

Comparison of the effects of gestational weight gain on pregnancy outcomes between non-diabetic and diabetic women

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Objective

Appropriate gestational weight gain (GWG) is important in diabetic women. Current GWG guideline is for US general population, but not specific for diabetic women. We compared the effect of GWG on perinatal outcomes between diabetic and non-diabetic women.

Methods

Fifty two hundred and twelve women who delivered live singleton infants at Korea University Medical Center from January 2009 to December 2013 were included. One hundred twenty-nine overt diabetes women and 322 gestational diabetes women were categorized as diabetic women, and the others were categorized as none-diabetic women. 5,212 women were categorized by GWG (low 1,081; adequate 2,102; or high 2,029; according to the 2009 Institute of Medicine guidelines), and each of the 3 GWG groups was categorized into 2 groups; diabetic or non-diabetic women. And then, we compared perinatal outcomes between diabetic and non-diabetic groups.

Results

In each 3 GWG groups, primary cesarean section delivery, high birth weight, and large for gestational age rates were significantly higher in diabetic women than non-diabetic women. Only in adequate GWG group, preterm birth rate was significantly higher in diabetic women than non-diabetic women.

Conclusion

Our study shows that diabetic women had higher rates of adverse perinatal outcomes than non-diabetic women, although they achieved same GWG. It suggests that current GWG guideline may not be adequate for diabetic women, and that diabetic women may need more strict GWG control than normal population.

Keywords: Diabetes; Gestational diabetes; Gestational weight gain

Introduction

It is known that maternal pre-pregnancy body mass index (BMI), rate of weight gain, and total gestational weight gain (GWG) are closely associated with maternal and neonatal outcomes [1-3]. High GWG increases maternal risk of cesarean section (CS) delivery, pregnancy-related hypertension, and gestational diabetes (GDM) [2]. GWG also impacts the neonate, as high GWG increases risk of high birth weight (HBW), and, as low GWG increases risk of low birth weight (LBW), and preterm birth.

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In 2009, the Institute of Medicine (IOM) revised their GWG guidelines, and these guidelines have been used as a current recommendation of GWG [3]. IOM 2009 guidelines are for the general obstetrical population in the United States, but application on subpopulations of pregnant women has not been evaluated extensively, especially diabetic women.

Diabetes during pregnancy, GDM or pregestational diabetes (overt diabetes), is associated with adverse obstetrical outcomes [4-13]. Diabetes increases the risk of preeclampsia, CS delivery, congenital malformation, spontaneous abortion, macrosomia, polyhydramnios, and preterm birth. Adequate GWG is important for diabetic women to prevent adverse outcomes [14,15], which is one of the goals of medical nutritional therapy and exercise.

Maternal diabetes and obesity are independently associated with adverse pregnancy outcomes, and their combination has a greater impact than either disorder alone [8,16]. IOM 2009 GWG guidelines, however, are for general obstetrical population, but applicability of these GWG targets to diabetic women is controversial [3].

To examine whether current GWG guidelines are sufficient for diabetic women, we compared the effect of GWG on perinatal outcomes between diabetic and non-diabetic women.

Materials and methods

1. Study population and study design

We conducted a retrospective cohort study of 5,840 women who delivered a live singleton infant at the Korea University Medical Center from January 2009 to December 2013. Patients who did not know their pre-pregnancy weight, who did not take glucose tolerance test, or who did not receive proper antenatal care were excluded. After exclusion, data of total 5,212 women were analyzed. Fifty two hundred and twelve women consisted of 322 GDM patients, 129 overt diabetes patients, and 4,761 non-diabetic patients. GDM and overt diabetes patients were categorized in "diabetic women" group, and the others were categorized in "non-diabetic women" group. All women were categorized into 3 GWG groups according to IOM 2009 guidelines; low GWG, adequate GWG, and high GWG.

Each of the 3 GWG groups was categorized into 2 subgroups; diabetic or non-diabetic women. And then, we compared perinatal outcomes between the 2 subgroups. In 1,081

low GWG women, 126 diabetic women were compared with 955 non-diabetic women. In 2,102 adequate GWG women, 137 diabetic women were compared with 1,965 non-diabetic women. In 2,029 high GWG women, 188 diabetic women were compared with 1,841 non-diabetic women.

