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Impact of maternal obesity on preterm delivery in patients with cervical cerclage

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BACKGROUND: Maternal obesity has risen in the United States in recent decades.

OBJECTIVE: This study aimed to evaluate the impact of maternal obesity on the risk for spontaneous preterm delivery and the risk for overall preterm delivery among patients with cervical cerclage placement.

STUDY DESIGN: This was a retrospective study in which data from the California Office of Statewide Health Planning and Development linked birth file from 2007 to 2012 were used, yielding a total of 3654 patients with and 2,804,671 patients without cervical cerclage placement. Exclusion criteria included patients with missing information on body mass index, multiple gestation, anomalous pregnancies, and gestations <20 weeks or >42 weeks. Patients in each group were identified and were further categorized based on body mass index with the nonobese group defined as having a body mass index of <30 kg/m², the obese group defined as having a body mass index of 30 to 40 kg/m², and the morbidly obese group defined as having a body mass index with obesity or those with morbid obesity patients. The analysis was stratified by cerclage placement.

RESULTS: Among patients who underwent cerclage placement, the risk for spontaneous preterm delivery was not significantly different in the obese and morbidly obese group when compared with the nonobese group (24.2% vs 20.6%; adjusted odds ratio, 1.18; 95% confidence interval, 0.97-1.43; and 24.5% vs 20.6%; adjusted odds ratio, 1.12; 0.78-1.62, respectively). However, among patients without cerclage placement, the obese and morbidly obese groups had a higher risk for spontaneous preterm delivery than the nonobese group (5.1% vs 4.4%; adjusted odds ratio, 1.04; 1.02-1.05; and 5.9% vs 4.4%; adjusted odds ratio, 1.03; 1.00-1.07, respectively). The risks for overall preterm delivery at <37 weeks' gestation were higher for the obese and morbidly obese groups than for the nonobese group among patients with cerclage (33.7% vs 28.2%; adjusted odds ratio, 1.23; 1.03-1.46; and 32.1% vs 28.2%; adjusted odds ratio, 1.01; 0.72-1.43, respectively). Similarly, among patients without cerclage placement, the risks for preterm delivery at <37 weeks' gestation were higher for the obese and morbidly obese groups than for the nonobese group among patients with cerclage groups than for the nonobese group (7.9% vs 6.8%; adjusted odds ratio, 1.05; 1.04-1.06; and 9.3% vs 6.8%; adjusted odds ratio, 1.10; 1.08 - 1.13, respectively).

CONCLUSION: Among patients who received a cervical cerclage for the prevention of preterm birth, obesity was not associated with an increased risk for spontaneous preterm delivery. However, it was associated with an overall increased risk for preterm delivery.

Key words: body mass index, cerclage, cerclage failure, morbid obesity, obesity, preterm delivery

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Introduction

The prevalence of obesity among pregnant women was estimated to be between 18.5% and 38.3%.¹⁻⁴ Maternal obesity is a preventable and modifiable risk factor for many adverse pregnancy outcomes, including hypertensive disorders, gestational diabetes, thromboembolic events, stillbirths, cesarean deliveries,⁵⁻¹¹ and preterm delivery.¹¹⁻¹³

Spontaneous preterm birth contributed to approximately 60% of all preterm births,^{14,15} whereas medically indicated preterm birth contributed to the remaining 30% to 40% of all preterm births.^{16,17} The effect of obesity on each type of preterm delivery has been shown to be variable. Previous studies have demonstrated that obesity could be a protective factor against spontaneous preterm delivery.^{18,19} Existing laboratory data suggest that obesity may be associated with higher endogenous ghrelin production, which may inhibit oxytocin receptor activity and lead to suppression of spontaneous labor.²⁰ Conversely, previous data also demonstrated that the presence of inflammatory biomarkers associated with obesity may play a role in the increased risk for spontaneous preterm birth.²¹ Meanwhile, other studies have shown that obesity is associated with an increased risk for medically indicated preterm delivery.^{22,23} One study demonstrated that as maternal body mass index (BMI) increased, the risks for both medically indicated and spontaneous preterm deliveries increased accordingly.²⁴

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Why was this study conducted?

Current literature on the risk for preterm delivery overall and for spontaneous preterm delivery among patients with obesity and morbid obesity with cervical cerclage is limited and contradictory, necessitating further elucidation.

Key findings

Among patients with cervical cerclage, obesity does not increase the risk for spontaneous preterm delivery. However, it is associated with an increased risk for overall preterm delivery.

What does this add to what is known?

