



## Diabetes in pregnancy

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# **Diabetes in Pregnancy: 2 Categories**

Pregestational diabetes	Gestational diabetes
Pregnancy in pre-existing diabetes  • Type 1 diabetes  • Type 2 diabetes	Diabetes diagnosed in pregnancy



nutritional therapy

1-2 weeks

physical activity



## Nutritional therapy

- Achieve normoglycemia
- Prevent ketosis
- Provide adequate gestational weight gain based on maternal body mass index (BMI)
- (70 to 85 percent) can achieve normoglycemia
- 40%fat,40%carbohydrate,20%protein
- • FBS: <95 mg/dL (5.3 mmol/L)
- One-hour postprandial : <140 mg/dL (7.8 mmol/L)</li>
- • Two-hour postprandial : <120 mg/dL



## PHARMACOLOGIC THERAPY

INSULIN

ORAL: metformin

glyburide





- ACOG and ADA:
- prefer use of insulin for treatment of diabetes during pregnancy
- FDA didn't approve oral drug
- ACOG:
- oral agent for women who decline insulin therapy recommends metformin over glyburide as the preferred oral antihyperglycemic agent





Society of Maternal-Fetal Medicine

 concluded metformin is safe first-line pharmacologic alternative to insulin





- FIGO recommends
- insulin as the first-line
- before 20 weeks of gestation
- pharmacologic therapy is needed >30 weeks,
- fasting blood glucose is >110 mg/dL
- one hour postprandial glucose is >140 mg/dL
- pregnancy weight gain is >12 kg





# **Insulin Therapy**

#### TABLE 57-8. Action Profiles of Commonly Used Insulins

Insulin Type	Onset	Peak (hr)	Duration (hr)
Short-acting (SC)			
Lispro	<15 min	0.5-1.5	3-4
Glulisine	<15 min	0.5 - 1.5	3-4
Aspart	<15 min	0.5 - 1.5	3-4
Regular	30-60 min	2-3	4-6
Long-acting (SC)			
Detemir	1–4 hr	Minimala	Up to 24
Glargine	1-4 hr	Minimala	Up to 24
NPH	1-4 hr	6–10	10–16

Minimal near activity



## Insulin therapy

- four-times-per-day regimen
- Short-acting:4 to 8 U before meals

NPH before breakfast:6 to 8 U(0.2 u/kg)

If FBS>90 to 95, NPH 4 to 6 U at 11PM





#### Pharmacokinetics of the most commonly used insulin preparations

Insulin type	Approximate onset of action	Peak effect	Approximate duration *
Lispro, aspart, faster aspart, glulisine	3 to 15 minutes	45 to 75 minutes	2 to 4 hours
Regular	30 minutes	2 to 4 hours	5 to 8 hours
NPH	2 hours	4 to 12 hours	8 to 18 hours, with usual duration of action around 12 hours
Insulin glargine	2 hours	No peak	20 to >24 hours
Insulin detemir	2 hours	3 to 9 hours	6 to 24 hours¶
NPL	2 hours	6 hours	15 hours
Insulin degludec	2 hours	No peak	>40 hours

NPH: neutral protamine hagedorn; NPL: neutral protamine lispro.

# eral antihyperglycemic agents

- Metformin did have some benefits:
- Lower mean birth weight Less gestational weight gain
- Less composite neonatal death or serious morbidity
- Women should be informed that it crosses the placenta

 Begins 500mg twice daily q 12 h to maximum of 2500mg/d





## glyburide

- frequency of treatment failure (lack of glycemic control) is similar for glyburide and metformin in most trials.
- Failure rate: 16 to 17%
- 2.5 mg every 12 hours
- Maximum dose:20 mg/d
- peaked at 2 to 3 hours
- returned to baseline by 8 to 10 hours.

# Pregestational diabetes

- 1-Women on medical nutritional therapy prior to pregnancy
- 2-Women on oral anti hyperglycemic agents prior to pregnancy
- insulin therapy is the only means of achieving the degree of glycemic control desirable
- Throughout pregnancy in women with type 1 and type 2 diabetes
- Type 2 diabetes have excellent glycemic control on an oral antihyperglycemia
- Drug such as metformin at conception
- 3-on multiple daily injection therapy prior to pregnancy



