



2-2023

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### How to Cite This Article

Elomda, Fahd Abd elaal; kheder, Mohamad Abdel Hameed; Said, Ahmed Mohamed; and Elsead, Mohamad Yousef (2023) "Prediction of Fetal Macrosomia in Patients with Gestational Diabetes Mellitus through measuring the umbilical cord thickness and glycated hemoglobin (HbA1c) levels," *Al-Azhar International Medical Journal*: Vol. 4: Iss. 2, Article 2.

DOI: <https://doi.org/10.58675/2682-339X.1649>

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## ORIGINAL ARTICLE

# Prediction of Fetal Macrosomia in Patients with Gestational Diabetes Mellitus Through Measuring the Umbilical Cord Thickness and Glycated Hemoglobin Levels

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## Abstract

**Background:** Diabetes mellitus, one of the most prevalent medical issues, has emerged as a serious concern to pregnant women. It is linked to a number of maternal and fetal issues, including delivery traumas, perinatal death, shoulder dystocia, macrosomia, and operating room interference. Early prediction of fatal complications will improve outcome and reduce perinatal mortality.

**Aim and objectives:** To determine whether glycated hemoglobin and umbilical cord thickness in pregnant women with diabetes can accurately predict fetus's macrosomia.

**Patients and methods:** A 6-month prospective observational research study was performed on 100 women with gestational diabetes who were 28–29-week pregnant at Al-Hussein University Hospital. Patients had a thorough medical history review, an ultrasound assessment, and an ultrasonography examination.

**Result:** Glycated hemoglobin and Umbilical Cord Thickness (UCT) at 27–28 weeks and 36–37 weeks of gestation were significantly different across the three groups that were examined.

**Conclusion:** Severe obstetric problems, including shoulder dystocia and postpartum hemorrhage, are caused by macrosomia. There are times when it is difficult to foresee shoulder dystocia. The group most at risk for these issues may be found, however, using macrosomia prediction. There have been documented studies using sonographic measurement for predicting fetal macrosomia. Fetal macrosomia is well predicted by the thickness of the umbilical cord and the fetal fat layer.

**Keywords:** Diabetes, Glycated hemoglobin, Macrosomia

## 1. Introduction

The umbilical cord controls the flow of blood between the mother and the fetus. It typically consists of two arteries that carry venous blood and a vein that carries arterial blood, both cushioned by remains of the allantoides and a unique kind of mucous connective tissue called Wharton's jelly.<sup>1</sup>

Macrosomia is defined by the American College of Obstetricians and Gynecologists as a birth weight

that is more than 4000 g regardless of weeks of gestation, or that is greater than the 90th percentile for gestational age after adjusting for neonatal sex and ethnicity.<sup>2</sup> Predictions of the risk of macrosomia during pregnancy have been made by medical experts using glycated hemoglobin (HbA1c) values and estimated fetus weight obtained from ultrasounds.<sup>3</sup>

Gestational diabetes mellitus (GDM) is associated with several unfavorable pregnancy outcomes, including macrosomia and cesarean section

Accepted 15 September 2022.

Available online 15 May 2023

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<https://doi.org/10.58675/2682-339X.1649>

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delivery.<sup>4</sup> Macrosomia is a well-known sign of maternal diabetes in fetus, which is highly connected with preterm, respiratory distress syndrome, birth trauma, fetal mortality, and bad mother outcome.<sup>5</sup>

Overall, 2–6% of pregnant women have GDM, which increases the risk of serious negative perinatal results such macrosomia and birth injuries.<sup>6</sup> Macrosomic fetal birth has been linked to unfavorable outcomes for both mother and baby. It is possible to have shoulder dystocia during birth and a resulting persistent brachial plexus damage. When compared with babies of normal weight, macrosomic fetuses have greater rates of newborn death and morbidity.<sup>7</sup>

