Gestational DiabetesThere Are Some Frustrations



Most cases identified are mild in nature Patients don't like the **OGCT or 3-hr GTT** Really counseling patients is time-consuming Patients are reluctant to check BGs Do we make a difference?

MFMU Network Randomized Treatment Trial of Mild GDM

- Multicenter randomized trial of women with
 - Abnormal 50 g OGC
 - -3-hr GTT \rightarrow GDM, but
 - Normal FBS on 3-hr GTT
- Subjects randomized to
 - Usual care (GTT results not available)
 - Dietary intervention, SBGM, and insulin if required

Effect of Treatment

Outcome	Treated (N=485)	Routine Care (N= 473)	P value
Death	0	0	
Hyperbili- rubinemia	43 (10%)	54 (13%)	0.12
Hypoglycemia	62 (16%)	55 (15%)	0.75
Elevated cord C-peptide	75 (18%)	92 (23%)	0.07
Birth trauma	3 (<1%)	6 (1%)	0.33
Composite	149 (32%)	163 (37%)	0.14

Landon et al, N Engl J Med 2009; 361:1339

Effect of Treatment

Outcome		Routine Care (N=473)	
Birthweight	$3,302 \pm 50$	$23,408 \pm 589$	<0.001
LGA	34 (7.1%)	66 (14.5%)	<0.001
Macrosomia	28 (5.9%)	65 (14.3%)	<0.001
Fat Mass (g)	427 ± 198	464 ±	<0.003

Landon et **222** ngl J Med 2009; 361:1339

Effect of Treatment

Outcome	Treated (N=485)	Routine Care (N= 473)	P value
GHTN - PE	41 (9%)	62 (14%)	0.01
Cesarean	128 (27%)	154 (34%)	0.02
Shoulder dystocia	7 (1.5%)	18 (4%)	0.02

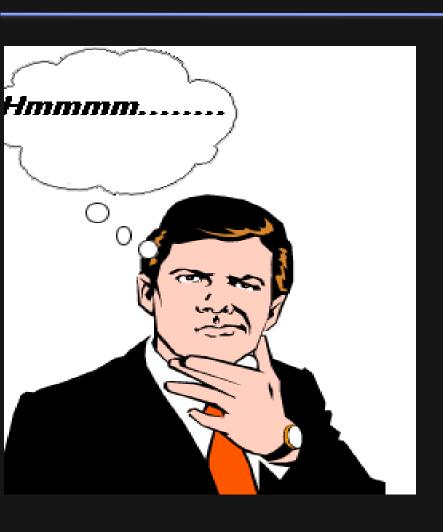
Landon et al, N Engl J Med 2009; 361:1339

MFMU Network Randomized Treatment Trial of Mild GDM

Outcome	Number Needed to Treat
Macrosomia	12
Cesarean Delivery	14
Shoulder Dystocia	40
PE+GHTN	20

Landon et al, Am J Obstet Gynecol 2009; 199:S2

Diabetes in Pregnancy Do we make a difference?



Offspring → childhood obesity and DM
Mothers → type 2 DM



Childhood Follow-up

- Follow-up of 500 (55%) children (ages 5-10) from MFMUN treatment trial
 - Physical parameters and BP
 - Fasting glucose and insulin
 - Triglycerides
 - HDL cholesterol

Childhood Follow-up

Group	Treated (N=264)	Untreated (N=236)	Adj RR (95% CI)
BMI >85 percentile for age and sex	86 (32.6%)	91 (38.6%)	0.88 (0.71-1.10)
Waist circ >90 th percentile for age/sex/ethnicity	31 (11.7%)	27 (11.4%)	1.05 (0.65-1.69)
Impaired FBG	12 (5.7%)	13 (7.2%)	0.76 (0.36-1.62)
Elevated Triglycerides	38 (18.2%)	29 (16.0%)	1.11 (0.71-1.72
Low HDL cholesterol	27 (13.0%)	22 (12.2%)	1.03 (0.61-1.76)
HBP >95 th percentile for age/sex/height	30 (11.5%)	23 (9.8%)	1.23 (0.74-2.05)

Maternal Follow-up

- Follow-up of 457 (50%) mothers from MFMUN treatment trial (at 4.5-10 yrs)
 - Physical parameters and BP
 - Fasting glucose and insulin
 - Triglycerides
 - HDL cholesterol
 - OGTT (75 g)

Maternal Follow-up

Group	Treated (N=229)	Untreated (N=201)	P Value or Adj RR (95% CI)
ВМІ	29.4 (28.6-30.3)	29.1 (28.2-30.0)	0.59
Waist circ (cm)	96.2 (94.3-96.1)	94.2 (92.2-96.3)	0.11
Hip circ (cm)	108.4 (106.7- 110.1)	106.8 (105.1- 108.7)	0.16
Diabetes	21 (9%)	17 (8%)	1.03 (0.56-1.92)
Metabolic syndrome	73 (32%)	69 (34%)	0.96 (0.73-1.26)

Casey et al. SMFM, 2015

Gestational Diabetes Do we make a difference?

