

#### ORIGINAL RESEARCH

# Correlation Between Postpartum Myometrial Elasticity and Obstetric Complications in Pregnant Women with Gestational Diabetes Mellitus

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Objective: To investigate postpartum myometrial elasticity in pregnant women with gestational diabetes mellitus (GDM) using shear wave elastography (SWE) and analyze the correlation between myometrial SWE values and obstetric complications.

Methods: Clinical data of women who gave birth at Affiliated Hospital of Yangzhou University from August 2022 to July 2023 were retrospectively analyzed and divided into two groups based on GDM diagnosis: the GDM group and the healthy control group. SWE was used to measure the elasticity values of the anterior and posterior myometrial walls in both groups. Differences in placental attachment position and SWE values at the placental attachment site were compared between the two groups. Spearman correlation analysis was utilized to evaluate the correlation between SWE values and obstetric complications.

Results: Glycated hemoglobin (HbA1c), homeostasis model assessment of insulin resistance (HOMA-IR), fasting plasma glucose (FPG), 2-hour postprandial glucose (2h PG), triglycerides (TG), and total cholesterol (TC) levels were higher in the GDM group than in the healthy control group (P<0.05). There was a statistically significant difference in placental attachment position between the two groups (P<0.05). In both the GDM (17.52±0.42 vs 25.29±0.74, P=0.001) and control groups (14.06±5.01 vs 22.20±6.34, P=0.001), mean SWE values were significantly lower for anterior versus posterior placental attachment, and mean SWE values were also significantly higher in the GDM versus control group for both anterior (17.52±0.42 vs 14.06±5.01, P=0.001) and posterior placental attachment (25.29±0.74 vs 22.20±6.34, P=0.001). Spearman correlation analysis showed that postpartum hemorrhage (r=0.632, P=0.017), preeclampsia (r=0.818, P=0.014), premature rupture of membranes (r=0.710, P=0.012), placental abruption (r=0.732, P=0.031), and ketoacidosis (r=0.729, P=0.022) were negatively correlated with average myometrial SWE values in the GDM group (P < 0.05).

Conclusion: SWE values at the placental attachment site were higher in GDM patients than in healthy pregnant women, and myometrial elasticity was positively correlated with obstetric complications.

Keywords: elastography, gestational diabetes, myometrium, pregnancy complications

## Introduction

Gestational diabetes mellitus (GDM) refers to glucose intolerance of variable severity with onset or first recognition during pregnancy, occurring in 1-10% of gravidas. The prevalence of GDM has progressively risen in tandem with elevated living standards in recent years.<sup>2</sup> If not controlled in a timely manner, GDM can easily lead to adverse pregnancy outcomes for pregnant women such as polyhydramnios, hypertensive disorders in pregnancy, dystocia, cesarean section, macrosomia, respiratory distress and hypoglycemia in newborns, posing serious threats to the health of mothers and infants.<sup>3</sup> Excessive glucose from pregnant women entering the fetus during pregnancy can lead to macrosomia, increasing the difficulty of delivery, and would also pose serious threats to the safety of mothers and

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infants. It has been reported that the placental structural and functional tissues in GDM pregnant women undergo pathological changes with the occurrence and development of GDM, leading to increased placental hardness. 5,6 Also, the dysregulation of various hormones and cytokines caused by GDM can lead to placental and myometrial dysfunction, increasing the risks of adverse pregnancy outcomes and long-term maternal and infant complications. <sup>7,8</sup>

Shear wave elastography (SWE) is an imaging method used to evaluate tissue stiffness. Studies have reported that SWE can be utilized to evaluate endometrial lesions, preeclampsia, placental elasticity, and post-abortion myometrial elasticity. 11 Research has been conducted on the elasticity of the myometrial measured by SWE of GDM patients during mid-pregnancy and its association with obstetric complications, however, there is currently a lack of studies on the elasticity of the myometrial postpartum in this population. Therefore, this study aims to explore the correlation between postpartum myometrial elasticity and obstetric complications.

