontaneous labor and achieving normal labor progress seen in women with obesity corroborate our findings that labor induction can be longer and may be more difficult for women with obesity.

Higher doses of prostaglandins and oxytocin were required for women with obesity compared with women of normal weight in all studies included in this review except for one in which dinoprostone was used.²⁷ O'Dwyer et al²⁷ reported a shorter labor duration for all women than the other oxytocin studies included in this review. This discrepancy is likely due to the investigators' labor management protocol, which included straight-line partograms to define active-phase labor, a method that was more conservative than other included studies' definitions of labor. Perhaps the more frequent administration schedule of misoprostol and oxytocin compared with 12-hour dinoprostone allowed us to better observe the influence of maternal BMI on medication requirements.

We also found evidence that certain cervical ripening agents may be better than others for women with obesity.³¹ Misoprostol for cervical ripening may also be more effective in women with obesity because of physiologic changes in prostaglandin expression that may decrease the response to dinoprostone in some women.^{31,32} Alternatively, there may be some pharmacokinetic or pharmacodynamic differences in how these prostaglandins function in women with obesity. More investigations comparing these methodologies in women with obesity are needed to further elucidate the relationship between maternal obesity and labor induction success using different doses and choice of cervical ripening agents.

Transcervical catheters for labor induction were not included in most of the studies of this review, and those that did include this method had small sample sizes. However, transcervical catheters appeared to be more successful than misoprostol for helping women with obesity complete cervical ripening, attain an active phase of labor, and end induction with vaginal birth.²² More research is needed on the use transcervical catheters for induction of labor among women with obesity. Future studies including the use of a combination of transcervical catheters and misoprostol in women with obesity are also needed, as this combination has shown success in mixed-weight groups of women³³ but has not been studied in women with obesity.

Additionally, we found evidence that gestational weight gain sufficient to move a woman from one BMI category to the next is also problematic.²² BMI at the time of labor and birth was the most common timing used to measure BMI in the studies examined in this review. The Beckwith et al study used both a prepregnancy BMI and a BMI at the time of birth. These authors found a greater correlation between BMI and unsuccessful cervical ripening among women with obesity at the time of birth compared with women of normal weight. This may be due to a greater distribution volume at the time of birth compared with prepregnancy BMI. Given these findings, we suggest using the BMI at the time of hospital admission for labor management decisions.

Changes in practice during the period of studies included in this review may have influenced outcomes for women with obesity. In the Safe Prevention of the Primary Cesarean Delivery,³⁴ the American College of Obstetricians and Gynecologists and the Society for Maternal-Fetal Medicine recommended that clinicians use 6 cm of cervical dilatation as the start of active-phase labor, in contrast to previous definitions of active phase starting at 4 cm of cervical dilatation.³⁵ It is possible that the significant differences we observed in this metaanalysis between maternal BMI and cesarean birth following labor induction would not be seen, or would not be as strong, since the Safe Prevention guidelines were published. Women with obesity are known to show slowest labor progress between 4 and 6 cm dilatation compared with women with normal range BMIs.³¹ Therefore, the new recommendation in the Safe Prevention guidelines could be particularly important for future clinical outcomes in women with obesity.

This systematic review is limited by the studies available on this subject in the published literature. We found considerable differences between studies with regard to the way maternal obesity was categorized and the standardization of labor induction processes. Also, some studies included in this review adjusted for a range of maternal factors when calculating odds of cesarean birth and other outcomes following labor induction, whereas others presented unadjusted odds or raw numbers only. Despite these differences in statistical adjustment, the overall ORs for cesarean birth following labor induction by maternal BMI were similar across the studies included in this review. This similarity suggests maternal BMI may have a clinically important influence on labor induction processes and outcomes.

The influences of maternal BMI were calculated using ORs rather than risk ratios, as ORs were reported in the original studies. The OR may overinflate the odds of an occurrence when the occurrence is common in a cohort.³⁶ Future studies examining risks associated with maternal BMI may consider reporting risk ratios instead.