This study provides further clarity on the risks for overall and spontaneous preterm delivery among patients with obesity with cervical cerclage. The findings of this study should be used in the counseling of patients with obesity who require cerclage placement and provide reassurance on the efficacy of cerclage in women with obesity. This study also highlights the risks for preterm delivery associated with obesity secondary to other medical indications.

Among patients with preexisting risks for preterm delivery that necessitate cervical cerclage placement, data on whether obesity affects the efficacy of cerclage placement are limited and conflicting. Among women with cerclage, it was shown that maternal obesity did not seem to impact the rates of overall preterm delivery²⁵ and spontaneous preterm delivery.²⁶ However, a retrospective study demonstrated that there was an inverse correlation between BMI and gestational age at delivery among patients with cervical cerclage.²⁷ Some researchers theorized that the increased abdominal pressure on the cerclage may have contributed to cerclage failure.^{27,28}

Thus far, the data on maternal obesity and preterm delivery among patients with cerclage are sparse and conflicting, necessitating further elucidation. Therefore, we aimed to evaluate the risk for preterm delivery associated with obesity among patients with cerclage using a large population-based cohort study. We hypothesized that maternal obesity increased the risks for spontaneous preterm delivery and overall preterm delivery among patients with cervical cerclage.

Materials and Methods

This was a retrospective study in which data from the California Office of Statewide Health Planning and Development linked birth file from 2007 to 2012 were used, which is a birth registry database that was linked to hospital discharge International Classification of Diseases, Ninth Revision (ICD-9), billing codes. Cohort exclusion criteria included patients with missing data on BMI, multiple gestation, anomalous pregnancies, and gestations <20 weeks or >42 weeks (Figure 1). Patients with cerclage placement were identified using hospital birth records. Patients in each group were identified and were further categorized based on maternal prepregnancy BMI, with the nonobese group defined as having a BMI of <30 kg/m², the obese group as having a BMI of 30 to 40 kg/m², and morbidly obese group as having a BMI >40 kg/m². The primary outcome of interest was preterm delivery, defined as any delivery between 20 +0/7 weeks and 36+6/7 weeks' gestation. Furthermore, we evaluated the risk for severe prematurity, defined as deliveries up to 34+6/7 weeks' gestation. The outcomes of interest were further delineated as deliveries following spontaneous labor based on ICD-9 code 644.0 to 644.2. The risk for preterm delivery associated with obesity was estimated separately for pregnancies that required cervical cerclage and pregnancies that did not require cerclage.

The demographics of interest are included Table 1. Categorical variables were compared between the 2 groups using the chi-square test. A 2-tailed P

value of <.05 was used as the threshold for statistical significance. Logistic regression analysis was used to estimate the association between BMI group and the risk for preterm delivery. Odd ratios (ORs) with 95% confidence intervals (CIs) were calculated and adjusted ORs (aORs) were adjusted for race, insurance type, hypertension, preeclampsia, pregestational diabetes, gestational diabetes, history of preterm delivery, and maternal age. Bootstrapping was performed 50 times in the full sample to evaluate the risk for spontaneous preterm delivery given the relatively limited cohort of patients with a cerclage with morbid obesity. Statistical analyses were performed using Stata 14 (StataCorp, College Station, TX). This study was approved by the California State University - Fullerton Institutional Review Board (IRB# HSR-19-20-548).

Results

This retrospective cohort study included 3654 pregnant patients with cerclage placement among which there were 184 patients with morbid obestiy, 832 patients with obesity, and 2638 patients without obesity. The group of 2,784,882 patients without cervical cerclage included 81,700 patients with morbid obesity, 489,536 patients with obesity, and 2,233,435 patients without obesity. The demographics of all patients in each BMI group are shown in Table 1. Compared with the population without obesity, the groups with obesity and morbid obesity demonstrated higher rates of pregestational and gestational complications including chronic hypertension, preeclampsia, pregestational diabetes, gestational diabetes, and higher rates of previous preterm delivery, and a history of cesarean delivery.

