

# In Defense of Carbs:

## Challenging the Low Carbohydrate Diet for Gestational Diabetes

### Implications for Long-term Metabolic Health



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- Janssen Research and Development
- No Conflicts of Interest



Investigations in the  
Gestational Origins of Lifelong  
Development

[www.infantgoldresearch.org](http://www.infantgoldresearch.org)

# Objectives

1. Make the case for flexibility in dietary carbohydrate content for Gestational Diabetes (GDM)
2. Present highlights of clinical trial results from the CHOICE diet parallel randomized controlled trial
  - *A Colorado approach to prevention*
3. Discuss implications for long-term metabolic health



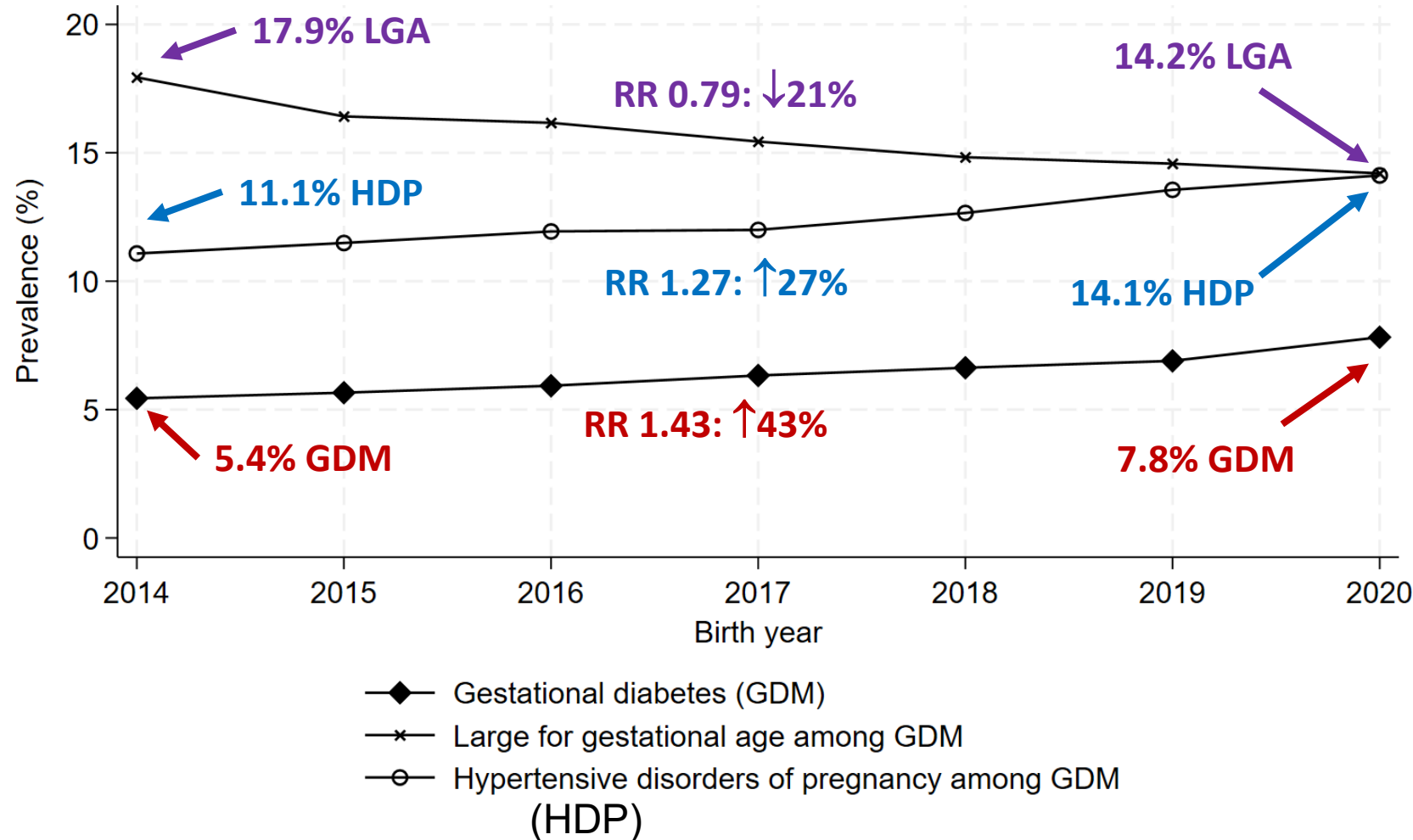


Lake Isabelle  
North of Boulder, Colorado  
August 2023

# **The Case for Flexibility in Dietary Carbohydrate Content ...**

**for Gestational Diabetes**

# Increasing Prevalence of GDM 2014-2020 with Decreased Large-for-Gestational Age



N= 25,437,281 total births

-National Center for Health Statistics, birth certificates  
-n=1,620,820 (6%) with GDM

-Despite lower LGA over time, LGA prevalence was 15.5% in GDM vs. 9.4% in controls



RR = risk ratio

Valent, A; Garg, B; and Hernandez TL, 2024, in preparation

# Nutrition Therapy in GDM: Doomed from the Start?

## Anxiety, Rigid Adherence, and Unintended Consequences

### Until GDM diagnosis, pregnancy was “normal”...

- Suddenly: High risk pregnancy label, “diet”, ↑rigid control of glucose, ↑surveillance, medications
- Anxiety, fear, depression

### Psychology related to nutrition therapy in GDM<sup>1-5</sup>

- Focus: rigid restriction of carbohydrate
- Rapid adaptation in late pregnancy is challenging; food selection is mentally taxing
- Infringement on cultural/social beliefs
- Feel confined by the diet
  - Narrow range of “acceptable foods,” limited food choices
- Rigid diet control: the most difficult component to treatment
  - **Unintended consequences: ↑↑fat intake**
- **Medicalization of eating**
  - “It feels like medically promoted disordered eating”



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2. Hui AL, 2014, Diabetes Educ, 40(5); 668

3. Teixeira PJ, 2015, BMC Med, 13:84

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University of Colorado Hospital, circa 2002



# A Low-Carbohydrate Diet in GDM is Conventional Therapy

- **Nutrition therapy is first line treatment for GDM across diagnostic criteria**
- Historically, diet designed to blunt post-prandial glycemia<sup>1</sup>
  - To prevent fetal overgrowth and large-for-gestational age (LGA)—as birthweight, or excess adiposity
  - 1990: 40% CHO; 45% fat; 15% protein
  - Pre-insulin era: Joslin diet was 2% carbohydrate, 70-80% fat
- The current evidence does not support one type of diet prescription for primary treatment of GDM and the quality of evidence is low<sup>2</sup>
- Evidence supports: ANY diet modification that *improves nutritional pattern once the diagnosis is made* reduces maternal glucose and birth weight (BW)<sup>3,4</sup>
  - Meta-analysis of 18 RCTs (n=1151 women). Fasting glucose ↓4mg/dL, postprandial glucose ↓8 mg/dL. Lower need for adjunctive medication (35%), birth weight ↓ 171g
- Infants born LGA or with ↑adiposity have higher risk for childhood overweight, insulin resistance, hypertension, some leukemias, and later diabetes and cardiovascular disease<sup>5</sup>