GDM was defined as diabetes which is diagnosed in pregnancy by two step oral glucose tolerance test (OGTT) (50 g OGTT for screening test, and 100 g OGTT for diagnostic test). GDM was diagnosed if two or more plasma glucose levels met or exceeded the following thresholds: fasting glucose concentration of 95 mg/dL, 1-hour glucose concentration of 180 mg/dL, 2-hour glucose concentration of 155 mg/dL, or 3-hour glucose concentration of 140 mg/dL. Overt diabetes was defined as a history of diabetes. BMI was calculated by dividing weight in kilograms by the square of height in meters. Pre-pregnancy BMI was categorized as underweight (BMI <18.5 kg/m²), normal weight (18.5 ≤ BMI <25 kg/m²), overweight (25 ≤ BMI <30 kg/m²), or obese (BMI ≥30 kg/m²) by World Health Organization classification. Adequacy of GWG was defined according to the maternal pre-pregnancy BMI status and the IOM 2009 GWG recommendations; 12.5 to 18 kg total weight gain, or 0.44 to 0.58 kg/wk rate of weight gain in underweight women, 11.5 to 16 kg total weight gain, or 0.35 to 0.50 kg/wk rate of weight gain in normal weight women, 7 to 11.5 kg total weight gain, or 0.23 to 0.33 kg/wk rate of weight gain in overweight women, and 5 to 9 kg total weight gain, or 0.17 to 0.27 kg/wk rate of weight gain in obese women. We used the guidelines of weight gain rate (kg/wk) to correct the effect of gestational age (GA) to GWG.

Perinatal outcomes assessed were primary CS delivery, preterm labor, preterm premature rupture of membranes (PPROM), and preterm birth as the maternal outcomes, and LBW, HBW, small for gestational age (SGA), large for gestational age (LGA), low Apgar score, and admission to neonatal intensive care unit (NICU) as the neonatal outcomes. Preterm labor was defined as presence of uterine contractions of sufficient frequency and intensity to enable progressive cervical effacement and dilation prior to term gestation. PPRM was defined as membrane rupture prior to term gestation. LBW was defined as birth weight less than 2,500 g, and HBW was defined as birth weight more than 4,000 g. SGA was defined as birth weight below the 10th percentile for GA, and LGA was defined as birth weight greater than the 90th percentile for GA. GA was estimated by ultrasound measurement of crown-rump length in first trimester. When we diagnosed

SGA or LGA, GA was rounded off to the nearest weeks to use birth weight percentile table made by Doubilet et al. [17]. Low Apgar score was defined as below 7 in 1-minute or 5-minutes Apgar score.

2. Statistical analysis

Data are expressed as the mean±standard deviation for continuous variables, and data for the categorical variables are expressed as the number and the percentage of patients. The chi-square test was used for the categorical variables of the two groups, and the Mantel-Haenszel chi-square test was used to adjust potential confounding factors. The independent samples *t*-test was used to compare continuous variables between the two groups. A *P*-value <0.05 was considered statistically significant.

Results

In the study population (n=5,212), diabetic women were 451, and non-diabetic women were 4,761 (Table 1). Mean age was older in the diabetic group than the non-diabetic group (33.6±4.3 vs. 32.0±4.5, *P*<0.001). The diabetic group have higher rate of multiparous pregnancy (50.8 vs. 45.9, *P*=0.049), and previous history of CS (31.7 vs. 18.3, *P*<0.001). Pre-pregnancy height was not significantly different between the two groups, but pre-pregnancy weight and BMI were higher in the diabetic group. GWG was higher in the non-diabetic group, but weight at delivery was higher in the diabetic group. Using IOM targets as reference, rate of adequate GWG was similar between the two groups, but fetal birth weight was higher in the diabetic group than the non-diabetic group (3.01±0.77 vs. 2.90±0.72, *P*=0.001). Mean GA was shorter in the diabetic group (259.7±20.2 vs. 262.9±22.2, *P*=0.002).

