

**In the name of God**

# **CLASSIFICATION AND DIAGNOSIS OF DIABETES MELLITUS**

## CLASSIFICATION OF DIABETES MELLITUS :

Diabetes can be classified into the following general categories:

- 1. Type 1 diabetes** (due to autoimmune B-cell destruction, usually leading to absolute insulin deficiency)
- 2. Type 2 diabetes** (due to a progressive loss of B-cell insulin secretion frequently on the background of insulin resistance)
- 3. Gestational diabetes mellitus (GDM)** (diabetes diagnosed in the second or third trimester of pregnancy that was not clearly overt diabetes prior to gestation)
- 4. Specific types of diabetes** due to other causes, e.g., monogenic diabetes syndromes (such as neonatal diabetes and maturity-onset diabetes of the young [MODY]), diseases of the exocrine pancreas (such as cystic fibrosis and pancreatitis), and drug- or chemical-induced diabetes (such as with glucocorticoid use, in the treatment of HIV/AIDS, or after organ transplantation)

# Criteria for the diagnosis of diabetes :

**FPG  $\geq 126$  mg/dL** Fasting is defined as no caloric intake for at least 8 h.\*

OR

**2-h PG  $\geq 200$  mg/dL during OGTT.** The test should be performed as described by the WHO, using a glucose load containing the equivalent of 75-g anhydrous glucose dissolved in water.\*

OR

**A1C  $\geq 6.5\%$ .** The test should be performed in a laboratory using a method that is NGSP certified and standardized to the DCCT assay.\*

OR

In a patient with classic symptoms of hyperglycemia or hyperglycemic crisis, a **random plasma glucose  $\geq 200$  mg/dL.**

\*In the absence of unequivocal hyperglycemia, diagnosis requires two abnormal test results from the same sample or in two separate test samples.

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# Criteria for testing for diabetes or prediabetes in asymptomatic adults

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## Criteria for testing for diabetes or prediabetes in asymptomatic adults

1. Testing should be considered in **overweight or obese** (BMI  $\geq 25$  kg/m<sup>2</sup> or  $\geq 23$  kg/m<sup>2</sup> in Asian Americans) adults who have one or more of the following **risk factors**:

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\* **First-degree relative with diabetes**

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\* First-degree relative with diabetes

\* **High-risk race/ethnicity (e.g., African American, Latino, Native American, Asian American, Pacific Islander)**

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- \* First-degree relative with diabetes
- \* High-risk race/ethnicity (e.g., African American, Latino, Native American, Asian American, Pacific Islander)

\* **History of CVD**

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- \* High-risk race/ethnicity (e.g., African American, Latino, Native American, Asian American, Pacific Islander)
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- \* **Hypertension** ( $\geq 140/90$  mmHg or on therapy for hypertension)

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- \* High-risk race/ethnicity (e.g., African American, Latino, Native American, Asian American, Pacific Islander)
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- \* Hypertension ( $\geq 140/90$  mmHg or on therapy for hypertension)
- \* **HDL cholesterol level  $<35$  mg/dL and/or a triglyceride level  $>250$  mg/dL**

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\* **Women with polycystic ovary syndrome**

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- \* Women with polycystic ovary syndrome
- \* **Physical inactivity**

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- \* Women with polycystic ovary syndrome
- \* Physical inactivity
- \* **Other clinical conditions associated with insulin resistance (e.g., severe obesity, acanthosis nigricans)**

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**2. Patients with prediabetes (A1C  $\geq 5.7\%$  , IGT, or IFG) should be tested yearly.**

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- \* High-risk race/ethnicity (e.g., African American, Latino, Native American, Asian American, Pacific Islander)
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- \* Hypertension ( $\geq 140/90$  mmHg or on therapy for hypertension)
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- \* Other clinical conditions associated with insulin resistance (e.g., severe obesity, acanthosis nigricans)

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**3. Women who were diagnosed with GDM should have lifelong testing at least every 3 years.**

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- \* High-risk race/ethnicity (e.g., African American, Latino, Native American, Asian American, Pacific Islander)
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**4. For all other patients, testing should begin at age 45 years.**

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**2.** Patients with prediabetes (A1C  $\geq 5.7\%$  , IGT, or IFG) should be tested yearly.