## **Materials and Methods**

## General Information

Clinical data of 189 women delivering at Affiliated Hospital of Yangzhou University Obstetrics 8/2022-7/2023 were selected after retrospective review. Those with GDM (n=58) were assigned to the GDM group, with healthy cases without GDM (n=131) comprising the control group. Inclusion criteria: (1) singleton pregnancy, no history of abortion or termination; (2) placenta attached to the anterior or posterior myometrial wall; (3) no fetal malformation or intrauterine infection detected during pregnancy; (4) successful acquisition of myometrial SWE results; (5) GDM diagnostic criteria:  $^{12}$  FPG  $\geq 5.1$  mmol/L or 2-hour postprandial glucose  $\geq 8.5$  mmol/L after a 75g oral glucose tolerance test. Exclusion criteria: (1) hypertension, hyperlipidemia, endocrine/metabolic diseases, heart/liver/kidney comorbidities; (2) polyhydramnios or other comorbidities; (3) Patients with incomplete clinical information. All participants provided signed informed consent forms approved by the hospital's ethics committee.

## **Examination Methods**

#### **SWE Measurement**

Examinations were performed using a Resona 9 color Doppler ultrasonography system (Mindray, China) with probe frequencies set at 1-6 MHz. The participants were instructed to lie supine and breathe steadily. Following inspection of the myometrial position, the ultrasound transducer was set to SWE mode and applied over the central and peripheral regions of the uterus. Pregnant women were instructed to hold their breath at the end of exhalation in a calm state for imaging. When scanning, care was taken to evade myometrial vascular sinuses and sites of heavy calcification, ensuring complete, homogeneous and consistent color saturation of the elastography patterns. Target imaging criteria: 100% credibility index and 5 star M-STB rating. The scanning protocol was repeated to acquire 3 comparable measurements, of which the mean was calculated. After measurement of the anterior myometrial wall, the posterior myometrial wall was measured using the same method. The subjects were examined by a physician with more than five years of experience in the ultrasound department of our hospital, and this study was kept confidential among the various operators.

## **Blood Glucose Measurement**

Blood glucose levels were measured using a glucose assay kit (GLU(HK) kit, cat:1000020106). Oral glucose tolerance test: Subjects at 24 weeks of gestation were instructed to fast for at least 8 hours overnight and then ingest 250 mL of water containing 75 g of glucose the next morning within 5 minutes. Venous blood samples were collected 1 hour before and 1 and 2 hours after the first glucose ingestion to measure blood glucose and insulin levels.

## Observation and Evaluation Indices

#### Participant Data

Body mass index (BMI), glycated hemoglobin (HbA1c), fasting plasma glucose (FPG), 2-hour postprandial glucose (2h PG), triglycerides (TG), total cholesterol (TC), and homeostasis model assessment of insulin resistance (HOMA-IR) were collected for participants in both groups. Obstetric complications recorded included postpartum hemorrhage, preeclampsia, premature rupture of membranes, preterm birth, and ketoacidosis.

#### Insulin Resistance

Homeostasis model assessment of insulin resistance (HOMA-IR)<sup>13</sup> was calculated using the following formula: fasting blood glucose (FPG, mmol/L) x fasting insulin (FINS,  $\mu$ U/mL)/22.5.

## Statistical Analysis

Data were analyzed using SPSS 22.0. Normal distributed data were presented as mean  $\pm$  standard deviation (x $\pm$ s) and groups were compared by independent t-tests. Categorical and ordinal variables were compared by chi-square test or Fisher's exact test. Spearman correlation analysis was used to evaluate associations between SWE and variables. P <0.05 indicates statistically significant differences.