A greater incidence of surgical births, postpartum hemorrhages, birth injuries in vaginal delivery, and newborn hypoglycemia are all linked to fetal macrosomia. Only 40% of mothers who give birth to macrosomic kids had known maternal risk factors recognized.<sup>8</sup> Additionally, macrosomia raises the chance of developing some malignancies.<sup>9</sup> The macrosomia morbidity ranges from 7 to 10%. Therefore, monitoring abnormal birth weight and associated risk variables has substantial consequences for public health.<sup>10</sup> According to studies, obesity and metabolic diseases are more likely to occur in newborns who are macrosomic.<sup>11</sup>

The care of people with type 2 diabetes includes measures to reduce hyperglycemia, which is a significant aspect of diabetes as a serious public health problem. The HbA1c test determines hyperglycemia by determining the median blood glucose value over the prior 60–120 days. It is advised for diabetic individuals to aim for a HbA1c of 6.5% (48 mmol/mol).<sup>12</sup> Overall, 3.5–5.5% is the range for a nondiabetic normal HbA1c result. A healthy range for diabetics is between 6.5 and 7%. The amount of this HbA1c in those with poorly managed diabetes is much greater than in healthy individuals.<sup>13</sup>

Studies have revealed that HbA1c values and umbilical cord components may be used to predict macrosomia with greater accuracy.<sup>3</sup> Several studies have shown that HbA1c values can be employed to predict macrosomia, nevertheless.<sup>14</sup>

The purpose of this research was to evaluate how well HbA1c and umbilical cord thickness predict fetal macrosomia in diabetic pregnant women.

## 2. Patients and methods

### 2.1. Ethical considerations

The study protocol was submitted for approval by the Institutional Review Board, Al-Azhar University. Each individual who participated in the research provided informed verbal permission. At every

stage of the research, confidentiality, and personal privacy were protected.

In this prospective, observational study conducted over a 6-month period at Al-Hussein University Hospital, 100 pregnant women with gestational diabetes, 28–29 weeks of gestation, received routine prenatal treatment as inpatients or at outpatient clinics.

Women who were between 28- and 29-week pregnant, experienced gestational diabetes, and had a singleton pregnancy with normal umbilical morphology met the inclusion criteria. All patients with age over of 40 years, repeated fetal pregnancies, gestational diabetes, maternal chronic illness, multiparity, and drug usage during pregnancy were excluded.

The patients received a comprehensive medical history, a general evaluation, an abdominal examination, and a local clinical diagnosis. The gestational age was established using Negele's rule and confirmed using ultrasonography. Ultrasound with a convex transabdominal probe was used to evaluate umbilical cord thickness beginning at 28–29 weeks of gestation and then every 2 weeks until 36–37 weeks of gestation (mm<sup>2</sup>).

The fetal anthropometric parameters including biparietal diameter, femur length, and estimated fetal weight, which were automatically computed using Hadlock's algorithm.

If a pregnant woman had a fasting plasma glucose level of 126 mg/dl (7.0 mmol/l), a random plasma glucose level of 200 mg/dl (11.0 mmol/l), and a HbA1c level of 6.5%, she was diagnosed with gestational diabetes. When the estimated fetal weight was greater than the 90th percentile for gestational age or over 4000 g, macrosomia was considered.

Monitoring of patients during delivery was done, including birth weight, birth method, and fetus sex. Population characteristics such as age, BMI, parity, mode of delivery, ultrasound-estimated birth weight, birth weight, HbA1c, and umbilical cord thickness were noted. The accuracy of the umbilical cord thickness and HbA1c value in predicting fetal macrosomia served as the primary indicators of the study's effectiveness.

Using the SPSS program, data input and analysis were performed (SPSS 20.0 Version, Chicago, USA). We computed the mean, proportion, and percentage.  $\chi^2$  test was used to see if there was a correlation.

## 3. Results

There was a statistically significant difference in maternal BMI and gravidity between the groups (Table 1).