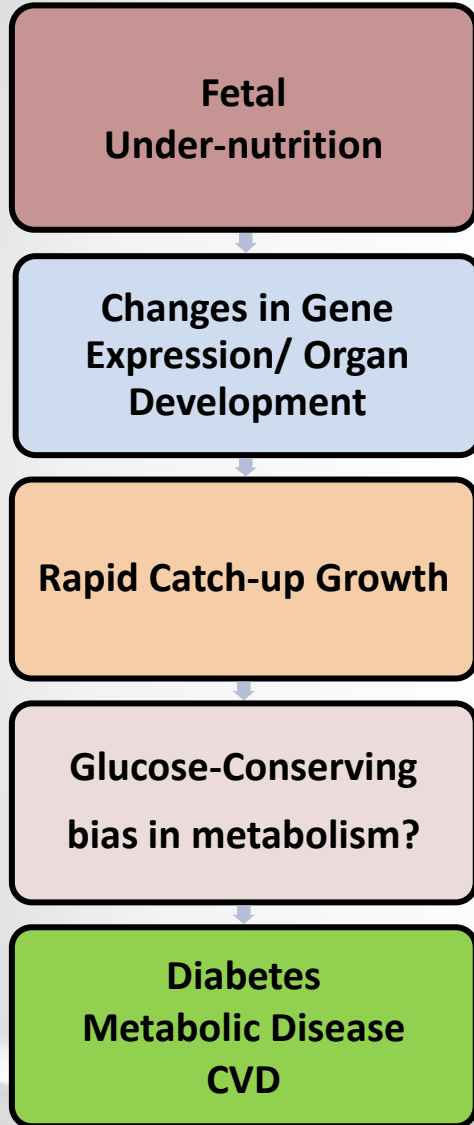
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2. Han, S, Cochrane Database Systematic Reviews, 2017, 2:CD009275

3. Yamamoto JM, 2018, Diabetes Care, 41:1346  
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5. Scifres, CM, 2021, Obstet Gynecol Clin N Am, 48:325



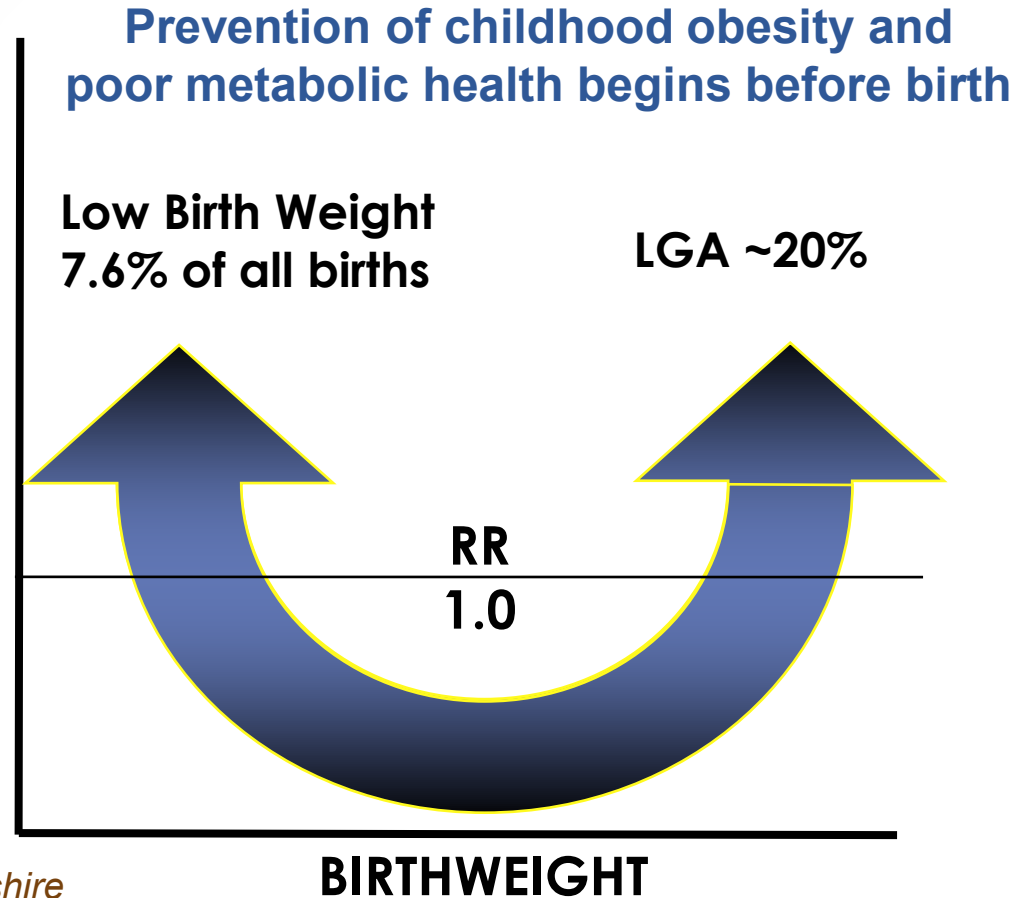
# Developmental Origins of Health and Disease:

## A “Thrifty” Phenotype

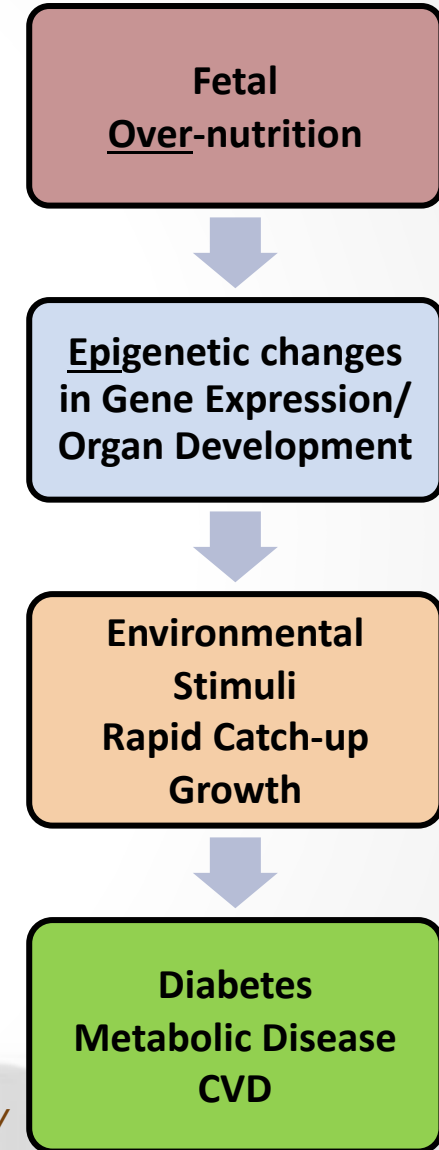


*Hertfordshire  
Helsinki  
Dutch Famine*

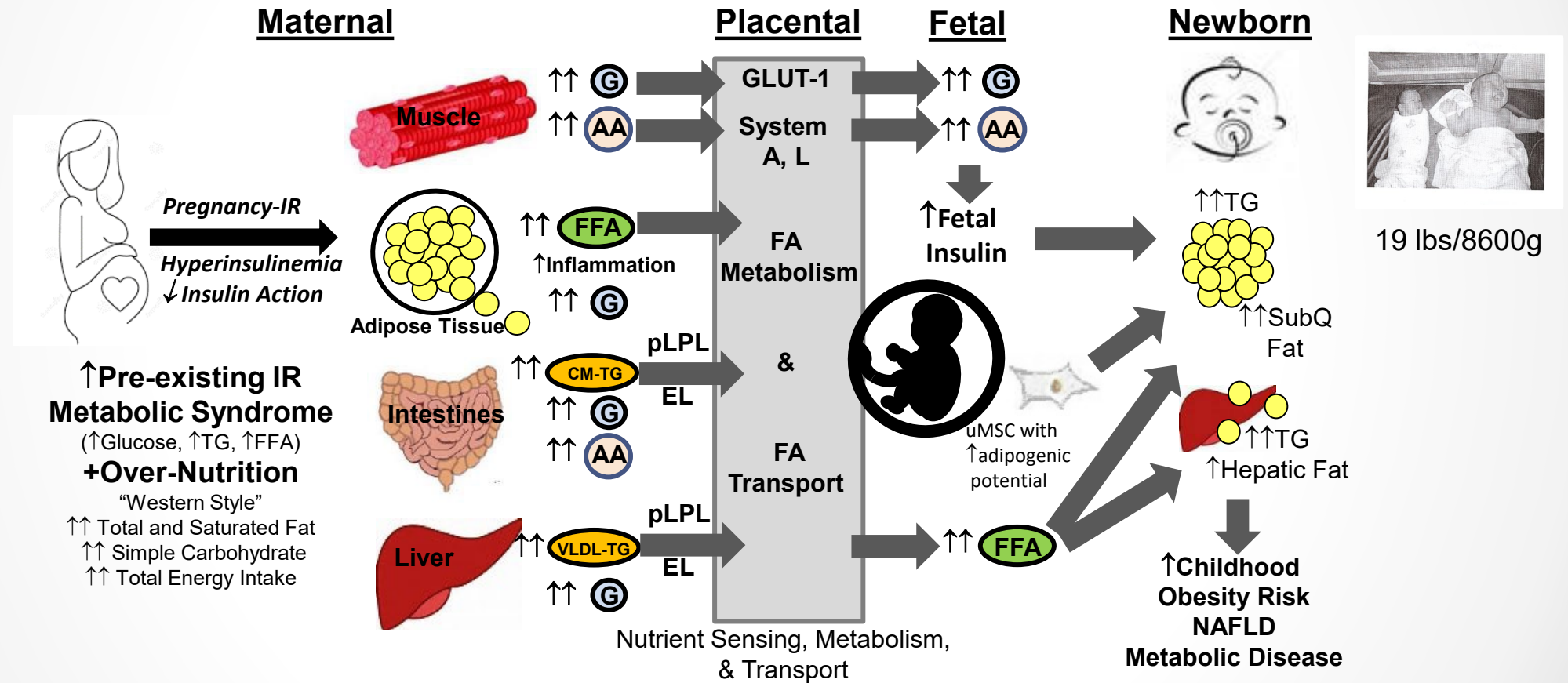
Hales & Barker, 1992, Diabetologia; 35:595  
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*Pima Indians  
NW University*



# Maternal Nutrition: Primary Intervention for Manipulation of Gestational Metabolism







Mom's garden  
Hoffman Estates, Illinois  
June 2024

# The CHOICE Diet Randomized Trial

Highlights of Clinical Trial Results

...

"...the placenta and  
the fetus develop in  
an incubation medium  
that is wholly derived  
from maternal fuels."  
-Norbert Freinkel  
1980 Banting Lecture

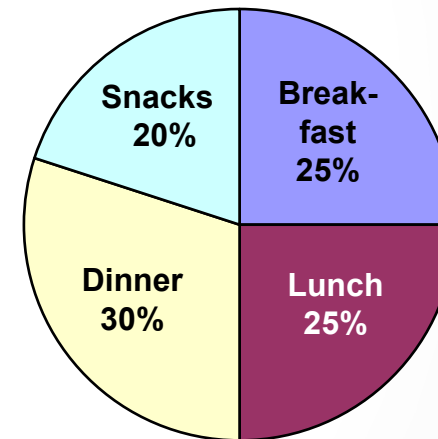


# Nutrition Arms:

## All Meals were Provided through CCTSI Bionutrition Core

- **CHOICE™ = Choosing Healthy Options  
In Carbohydrate Energy**
  - 60% carb, mostly complex
  - 25% fat
  - 15% protein
- **LC/CONV = Conventional Low Carbohydrate**
  - 40% carb
  - 45% fat
  - 15% protein
- **Both diets**
  - Eucaloric
  - SFA- 35-45%; MUFA- 35-45%; PUFA- 15-20%
  - **Simple Sugars: fixed at 70±5g in both diets**
  - Carbs are 'complex:' low-moderate glycemic index
  - Fiber is similar (~24g/day in LC, ~29g/day in CHOICE)

**Caloric Distribution**





# Hypothesis

7-8 weeks of CHOICE (60% carbohydrate, mostly complex) vs. a Conventional low carbohydrate diet (40%) would improve glycemia, insulin resistance, reduce free fatty acids (FFA), and reduce newborn adiposity



Hernandez TL, R01 DK101659

# Diabetes Care.



## Randomization to a Provided Higher-Complex-Carbohydrate Versus Conventional Diet in Gestational Diabetes Results in Similar Maternal 24-Hour Glycemia and Newborn Adiposity

Teri L. Hernandez, Sarah S. Farabi, Bailey K. Fosdick, Nicole Hirsch, Emily Z. Dunn, Kristy Roloff, John P. Corbett, Elizabeth Haugen, Tyson Marden, Janine Higgins, Jacob E. Friedman, and Linda A. Barbour