#### Results

## Comparison of Metabolic Indicators

The GDM group comprised individuals aged between 20 and 38 years, with a mean age of (29.34±3.57) years and an average BMI of 25.19±3.27 kg/m<sup>2</sup>. The healthy control group consisted of individuals aged between 24 and 39 years, with a mean age of (29.29±3.70) years and an average BMI of 24.38±4.51 kg/m<sup>2</sup>. In GDM group, 39 (67.24%) pregnant women had vaginal birth, 19 (32.76%) pregnant women had Caesarean section and number of deliveries was 1.40±0.50. In healthy group 97 (71.32%) pregnant women had vaginal birth, 37 (27.21%) pregnant women had Caesarean section and number of deliveries was 1.44±0.55. Statistical analysis revealed no significant differences between the two groups (P>0.05).

The GDM group exhibited higher HbA1c (5.29±1.31 vs 5.07±0.67), HOMA-IR (2.8±1.29 vs 1.3±0.50) levels, FBG (5.45±2.76 vs 4.79±2.14), 2h PG (9.21±3.31 vs 6.45±2.76), TG (3.72±2.02 vs 1.65±0.57), and TC (6.42±1.13 vs 5.14 ±2.12) levels, with statistically significant differences (P<0.05). as demonstrated in Table 1. The routine examinations confirmed the GDM status of enrolled subjects in corresponding cohort. These demographic details contribute to the robustness of our study, ensuring a comprehensive portrayal of the participant characteristics.

## Comparison of SWE Values by Placental Location

In the GDM group, 34 (58.62%) cases exhibited placental attachment to the anterior uterine wall, while 24 (41.38%) cases displayed attachment to the posterior uterine wall. Conversely, In the healthy control group, 65 (47.79%) cases demonstrated placental attachment to the anterior uterine wall, and 71 cases (52.21%) to the posterior uterine wall. The disparity between the two groups was statistically significant (P<0.05).

Table 1 Chilical Characteristics of the Study Groups					
	GDM Group (58 Cases)	Healthy Control Group (136 Cases)	χ²/t	P	
Age	29.34±3.57	29.29±3.70	9.923	0.730	
HbAIc (mmol/L)	5.29±1.31	5.07±0.67	10.347	0.021	
HOMA-IR	2.8±1.29	1.3±0.50	12.046	0.017	
BMI (Kg/m <sup>2</sup> )	25.19±3.27	24.38±4.51	8.912	0.052	
FBG (mmol/L)	5.45±2.76	4.79±2.14	7.629	0.039	
2h PG (mmol/L)	9.21±3.31	6.45±2.76	6.701	0.023	
TG (mmol/L)	3.72±2.02	1.65±0.57	2.904	0.027	
TC (mmol/L)	6.42±1.13	5.14±2.12	3.901	0.032	
Delivery type					
Vaginal birth	39 (67.24)	97 (71.32)	13.781	0.057	
Caesarean section*	19 (32.76)	37 (27.21)			

Table I Clinical Characteristics of the Study Groups

1.40±0.50

Notes: \*The caesarean section information of 2 patients in healthy control group was missing.

Abbreviations: HbA1c, glycated hemoglobin; HOMA-IR, homeostatic model assessment of insulin resistance; GDM, gestational diabetes mellitus; SWE, shear wave elastography; BMI, body mass index; FBG, fasting blood glucose; 2hPG, 2-hour postprandial glucose; TG, triglycerides; TC, total cholesterol.

1.44+0.55

Number of deliveries

0.672

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Both in the GDM (17.52±0.42 vs 25.29±0.74, P=0.001) and control groups (14.06±5.01 vs 22.20±6.34, P=0.001), mean SWE values were significantly lower for anterior placental attachment compared to posterior attachment. Additionally, mean SWE values were significantly higher in the GDM group compared to the control group for both anterior (17.52±0.42 vs 14.06±5.01, P=0.001) and posterior placental attachment (25.29±0.74 vs 22.20±6.34, P=0.001). Refer to Table 2 and Figures 1 and 2 for detailed information.

## Comparison of Obstetric Complications Between Both Groups

Following the computation of the mean SWE values for the anterior and posterior uterine walls in both groups, we assessed the incidence of obstetric complications between them. Results indicated a significantly higher occurrence of postpartum hemorrhage ( $\chi$ 2=9.328, P=0.004), preeclampsia ( $\chi$ 2=10.457, P=0.027), premature rupture of membranes ( $\chi$ 2=21.794, P=0.012), placental abruption ( $\chi$ 2=16.674, P=0.038), and ketoacidosis ( $\chi$ 2=6.652, P=0.023) in the GDM group compared to the healthy control group, with statistically significant differences (P<0.05), refer to Table 3 for comprehensive details.