Table 1. Characteristics between non-diabetic and diabetic women (n=5,212)

Variable	Non-diabetes (n=4,761)	Diabetes (n=451)	P-value
Age (yr)	32.0±4.5	33.6±4.3	<0.001
GA at delivery (day)	262.9±22.2	259.7±20.2	0.002
Parity (multiparous)	2,187 (45.9)	229 (50.8)	0.049
Previous CS	870 (18.3)	143 (31.7)	<0.001
Pre-pregnancy height (cm)	160.5±6.6	162.9±65.5	0.436
Pre-pregnancy weight (kg)	55.8±9.7	64.4±14.4	<0.001
Pre-pregnancy BMI (kg/m ²)	21.7±5.3	25.3±6.3	<0.001
Underweight	683 (14.3)	23 (5.1)	<0.001
Normal weight	3,425 (71.9)	225 (49.9)	
Overweight	510 (10.7)	120 (26.6)	
Obese	143 (3.0)	83 (18.4)	
Hypertensive disease			<0.001
Normotensive	4,375 (91.9)	364 (80.7)	
Chronic hypertension	96 (2.0)	39 (8.6)	
Preeclampsia	277 (5.8)	47 (10.4)	
Eclampsia	13 (0.3)	1 (0.2)	
Weight at delivery (kg)	68.5±18.5	75.4±15.4	<0.001
Weight change (kg)	12.8±15.5	10.9±6.4	0.013
Inadequate GWG	955 (20.1)	126 (27.9)	0.189
Adequate GWG	1,965 (41.3)	137 (30.4)	
Excessive GWG	1,841 (38.7)	188 (41.7)	
Birth weight (kg)	2.90±0.72	3.01±0.77	0.001

Values are presented as mean±standard deviation or number (%).

GA, gestational age; CS, cesarean section; BMI, body mass index; GWG, gestational weight gain.

Table 2. Perinatal outcomes among GWG group in diabetic women (n=451)

Variable	Low GWG (n=126)	Adequate GWG (n=137)	High GWG (n=188)	P-value
Primary cesarean section	37 (40.2)	45 (48.4)	72 (58.5)	0.027
Preterm labor	18 (14.3)	9 (6.6)	14 (7.4)	0.055
PPROM	7 (5.6)	11 (8.0)	14 (7.4)	0.715
Preterm birth	32 (25.4)	47 (34.3)	49 (26.1)	0.181
Low birth weight	32 (25.4)	34 (24.8)	27 (14.4)	0.021
High birth weight	4 (3.2)	9 (6.6)	19 (10.1)	0.061
Small for gestational age	15 (12.0)	15 (10.9)	9 (4.8)	0.046
Large for gestational age	9 (7.2)	22 (16.1)	41 (21.9)	0.002
Low 1 min Apgar	20 (15.9)	24 (17.5)	33 (17.6)	0.915
Low 5 min Apgar	7 (5.6)	6 (4.4)	6 (3.2)	0.589
NICU admission	35 (27.8)	55 (40.1)	58 (30.9)	0.077

Values are presented as number (%).

GWG, gestational weight gain; PPRM, preterm premature rupture of membranes; NICU, neonatal intensive care unit.

We categorized the study population into three GWG groups; low, adequate, and high GWG by IOM 2009. First, we compared perinatal outcomes among GWG groups in diabetic women (Table 2). The results showed tendency that higher GWG group had higher rates of primary CS delivery and LGA, and that lower GWG group had higher rates of LBW and SGA. However, the other outcomes were not significantly different among GWG groups.

Then perinatal outcomes were compared between diabetic and non-diabetic women in each GWG groups, respectively (Table 3). We found that primary CS delivery rate was higher in diabetic women, even though both diabetic and non-diabetic women belonged to same GWG group; 40.2% vs. 29.4% in low GWG group, 48.4% vs. 33.9% in adequate GWG group, 58.5% vs. 43.7% in high GWG group, respectively. The results were similar after adjustment of maternal age and parity. The other perinatal outcomes were also compared. In each 3 GWG groups, rates of HBW and LGA were significantly higher in diabetic women than non-diabetic women. In the low GWG group, NICU admission rate was higher in non-diabetic women. In the adequate GWG group, however, NICU admission rate was higher in diabetic women. In the high GWG group, there was no significant difference between non-diabetic and diabetic women. In the adequate GWG group, PPRM, preterm birth, and low 1 minute Apgar score rates were higher in diabetic women. In the high GWG group, PPRM rate was higher in diabetic women, but LBW and SGA rates were higher in non-diabetic women. The

other outcomes were not significantly different between non-diabetic and diabetic women. The results were similar after adjustment of maternal age and parity.

To adjust for pre-pregnancy BMI as potential confounding factor, we also compared perinatal outcomes between non-diabetic and diabetic women in normal BMI group (Table 4). In the low GWG group, preterm birth and NICU admission rates were higher in non-diabetic women. In the adequate GWG group, primary CS delivery, preterm birth, HBW, and LGA rates were higher in diabetic women. In the high GWG group, primary CS delivery, preterm labor, and LGA rates were higher in diabetic women. The results were similar after adjustment of maternal age and parity.