The risks for spontaneous preterm deliveries were not different between those who received a cerclage in the obese and morbidly obese groups and those in the nonobese group (24.2% vs 20.6%; aOR, 1.18; 0.97–1.43; and 24.5% vs 20.6%; aOR, 1.12; 0.78–1.62, respectively) (Table 2). However, among patients without cerclage placement, the obese and morbidly obese groups had a

FIGURE 1 The cohort selection Initial Cohort = 3,186,553 260,813 Missing BMI excluded 95,053 Multiple gestations excluded 8,597 Anomalous pregnancies excluded 13,765 Gestations < 20 weeks or > 42 weeks excluded Final Cohort = 2,808,325

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higher risk for spontaneous preterm delivery than the nonobese group (5.1% vs 4.4%; aOR, 1.04; 1.02–1.05; and 5.9% vs 4.4%; aOR, 1.03; 1.00–1.07, respectively). After bootstrapping was performed, the risks for spontaneous preterm delivery remained the same (Supplementary Table 1).

TABLE 1 Demographics of patients in the nonobese, obese, and morbidly obese groups

Demographics	Nonobese	Obese (BMI 30.0–40.0 kg/m²)	Morbidly obese (BMI \geq 40 kg/m ²)	<i>P</i> value
	2,236,073	490,368	81,884	
Maternal age, mean \pm SD	28.2±6.4	28.5±6.0	28.8±5.6	<.0001
Gestational age at delivery	39 (38-40)	39 (38-40)	39 (38-40)	<.0001
Race				
White	41	53.3	46.3	<.001
Black	1	1.6	2.4	
Hispanic	54.5	41.2	47.7	
Asian	0.9	0.5	0.5	
Mix	1.6	2.3	2	
Other	0.9	1.1	1.1	
Chronic hypertension	1	1.8	3.1	<.001
Preeclampsia	2.6	4.9	7.6	<.001
Pregestational diabetes mellitus	0.04	0.2	0.5	<.001
Gestational diabetes mellitus	1.7	3.7	5.6	<.001
Private insurance	50.7	40.8	40.8	<.001
History of preterm	0.3	0.4	0.6	<.001
History of cesarean delivery	13.2	21.6	28.4	<.001
BMI, body mass index.				

Among patients with cerclage placement, the rates of overall preterm delivery at <37 weeks' gestation in the obese and morbidly obese groups were higher than those in the nonobese group (33.7% vs 28.2%; aOR, 1.23; 1.03-1.46; and 32.1% vs 28.2%; aOR, 1.01; 0.72 -1.43, respectively) as demonstrated in Table 3. Similarly, the rates of preterm delivery at ≤ 34 weeks' gestation in the obese group was higher than those in the nonobese group (21% vs 16.9%; aOR, 1.33; 1.08-1.63). The morbidly obese group carried the highest rate of preterm delivery at ≤34 weeks' gestation when compared with the nonobese group (25% vs 16.9%; aOR, 1.54; 1.06 -2.23) (Figure 2).

Among patients without cerclage placement, the obese and morbidly obese groups continued to demonstrate increased risks for preterm delivery when compared with the nonobese group, similar to the trend seen in the population of patients with cerclage (Figure 2). The rates for preterm delivery at <37 weeks' gestation in the obese and morbidly obese groups were higher than that in the nonobese group (8.1% vs 6.7%; aOR, 1.04; 1.03-1.05; and 9.5% vs 6.7%; aOR, 1.07; 1.04-1.10, respectively). Similarly, the rates of preterm delivery <34 weeks' gestation in the obese and morbidly obese group were higher than that in the nonobese group, (3.3% vs 2.5%; aOR, 1.11; 1.09-1.14; and 3.9% vs 2.5%; aOR, 1.13; 1.08-1.17, respectively).

Discussion Principal findings

Among patients with cervical cerclage, there was no difference in the risk for spontaneous preterm delivery among all of the BMI groups. However, maternal obesity was associated with an overall increased risk for preterm delivery. Specifically, the risk for overall preterm delivery by 34 weeks' gestation was 33% higher among women with obesity and 54% higher among women with morbid obesity. Together, these findings suggest that the primary reason for the observed increase in preterm delivery associated with obesity was because of medical indications. TABLE 2

Cerclage groups	Obesity type	Spontaneous preterm delivery ^a	OR (95% CI)	Adjusted OR ^b (95% CI)
Cerclage placed	Nonobese ^c	544 (20.6)	Ref	Ref
	Obese	201 (24.2)	1.23 (1.02-1.47)	1.18 (0.97-1.43)
	Morbidly obese	45 (24.5)	1.25 (0.88–1.77)	1.12 (0.78–1.62)
Without cerclage	Nonobese	99,125 (4.4)	Ref	Ref
	Obese	25,102 (5.1)	1.16 (1.15—1.18)	1.04 (1.02-1.05)
	Morbidly obese	4802 (5.9)	1.34 (1.31-1.39)	1.03 (1.00-1.07)

BMI, body mass index; CI, confidence interval; OR, odds ratio.