Another limitation is our decision to use a fixed-effect method for the meta-analysis.¹⁸ Fixed-effect meta-analyses presume that studies fundamentally share a common effect and therefore only include within-study variance in the model, with larger studies given greater weight. Random-effects meta-analyses do not presume this and include both within-study and between-study variance, with studies weighted more equally (ie, smaller studies weighted upward and larger studies weighted downward) in the model. At the core, choosing between fixed-effect and random-effects meta-analyses is a determination of whether selected studies vary substantially in their methodologies, populations, or in other ways that affect outcomes. Typically, a random-effects analysis is justified, as most studies vary substantially in meaningful ways. Borenstein et al¹⁸ recommend, however, that when there are few studies available, such as in our analysis, a fixed-effect analysis is appropriate with a caveat: The fixed-effect meta-analysis is for descriptive purposes only; generalizing to studies beyond those described is not possible.

These results gathered from a range of well-conducted studies performed over the past few decades in a variety of countries lend credence to the observation that maternal obesity increases a woman's risk for cesarean birth following labor induction. That said, our

inclusion of only observational studies (2 of which were a secondary analysis of a trial^{20,21} and one of which was a secondary analysis of an observational study¹⁹) is another limitation of this review, as observational studies do not include the same level of control for bias as randomized trials. Our decision to only include observational studies in this review if they scored high in methodologic quality offsets this limitation.

This review is also limited by our inability to present separate estimates of the influence of maternal obesity on labor induction outcomes and processes by parity. Labor progress and outcomes appear to be most altered in women who are obese and nulliparous.^{37,38} However, a sufficient number of investigations in the existing literature that allowed presentation of results for both nulliparous and multiparous women separately could not be found. Finally, this analysis was not able to compare longer-term outcomes for women or their neonates following labor induction.

Despite these limitations, this study has a number of strengths. First, study authors conducted a thorough search of existing literature, using duplicate searches by the primary author and a trained university librarian. Selected studies met a strict set of inclusion and exclusion criteria, including high standards for methodologic quality using established tools. Additionally, by including women of mixed parity, we were able to better isolate the influence of maternal obesity on labor induction processes and outcomes.¹⁰ This review focused on contemporary labor induction practices by limiting the inclusion years of chosen studies. Our use of meta-analysis demonstrating minimal heterogeneity between studies is a final strength of this investigation. More information is needed, but this first meta-analysis on this subject is an important step toward guiding the format of future investigations.

Clinical Implications

Clinicians should consider maternal BMI as they undertake labor induction. As shown in this review, women with obesity are more likely than women of normal weight to have a cesarean birth following induction of labor. Therefore, clinicians should carefully discuss the risks and benefits of labor induction in women with obesity, along with maternal or fetal indications for expedited birth.

Maternal BMI near the time of labor should be calculated and used to determine the best agents for cervical ripening. Hospital admission BMI, not prepregnancy BMI, was used by most investigators considering the influence of maternal BMI on labor processes and outcomes.^{22,39–41} Although prepregnancy BMI may be helpful for guiding antenatal care, maternal BMI at the end of pregnancy may better reflect a woman's metabolic condition near the time of labor. In an investigation reporting cesarean rates by maternal BMI category, Kominiarek and colleagues found that women's risk of cesarean increased by 30% with each increase in hospital admission BMI category.⁴¹ Although having a BMI in obese ranges at the time of labor admission may be normal in many women achieving full-term pregnancy, it is nevertheless important that clinicians use the most accurate measurement of BMI to guide their decisions on optimal labor induction strategies for individual women.

This review suggests that transcervical catheters may be more effective than prostaglandin agents at achieving cervical ripening in women who are obese. Clinicians should anticipate

that women with obesity are more likely than women of normal weight to require repeated doses of misoprostol and/or longer administration, as well as higher doses of synthetic oxytocin, to attain active-phase labor. Clinicians should also expect that induction of labor in a woman with obesity may take longer than in a woman of normal weight and should prepare the clinical team, the labor support person, and the woman for this possibility.

This review supports the need for research to better describe optimal labor induction practices for women with obesity. Prospective investigations using standard BMI categories, standard labor induction regimens, and standard definitions of active labor onset will provide more accurate and precise information for use by clinicians when inducing the labor of women who are obese. Further research should focus on the effectiveness of individual induction agents for women with obesity, as well as that of agents used in combination. Also, researchers in this area should strive to include women with wide variations in BMI as participants, thereby better elucidating changes in labor induction success with different degrees of maternal obesity.