## Correlation Between Obstetric Complications and SWE Values in the GDM Group

Following the computation of mean SWE values at the placental attachment site in the GDM group, we conducted an analysis to examine the correlation between maternal obstetric complications and the mean uterine SWE values. Spearman correlation analysis showed that postpartum hemorrhage (r=-0.632, P=0.017), preeclampsia (r=-0.818, P=0.014), premature rupture of membranes (r=-0.710, P=0.012), placental abruption (r=-0.732, P=0.031), and ketoacidosis (r=-0.729, P=0.022) exhibited a negative correlation with mean uterine SWE values (P<0.05). Comprehensive details are available in Table 4.

## **Discussions**

The uterus plays a vital role in fostering fetal development, and maternal and uterine disorders can impact delivery outcomes. Clinical observations have found that changes in myometrial elasticity in mid to late pregnancy are related to pregnancy complications, although literature on postpartum myometrial elasticity and obstetric complications remains scarce.

While there have been studies focusing on the measurement of uterine muscle layer elasticity using SWE, the unique aspect of this research lies in dynamically observing the patterns of uterine contractions. <sup>14</sup> To date, there is a lack of research specifically addressing the measurement of uterine muscle elasticity during contraction and relaxation using SWE. SWE is an imaging technique for assessing tissue stiffness by using ultrasound radiation force to induce shear waves in tissues. Stiffer tissues exhibit faster shear velocities and higher Young's moduli. Yuan Shengmei et al. <sup>15</sup> performed SWE examinations on mid-pregnancy singleton placentas and found that the placenta elasticity values in the early-onset preeclampsia group were significantly higher than the normal pregnancy group, thereby identifying SWE as useful for evaluating placental function. This study used SWE to assess myometrial elasticity and found that the mean SWE values at the placental attachment site were significantly higher in GDM patients than the healthy controls. SWE values are thought to be related to myometrial and placental tissue changes during pregnancy in GDM patients. Focal thickening of the chorionic basement membrane and patchy necrosis of trophoblastic cells are accompanied by significant

Table 2 SWE Measurements Results

	GDM Group (58 Cases)	Healthy Control Group (136 Cases)	χ²lt	P
Placental location				
Anterior wall of uterus	34 (58.62)	65 (47.79)	21.791	0.018
Posterior wall of uterus	24 (41.38)	71 (52.21)	27.238	0.0012
SWE value of placenta attachment (kPa)				
Anterior wall of uterus	17.52±0.42	14.06±5.01	15.294	0.001
Posterior wall of uterus	25.29±0.74	22.20±6.34	28.372	0.001