Table 1. Demographic and clinical features of the examined groups.

	No-macrosomia (N = 82)	Macrosomia (N = 18)	t	P
Age (years)				
Mean ± SD	30.11 ± 4.73	28.56 ± 4.12	1.29	0.201
BMI (kg/m <sup>2</sup> )				
Mean ± SD	26.34 ± 2.39	28.6 ± 2.88	3.49	0.001
Parity				
Mean ± SD	2.42 ± 1.15	2.85 ± 1.27	1.41	0.162
Gravidity				
Mean ± SD	2.73 ± 1.32	3.51 ± 1.44	2.23	0.028

Table 2. Neonatal characteristics and clinical data between the two studied groups.

	No-macrosomia (N = 82)	Macrosomia (N = 18)	t/χ <sup>2</sup>	P
GA (weeks)				
Mean ± SD	37.25 ± 1.86	36.76 ± 2.41	0.957	0.341
Birth weight (kg)				
Mean ± SD	3.18 ± 0.609	3.56 ± 0.863	2.21	0.029
Neonatal sex [n (%)]				
Male	36 (43.9)	7 (38.9)	0.151	0.697
Female	46 (56.1)	11 (61.1)		
Apgar at 1 min				
Mean ± SD	6.84 ± 2.26	7.2 ± 1.23	0.653	0.515
Apgar at 5 min				
Mean ± SD	9.25 ± 1.38	9.7 ± 1.09	1.29	0.198

Table 3. Fetal biometry of the two studied groups.

	No-macrosomia (N = 82)	Macrosomia (N = 18)	t	P
Biparietal diameter				
Mean ± SD	75.42 ± 4.53	77.15 ± 3.86	1.5	0.136
Abdominal circumference				
Mean ± SD	24.11 ± 3.92	25.29 ± 3.68	1.17	0.245
Femur length				
Mean ± SD	55.18 ± 3.98	56.9 ± 3.49	1.69	0.093

Table 4. Mode of delivery distribution among the studied groups.

	No-macrosomia (N = 82)	Macrosomia (N = 18)	χ <sup>2</sup>	P
Cesarean section	48 (58.5%)	13 (72.2%)	1.16	0.282
Vaginal delivery	34 (41.5%)	5 (27.8%)		

There was a significant difference between the groups regarding birth weight (Table 2).

Regarding biparietal diameter, belly circumference, and femur length, there was no significant difference between the two study groups (Table 3).

There was no statistically significant difference between the groups in term of mode of delivery (Table 4).

There was a significant difference between the three studied groups regarding HbA1c and Umbilical Cord

Table 5. Glycated hemoglobin and the umbilical cord area of the two studied groups.

	No-macrosomia (N = 82)	Macrosomia (N = 18)	t	P
HbA1c (%) (mean ± SD)				
27–28 weeks	6.42 ± 0.418	6.15 ± 0.322	2.57	0.012
36–37 weeks	5.93 ± 0.493	6.58 ± 0.426	5.18	<0.001
UCT (mm <sup>2</sup> ) (mean ± SD)				
27–28 weeks	202.24 ± 2.68	211.8 ± 3.19	13	<0.001
36–37 weeks	217.12 ± 4.27	235.23 ± 3.58	16	<0.001

HbA1c, glycated hemoglobin.

Table 6. Correlation of birth weight with glycated hemoglobin and UCT among macrosomia group.

	Birth weight	
Macrosomia group	r	P
UCT	0.654	<0.001
HbA1c	0.281	0.092

HbA1c, glycated hemoglobin.

Thickness (UCT) at 27–28 weeks and 36–37 weeks of pregnancy (Table 5).

Within the macrosomia group, there was a substantial positive connection between birth weight and UCT (Tables 6 and 7).

UCT achieved significance for predicting fetal macrosomia with sensitivity of 91.8%, specificity of 93.6%, positive predictive value of 89%, and negative predictive value of 96%. However, HbA1c did not achieve significance for predicting fetal macrosomia.