CONCLUSION

Maternal obesity and labor induction are now normal in contemporary clinical practice. Until clinicians have better information on the risks of labor induction and techniques to optimally implement labor induction in women who are obese, this population could see increases in rates of cesarean birth and other poor outcomes because of unsuccessful or prolonged labor induction. Our review supports the need for new labor induction protocols that are individualized by the degree of maternal obesity in both the timing and choice of induction agents. With patience and time, many more women with obesity might achieve normal labor outcomes following a safe and effective labor induction.

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Quick Points

- Women with obesity are more likely to end labor induction with cesarean birth.
- Women with obesity need higher doses and a longer duration of exposure to prostaglandins to complete labor initiation and birth compared with women of normal weight.
- Women with obesity need higher doses of synthetic oxytocin to complete labor initiation and birth compared with women of normal weight.

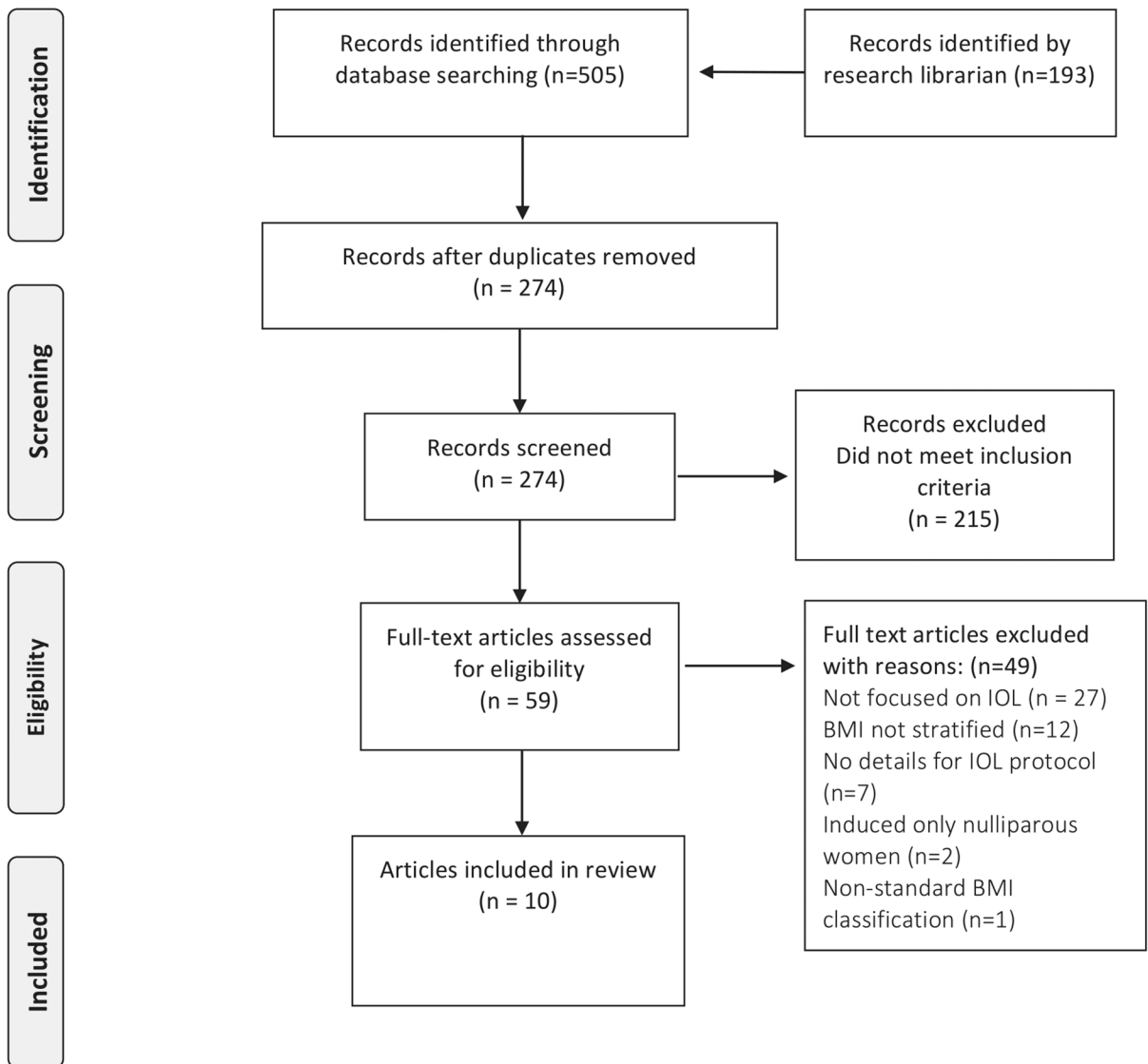


Figure 1.
Flow Chart Describing Literature Extraction Process
Abbreviation: OR, odds ratio.

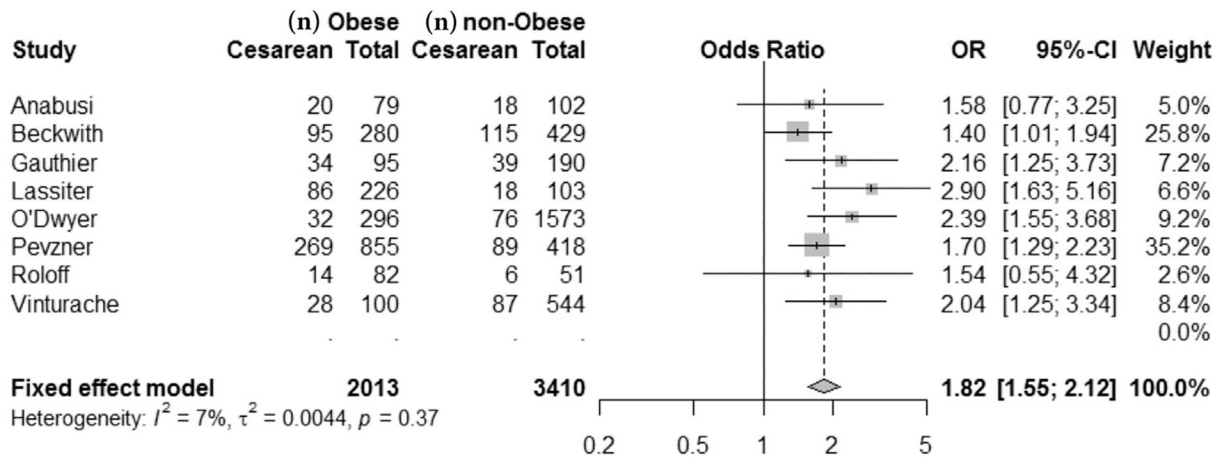


Figure 2.
Forest Plot of Mantel-Haenszel Fixed-Effect Meta-Analysis Displaying Odds of Cesarean Birth

Abbreviations: BMI, body mass index; IOL, induction of labor.

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Table 1.