Diabetes Care 2023;46(11):1–10 | <https://doi.org/10.2337/dc23-0617>

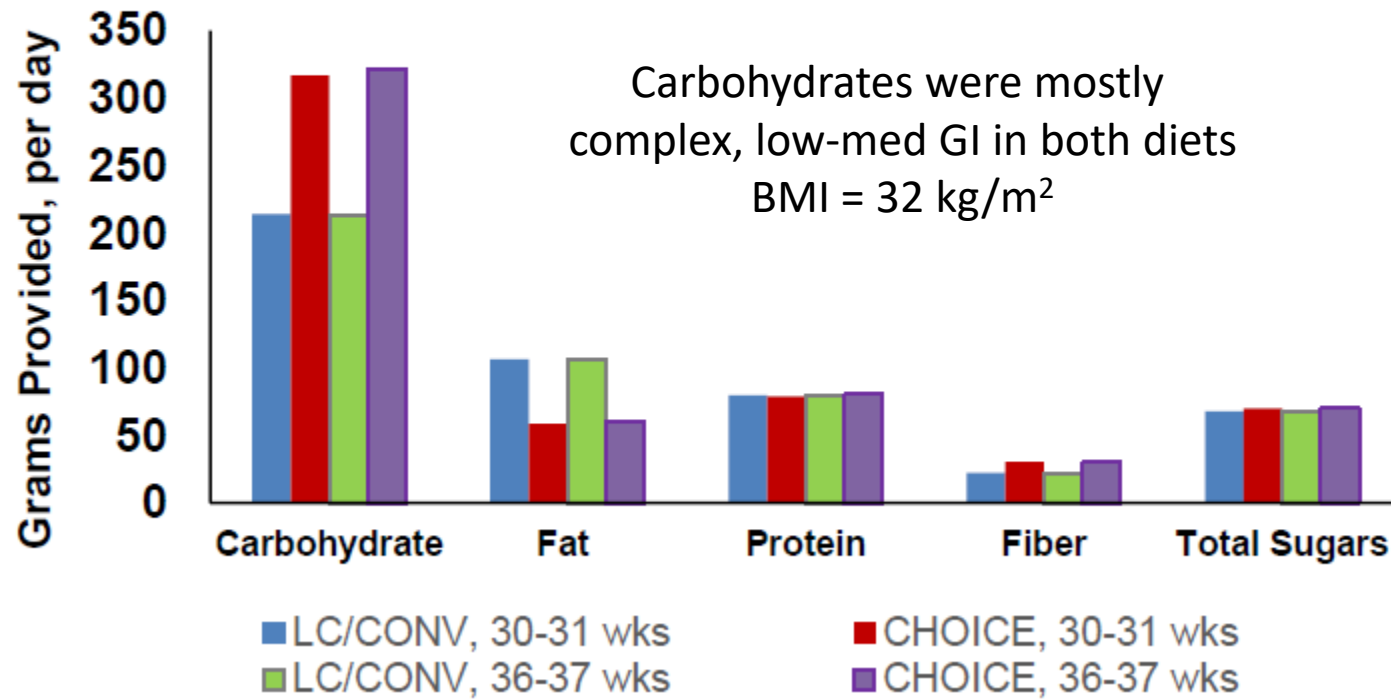


*Newborn body  
Composition  
by PEAPOD*

# RCT: 7-8 wks, All Meals Provided

## Lower Carbohydrate vs. Higher Carbohydrate

**Conventional: 40% CHO, 45% Fat, 15% Pro**  
**CHOICE: 60% CHO, 25% Fat, 15% Pro**



CHO = Carbohydrate

	40% CHO	60% CHO
Carbohydrate, g/d	214	316
Fat, g/d	106	59
Protein, g/d	80	79
Calories, kcal/d	2101	2098
Saturated Fat, g/d	34	18
Sugars, g/d	68	70