#### 4. Discussion

The umbilical cord is in charge of the flow of mother and fetal blood. In the first and early second trimesters of pregnancy, there was a significant difference between fetuses with umbilical cord thickness below the fifth percentile (lean cord) and those with thickness above the fifth percentile (non-lean cord) in terms of average gestation age, mode of delivery, birth weight, and adverse perinatal outcomes.<sup>1</sup>

GDM is linked to a number of unfavorable pregnancy outcomes, including macrosomia and cesarean section delivery. A greater prevalence of surgical births, postpartum hemorrhages, birth injuries during natural delivery, and newborn hypoglycemia are all linked to fetal macrosomia. Only 40% of mothers who gave birth to macrosomic infants had known maternal risk factors identified. It has been proposed that one of the potential risk factors for obesity is macrosomia.<sup>14</sup>

Table 7. Validity of glycated hemoglobin and UCT.

Variables	AUC	SE	Significance	95% CI	Sensitivity	Specificity	PPV	NPV
UCT	0.928	0.034	<0.001*	0.861–0.995	91.8%	93.6%	89.2%	96%
HbA1c	0.615	0.074	0.126	0.471–0.761	58.1%	87.2%	33%	89%

AUC, area under the curve; CI, confidence interval; HbA1c, glycated hemoglobin; NPV, negative predictive value; PPV, positive predictive value.

There was a significant difference between the groups (no-macrosomia and macrosomia) regarding maternal BMI and gravidity.

Our results were in agreement with a study of Fayeze Mohamed Fathi et al.<sup>15</sup> as they reported that there was a significant difference between their investigated groups (no-macrosomia and macrosomia) regarding maternal BMI and gravidity.

Similarly, Ismail et al.<sup>16</sup> revealed that both groups varied considerably in terms of gravidity, where in group 1 (macrosomia), women had a mean of  $3.6 \pm 1.3$  compared with  $2.5 \pm 1.2$  for group 2 (non-macrosomia), although neither group varied in terms of parity. Five (33%) women of the 15 patients who delivered macrosomic fetuses were obese, eight (54%) were overweight, and two (13%) were normal, with a significant difference between the two groups.<sup>16</sup>

The present study showed that there was no significant difference between the groups in term of mode of delivery.

In contrary to our results, a study by Ismail et al.<sup>16</sup> reported that there was significant difference between their studied groups regarding mode of delivery.

The current study showed that there was a significant difference between the groups regarding birth weight. There was no significant difference among the three studied groups regarding biparietal diameter, abdominal circumference, and femur length.<sup>16</sup>

In accordance with our results, a study by Fayeze Mohamed Fathi et al.<sup>15</sup> reported that both groups' delivered fetuses had different birth weights; group 1 (macrosomic) had a mean birth weight of  $3924.9 \pm 418.3$  gm for 15 delivered fetuses, which was significantly higher ( $P < 0.0001$ ) than that in group 2 ( $3332.3 \pm 296.1$  g) for 85 delivered fetuses.<sup>15</sup>

Furthermore, Ismail et al.<sup>16</sup> revealed that there was a significant difference between the groups regarding birth weight.<sup>16</sup>

In our study, there was a significant difference between the two investigated groups regarding HbA1c and UCT at 27–28 weeks and 36–37 weeks of gestation.