^a The data are presented as number (percentage); ^b ORs were adjusted for maternal age, race, insurance type, hypertension, preeclampsia, pregestational diabetes mellitus, gestational diabetes mellitus, history of preterm delivery, and history of cesarean delivery; ^c ^cPeople without obesity are those with a BMI <30 kg/m², people with obesity are those with a BMI of 30 to 40 kg/m², and those with morbid obesity have a BMI >40 kg/m².

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Results in the context of what is known

Obesity continues to serve as one of the major risk factors for many adverse pregnancy complications. In existing literature, the association between obesity and preterm delivery in patients with cerclage has been conflicting. In one study, maternal obesity was observed to not have an impact on the gestational age at delivery among women with a cerclage,²⁵ whereas another study showed that BMI was a significant predictor of preterm delivery.²⁷

Our study has further elucidated the risks for preterm delivery. In the patient population at risk for preterm delivery with a need for cerclage placements, this study demonstrated that obesity was not associated with a higher risk for preterm delivery secondary to the onset of spontaneous labor. This finding reassuringly suggests that cerclage is as efficacious in preventing spontaneous preterm delivery in women with obesity as it is in women without obesity. However, the overall risk for preterm delivery was higher in the obese and morbidly obese groups. These findings suggest that medically indicated preterm deliveries may have been a major contributor to the observed increase in the overall preterm delivery in our study and in other published studies.²⁷

In the population of pregnant patients without cervical cerclage, the mechanisms

of how obesity impacts spontaneous preterm delivery is thought to be a consequence of the increased levels of cytokines causing prostaglandin production and matrix-degrading enzymes, which is associated with cervical ripening, weakening of the membranes, and preterm contractions.^{29,30} The results of this study demonstrated that among patients with cerclage, the impact of obesity and morbid obesity on spontaneous preterm delivery was mitigated such that the risks were the same as those of patients without obesity. Increased adipokines from visceral fats in patients with obesity have been shown to increase systemic inflammation,^{31,32} contributing to many obesity-related maternal complications, such as endothelial damage and insulin resistance seen in preeclampsia and gestational diabetes, respectively, among other gestational disorders.^{33–35} The results from this study of patients with obesity and morbid obesity with cerclage align with the current literature on their counterparts without cerclage, demonstrating the role of obesity in increasing the overall preterm delivery risks associated with other gestational disorders.

Other factors surrounding the cervical cerclage placement procedure could potentially play a role in cerclage effectiveness. Based on current knowledge, cerclage suture materials have been proposed to have differential effects on pregnancy loss. In a prospective cohort study, it was noted that the risk for pregnancy loss and preterm birth when using a monofilament suture was lower than when braided suture was used (7% vs 18%; relative risk, 0.34; 95% CI, 0.18-0.63; and 17% vs 28%; P=.0006, respectively).³⁶ However, a recent, large randomized control trial (the Cerclage Suture Type for an Insufficient Cervix and its Effects on Health Outcomes trial) in which monofilament and braided suture for cervical cerclage were compared showed no difference in the primary outcome of pregnancy loss.³⁷ The effects of different suture materials on preterm birth in obese and nonobese populations deserve further elucidation. Physicians who perform cerclage could also face technical challenges during placement because of obesity status, requiring expert skills. Furthermore, a retrospective study in which the effectiveness of the McDonald vs Shirodkar cerclage type was investigated suggested that in normal patients, when compared with patients who were overweight, cerclage type did not seem to impact pregnancy duration. However, when patients with obesity who received the McDonald cerclage were compared with those who received the Shirodkar cerclage, they had significantly longer pregnancies.³⁸ The Shirodkar cerclage presented a more technically challenging procedure. Similar to any other

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The risks for preterm delivery at <37 weeks and <34 weeks of gestation among patients with obesity and morbid obesity and with and withou	cerclage	Adiusted OR ^b