Discussion

Currently IOM guidelines are often used as ideal GWG, but there are no weight gain guidelines specific for diabetic women [18-21]. So, currently, the GWG goal of diabetic women was same as non-diabetic women. We found that rates of some adverse perinatal outcomes were higher in high GWG or low GWG groups (primary CS delivery, LBW, SGA, and LGA). These results suggest that adequate GWG is not only important for non-diabetic women, but also for diabetic women. But, we found that primary CS delivery, HBW, and LGA rates were significantly higher in diabetic women than non-diabetic women, although diabetic women achieved their

Table 3. Perinatal outcomes between non-diabetic and diabetic women (n=5,212)

Variable	Low GWG (n=1,081)		Adequate GWG (n=2,102)		High GWG (n=2,029)				
	Non diabetes (n=955)	Diabetes (n=126)	P-value	Non diabetes (n=1,965)	Diabetes (n=137)	P-value	Non diabetes (n=1841)	Diabetes (n=188)	P-value
Primary CS	230 (29.4)	37 (40.2)	0.033	551 (33.9)	45 (48.4)	0.004	647 (43.7)	72 (58.5)	0.001
Preterm labor	162 (17.0)	18 (14.3)	0.448	212 (10.8)	9 (6.6)	0.120	183 (9.9)	14 (7.4)	0.271
PPROM	51 (5.3)	7 (5.6)	0.920	76 (3.9)	11 (8.0)	0.018	75 (4.1)	14 (7.4)	0.031
Preterm birth	324 (33.9)	32 (25.4)	0.056	473 (24.1)	47 (34.3)	0.007	513 (27.9)	49 (26.1)	0.599
LBW	300 (31.4)	32 (25.4)	0.169	443 (22.5)	34 (24.8)	0.539	434 (23.6)	27 (14.4)	0.004
HBW	8 (0.8)	4 (3.2)	0.041	45 (2.3)	9 (6.6)	0.007	74 (4.0)	19 (10.1)	<0.001
SGA	130 (13.6)	15 (12.0)	0.613	257 (13.1)	15 (10.9)	0.470	187 (10.2)	9 (4.8)	0.019
LGA	24 (2.5)	9 (7.2)	0.010	62 (3.2)	22 (16.1)	<0.001	120 (6.5)	41 (21.9)	<0.001
Low 1 min Apgar	139 (14.6)	20 (15.9)	0.695	228 (11.6)	24 (17.5)	0.039	261 (14.2)	33 (17.6)	0.210
Low 5 min Apgar	44 (4.6)	7 (5.6)	0.637	69 (3.5)	6 (4.4)	0.630	80 (4.3)	6 (3.2)	0.454
NICU admission	615 (33.4)	58 (30.9)	0.012	602 (30.6)	55 (40.1)	0.020	615 (33.4)	58 (30.9)	0.475

Values are presented as number (%).

GWG, gestational weight gain; CS, cesarean section; PPROM, preterm premature rupture of membranes; LBW, low birth weight; HBW, high birth weight; SGA, small for gestational age; LGA, large for gestational age; NICU, neonatal intensive care unit.

Table 4. Perinatal outcomes between non-diabetic and diabetic women in normal BMI group (n=3,650)