Our results were in accordance with Ismail et al.,<sup>16</sup> who assessed the association between umbilical cord components, HbA1c, and baby macrosomia at 27–28 weeks of gestation. Fetuses with and without macrosomy were compared. The median umbilical

cord area of macrosomic fetuses was  $213.1 \pm 2.8$  mm<sup>2</sup> compared with  $204.2 \pm 2.1$  mm<sup>2</sup> for the nonmacrosomic group, and there were statistically distinct results for Wharton's jelly for each group. At this gestational stage, there were no statistically significant differences in the cord diameter, umbilical artery, or vein area measurements between groups. When compared with the levels obtained at 27–28 gestational weeks, neither group's HbA1c levels differed substantially. Although the macrosomic group had a greater HbA1c than group 2 at 36–37 weeks of gestation ( $6.4 \pm 0.3$  vs.  $5.8 \pm 0.4\%$ , respectively), this difference was extremely statistically substantial.<sup>16</sup>

Furthermore, Fayeze Mohamed Fathi and colleagues, demonstrated that HbA1c and UCT were substantially different between the two study groups at 27–28 weeks and 36–37 weeks of pregnancy, respectively.

In our research, macrosomic babies were delivered by six (14.6%) of 41 women with GDM or pre-GDM, compared with five (10%) of 50 individuals who were not diabetic. The diabetic group's relative risk of macrosomia was found to be 1.5 times greater.

Naylor et al.<sup>17</sup> reported that in individuals with GDM, the frequency of macrosomia was 16–29% compared with 10% in the general population. In those with diabetes, the relative risk of macrosomia might be 1.5 to three times greater. Additionally, Naylor et al.,<sup>17</sup> reported that moms with GDM had a 30% cesarean section rate compared with a 20% rate in the control group.

Our results showed that among the macrosomia group, there was a substantial positive connection between birth weight and UCT. Using ROC curve, UCT achieved significance for predicting fetal macrosomia with sensitivity of 91.8% and specificity of 93.6%, with positive predictive value of 89% and negative predictive value of 96%.

However, a study by Ismail et al.<sup>16</sup> reported that the umbilical cord area was shown to be more accurate in predicting fetal macrosomia at the correct criteria when the ROC curves of both the umbilical cord area and the HbA1c were compared. It was determined that the difference in predicting effectiveness between the two factors was statistically very substantial. The HbA1c did not significantly or



strongly correlate with birth weight, nor did measurements taken at 27–28 weeks of gestation or 36–37 weeks of gestation.<sup>16</sup>

In the study by Fayez Mohamed Fathi et al.,<sup>15</sup> when the umbilical cord area and birth weight in group 1 (macrosomic fetuses) were correlated, it was discovered that there was a strong, dependent, and positive (direct) connection between the two parameters, whether measurement at 27–28 weeks or measurement at 36–37 weeks of gestation ( $r = 0.7340$  and  $0.7483$ , respectively). Additionally, it was discovered that these connections were statistically very substantial ( $P = 0.0002$  and  $0.0001$ , respectively). HbA1c measurements at 27–28 weeks of gestation and 36–37 weeks of gestation did not significantly or strongly correlate with birth weight.<sup>15</sup>

#### 4.1. Conclusion

Severe obstetric problems, including shoulder dystocia, delivery hypoxia, and postpartum hemorrhage, are caused by macrosomia. There are times when it is difficult to foresee shoulder dystocia. The group most at risk for these issues may be found, however, using macrosomia prediction. There have been documented studies using sonographic measurement for predicting fetal macrosomia. Fetal macrosomia is well predicted by the thickness of the umbilical cord and the fetal fat layer.

The clinical risk factors must be taken into account in addition to the ultrasonographic data when determining the risk of macrosomia. Further research is required to assess the therapeutic utility of using these soft tissue measures in fetal weight estimate algorithms.

#### Conflict of interest

There are no conflicts of interest.