Cerclage groups	Obesity type	z	Preterm <34 wk ^a	<i>P</i> value	or (95% CI)	Adjusted OR ^b (95% Cl)	Preterm <37 wk	<i>P</i> value	OR (95% CI)	Adjusted OR (95% Cl)
Cerclage placed	Nonobese ^c	2638	445 (16.9)	.001	Ref	Ref	745 (28.2)	600'	Ref	Ref
	Obese	832	175 (21.0)		1.31 (1.08–1.60)	1.33 (1.08–1.63)	280 (33.7)		1.29 (1.09–1.52)	1.23 (1.03–1.46)
	Morbidly obese	184	46 (25.0)		1.64 (1.16–2.33)	1.54 (1.06–2.23)	59 (32.1)		1.20 (0.87-1.65)	1.01 (0.72-1.43)
Without cerclage	Nonobese	2,233,435	56,815 (2.5)	<.0001	Ref	Ref	153,620 (6.7)	<.0001	Ref	Ref
	Obese	489,536	16,077 (3.3)		1.30 (1.28–1.32)	1.11 (1.09—1.14)	39,417 (8.1)		1.18 (1.17–1.20)	1.04 (1.03-1.05)
	Morbidly obese	81,700	3195 (3.9)		1.56 (1.50-1.62)	1.13 (1.08–1.17)	7724 (9.5)		1.41 (1.38–1.45)	1.07 (1.04–1.10)
BMI, body mass inde>	BMI, body mass index; CI, confidence interval; OR, odds ratio.	OR, odds ratio.								

People without obe-The data are presented as number (percentage)." ORs were adjusted for maternal age, race, insurance type, hypertension, preeclampsia, pregestational diabetes, gestational diabetes, history of preterm delivery, and history of cesarean delivery. sity are those with a BMI < 30 kg/m², people with obesity are those with a BMI of 30 to 40 kg/m², and those with morbid obesity have a BMI > 40 kg/m²

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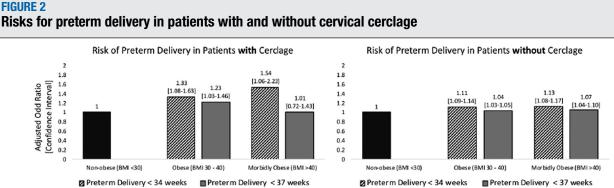
surgical procedures, cerclage effectiveness, in large part, depend on the skills of the practitioners.

Clinical implication

Based on the results of this study, patients who meet the criteria for history-indicated cerclage should be counseled on the risk for preterm delivery associated with obesity and its link with medical indications and adverse pregnancy outcomes. Reassurance should be provided to patients with obesity because the procedure itself seemed to be equally efficacious in patients with obesity as in patients without obesity. More importantly, the results of this study could provide further information on risk stratification for pregnancies affected by obesity during preconception counseling and initial prenatal visits. As such, during preconception counseling, patients with a history of cerclage placement in a previous pregnancy should be encouraged to optimize their BMI before a future pregnancy to further lower their risk for preterm delivery.

Strengths and limitations

One of the main strengths of this study was the large cohort of patients with cerclage placement during pregnancy, which further allowed us to study the risks for preterm delivery in a relatively small subgroup of women with cerclage and a BMI of >40 kg/m². The large cohort of patients with and without cerclage allowed us to evaluate the overall risk for preterm delivery associated with maternal obesity and to adjust for many of the common medical conditions that lead to medically indicated preterm deliveries. Furthermore, in this study, we were able to separately analyze the impact of obesity on the preterm delivery risks in the general population without cerclage placement to comprehensively evaluate whether these risks could persist in patients with cerclage. This study evaluated the impact of obesity on the risk for spontaneous preterm delivery while further assessing the



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overall risk for preterm delivery in this particular population.

We acknowledge the limitations of this study. Because of the nature of this study in that a population-based data registry was used, it is subject to data entry and coding errors. Furthermore, patients with history-indicated, ultrasound-indicated, or physical examination-indicated cerclage were grouped together, and the indications leading to cerclage placement and the appropriateness of cerclage placement could not be stratified further and evaluated because of the lack of specification in the data registry. In addition, we acknowledge the limited sample size of patients with morbid obesity with a cervical cerclage and the requirement for performing bootstrapping. We cannot fully determine the risk in this unique group and more studies would be needed to further elucidate this. Lastly, given the retrospective cohort study design, the findings should be interpreted within its limitation. As such, the findings may not necessarily establish causation between obesity and preterm delivery in the cerclage population, and thus, further randomized control trials are needed to evaluate this role.

Conclusion

Maternal obesity is not associated with an increased risk for spontaneous preterm delivery. Patients with obesity and morbid obesity should be reassured that cerclage does not seem to be differentially efficacious in preventing spontaneous preterm labor in this group when compared with cerclage in their counterparts without obesity, but they should be aware that they are still at risk for overall preterm delivery because of other gestational complications.

Supplementary materials

Supplementary material associated with this article can be found in the online version at doi:10.1016/j.xagr.2023.10 0211.

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