**3.** Women who were diagnosed with GDM should have lifelong testing at least every 3 years.

**4.** For all other patients, testing should begin at age 45 years.

**5. If results are normal, testing should be repeated at a minimum of 3-year intervals, with consideration of more frequent testing depending on initial results and risk status.**

# A1C :

- 2.1 To avoid misdiagnosis or missed diagnosis, **the A1C test should be performed** using a method that is certified by NGSP and **standardized** to the Diabetes Control and Complications Trial (DCCT) assay. B
- 2.2 Marked **discordance between measured A1C and plasma glucose** levels should raise the possibility of A1C assay interference due to hemoglobin variants (i.e., **hemoglobinopathies**) and consideration of using an assay without interference or plasma blood glucose criteria to diagnose diabetes. B
- 2.3 In conditions associated with an altered relationship between A1C and glycemia, such as **sickle cell disease, pregnancy** (second and third trimesters and the postpartum period), **glucose-6-phosphate dehydrogenase deficiency, HIV, hemodialysis, recent blood loss or transfusion, or erythropoietin therapy**, only plasma glucose criteria should be used to diagnose diabetes. B

# Type 1 Diabetes :

2.4 Plasma **blood glucose rather than A1C** should be used to diagnose the acute onset of type 1 diabetes in individuals with symptoms of hyperglycemia. E

2.5 **Screening** for type 1 diabetes risk with a panel of autoantibodies is currently recommended only in the setting of a research trial or in first-degree family members of a proband with type 1 diabetes. B

2.6 Persistence of **two or more autoantibodies** predicts clinical diabetes and may serve as an indication for intervention in the setting of a clinical trial. B

# Prediabetes and Type 2 Diabetes (1) :

- 2.7 Screening** for prediabetes and type 2 diabetes with an informal assessment of **risk factors** or validated tools should be considered in asymptomatic adults **B**
- 2.8** Testing for prediabetes and/or type 2 diabetes in asymptomatic people should be considered in adults of any age who are **overweight or obese** (BMI  $\geq 25$  kg/m<sup>2</sup> or  $\geq 23$  kg/m<sup>2</sup> in Asian Americans) and who have one or more additional **risk factors** for diabetes (**Table 2.3**) **B**
- 2.9** For all people, testing should **begin at age 45 years** **B**
- 2.10** If tests are normal, repeat testing carried out at a minimum of **3-year** intervals is reasonable **C**

## Prediabetes and Type 2 Diabetes (2) :

- 2.11 To test for prediabetes and type 2 diabetes, fasting plasma glucose, 2-h plasma glucose during 75-g oral glucose tolerance test, and A1C are **equally** appropriate **B**
- 2.12 In patients with prediabetes and type 2 diabetes, identify and, if appropriate, treat **other cardiovascular disease risk factors** **B**
- 2.13 Risk-based screening for prediabetes and/or type 2 diabetes should be considered after the onset of puberty or after 10 years of age, whichever occurs earlier, in **children** and adolescents who are **overweight** (BMI  $\geq 85^{\text{th}}$  percentile) or **obese** (BMI  $\geq 95^{\text{th}}$  percentile) and who have **additional risk factors** for diabetes (see **Table 2.4** for evidence grading of risk factors)

### Table 2.4—Risk-based screening for type 2 diabetes or prediabetes in asymptomatic children and adolescents in a clinical setting

Testing should be considered in youth\* who are overweight ( $\geq 85\%$  percentile) or obese ( $\geq 95\%$  percentile) **A** and who have one or more additional risk factors based on the strength of their association with diabetes:

- Maternal history of diabetes or GDM during the child's gestation **A**
- Family history of type 2 diabetes in first- or second-degree relative **A**
- Race/ethnicity (Native American, African American, Latino, Asian American, Pacific Islander) **A**
- Signs of insulin resistance or conditions associated with insulin resistance (acanthosis nigricans, hypertension, dyslipidemia, polycystic ovary syndrome, or small-for-gestational-age birth weight) **B**

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\*After the onset of puberty or after 10 years of age, whichever occurs earlier. If tests are normal, repeat testing at a minimum of 3-year intervals, or more frequently if BMI is increasing, is recommended.



# ARE YOU AT RISK FOR

# TYPE 2 DIABETES?



## Diabetes Risk Test

**1 How old are you?**

- Less than 40 years (0 points)
- 40—49 years (1 point)
- 50—59 years (2 points)
- 60 years or older (3 points)

Write your score in the box.




