# **Original Article**

# Gestational Diabetes and Risk of Developing Postpartum Type 2 Diabetes: How to Improve Follow-up?

#### Abstract

**Background:** Gestational diabetes mellitus (GDM) is defined by World Health Organization as glucose intolerance diagnosed for the first time during pregnancy; GDM affects 7% of pregnancies. Women with earlier GDM have higher risk to develop type 2 diabetes (T2D). The aim of our study was to evaluate the outcomes of GDM and to assess the impact of recalling patients in the postpartum stage by phone, the target was to assess T2D or impaired glucose tolerance in women with a history of GDM. **Methods:** This prospective study included 200 patients with GDM that have received education sessions regarding the major interest of screen T2D using 75 g of oral glucose tolerance testing in the 3<sup>rd</sup> month after birth. **Results:** Only 22.5% (n = 45) women spontaneously complied to assess T2D. About 15% have had developed T2D and 28% prediabetes. Risk factors of T2D onset were younger gestational age at the occurrence GD, higher fasting blood glucose, and frequent use of insulin. **Conclusions:** Women with GD history demonstrated high risk of developing T2D. Simple changes of lifestyle were shown to be an efficient prevention protocol. Despite therapeutical education, few women spontaneously complied with T2D screening. The telephone reminders could improve the screening observance therefore patient's outcome.

Keywords: Diabetes mellitus, follow-up, gestational diabetes

## Introduction

According World Health to Organization (WHO), gestational diabetes (GD) is a disorder of glucose tolerance that is diagnosed for the first time during the pregnancy.<sup>[1]</sup> It is a public health problem since diagnosed in 7% of pregnancies with shorter and longer term impact on maternal and fetal health.<sup>[1,2]</sup> Former pregnant women with GD have higher risk to developing a diabetes mellitus, especially type 2 diabetes (T2D), metabolic syndrome, and obesity.<sup>[3-5]</sup> Appropriate care has significantly reduced the maternal and fetal morbidity.<sup>[5]</sup> The compliance to screening was demonstrated to be low; fewer patients adjust their lifestyle to prevent later occurrence of T2D.[3]

The mail reminders have allowed increasing T2D screening compliance in women with earlier GD; however, the impact of telephone reminder was never assessed.<sup>[6]</sup>

The goal of our study was to assess the prospect of GD, to highlight the predictive factors of T2D occurrence, and to evaluate the impact of phone reminder on the

compliance to T2D screening in postpartum in women with earlier GD.

## Methods

This prospective was carried along 5 years between January of 2010 and 2015. The study included 200 patients followed for GD in the Department of Endocrinology and Diabetology of University hospital of Fez, Morocco.

According to diagnosis criteria of "Collège national des gynécologues et obstétriciens français" established in 2010; any FBG higher than 0.92 g/L or oral glucose tolerance test (OGTT) achieved between 24<sup>th</sup> and 28<sup>th</sup> week of amenorrhea would allow establishing the diagnosis of GD.

In our study, the diagnosis of GD was based on any of both assessing fasting blood glucose (FBG) in the first trimester of pregnancy. All patients have had benefited of therapeutical education, this emphasized the importance of follow-up of diabetes using OGTT with 75 g of carbohydrates after the thirds month of the postpartum. The statistical analysis was realized by SPSS Inc. Released 2009. PASW Statistics for Windows, Version 18.0. Chicago, USA.

How to cite this article: El Ouahabi H, Doubi S, Boujraf S, Ajdi F. Gestational diabetes and risk of developing postpartum type 2 diabetes: How to improve follow-up? Int J Prev Med 2019;10:51.

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#### Results

The average age of patients was  $33.5 \pm 5.5$  years old, and their middleweight before pregnancy was  $70 \pm 9$  kg. The average gestational age of GD assessment was 19.36 weeks of amenorrhea. GD screening in our study was focused on risk factors in 80% of cases and systematic in 20% of cases. These risk factors consisted of being older than 30 years-old found in 71% of patients, presence of maternal heredity of T2D, history of polymicrocyst in ovaries, and history of high arterial blood pressure, *in utero* fetal death, former GD as well as of macrosomia [Figure 1].

The postpartum consultation was spontaneously achieved in 22.5% (n = 45) of patients. About 77.5% of patients were followed up by phone reminder and 51.5% of called responded positively; while 26% (n = 52 patients) were not reachable despite multiple recall [Table 1].

The "self-dismiss" of follow-up was mainly due to being busy with baby care, negligence, and fear of discovering diabetes, despite the follow-up of patients lasted 6 months.

Screening protocol consisted of achieving OGTT in 67.6% of patients, followed by glycosylated hemoglobin test (HbA1C) achieved in 25.7% of cases and FBG realized in 6.7% of cases. The upcoming of our GD patients is summarized in Figure 2.

The risk factors to develop T2D after delivery were marked by the lower gestational age at the GD diagnosis associated with higher FBG, requiring insulin treatment in 86% of cases. The obesity was shown in 9% of cases.

Neonates were females in 58% of cases and benefited of an exclusive breastfeeding in 83% of cases along an average duration of 6 months.