Variable	Low GWG (n=672)		Adequate GWG (n=1,581)		High GWG (n=1,397)				
	Non diabetes (n=605)	Diabetes (n=67)	P-value	Non diabetes (n=1,493)	Diabetes (n=88)	P-value	Non diabetes (n=1,327)	Diabetes (n=70)	P-value
Primary CS	150 (30.5)	20 (37.0)	0.329	418 (33.8)	30 (50.0)	0.010	465 (43.1)	30 (61.2)	0.012
Preterm labor	97 (16.0)	8 (11.9)	0.381	157 (10.5)	6 (6.8)	0.268	129 (9.7)	1 (1.4)	0.020
PPROM	28 (4.6)	1 (1.5)	0.346	56 (3.8)	7 (8.0)	0.082	52 (3.9)	5 (7.1)	0.203
Preterm birth	204 (33.7)	14 (20.9)	0.033	349 (23.4)	31 (35.2)	0.011	353 (26.6)	15 (21.4)	0.338
LBW	183 (30.2)	16 (23.9)	0.279	321 (21.5)	25 (28.4)	0.128	300 (22.6)	10 (14.3)	0.102
HBW	6 (1.0)	1 (1.5)	0.522	31 (2.1)	6 (6.8)	0.014	45 (3.4)	5 (7.1)	0.101
SGA	75 (12.4)	9 (13.4)	0.816	190 (12.7)	10 (11.4)	0.707	134 (10.1)	5 (7.1)	0.419
LGA	15 (2.5)	3 (4.5)	0.411	43 (2.9)	12 (13.6)	<0.001	71 (5.4)	12 (17.1)	0.001
Low 1 min Apgar	83 (13.7)	8 (11.9)	0.686	166 (11.1)	15 (17.0)	0.090	175 (13.2)	8 (11.4)	0.671
Low 5 min Apgar	29 (4.8)	1 (1.5)	0.348	52 (3.5)	6 (6.8)	0.133	49 (3.7)	3 (4.3)	0.742
NICU admission	234 (38.7)	16 (23.9)	0.017	440 (29.5)	34 (38.6)	0.068	426 (32.1)	16 (22.9)	0.104

Values are presented as number (%).

GWG, gestational weight gain; CS, cesarean section; PPROM, preterm premature rupture of membranes; LBW, low birth weight; HBW, high birth weight; SGA, small for gestational age; LGA, large for gestational age; NICU, neonatal intensive care unit.

recommended (adequate) GWG. We also found similar results in pre-pregnancy normal BMI women. These results suggest that diabetic condition is independently associated with increased risk of adverse perinatal outcomes, and that current IOM GWG goal is insufficient for diabetic women.

Although it was not constant trend, some other adverse perinatal outcomes were increased in diabetic women than non-diabetic women. But, in low GWG group, NICU admission rate was higher in non-diabetic women than diabetic women. This result may be associated with fetal birth weight. Previous study showed that singletons born to women with low total GWG have higher risks of preterm birth and LBW [22]. In low GWG group, mean fetal birth weight of diabetic women was 2.84 kg, and non-diabetic women was 2.72 kg, but it was not significantly different. Mean fetal birth weight of total low GWG group (diabetic and non-diabetic women) was 2.73 kg, and it was significantly lower than high or adequate GWG groups (2.97 and 2.92 kg). We used multivariate logistic regression to analyze risk factors of NICU admission in low GWG group. Multivariate binary logistic regression analysis showed that preterm birth ($P<0.001$), LBW ($P<0.001$), and maternal diabetes ($P=0.049$) were significantly associated with increased risk of NICU admission, but SGA, HBW, and LGA were not significantly associated. Although it was not significantly different, mean fetal birth weight was lower and LBW rate was higher in non-diabetic women. These may be associated with increased NICU admission rate of non-diabetic women in low GWG group. These results also imply that specific weight gain guideline may be needed for diabetic women.

In addition, we evaluate the risk of primary CS along the whole range of maternal weight gain to establish optimal GWG for normal weight BMI diabetic women by using similar method with previous study [23]. The number of individuals in each weight gain class was compared with the number of individuals in all other weight gain classes, and optimal weight gain range was based on significant risk estimates below 1 for each maternal weight gain. According to the results, possible optimal weight gain range in normal weight BMI diabetic women was 9 to 13 kg, but estimates of 95% confidence intervals were not statistically significant. The other BMI groups of this study could not be evaluated because study population number was too small to do this analysis, and it was our limitation of the study. Currently IOM recommended GWG in normal weight BMI group is 11.5 to 16.0 kg. Although it

was not statistically significant meaningful, this possible optimal weight gain range (9 to 13 kg) may suggest that diabetic women need stricter GWG control than normal population in spite of same BMI. Thus, further large population studies are needed to evaluate and estimate adequate GWG goals in diabetic women.

Limitation of this study is that type of therapy of diabetes and degree of diabetes control may be associated with perinatal outcomes. However, the purpose of this study was to suggest the necessity of specific GWG guidelines for diabetic women. Considering the impact of specific situation may be needed, but that may also make it unnecessarily complicated to make guidelines for diabetic women.

In summary, our study shows that diabetic women had higher primary CS delivery, HBW and LGA rate than non-diabetic women, although they achieved same GWG. It suggests that current GWG guideline may be not adequate for diabetic women, and that diabetic women may need stricter GWG control than general population.

Conflict of interest

No potential conflict of interest relevant to this article was reported.

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