At Randomization, ~31 wks

Eucaloric Diets

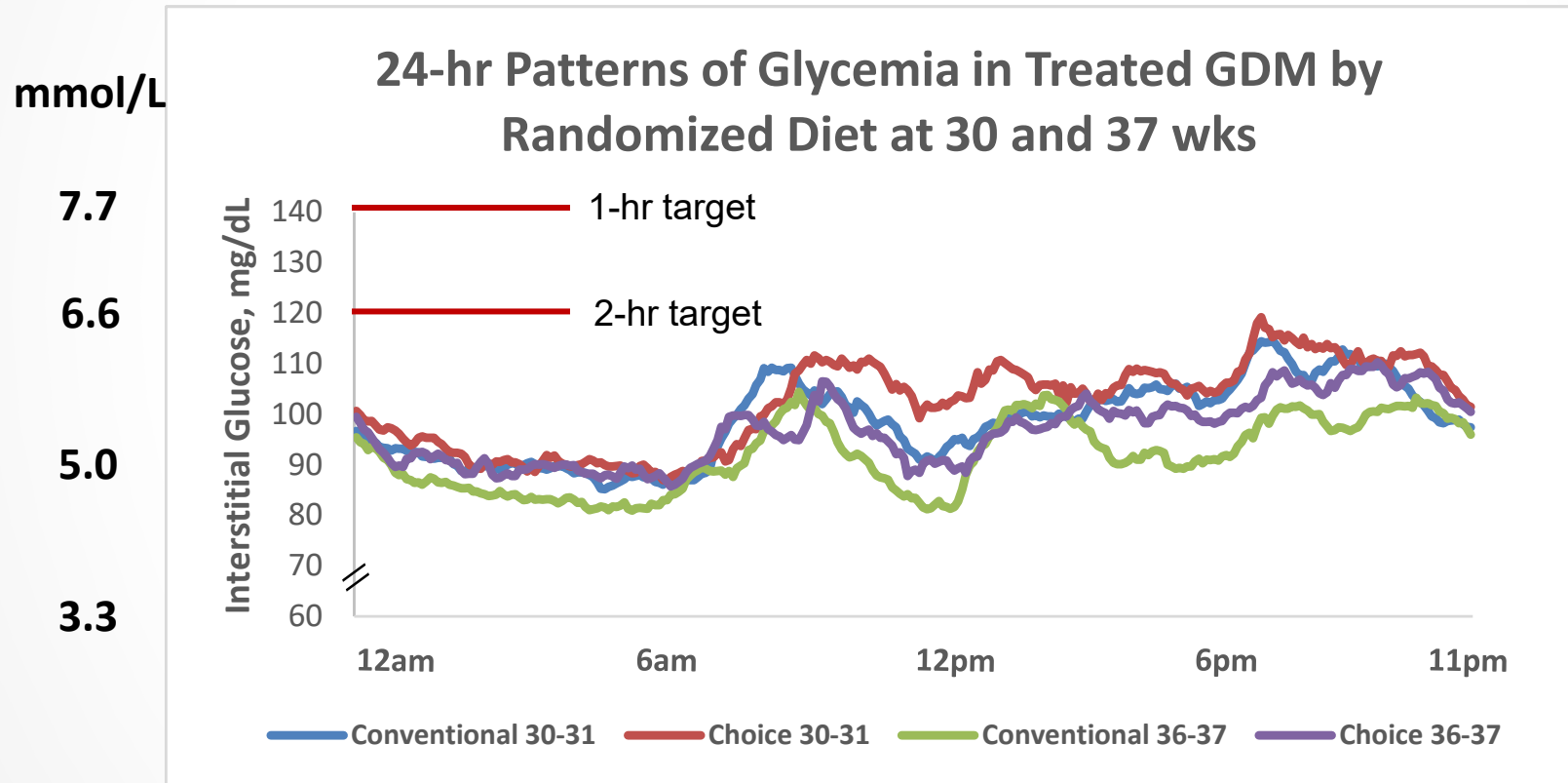
**Adherence >90%, both groups**



# Participant Characteristics

Women with A1 GDM, Baseline		Mean±SEM
	CONVENTIONAL	CHOICE
N=59 randomized 	28	31
Required Medication/Pregnancy Complication Exclusion	2/3	1/6
Completed the study 	23	23
Caucasian/Hispanic/Asian/Black, %	78/17/13/9	74/17/17/0
Weeks gestation, study baseline	31±0.1	32±0.1*
Age, yrs	32±1	33±1
BMI, pre-pregnancy, kg/m <sup>2</sup>	28±1	30±1
BMI, kg/m <sup>2</sup> , study baseline 	32±1	32±1
Gravida/Para	2/1	2/1
Characteristics at Delivery		
Gestational Age, delivery, wks 	39±0.2	39±0.3
Vaginal/C-Section delivery, %	78/4	61/17
BMI at delivery	32±1	34±1
Delta weight (delivery-study baseline), kg 	1.8±0.3	2.0±0.4
Total Gestational Weight Gain, kg	11±1.2	10±0.9
Days on Diet 	54±2	48±2

# No Between-Group Differences in 24-hour Glycemia



**40% CHO diet:**  
(214g/day)

**60% CHO diet:**  
(316g/day)

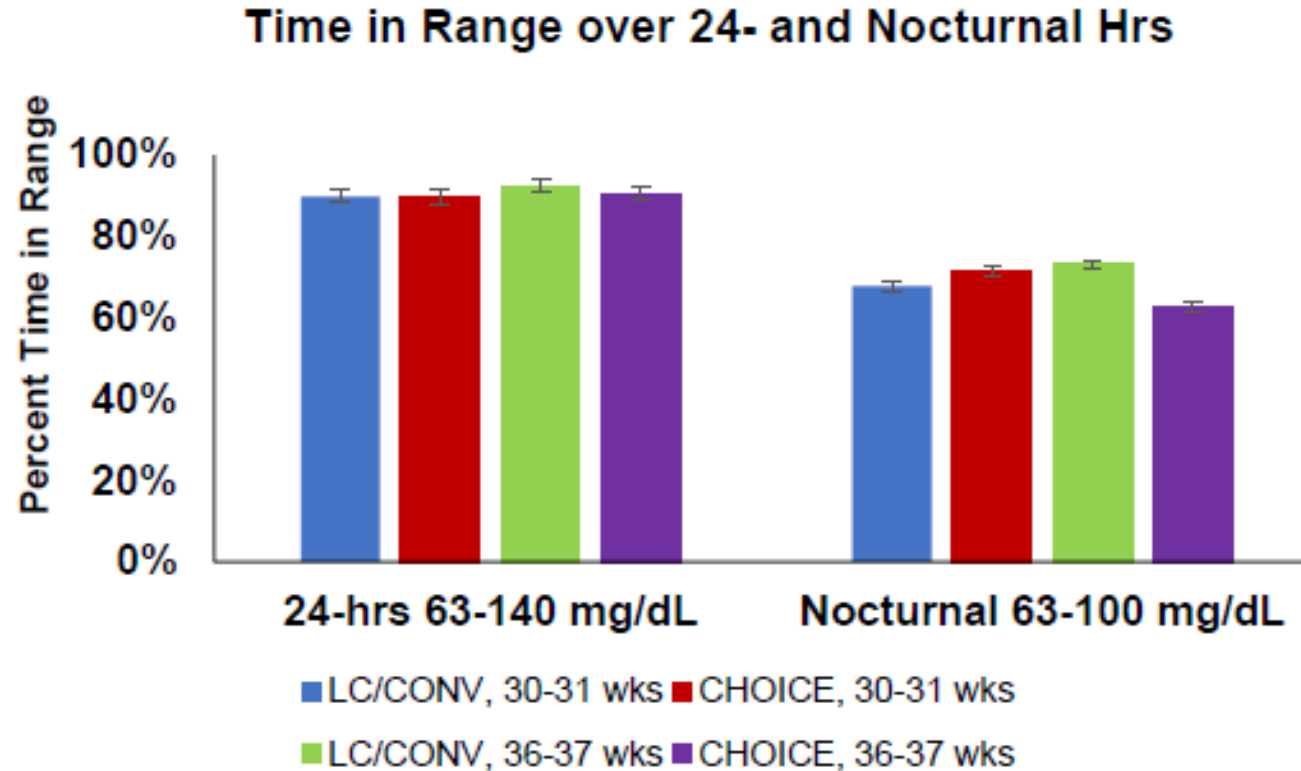
Mean  $\pm$  SEM  
All between-group  
Comparisons,  $>0.05$

No increases in glycemic  
measures over time with  
 $\uparrow$ insulin resistance