The postpartum contraception was established in 52.5% of cases with an intrauterine contraceptive device in 13.5%, estrogen-progesterone microdose in 27%, estrogen-progesterone minidose in 2%, and mechanical contraception in 10%.

In postpartum, patients have had benefited of care mainly based on rigorous healthy lifestyle with particular rejection of rapid sugars, a balanced diet, and a regular physical activity.

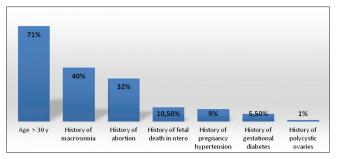


Figure 1: Risk factors of developing a diabetes mellitus

Regarding cardiovascular risk factors, patients have established an education taking into account regular control follow-up of glycemia and glycosuria, proteinuria, and the blood pressure.

T2D was treated by insulin during the feeding stage, which was substituted by oral antidiabetics. Metformin was indicated for patients with prediabetes when associating obesity when they are not successful in modifying their lifestyle.

## Discussion

"Prediabetes" is an intermediate state between a normal homeostasis of glucose and confirmed T2D and includes two clinical entities: (i) moderate hyperglycemia of fasting individual and (ii) glucose intolerance (GI). According to WHO, the fasting hyperglycemia (FG) is defined by FBG ranging between 1.10 and 1.26 g/L. GI is defined by a glycemia measured 2 h after an oral load of 75 g of glucose with values ranging between 1.40 and 2 g/L.

Despite the therapeutical education, only 22.5% (n = 45) of cases spontaneously complied to T2D screening. Most patients required phone reminder. Among 148 patients that were included in our study, 15% have developed T2D and 28% prediabetes. The risk factors of occurrence of T2D include a younger gestational age at the assessment of GD, higher FBG, and a more frequent use of insulin.

The GD is defined by WHO as a disorder of sugar tolerance, diagnosed for the first time during the pregnancy.<sup>[1]</sup> GD includes preexisting diabetes, as well as glycoregulation disorders initiated by the pregnancy, and might evolve and/or vanish after delivery. GD screening is based on risk factors, rather than a universal screening.<sup>[5]</sup> Indeed, the assessment of FBG in the first trimester of pregnancy in higher risk patients would allow diagnosing of any unrevealed T2D.<sup>[5]</sup>

The benefits of diabetological care were demonstrated with a significant reduction of the maternal and fetal morbidity.<sup>[5]</sup> Although the sugar tolerance quickly

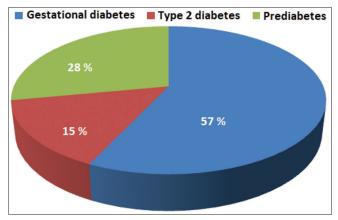


Figure 2: Reclassification of becoming of former gestational diabetes after birth child

normalizes after the delivery in most women with history of GD, the risk of developing later diabetes or an intolerance to glucose is 43% in our study [Table 2].<sup>[7,8]</sup> A history of GD increases the risk of T2D by 7 times, metabolic syndrome by 2–5 times, and cardiovascular diseases by 1–7 times.<sup>[9]</sup> The recurrence of GD is estimated between 30% and 84% and even more frequent in GD cases with insulinotherapy.<sup>[5]</sup> The outcome of gestational diabetic patients after delivery is summarized in Table 2.<sup>[10-14]</sup>

The American College of Obstetricians and Gynecologists<sup>[15]</sup> and the American Diabetes Association (ADA)[16] recommend systematic screening within 6th and 12th week after delivery. The feeding and contraception do not justify delaying diabetes assessment. Indeed, the screening could be achieved using FBG or OGTT.<sup>[17]</sup> The OGTT with 75 g of glucose has the highest sensitivity of 100%.<sup>[17]</sup> The FBG has lower sensitivity compared to OGTT but HbA1C could supplant these assessments.<sup>[3]</sup> Indeed, FBG allows assessing moderate hyperglycemia and diabetes in fasting patients, but it is not sensitive to GI. According to ADA recommendations,<sup>[16]</sup> a HbA1C indicates increased risk of diabetes requiring a greater supervision, but it does not allow a positive diagnosis in the view of insufficient predictive value.<sup>[18]</sup> The collaborative work of ATLANTIC DIP group has used a threshold value of HbA1C that was fixed at 5.9%.<sup>[18]</sup> In our series, OGTT of 75 g glucose were prescribed to patients at the end of pregnancy, which has increased the achievement rate [Table 3].[12,19,20]

ADA recommends performing a T2D assessment every 3 years in each GD patient with a first negative screening in postpartum and yearly assessments in cases presenting CHI, a moderate fasting hyperglycemia or both abnormalities.<sup>[21]</sup>

Table 1: Postpartum follow-up							
	Patient's followed up spontaneously	Patient's followed up phone call	Patients lost at the follow-up				
<i>n</i> (%)	45 (22.5)	103 (51.5)	52 (26)				
Patient's motivations	Self-motivation Heredity of	Busy with the baby	False phone number				
	diabetes Education impact	Forgotten Carelessness	Changed phone number				
		Fear of	Unreachable				
		discovering T2D	patient				