#### References

1. Marino L, Castaldi MA, Rosamilio R, et al. Mesenchymal stem cells from the Wharton's jelly of the human umbilical cord: biological properties and therapeutic potential. *Int J Stem Cells*. 2019;12:218–226.
2. Mohammadbeigi A, Farhadifar F, Zadeh NS, Mohammadsalehi N, Rezaiee M, Aghaei M. Fetal macrosomia: risk factors, maternal, and perinatal outcome. *Ann Med Health Sci Res*. 2013;3:546–550.
3. Binbir B, Yeniel AO, Ergenoglu AM, Kazandi M, Akercan F, Sagol S. The role of umbilical cord thickness and HbA1c levels for the prediction of fetal macrosomia in patients with gestational diabetes mellitus. *Arch Gynecol Obstet*. 2012;285:635–639.
4. Alfadhli EM. Maternal obesity influences birth weight more than gestational diabetes. *BMC Pregnancy Childbirth*. 2021;21:1–7.
5. Picón-César MJ, Molina-Vega M, Suárez-Arana M, et al. Metformin for gestational diabetes study: metformin vs insulin in gestational diabetes: glycemic control and obstetrical and perinatal outcomes: randomized prospective trial. *Am J Obstet Gynecol*. 2021;225:517e1.
6. Beta J, Khan N, Khalil A, Fiolna M, Ramadan G, Akolekar R. Maternal and neonatal complications of fetal macrosomia: systematic review and meta-analysis. *Ultrasound Obstet Gynecol Off J Int Soc Ultrasound Obstet Gynecol*. 2019;54:308–318.
7. Campbell S. Fetal macrosomia: a problem in need of a policy. *Ultrasound Obstet Gynecol*. 2014 Jan;43(1):3–10. <https://doi.org/10.1002/uog.13268>. PMID: 24395685.
8. Auger N, Park AL, Zoungana H, Sing MF, Lo E, Luo Z. Widening inequality in extreme macrosomia between Indigenous and non-Indigenous populations of Québec, Canada. *Aust N Z J Publ Health*. 2013;37:58–62.
9. Sprehe MR, Barahmani N, Cao Y, et al. Comparison of birth weight corrected for gestational age and birth weight alone in prediction of development of childhood leukemia and central nervous system tumors. *Pediatr Blood Cancer*. 2010;54:242–249.
10. Shan X, Chen F, Wang W, et al. Secular trends of low birth-weight and macrosomia and related maternal factors in Beijing, China: a longitudinal trend analysis. *BMC Pregnancy Childbirth*. 2014;14:1–9.
11. Dyer JS, Rosenfeld CR, Rice J, Rice M, Hardin DS. Insulin resistance in Hispanic large-for-gestational-age neonates at birth. *J Clin Endocrinol Metab*. 2007;92:3836–3843.
12. Masuch A, Friedrich N, Roth J, Nauck M, Müller UA, Petersmann A. Preventing misdiagnosis of diabetes in the elderly: age-dependent HbA1c reference intervals derived from two population-based study cohorts. *BMC Endocr Disord*. 2019;19:20.
13. Suchitra MR, Jaiganesh K, Parthasarathy S. Diabetic profile-screening of HBA1C-a random community assessment. *J Clin Diagn Res*. 2013;7:2200.
14. Katon J, Reiber G, Williams MA, Yanez D, Miller E. Antenatal haemoglobin A1c and risk of large-for-gestational-age infants in a multi-ethnic cohort of women with gestational diabetes. *Paediatr Perinat Epidemiol*. 2012;26:208–217.
15. Fayez Mohamed Fathi M, Mohammed Moustafa T, Ramadan Alsawy Rady I. The role of umbilical cord thickness and glycated hemoglobin (HBA1C) levels for prediction of fetal macrosomia in patients with gestational diabetes mellitus. *Al-Azhar Med J*. 2020;49:1741–1752.
16. Ismail OEA, Rady IRA, Fathi MFM. The role of umbilical cord thickness and glycated hemoglobin (HbA1c) levels for prediction of fetal macrosomia in patients with gestational diabetes mellitus. *Egypt J Hosp Med*. 2019;77:4906–4912.
17. Naylor CD, Sermer M, Chen E, Sykora K. Cesarean delivery in relation to birth weight and gestational glucose tolerance: pathophysiology or practice style? *JAMA*. 1996;275:1165–1170.