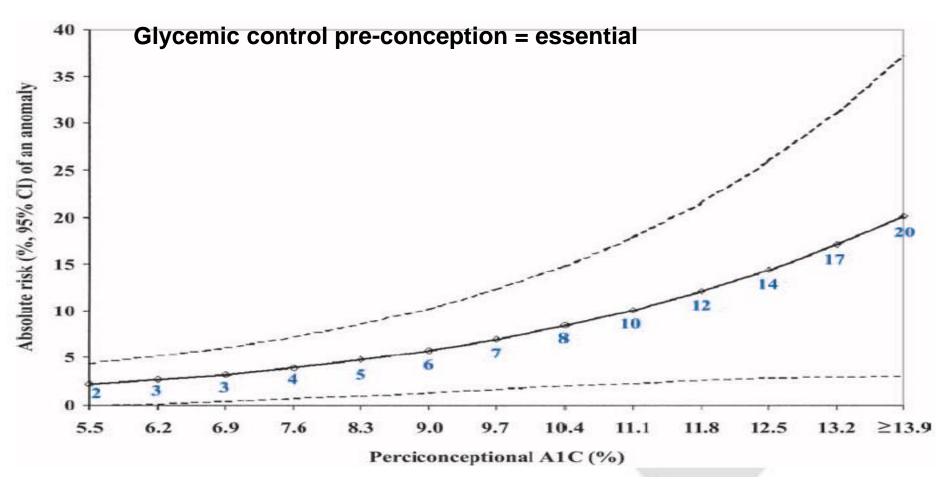


## **Treatment**

#### Target Plasma Glucose Levels in Pregnancy

Time	Glucose Level (mg/dL)
Before breakfast	60-90
Before lunch, supper, bedtime snack	60-105
One hour after meals	≤140
Two hours after meals	≤120
2 AM to 6 AM	>60

# Risk of Fetal Anomaly Relative to Periconceptional A1C







ADA and CDA: Hb A1c

• NICE: HbA1c<6.5 %



### macrosomia

Above 4000,4500 gram or greater than 90<sup>th</sup> or 95th centile

- 40-60% of pregnancies with pregestational diabet
- Around 26 weeks
- Tight control in the first trimester
- Correlation between1-hour postprandial glucose level and HbA1c
- Fasting glucose between 29 and 32 weeks



## stillbirth

- Higher in type 1 and 2 rather than GDM
- Maternal obesity
- Higher in macrosomia
- Possible reasons is hypoxia due to
- acidosis, cardiac arrhythmias, hypokalemia
- Higher in woman with moderate untreated GDM





# Fetal cardiomegaly

- Myocardial hypertrophy
- Hypertrophy of ventricular walls and septum
- Subaortic stenosis
- Secondary mitral insufficiency





# polyhydramnious

- Higher in patient with poor glycemic control and LGA fetuses, with higher A1c
- Fetal hyperglycemia leading to polyuria
- Decreased fetal swalloing





#### Antepartum Fetal Surveillance in Low-Risk Insulin-Dependent Diabetes Mellitus\*

Study	Indicated
Ultrasonography at 4- to 6-wk intervals	Yes
Maternal assessment of fetal activity, daily at	Yes
28 wk	
NST at 32 wk; BPP or CST if NST is nonreactive	Yes
Amniocentesis for lung profile	Yes, if elective delivery is planned before
	39 wk

BPP, biophysical profile; CST, contraction stress test; NST, nonstress test.

<sup>\*</sup> Excellent control, no vasculopathy (classes B, C), no stillbirth.





#### Antepartum Fetal Surveillance in High-Risk Insulin-Dependent Diabetes Mellitus\*

Study	Indicated
Ultrasonography at 4- to 6-wk intervals	Yes
Maternal assessment of fetal activity daily at 28 wk	Yes
NST; BPP or CST if NST is nonreactive	Initiate at 28-30 wk
Consider amniocentesis for lung profile prior to 38 wk	

BPP, biophysical profile; CST, contraction stress test; NST, nonstress test.

<sup>\*</sup> Poor control (macrosomia, hydramnios), vasculopathy (classes D, F, R), prior stillbirth.





#### Labor and delivery

#### Timing of birth:

- Offer elective delivery at 37<sup>+0</sup>–38<sup>+6</sup> weeks in women with preexisting diabetes
- Offer elective delivery at no later than 40<sup>+6</sup> weeks in women with GDM with no complications and whose diabetes is well controlled
- Discuss the potential for shoulder dystocia with women before labor; consider elective CS if estimated fetal weight ≥ 4500 g





#### TABLE 57-10. Insulin Management During Labor and Delivery

- Usual dose of intermediate-acting insulin is given at bedtime.
- · Morning dose of insulin is withheld.
- Intravenous infusion of normal saline is begun.
- Once active labor begins or glucose levels decrease to less than 70 mg/dL, the infusion is changed from saline to 5% dextrose and delivered at a rate of 100–150 cc/h (2.5 mg/kg/min) to achieve a glucose level of approximately 100 mg/dL.
- Glucose levels are checked hourly using a bedside meter allowing for adjustment in the insulin or glucose infusion rate.
- Regular (short-acting) insulin is administered by intravenous infusion at a rate of 1.25 U/h if glucose levels exceed 100 mg/dL.

Data from Coustan DR. Delivery: timing, mode, and management. In: Reece EA, Coustan DR, Gabbe SG, editors. Diabetes in women: adolescence, pregnancy, and menopause. 3rd ed. Philadelphia (PA): Lippincott Williams & Wilkins; 2004; and Jovanovic L, Peterson CM. Management of the pregnant, insulin-dependent diabetic woman. Diabetes Care 1980;3:63–8.