Hernandez, TL, Diabetes Care,  
2023, 46:11 1-10



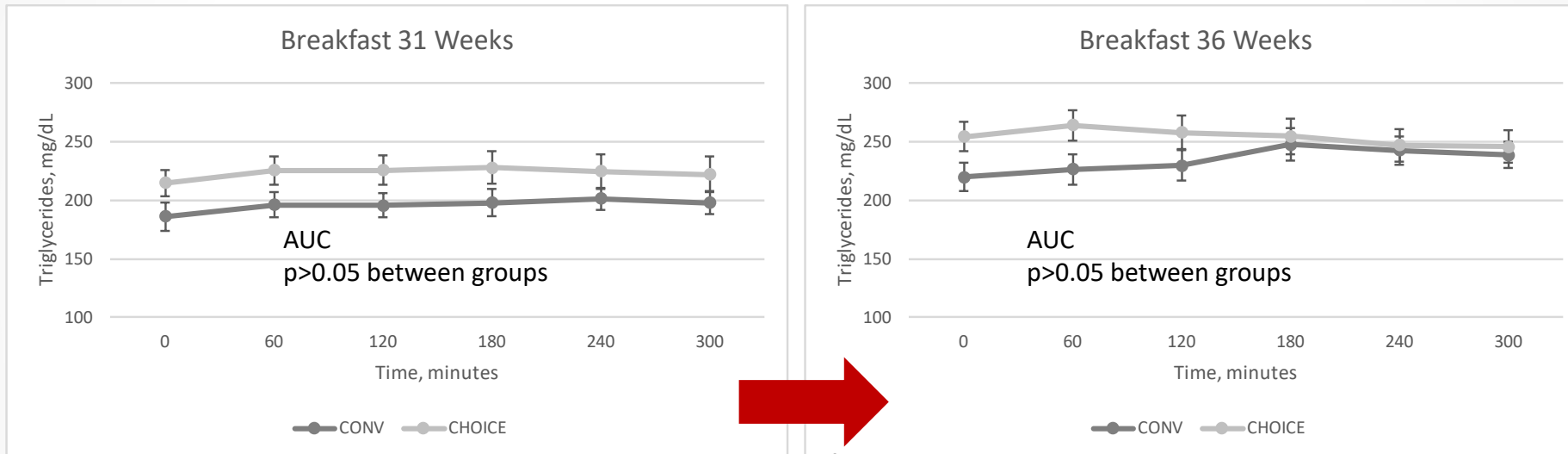
# No Between-Group Differences: %Time in Range



**40% CHO-45% Fat  
(214g/d)  
vs.  
60% CHO-25% fat  
(316g/d)**

# Similar Increase in Triglycerides on both Diets

## 40% CHO-45% Fat vs. 60% CHO-25% fat



Fasting TG:  
CONV: 186  
CHOICE: 217

Women on CHOICE  
at higher TG at study baseline

7-8 wks

Fasting TG:  
CONV: 219  
CHOICE: 257

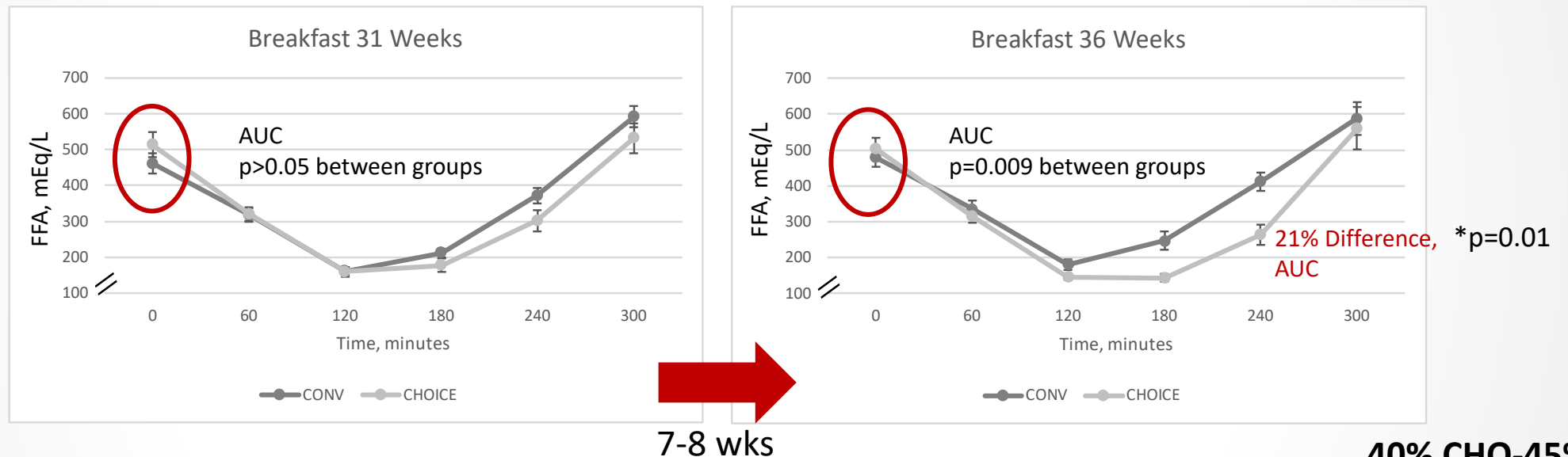
Women on CONV had 5% higher  
Increase in TG over time ( $p>0.05$ )  
—*clinically significant?*

Breakfast test meals  
30% of total energy intake  
Mean $\pm$ SEM

Fasting TG increased by ~33-40 mg/dL  
in both groups ( $p<0.001$  for both, within group change)

# Lower Postprandial FFA

## Higher Complex Carbohydrate Diet after 7-8 wks



Breakfast test meals  
30% of total energy intake  
Mean  $\pm$  SEM

FFA AUC increased on the Conventional diet and decreased on CHOICE, resulting in 21% decreased FFA exposure on Choice compared to CONV

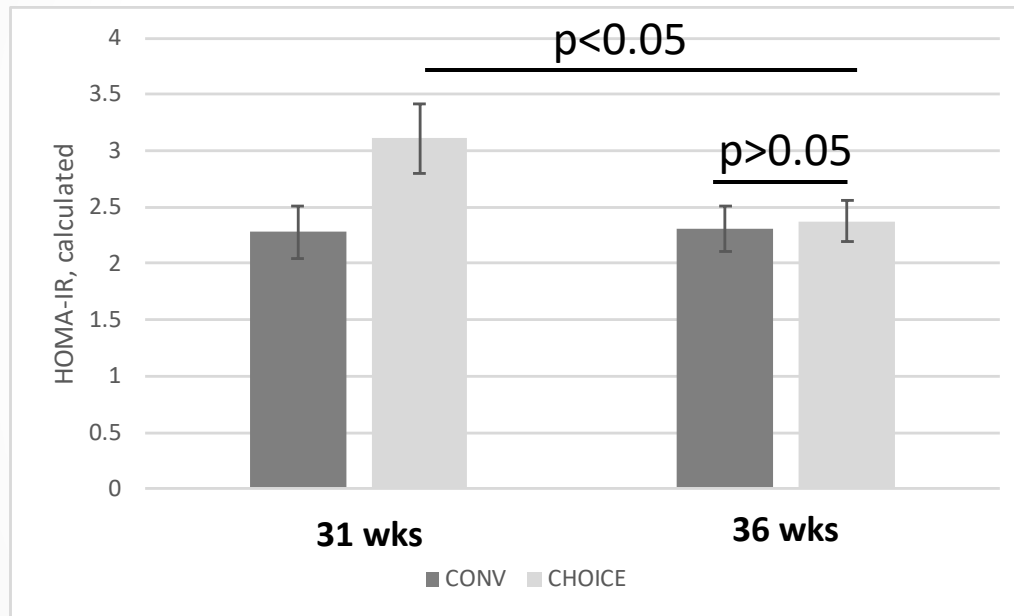
**40% CHO-45% Fat  
(214g/d)  
vs.  
60% CHO-25% fat  
(316g/d)**