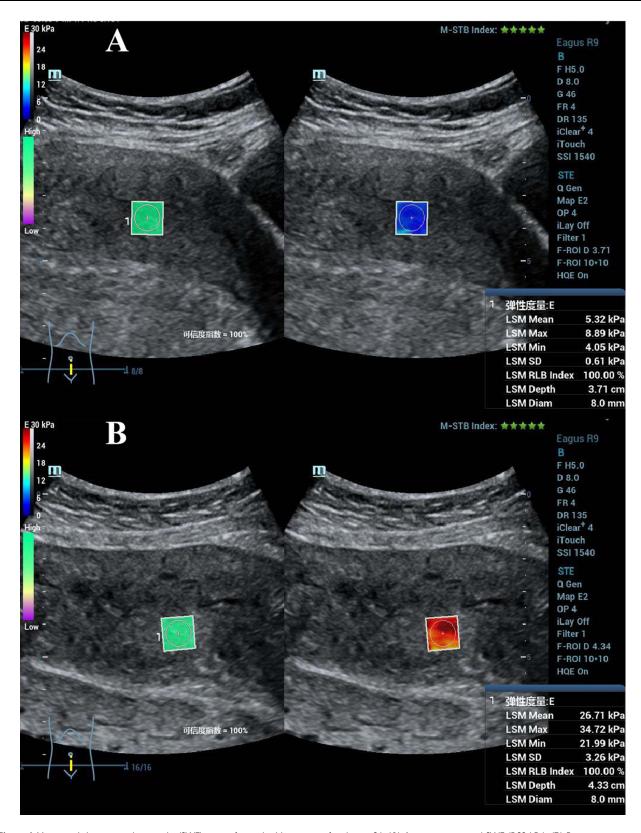


Figure 1 Myometrial shear wave elastography (SWE) images from a healthy pregnant female, age 36. (A) Anterior myometrial SWE (5.32 kPa). (B) Posterior myometrial SWE (26.71 kPa).

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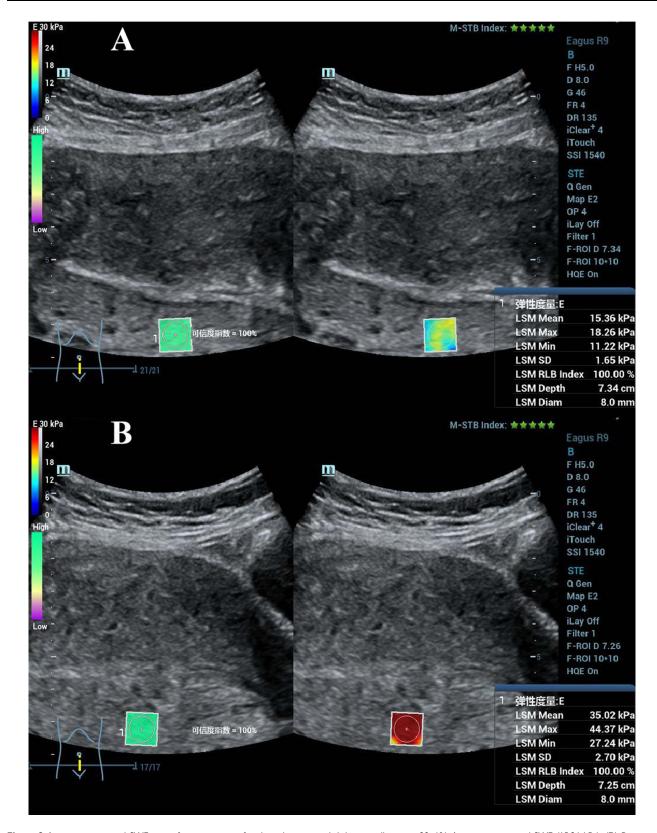


Figure 2 Anterior myometrial SWE image from a pregnant female with gestational diabetes mellitus, age 29. (A) Anterior myometrial SWE (15.36 kPa). (B) Posterior myometrial SWE (35.02 kPa).

Table 3 Obstetric Complications in the Study Groups

Group	GDM Group (58 Cases)	Healthy Control Group (131 Cases*)	χ²	P value
Postpartum hemorrhage	5(8.62)	7(5.34)	9.328	0.004
Preeclampsia	7(12.07)	2(1.53)	10.457	0.027
Premature rupture of membranes	11(18.97)	13(9.92)	21.794	0.012
Placental abruption	7(12.07)	8(6.11)	16.674	0.038
Preterm birth	6(10.39)	14(10.69)	19.353	0.326
Ketoacidosis	3(5.17)	2(1.53)	6.652	0.023

Notes: \*The obstetric information of 5 patients in healthy control group is missing. GDM stands for gestational diabetes mellitus.