**2 Are you a man or a woman?**

- Man (1 point)    Woman (0 points)

**3 If you are a woman, have you ever been diagnosed with gestational diabetes?**

- Yes (1 point)    No (0 points)

**4 Do you have a mother, father, sister, or brother with diabetes?**

- Yes (1 point)    No (0 points)

**5 Have you ever been diagnosed with high blood pressure?**

- Yes (1 point)    No (0 points)

**6 Are you physically active?**

- Yes (0 points)    No (1 point)

**7 What is your weight status? (see chart at right)**

Height	Weight (lbs.)		
4' 10"	119-142	143-190	191+
4' 11"	124-147	148-197	198+
5' 0"	128-152	153-203	204+
5' 1"	132-157	158-210	211+
5' 2"	136-163	164-217	218+
5' 3"	141-168	169-224	225+
5' 4"	145-173	174-231	232+
5' 5"	150-179	180-239	240+
5' 6"	155-185	186-246	247+
5' 7"	159-190	191-254	255+
5' 8"	164-196	197-261	262+
5' 9"	169-202	203-269	270+
5' 10"	174-208	209-277	278+
5' 11"	179-214	215-285	286+
6' 0"	184-220	221-293	294+
6' 1"	189-226	227-301	302+
6' 2"	194-232	233-310	311+
6' 3"	200-239	240-318	319+
6' 4"	205-245	246-327	328+
	(1 Point)	(2 Points)	(3 Points)

You weigh less than the amount in the left column (0 points)

Add up your score.




**If you scored 5 or higher:**

You are at increased risk for having type 2 diabetes. However, only your doctor can tell for sure if you do have type 2 diabetes or prediabetes (a condition that precedes type 2 diabetes in which blood glucose levels are higher than normal). Talk to your doctor to see if additional testing is needed.

Adapted from Bang et al., Ann Intern Med 151:775-783, 2009. Original algorithm was validated without gestational diabetes as part of the model.

## Table 2.5—Criteria defining prediabetes\*

FPG 100 mg/dL (5.6 mmol/L) to 125 mg/dL (6.9 mmol/L) (IFG)

OR

2-h PG during 75-g OGTT 140 mg/dL (7.8 mmol/L) to 199 mg/dL (11.0 mmol/L) (IGT)

OR

A1C 5.7–6.4% (39–47 mmol/mol)

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\*For all three tests, risk is continuous, extending below the lower limit of the range and becoming disproportionately greater at the higher end of the range.

# Gestational Diabetes Mellitus (1) :

- 2.14 Test for undiagnosed diabetes **at the first prenatal visit** in those with risk factors using standard diagnostic criteria **B**
- 2.15 Test for gestational diabetes mellitus **at 24-28 weeks of gestation** in pregnant women not previously known to have diabetes **A**
- 2.16 Test women with gestational diabetes mellitus for prediabetes or diabetes **at 4-12 weeks postpartum**, using the 75-g oral glucose tolerance test and clinically appropriate nonpregnancy diagnostic criteria **B**

## Gestational Diabetes Mellitus (2) :

**2.17** Women with a history of gestational diabetes mellitus should have **lifelong screening** for the development of diabetes or prediabetes at least **every 3 years** B

**2.18** Women with a history of gestational diabetes mellitus found to have **prediabetes** should receive intensive **lifestyle interventions** or **metformin** to prevent diabetes A

# DIAGNOSIS OF GDM :

GDM diagnosis can be accomplished with either of two strategies:

1. “**One-step**” 75-g OGTT

or

2. “**Two-step**” approach with a 50-g (non fasting) screen followed by a 100-g OGTT for those who screen positive

# Screening for and DIAGNOSIS OF GDM

## One-step strategy

Perform a **75-g OGTT** , with plasma glucose measurement when patient is fasting and at 1 and 2 h, at **24–28 weeks** of gestation in women not previously diagnosed with diabetes.

The OGTT should be performed in the morning after an overnight fast of at least 8 h.