R01 DK101659



# Improved Insulin Resistance by HOMA-IR on Choice by 36-37 wks Weeks



Fasting plasma glucose/insulin, OGTT day

## Insulin Resistance by Matsuda Index

	31 Wks	36 Wks
CONV	3.1±0.2	3.1±0.3
CHOICE	2.5±0.2	2.8±0.2

Higher HOMA-IR indicates ↑ insulin resistance  
Higher Matsuda Index indicates ↑ insulin sensitivity

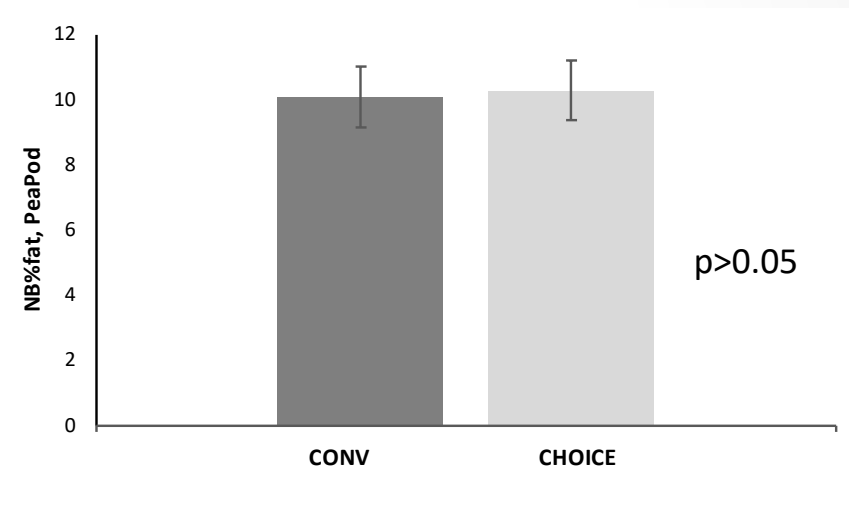


# Primary Outcome: Similar Neonatal Adiposity and Cord Blood Insulin between Diets

	Conventional	CHOICE
Birth weight, g	3303±470	3293±389
Body fat, %	10.25±4.5	10.8±4.3
LGA, %	17	13
SGA, %	4	0
Gender (M/F)	11/12	11/12
TG, mg/dL, cord	48±24	43±12
Glucose, mg/dL, cord	77±21	83±28
Insulin, uIU/L, cord	6.6±5	7.2±3
C-peptide, ng/mL, cord	0.69±0.27	0.83±0.35

Mean±SD

LGA=large-for-gestational age; SGA=small-for-gestational age



Primary Outcome

**These data reveal implications for eucaloric conditions**

# Implications ... for Long-Term Metabolic Health



“Top of the World”  
Pike’s Peak: 14,115 feet  
Colorado Springs, Colorado  
August 2023



# Choices for Women can support a *Food as Medicine* Approach

**TABLE 2**

## Recommended reproductive diet patterns

Name	Includes	Excludes	Benefits	Risks
Optimal diets				
Mediterranean diet	Plant-based foods—vegetables, fruits, whole grains, legumes, nuts, herbs, spices, olive oil, fish, poultry, and red wine Up to 40% calories from fat	Limits red meat a few times per month	Reduces the risk of CVD, mortality, cancers, and cognitive diseases	
Dietary approaches to stop hypertension	Balanced complex carbohydrates (58%), lower fat (28%), and moderate protein (18%) High in fiber, calcium, phosphorus, magnesium, and potassium	Low in cholesterol, fat, and sodium	Reduces weight, lowers BP and cholesterol, and reduces the risk of CVD and bone loss	Needs vitamin D supplementation
Flexitarian diet	Vegetarian most of the time, more vegetables, whole grains, plant-based or nonmeat proteins ("new meat"), dairy, and "sugar and spice" Focus on home prepared food with < 5 ingredients	Meat and dairy in moderation if at all	Lowers BP and cholesterol, reduces weight, and reduces the risk of heart disease, stroke, and diabetes mellitus	May need calcium, vitamin B12, and iron supplementation
Nordic diet	Fruits, vegetables, legumes, potatoes, whole grains, nuts, seeds, rye bread, fish, seafood, low-fat dairy, herbs, spices, and canola oil	Rare red meat and animal fats No sugar-sweetened beverages, added sugars, processed meats, and refined fast foods	Reduces weight and lowers BP and inflammatory markers	
Diets to avoid during pregnancy				
Atkins diet 20-40-100	Low carbohydrate (20 g), high fat, beef, pork, poultry, fish, eggs, cheese, and sources of fat	Limit starchy vegetables, grains, legumes, simple sugars, and milk		Needs vitamin C, B vitamins, folate, calcium, and magnesium
Paleo diet	Lean meats, fish, eggs, nuts, seeds, fruits, vegetables, and oils	Processed foods, wheat, other grains, legumes, dairy, potatoes, refined sugar, salt, and refined oils	Reduces weight and lowers the risk of diabetes mellitus, heart disease, and cancer	Needs calcium, B vitamins, and whole grain nutrients
Ketogenic diet	Extreme carbohydrate restriction ketosis, skin-on poultry, fattier beef, pork, fish, green leafy vegetables, oils, and solid fats	Avoid starchy root vegetables, bread, pasta, other grains, and fruit	Reduces weight	Not recommended in pregnancy because of altered neonatal brain development Needs vitamin C, B vitamins, folic acid, calcium, and fiber

BP, blood pressure; CVD, cardiovascular disease.

Marshall. Nutrition in pregnancy: lifelong consequences. Am J Obstet Gynecol 2021.