- Identification and treatment of GDM associated with
 - ➤ Modest reductions in birthweight / fat mass / LGA / macrosommia / GHTN-PE / shoulder dystocia ☺
 - > Possible reduction in CS ©
- Childhood and post-delivery maternal outcomes are not altered ⊗

- Recent controversy in GDM diagnosis:
 - IADPSG and ADA versus ACOG and NIH
 Consensus Panel

Screening Strategy for GDM

- All women "whether by ...medical history, clinical risk factors, or laboratory screening test...."
- Laboratory screening
 - 50 g glucose challenge → 1 hour venous glucose
 - Threshold of 135 or 140
 - 3 hour diagnostic OGTT

Screening Strategy for GDM

Status	Plasma or Serum Serum Glucose (Carpenter and Coustan)	Plasma Glucose (NDDG)
Fasting	95	105
1 hour	180	190
2 hour	155	165
3 hour	140	145

ACOG Practice Bulletin 137, August 2013

IADPSG Screening Strategy for GDM

- One-step approach
 - 75 g, 2 hour OGTT
- GDM diagnosis if any one threshold met or exceeded

Status	Threshold
	Glucose
Fasting	92
1 hour	180
2 hour	153

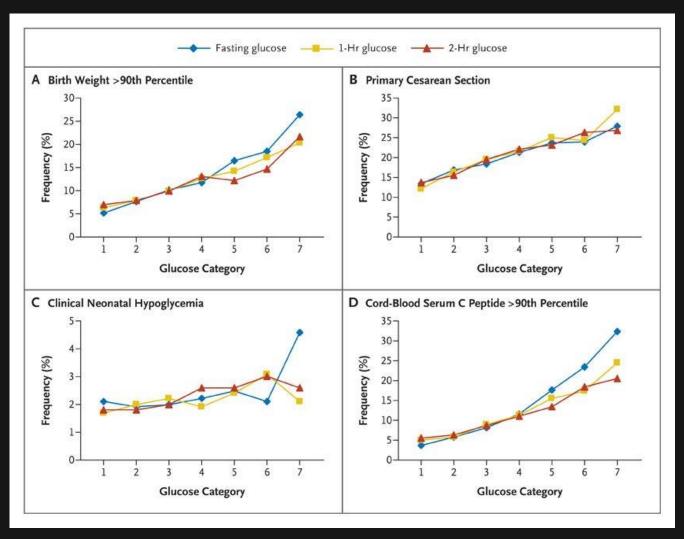
IADPSG Screening Strategy for Diabetes in Pregnancy

- FBG or HgbA1c at first prenatal visit
 - -≥126 or 6.5% → overt diabetes
 - $-92-125 \rightarrow GDM$
 - -<92 → 75 g OGTT at 24-28 weeks</p>

HAPO Study

- Prospective, observational study of 23,316 pregnant women
- 75 g OGTT > 7 glucose categories
- Primary outcomes
 - -LGA
 - -CS
 - Neonatal hypoglycemia
 - Cord C-peptide

Frequency of Primary Outcomes across the Glucose Categories



The HAPO Study Cooperative Research Group. N Engl J Med 2008;358:1991-2002



IADPSG Screening Strategy for GDM

- One-step approach
 - 75 g, 2 hour OGTT
- GDM diagnosis if any one threshold met or exceeded

Status	Threshold Glucose
Fasting	92
1 hour	180
2 hour	153

Based on glucose levels associated with 1.75-fold increased risk of LGA, neonatal body fat >90th%tile, & ⊕cord insulin

Implication

 HAPO→IADPSG recommendation would result in nearly 20% of the population being diagnosed with GDM!

Pros for IADPSG Recommendations

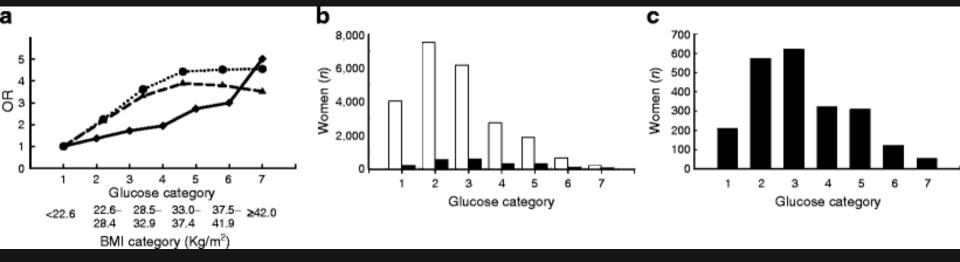
- Based on 2 randomized trials
 - ~30% to 60% reduction in adverse outcomes
- In HAPO (23,316 pregnancies), dx and treatment would
 - Prevent 140 cases of LGA, 21 SDs, and 16 birth injuries
- Potential for long-term health improvements related to patient education in pregnancy