Description of Included Studies

Study Year Country	Sample n	Methodology	Inclusion Criteria	Time of BMI Determination and Classification	Induction Method Used and Dosage Schedule	Labor Induction Protocol	Definition of Onset of Active Labor	Study Outcomes
Anabusi et al ²⁰ 2016 Israel	181	Cohort study	Singleton, cephalic, intact membranes, term (< 37 wk gestation), with an unfavorable cervix, planning induction	Hospital admission	Cook or Foley catheter Prostaglandins (if first attempt was unsuccessful)	Catheter remained in place for up to 12 h Ripening continued to Bishop score increase of 2 points or >3 cm dilatation	4 cm Successful cervical ripening Cesarean birth rate Maternal and neonatal adverse events	Time from device insertion to birth Successful cervical ripening Cesarean birth rate Maternal and neonatal adverse events
Beckwith et al ²² 2017 United States	709	Retrospective cohort	Singleton, live birth, nonanomalous fetus/newborn, induced labor	Hospital admission	Misoprostol 25 mcg Foley bulb inflated to 30 mL, accompanied by oxytocin	Ripening continued to Bishop score >5	6 cm	Primary: failure to achieve active labor Secondary: cesarean birth rate, doses of misoprostol used, need for protocol deviation
Gauthier et al ²³ 2012 France	285	Retrospective cohort with matching	Singleton, cephalic, live birth, term (>37 wk gestation), one previous cesarean, no contraindications to vaginal birth, Bishop score <6, >18 y of age	First prenatal visit	Dinoprostone 10 mg for 12 h if Bishop score <3 or 1 mg if Bishop score 4-6	Ripening to Bishop score of >6 Continuing for up to 3 days	Bishop score >6	Primary: first cervical ripening attempt unsuccessful
Lassiter et al ²⁴ 2016 United States	329	Retrospective cohort	All women undergoing induction at the research site, gestational age < 37 wk, Bishop score <5	Hospital admission	Misoprostol 25 mcg Oxytocin 1 millilunit per min, increased 1-2 millunits every 30 min	Ripening continued until favorable cervix; then oxytocin was started	Favorable cervix	Primary: time to birth Secondary: number of doses of misoprostol, duration of oxytocin, cesarean birth
Maeder et al ²⁸ 2017 United States	280	Retrospective cohort	Singleton, cephalic, documented weight and height at initiation of prenatal care and labor admission, labor induction, oxytocin	Hospital admission	Oxytocin at 1-2 millunits per min, increased every 15-30 min	Cervical ripening to 3-4 cm Oxytocin per protocol 1-2 millunits, maybe increased every 15-30 min	Not reported	Primary: total oxytocin Secondary: length of labor, method of birth
Melamed et al ²⁵ 2010 Israel	488	Retrospective cohort	Singleton, cephalic, one previous cesarean, no contraindication for vaginal birth	First prenatal visit	Dinoprostone 3 mg Oxytocin 2.5 millunits per min, increased by	Ripening continued until Bishop score >7	Bishop score 7	Primary: failure of cervical ripening with prostaglandin

Study Year Country	Sample n	Methodology	Inclusion Criteria	Time of BMI Determination and Classification	Induction Method Used and Dosage Schedule	Labor Induction Protocol	Definition of Onset of Active Labor	Study Outcomes
O'Dwyer et al ²⁷ 2013 Ireland	1927	Prospective cohort	Singleton pregnancy in the first trimester, Northern European race, > 18 y of age, no gestational diabetes mellitus	On enrollment (in the first trimester)	2.5 milliunits per min every 20 min Prostaglandin and oxytocin	Oxytocin if prostaglandin unsuccessful Ripening continued to favorable cervix Induction of labor with amniotomy then oxytocin	Favorable cervix	Primary: mode of birth, obstetric outcomes
Pevzner et al ²¹ 2009 United States	1273	Cohort study	Singleton pregnancy, < 36 wk gestation, < 18 y of age, low parity (< 3 previous births)	Hospital admission	Misoprostol 100 or 50 mcg Dinoprostone 10 mcg ⁴²	Ripening protocol continued for 24 h or until active labor	4 cm	Primary: active labor Secondary: total oxytocin for induction, birth in <24 h, type of birth
Roloff et al ²⁶ 2015 United States	413	Retrospective cohort	Viable pregnancy with a singleton, cephalic at term (37– 42 wk gestation)	Hospital admission	Cook or Foley catheter Prostaglandins (if first attempt was unsuccessful)	Cervical ripening agent administered every 4–6 h as needed at the discretion of the attending physician	6 cm	Primary: cumulative oxytocin needed for vaginal birth
Vinturache et al ¹⁹ 2014 Canada	1996	Retrospective cohort	Term singleton pregnancies, participation in All Our Babies cohort study	On enrollment (before 25 wk gestation)	Misoprostol 25 mcg Foley bulb inflated to 30 mL, accompanied by oxytocin use	No timing of interventions or Bishop score reported	Not reported	Primary: type of birth Secondary: obstetric outcomes

Abbreviation: BMI, body mass index.