**TABLE 3**

## Common ground for healthy dietary patterns

- Whole, unprocessed foods and beverages
- Rich in fruits and vegetables
- Whole grains and complex carbohydrates, including ancient grains
- Healthy fats (monounsaturated and polyunsaturated), including nuts and seeds
- Healthy fish
- Plant-based protein
- Drink more water
- Lean meats and dairy products

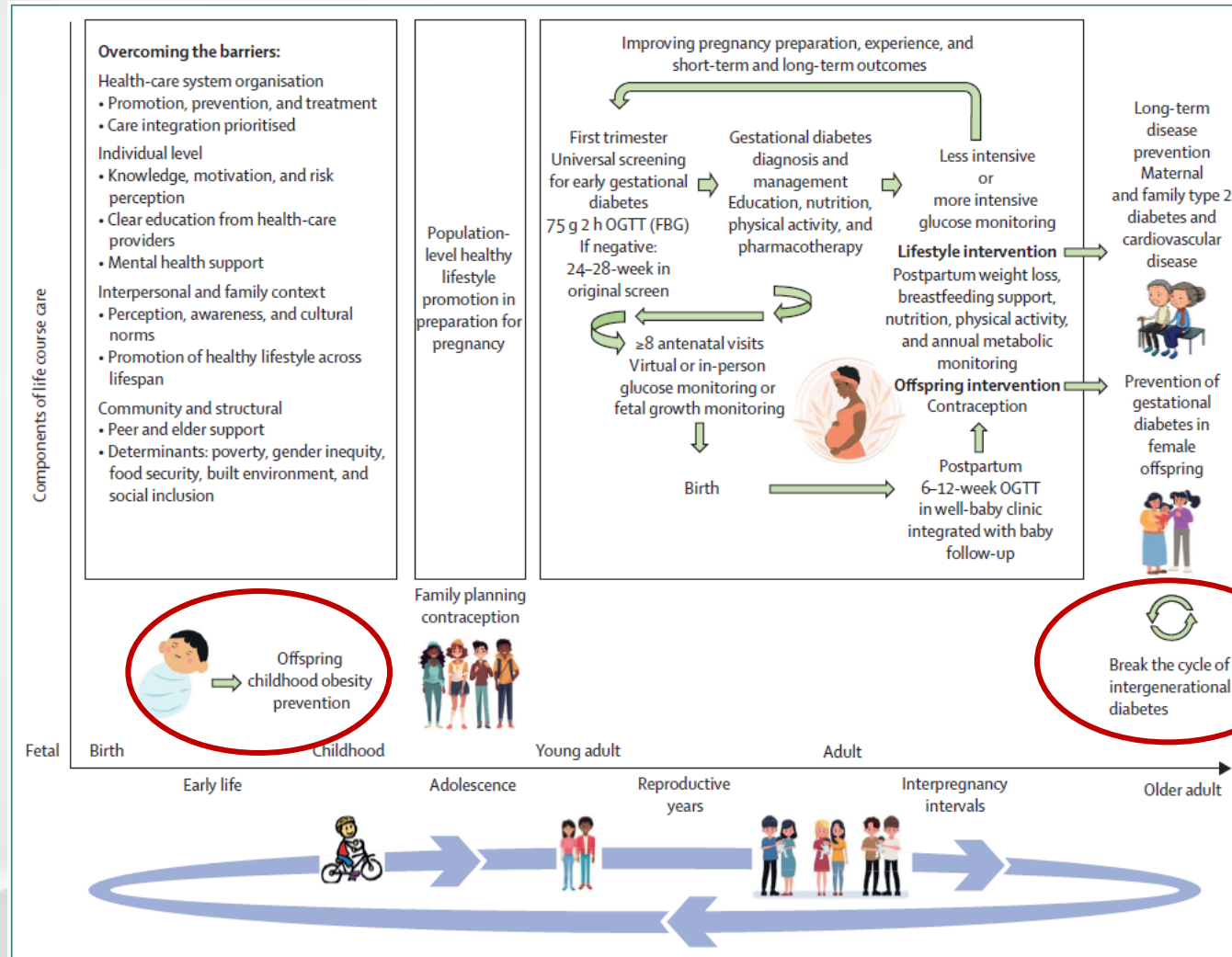
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# A Less Restrictive Carbohydrate Approach is Culturally Adaptive

**Table 1 – Local/traditional foods from varied geographical regions that qualify for a less carbohydrate-restricted dietary approach in GDM.**

Geographical region	Examples of traditional foods that qualify for less restricted GDM Nutrition therapy approach
North America	<ul style="list-style-type: none"> <li>– Whole grain breads, pasta, brown or parboiled rice, oats</li> <li>– Vegetables, fruits, beans, lentils</li> <li>– Low-fat dairy, lean poultry and fish</li> <li>– Occasional meats, cheese and nuts</li> </ul>
Latin America	<ul style="list-style-type: none"> <li>– Whole grains like amaranth, maize, quinoa, brown rice</li> <li>– Vegetables, fruits, beans</li> <li>– Lean poultry, fish, low-fat dairy</li> <li>– Occasional meats, nuts and cheese</li> </ul>
Mediterranean Region	<ul style="list-style-type: none"> <li>– Whole grain bread/pasta, brown rice, couscous</li> <li>– Vegetables, fruits, beans, lentils</li> <li>– White fish, lean poultry, low-fat dairy</li> <li>– Occasional nuts, cheese, meats and shellfish</li> </ul>
Africa	<ul style="list-style-type: none"> <li>– Whole grains like millets, sorghum, teff, parboiled rice</li> <li>– Vegetables, fruits, roots, tubers, beans</li> <li>– Fish, eggs, poultry</li> <li>– Occasional meats and dairy</li> </ul>
South Asia	<ul style="list-style-type: none"> <li>– Whole wheat, millets, barley, rye, buckwheat, parboiled rice, wheat rotis</li> <li>– Vegetables, roots, tubers, fruits</li> <li>– Beans, lentils, dals, low-fat dairy, lean poultry, fish</li> <li>– Occasional meats, nuts and cottage cheese (paneer)</li> </ul>
East Asia	<ul style="list-style-type: none"> <li>– Noodles and brown rice</li> <li>– Soybeans, fish, seafood, vegetables, wild plants, seaweed, mushrooms</li> <li>– Occasional lean meats, shellfish, dairy</li> </ul>

# A Lifecourse Approach to Prevention of Diabetes and Cardiovascular Disease is Urgently needed



**Consistent Risk Factors for childhood overweight/ Obesity can be addressed through nutrition in pregnancy<sup>1</sup>**

- ↑ pre-pregnancy BMI
- Prenatal tobacco exposure
- Excess GWG
- Higher neonatal birth weight
- Higher neonatal adiposity
- Accelerated infant weight gain

Healthy nutrition is a key component to lifecourse prevention of diabetes in mother, offspring and families

Remember the U-shaped curve



# In Defense of Carbs: Implications Moving Forward

In our Colorado approach to prevention of fetal overgrowth:

- First controlled data in humans to support that liberalization of higher complex carbohydrate results in:

- Glucose that meets treatment targets

- Lower FFA

- No difference in neonatal %fat or cord insulin levels

- Expanding nutrition options in GDM***

- **Liberalization of high-quality carbohydrates in GDM supports long-term healthy lifecourse development trajectories**

- For women, children, and families

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# Thank you!



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**Infant  
GOLD**