**Table 4** Correlations Between SWE Values and Obstetric Complications in the GDM Group

Group	SWE Results		
	r	P value	
Postpartum hemorrhage	-0.632	0.017	
Preeclampsia	-0.818	0.014	
Premature rupture of membranes	-0.710	0.012	
Placental abruption	-0.732	0.031	
Preterm birth	−0.25 I	0.329	
Ketoacidosis	-0.729	0.022	

**Notes**: GDM stands for gestational diabetes mellitus and SWE for shear wave elastography.

proliferation of trophoblastic cells, which can lead to increased placental hardness and higher SWE values. <sup>16</sup> However, since the placental attachment site is adjacent to the placenta, trophoblastic cell proliferation can also cause stiffening and increased SWE values. SWE has also been applied for the assessment of uterine tonicity after vaginal delivery. <sup>17</sup> For relative researches SWE using on myometrial elasticity, there have been studies in relation to ectopic pregnancy and the prediction of preterm birth probability, and myometrial elasticity presented to be a significant predictor for preterm delivery. <sup>18,19</sup>

The results of this study also showed that the incidence of postpartum hemorrhage, preeclampsia, premature rupture of membranes, low birth weight, macrosomia, neonatal asphyxia, and neonatal hypoglycemia in the GDM group was significantly higher than that in the healthy control group, and was negatively correlated with SWE values. The possible reason is that GDM pregnant women secrete excessive anti-islet substances and experience inflammatory reactions during pregnancy, while high expression of anti-islet substances and inflammatory mediators can affect maternal metabolic processes, causing vascular endothelial damage, uterine dysfunction, and thus lead to pathological histological changes in the uterus, resulting in increased tissue stiffness.<sup>20</sup>

There is a strong likelihood of a correlation between the thickness of the uterine myometrium and its elasticity. However, current research in this area is still lacking. Clinically, a decrease in myometrial thickness is associated with an increased risk of adverse pregnancy outcomes. Researchers have discovered<sup>21</sup> that uterine muscular thickness is associated with preterm delivery if there is no premature rupture of membranes. Guo et al<sup>22</sup> observed that spontaneous preterm birth patients exhibited markedly reduced anterior uterine muscular thickness in mid-gestation compared to full-term counterparts with intact membranes. Atarjavdan L et al<sup>23</sup> demonstrated pronounced anterior and lower myometrial thickening in preterm premature rupture of membranes cases, arising after lengthy latency. Thus, late gestational assessment of myometrial muscularity could predict adverse pregnancy outcomes. Some studies have shown that uneven uterine muscular thickness during delivery may lead to uncoordinated myometrial contractions, resulting in dystocia. For such pregnant women, early evaluation of the uterine muscular condition is instructive in choosing the delivery method,

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which helps reduce pregnancy and obstetric complications and lower the risks faced by fetuses and mothers. In subsequent studies, we will specifically investigate the connection between mid-pregnancy myometrial thickness and uterine elasticity. Additionally, at the current clinical stage, there is no conclusive evidence from clinical research indicating a difference in elasticity between the anterior and posterior walls of the uterus. When establishing our control group, we initially considered it as a potential clinical variable to control for. Surprisingly, our results did reveal a lower elasticity in anterior uterine wall than posterior walls. We speculate that this outcome may be related to the physiological structure of the uterus and its ligaments.

In summary, SWE emerges as a valuable tool for assessing uterine elasticity, providing insights into pregnancy complications, placental function, and potential predictive value for preterm birth, particularly in the context of conditions like gestational diabetes mellitus. The findings suggest a promising avenue for further research and clinical applications to reduce risks and complications for both fetuses and pregnant women. However, this study has certain limitations: it only explored the correlation between postpartum myometrial elasticity at the placental attachment site and the occurrence of obstetric complications, without an in-depth investigation of changes in the myometrial layer. In addition, the sample size included in the study was small, which may have led to bias. Future studies should use more samples to explore how to reduce pregnancy and obstetric complications, thereby lowering the risks faced by fetuses and pregnant women.

## **Conclusions**

SWE values measured at the placental attachment site were higher in GDM patients compared to healthy pregnant women and mean myometrial SWE values were strongly correlated with adverse pregnancy and delivery outcomes. Thus, SWE detection may be considered an effective means of myometrial functional assessment.

## **Disclosure**

The authors report no conflicts of interest in this work.

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