The diagnosis of **GDM** is made when **any** of the following plasma glucose values are met or exceeded:

\* Fasting: **92** mg/dL

\* 1 h: **180** mg/dL

\* 2 h: **153** mg/dL

International Association of the Diabetes and Pregnancy Study Groups (IADPSG)

Metzger BE, Lowe LP, Dyer AR, et al.; HAPO Study Cooperative Research Group. Hyperglycemia and adverse pregnancy outcomes. N Engl

J Med 2008;358:1991–2002

# Screening for and DIAGNOSIS OF GDM

## Two-step strategy

**Step 1:** Perform a **50-g GCT** (non fasting), with plasma glucose measurement at 1 h, at **24–28 weeks** of gestation in women not previously diagnosed with diabetes.

If the plasma glucose level measured 1 h after the load is **≥130** mg/dL, **135** mg/dL, or **140**mg/dL, proceed to a 100-g OGTT.

**Step 2:** The **100-g OGTT** should be performed when the patient is fasting.

The diagnosis of GDM is made if at least **two\*** of the following four plasma glucose levels (measured fasting and 1 h, 2 h, 3 h during OGTT) are met or exceeded:

	Carpenter-Coustan	or	NDDG
* Fasting	<b>95</b> mg/dL		105 mg/dL
* 1 h	<b>180</b> mg/dL		190 mg/dL
* 2 h	<b>155</b> mg/dL		165 mg/dL
* 3 h	<b>140</b> mg/dL		145 mg/dL

# Screening for and DIAGNOSIS OF GDM

## Two-step strategy

**Step 1:** Perform a **50-g GCT** (non fasting), with plasma glucose measurement at 1 h, at **24–28 weeks** of gestation in women not previously diagnosed with diabetes.

If the plasma glucose level measured 1 h after the load is **≥130** mg/dL, **135** mg/dL, or **140**mg/dL, proceed to a 100-g OGTT

The higher cutoff yielded sensitivity of 70–88% and specificity of 69–89%, while the lower cutoff was 88–99% sensitive and 66–77% specific.

**Step 2:** The **100-g OGTT** should be performed when the patient is at 24–28 weeks of gestation. The diagnosis of GDM is made if at least **two\*** of the following are met or exceeded:

	Carpenter-Coustan	or	NDDG
* Fasting	<b>95</b> mg/dL		105 mg/dL
* 1 h	<b>180</b> mg/dL		190 mg/dL
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# Screening for and DIAGNOSIS OF GDM

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**Step 1:** Perform a **50-g GCT** (nonfasting), with plasma glucose measurement at 1 h, at **24–28 weeks** of gestation in women not previously diagnosed with diabetes.

If the plasma glucose level measured 1 h after the load is  **$\geq 130$**  mg/dL, **135** mg/dL, or **140** mg/dL, proceed to a 100-g OGTT

**The use of A1C at 24–28 weeks of gestation as a screening test for GDM does not function as well as the GCT**

**Step 2:** The **100-g OGTT** should be performed when the patient is between 24 and 28 weeks of gestation. The diagnosis of GDM is made if at least **two\*** of the following four plasma glucose levels (measured fasting and 1 h, 2 h, 3 h during OGTT) are met or exceeded:

	Carpenter-Coustan	or	NDDG
* Fasting	<b>95</b> mg/dL		105 mg/dL
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## Two-step strategy

**Step 1:** Perform a 50-g GCT (non fasting), with plasma glucose measurement at 1 h, at 24–28 weeks of gestation in women not previously diagnosed with diabetes.

If the plasma glucose level measured 1 h after the load is  $\geq 130$  mg/dL, 135 mg/dL, or 140mg/dL, proceed to a 100-g OGTT.

**Step 2:** The **100-g OGTT** should be performed when the patient is fasting.

The diagnosis of GDM is made if at least **two\*** of the following four plasma glucose levels (measured fasting and 1 h, 2 h, 3 h during OGTT) are met or exceeded:

	Carpenter-Coustan	or	NDDG
* Fasting	<b>95</b> mg/dL		105 mg/dL
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\*American College of Obstetricians and Gynecologists (ACOG) notes that one elevated value can be used for diagnosis

Carpenter MW, Coustan DR. Criteria for screening tests for gestational diabetes. Am J Obstet Gynecol 1982;144:768–773

National Diabetes Data Group. Classification and diagnosis of diabetes mellitus and other categories of glucose intolerance. Diabetes 1979; 28:1039–1057

# Screening for and DIAGNOSIS OF GDM

## Two-step strategy

**Step 1:** Perform a 50-g GLT (nonfasting), with plasma glucose measurement at 1 h, at 24–28 weeks of gestation in women not previously diagnosed with diabetes.