T2D=Type 2 diabetes

However, literature reported half of GD patients achieved a screening after delivery.<sup>[11,12,22]</sup> In our series, we recorded higher rate of 74%, indeed this can be elucidated by the active reminding of former GD by phone in postpartum stage. Van Ryswyk *et al.* reported a study comparing results of a reminding system by messages sent to 276 women after GD episodes and delivery; randomized patients were split into two groups: the first group included 140 patients that were reminded by 6<sup>th</sup> week, 3<sup>rd</sup> and 6<sup>th</sup> month; while the second group enclosed 136 patients reminded only in the 6<sup>th</sup> month. The screening compliance and follow-up results were similar in both groups; hence, they suggested a single reminder by the 6<sup>th</sup> after delivery since sufficient to motivate the follow-up.<sup>[23]</sup>

The predictive factors of T2D development after a GD reported in our series agreed with earlier literature; these predictive factors includes obesity, the use of insulin during the pregnancy, higher FBG and HbA1C at the diagnosis, GD occurrence before 24th week of amenorrhea.<sup>[19,20,24-26]</sup>

Schaefer-Graf *et al.* calculated the prevalence and risk of diabetes attributed to each factor.<sup>[11]</sup> Hence, raised FBG during OGTT at the time of the diagnosis was the most discriminating since glycemia  $\leq 0.95$  g/l reflects minor risk of 0.5%. The risk rises to 36.7% with a glycemia over 1.21 g/L.<sup>[27]</sup> The gestational age at the diagnosis seems to be significant because the risk is 21.4% for GD diagnosed at 19<sup>th</sup> week, while dropping to 8.7% after the 31<sup>st</sup> week. The risk is 29% with existing former history of GD.<sup>[27]</sup>

The results of O'Sullivan tests are also a predictive factor; the risk of T2D is 2.6% when glycemia is  $\leq 1.55$  g/L and increases to 11.8% for glycemia ranging between 1.75 and 2.02 g/L and raises to 27.3% when glycemia is >2.02 g/L. Our series did not reveal that raised body mass index as a predictive factor, while other studies did.<sup>[10]</sup> The female gender of fetus seems to present higher risk factor of developing T2D; this was noticed in our study and also reported in earlier study involving 23363 patients.<sup>[27]</sup>

All these factors are easily detectable during the pregnancy and allow outlining females with higher risks of diabetes; therefore allows warning on the high requirement of a regular metabolic follow-up targeting a preventive or

Table 2: Outcome of gestational diabetic patients after delivery							
Outcome of glycemia after delivery	T2D (%)	Prediabetes (%)	GD (%)	Lost at follow-up (%)			
Notre série (200 cases)	15	28	57	26			
Cundy <i>et al</i> . <sup>[10]</sup> (1110 cases)	41		16				
Schaefer-Graf et al.[11] (4041 cases)	14	-	-	40			
Russell et al. <sup>[12]</sup> (344 cases)	8	-	-	55			
Vambergue <i>et al.</i> <sup>[13]</sup> (466 cases)	18	22	60				
Kjos <i>et al</i> . <sup>[14]</sup> (671 cases)	22						

T2D=Type 2 diabetes, GD=Gestational diabetes

Table 3: Outcome of glycemia assessment according to used diagnostic approach						
Diagnostic method of T2D (postpartum stage)	FBG	OGTT (75 g)	HbA1C			
Our series (148 cases)	6.7	67.6	25.7			
Morrison et al. <sup>[19]</sup> (1372 cases)	72.7	27.3	-			
El Feleh et al. <sup>[20]</sup> (73 cases)	-	82.2	-			
Russell et al. <sup>[12]</sup> (344 cases)	8	-	-			

HbA1C=Glycosylated hemoglobin, FBG=Fasting blood glucose, OGTT=Oral glucose tolerance testing

curative therapeutic care. Studies considering adjustment of lifestyle including regular physical activity and balanced diet were achieved in this population with higher risks of cardiovascular disorders and T2D; it showed the prevention efficiency.<sup>[23,28]</sup> The study Diabetes Prevention Program showed the efficiency of using metformin in the prevention of T2D.<sup>[29]</sup> After GD, women presenting of minor glycoregulation disorders and excess of weight have shown to half the risk of T2D if they do consider change in the lifestyle with metformin intake.<sup>[3]</sup>

#### Conclusions

This study established the GD prospective and defined risk factors of evolvement of GD toward T2D; it has also revealed motivations behind neglecting the follow-up. The women with GD have higher risk to become T2D. Indeed, the pregnancy offers an important opportunity to instigate very early therapeutical education on the interest of regular follow-up. Simple preventive strategies showed their efficiency to prevent T2D, therefore any associated complications. Reminding by phone besides to therapeutical education might improve the involvement of patients in T2D screening during the postpartum stage.

#### Financial support and sponsorship

Nil.

#### **Conflicts of interest**

There are no conflicts of interest.

Received: 02 Jan 17 Accepted: 19 Feb 18 Published: 06 May 19

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