Table 2. Primary Outcome-Cesarean Birth: Odds of Having a Cesarean Birth Among Women with Obesity Undergoing Labor Induction

Study, Year	Indication for Cesarean Birth	BMI Ranges, kg/m ²	Odds Ratio (95% CI)	P Value
Anabusi et al, 2016 ²⁰	Not reported	30	1	
		>30	1.58 (0.77–3.24) ^a	.21
Beckwith et al, 2017 ²²	Failure to reach active labor prior to 5 cm dilatation	<30	1	
		>30	1.4(1.01–1.94)	.04
Gauthier et al, 2012 ²³	Not reported	20–25	1	
		30	2.16 (1.25–3.73)	.005
Lassiter et al, 2016 ²⁴	Not reported	30	1	
		30–39.9	2.78 (1.52–5.10) ^a	<.001
O'Dwyer et al, 2013 ²⁷	Emergency cesarean	40	3.14 (1.58–6.25) ^a	.001
		20–25	1	
		>30	3.03 (1.89–4.86) ^a	<.001
Pevzner et al, 2009 ²¹	Arrest disorders	<30	1	
	Failure to progress			
	Failed labor induction	30–39	1.73 (1.28–2.35) ^b	.002
	Nonreassuring FHR pattern	>40	2.32 (1.58–3.42) ^b	<.001
	Malpresentation			
	Failure to progress	<30	1	
Roloff et al, 2015 ²⁶	Category 2–3 FHR tracing			
	Hemorrhage	>30	1.54 (0.55–4.31) ^a	.407
Vinturache et al, 2014 ¹⁹	Emergency cesarean	18.5–24.9	1	
		>30	2.2 (1.2–4.1) ^c	.011

Abbreviations: BMI, body mass index; FHR, fetal heart rate.

^aOdds ratios and significance values calculated using MedCalc; unadjusted odds ratios reported.

^b Adjusted for parity, race, and treatment group.

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Adjusted for maternal age, parity, preexisting health conditions (diabetes mellitus, hypertension, chronic heart disease, chronic renal disease), pregnancy complications (gestational diabetes, preeclampsia, eclampsia, placental abruption, placenta previa, prolonged rupture of membrane, intrauterine growth restriction), fertility treatments, previous cesarean birth.

Table 3. Influences of Maternal Obesity on Secondary Outcomes by Labor Inducing Agent

Study	Secondary Outcome	BMI Ranges, kg/m ²	Odds Ratio (95% CI)	P Value
Prostaglandins				
Beckwith et al, 2017 ²²	Failure to attain an active phase of labor ^a	Prepregnancy, <30	1	
		Prepregnancy, >30	1.56 ^b (1.18–2.05)	<.05
Gauthier et al, 2012 ²³	First cervical ripening attempt failure	Admission, <30	1	
		Admission, >30	3.23 ^b (1.15–9.09)	<.05
Melamed et al, 2010 ²⁵	Prostaglandin ripening failure	20–25	1	
		30	2.32 (1.37–4.00)	.0019
		21–25	1	
		26–30	5.75 ^c (1.92–23.27)	.006
Pevzner et al, 2009 ²¹	Failed to reach active labor	>30	6.22 ^c (3.73–41.03)	.004
		<30	1	
Anabusi et al, 2016 ²⁰	Cervical ripening failure ^e	30–39	2.31 ^d (1.28–4.16)	.0054
		>40	2.50 ^d (1.24–5.08)	.01
		30	1	
Beckwith et al, 2017 ²²	Failure to attain an active phase of labor ^e	>30	2.72 ^d (0.48–15.24)	.255
		Prepregnancy, <30	1	
		Prepregnancy, >30	1.20 ^b (0.74–1.96)	>.05
		Admission, <30	1	
Pevzner et al, 2009 ²¹	Oxytocin required for IOL	Admission, >30	1.61 ^b (0.70–3.69)	>.05
		20–25	1	
O'Dwyer et al, 2013 ²⁷	Oxytocin required for IOL	Overweight	0.88 ^d (0.69–1.13)	.35
		>30	1.56 ^d (1.18–2.06)	.0016
Pevzner et al, 2009 ²¹	Oxytocin required for IOL	>30	1	
		<30	1	

Study	Secondary Outcome	BMI Ranges, kg/m ²	Odds Ratio (95% CI)	P Value
		30–39	1.51 ^f (1.15–1.97)	<.05
		>40	2.24 ^f (1.57–3.18)	<.05
Roloff et al, 2015 ²⁶	Oxytocin required for IOL	<30	1	
		>30	5.33 ^d (1.03–27.5)	.04
Vinturache et al, 2014 ¹⁹	Oxytocin required for IOL	18.5–24.9	1	
		25–29.9	1.69 ^d (1.33–2.14)	<.0001
		30	2.36 ^d (1.74–3.21)	<.0001

Abbreviations: BMI, body mass index; IOL, induction of labor.