If the plasma glucose level measured 1 h after the load is  $\geq 130$  mg/dL, 135 mg/dL, or 140mg/dL, proceed to a 100-g OGTT.

**Step 2:** The **100-g OGTT** should be performed when the patient is fasting.

The diagnosis of GDM is made if at least **two\*** of the following four plasma glucose levels (measured fasting and 1 h, 2 h, 3 h during OGTT) are met or exceeded:

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## Two-step strategy

**Step 1:** Perform a 50-g GLT (nonfasting), with plasma glucose measurement at 1 h, at 24–28 weeks of gestation in women not previously diagnosed with diabetes.

If the plasma glucose level measured 1 h after the load is  $\geq 130$  mg/dL, 135 mg/dL, or 140mg/dL, proceed to a 100-g OGTT.

**Step 2:** The **100-g OGTT** should be performed when the patient is fasting.

The diagnosis of GDM is made if the following four plasma glucose levels (at 0, 1, 2, and 3 h during OGTT) are met or exceeded:

If the two-step approach is used, it would appear advantageous to use the lower diagnostic thresholds as shown in step 2

Carpenter-C

NDDG

* Fasting	<b>95</b> mg/dL	<b>105</b> mg/dL
* 1 h	<b>180</b> mg/dL	<b>190</b> mg/dL
* 2 h	<b>155</b> mg/dL	<b>165</b> mg/dL
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National Diabetes Data Group. Classification and diagnosis of diabetes mellitus and other categories of glucose intolerance. Diabetes 1979; 28:1039–1057

# Cystic Fibrosis-Related Diabetes

- 2.19 Annual screening for cystic fibrosis-related diabetes with an oral glucose tolerance test should begin by age 10 years in all patients with cystic fibrosis not previously diagnosed with cystic fibrosis-related diabetes B
- 2.20 A1C is not recommended as a screening test for cystic fibrosis-related diabetes B
- 2.21 Patients with cystic fibrosis-related diabetes should be treated with insulin to attain individualized glycemic goals A
- 2.22 Beginning 5 years after the diagnosis of cystic fibrosis-related diabetes, annual monitoring for complications of diabetes is recommended E

# Post transplantation Diabetes Mellitus

- 2.23 Patients **should be screened** after organ transplantation for hyperglycemia, with a formal diagnosis of post transplantation diabetes mellitus being best made once a patient is **stable** on an immunosuppressive regimen and **in the absence of an acute infection** **E**
- 2.24 The **oral glucose tolerance test** is the preferred test to make a diagnosis of post transplantation diabetes mellitus **B**
- 2.25 Immunosuppressive regimens shown to provide the best outcomes for patient and graft survival should be used, irrespective of post transplantation diabetes mellitus risk **E**

# Monogenic Diabetes Syndromes

The diagnosis of monogenic diabetes should be considered in **children and adults diagnosed with diabetes in early adulthood** with the following findings:

- Diabetes diagnosed within the **first 6 months** of life (with occasional cases presenting later, mostly *INS* and *ABCC8* mutations)
- Diabetes **without typical features of type 1 or type 2 diabetes** (negative diabetes-associated autoantibodies, nonobese, lacking other metabolic features especially with strong family history of diabetes)
- Stable, **mild fasting hyperglycemia** (100-150 mg/dL [5.5-8.5 mmol/L]), **stable A1C between 5.6 and 7.6%** (between 38 and 60 mmol/mol), especially if nonobese

# Monogenic Diabetes Syndromes

- 2.26** All children diagnosed with diabetes in the first 6 months of life should have immediate **genetic testing** for **neonatal diabetes** **A**
- 2.27** Children and adults, diagnosed in early adulthood, who have diabetes not characteristic of type 1 or type 2 diabetes that occurs in successive generations (suggestive of an autosomal dominant pattern of inheritance) should have **genetic testing** for **maturity-onset diabetes of the young** **A**
- 2.28** In both instances, consultation with a center specializing in diabetes genetics is recommended to understand the significance of these mutations and how best to approach further evaluation, treatment, and genetic counseling **E**



**Thank you**