Cons for IADPSG Recommendations

- High percentage of population diagnosed with GDM
 - But would <u>only</u> prevent 140 cases of LGA, 21 SDs, and 16 birth injuries
- Reproducibility of 75 g OGTT suboptimal
 - 25% could be reclassified
- "Treatment" in real-world may not be practical or effective

Cons for IADPSG Recommendations

- Unintended consequences of "overmedicalization"
 - Inductions and early delivery
- Costs > benefits?



Ryan EA. Diabetologia 2011;54:480

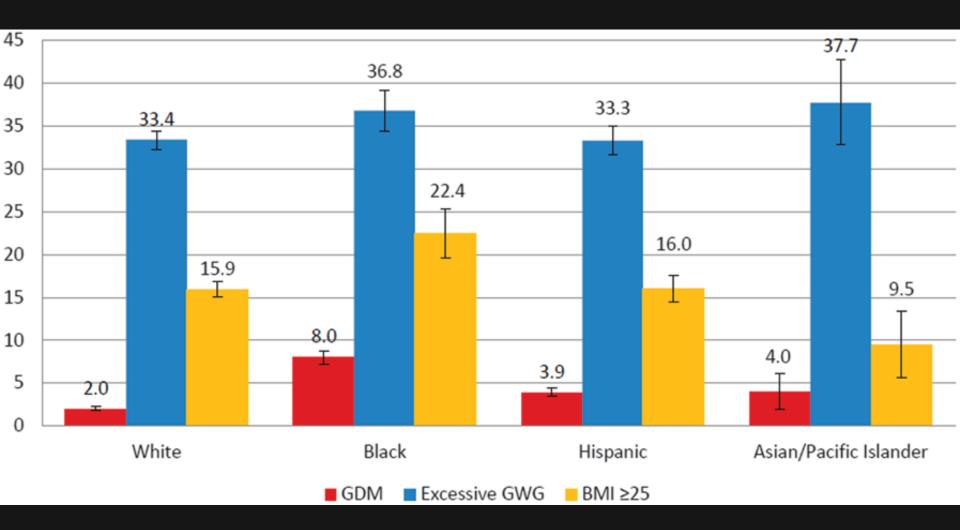
Diagnosing GDM?

- Both glucose and BMI predict LGA
- In HAPO, 78% of LGA infants were born of women without IADPSG GDM diagnosis!
- BMI is more relevant except at highest glucose levels

Obesity?



Prevalence of LGA for Births at 37-41 Weeks



Pre-Pregnancy BMI Category (Body Mass Index = BMI)	Recommende Gain During	the state of the s
BMI < 18.5 Underweight	12.5 – 18.0 kg	28.0 – 40.0 lbs
BMI 18.5 – 24.9 Normal Weight	11.5 – 16.0 kg	25.0 – 35.0 lbs
BMI 25.0 – 29.9 Overweight	7.0 – 11.5 kg	15.0 – 25.0 lbs
BMI ≥ 30 Obese	5.0 – 9.0 kg	11.0 – 20.0 lbs

Body Mass Index (BMI) = Weight (kg) / [Height (m)]²

Diet and Activity in GDM

- Carbohydrate intake [should] be limited to 33–40% of calories, with the remaining calories divided between protein (20%) and fat (40%)
- A moderate exercise program as part of the treatment plan for women with GDM is recommended

Diet and Carbs in GDM

Total Calories	Carbohydrate Calories (@40% of total) / g of carbs	Examples of Daily Carbohydrate Selection
2000 calories per day (1st 6 months of pregnancy)	800 calories 200 g carbohydrates	1 medium fruit, 1 average bread, 1 ½- ½- cup cereals/grains/potatoes per meal, along with assorted other carbs
2200 calories per day (last 3 months of pregnancy)	880 calories 220 g carbohydrates	Add 1-2 carb servings per day

Activity in GDM

- 30 minutes of moderate-to-vigorous intensity aerobic exercise at least 5 days a week (150 min per week)
 - You can talk but not sing
 - You cannot say more than a few words without pausing for a breath

Diabetes Management 101 (aka "Coaching")

- Insist on patient self blood glucose monitoring - "you manage what you measure"
- Review the results at every patient visit
- Suggest ways to improve coach!
- Establish reliable, outcome-oriented referral services