^aFailure to achieve an active phase of labor was defined as a cesarean birth performed at a cervical dilatation of 5 cm or less.

^bAdjusted for primiparity, gestational age at birth, cervix dilatation and effacement at induction initiation, fetal birth weight, and presence of hypertensive disorders of pregnancy.

^cAdjusted for gestational age, maternal age, BMI, parity, and cervical effacement.

^dOdds ratio and significance values were calculated using MedCalc; raw odds data presented.

^eCervical ripening failure calculation is based on the numbers for cervical ripening success, defined as second Bishop score increased by 2 or more points or cervical dilatation of 3 cm or more.

^fAdjusted for parity, race, and treatment group.

Table 4. Influences of Maternal Obesity on Time to Birth and Medication Dosage by Labor Inducing Agent

Study	BMI Ranges kg/m ²	Time to Birth h	P Value	Medication Dosage in Labor	P Value
Prostaglandins					
Gauthier et al, 2012 ²³	20–25	21.20 (17.18) ^{a,b}		1.2 (0.3) ^c	
	30	26.25 (19.22) ^{a,b}	.0258	1.4(0.6) ^c	.0126
Lassiter et al, 2016 ²⁴	<30	17.72 (7.3) ^{a,b}		1.59 ^d	
	30–0	20.01 (8.3) ^{a,b}		2.05 ^d	
	40	22.90 (11.6) ^{a,b}	.0001	2.32 ^d	.0003
O'Dwyer et al, 2013 ²⁷	20–29	6.10 ^{a,e}		2.5 ^f	
	Overweight	6.00 ^{a,e}		2.7 ^f	
	>30	6.60 ^{a,e}	NS	2.8 ^f	NS
Transcervical catheters					
Anabusi et al, 2016 ²⁰	30	16.00 ^g		Not applicable	
	>30	16.95 ^g	.092		
Oxytocin					
Lassiter et al, 2016 ²⁴	<30	17.72 (7.3) ^{a,b}		7.17 ^{a,h}	
	30–40	20.01 (8.3) ^{a,b}		8.54 ^{a,h}	
	>40	22.90 (11.6) ^{a,b}	.0001	10.39 ^{a,h}	.023
Maeder et al, 2017 ²⁸	<25	13.96 (8.10) ^{a,b}		6.37 (7.06) ^{a,h}	
	25–29.9	16.00 (7.54) ^{a,b}		5.98 (5.00) ^{a,h}	
	30	18.30 (8.65) ^{a,b}	.018	7.50 (6.49) ^{a,h}	.252
O'Dwyer et al, 2013 ²⁷	20–25	6.10 ^{a,e}		Not reported	
	Overweight	6.00 ^{a,e}			
	>30	6.60 ^{a,e}	NS		

Study	BMI Ranges kg/m ²	Time to Birth h	P Value	Medication Dosage in Labor	P Value
Pevzner et al, 2009 ²¹	<30	22.70 (11.4–39.6) ^{b,g}		2.6 (0.4–13.5) ^{g,h}	
	30–39	24.90 (12.2–43.0) ^{b,g}		3.5 (0.3–15.8) ^{g,h}	
	>40	27.00 (12.9–50.4) ^{b,g}	<.001	5.0 (0.4–20.9) ^{g,h}	<.001

Abbreviations: BMI, body mass index; NS, not significant.

^aMean (SD, if reported).

^bTime from labor induction to birth.

^cMean (SD) number of prostaglandin doses.

^dMean number of doses of misoprostol.

^eTime in first stage of labor for nulliparous women.

^fMean prostaglandin dose for nulliparous women.

^gMedian (10%–90% range, if reported).

^hTotal units of oxytocin.