

# **Delayed Lactogenesis II in Women with Gestational Diabetes Mellitus**

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# GDM Background: Maternal

- GDM has been increasing globally, overall rates ~10%
- Early postpartum, 5-10% develop type 2 diabetes (T2D)
- Relative risk of developing T2D among with GDM ranges in the USA from 3.9-6.6 (some countries have higher relative risks)
- Higher intensity and longer duration of breastfeeding associated with decreased risk of T2D in women with GDM for 2 years postpartum

# Background: Glucose Homeostasis

- Brain depends on glucose for fuel
- Early breastfeeding initiation ( $\leq 1$  hour of birth) supports infant glucose levels
- When infant is “stressed,” glucose is utilized too fast
- AAP recommends screening and managing high-risk infants in the first 24 hours (such as infants born to diabetic mothers). If borderline:
  - 0-4 hours 25-40 mg/dL consider management
  - 4-24 hours 35-45 mg/dL consider management
- Increased risk of maternal-infant separation in GDM dyads which precludes early, unrestricted breastfeeding

# GDM Background: Infant

- Maternal prenatal hyperglycemia →
- Fetal hyperinsulinism →
- Neonatal hyperinsulinism →
- Result: Neonatal hypoglycemia
  - Risk of complications: infant seizures, neurologic damage, coma

# Effects of early breastfeeding on glucose levels of infants born to GDM women

- **Methods:** Prospective pilot study of 76 infants born to GDM women comparing glucose outcomes in infants based upon feeding differences.
- **Conclusions:** Early breastfeeding facilitates glycemic stability in infants born to GDM women. Human milk is appropriate for infant feeding compared to formula for infants born to GDM women.

	Breastfed	Not Breastfed	P
Borderline hypoglycemia at mean 1.6 hours	10% (4)	28% (11)	0.05 (Fisher's)

# Review of Qualitative Literature: GDM & Pregnancy

10 primary qualitative research studies published in 2005-2011 examining women's perspectives into their experiences with GDM were reviewed (focus groups, interviews, and surveys).

- **Locations:** USA-6, Australia-2, Sweden-1, Canada-1
- **Sample:** Women with GDM (also studies that included GDM with type 2 diabetes)
  - Number of participants ranged from 8-228.
  - Ethnic diversity (including immigrant and minority women)

# Concern for Infant Specific to Diabetes

- **Pregnancy: concern for infant**
  - Anxiety, worry, and emotional distress over infant health
  - Self-blame, guilt
  - Medicalized prenatal care and delivery with intervention (induction, forceps, cesarean delivery, NICU admission)
- **Postpartum: concern for infant**
  - Guilt especially when infant needed additional monitoring, tests, care, and NICU admission
  - Concern about infant's current (hypoglycemia)
  - Concern about infant's future health (risk of diabetes)

# What do women with GDM experience with early breastfeeding?

- **Design:** Qualitative phenomenological study
- **Method:** Focus groups and interviews of 27 women who had GDM and had attempted to breastfeed
- **Purpose:** To examine the meaning of the lived experience of early breastfeeding for postpartum women who had GDM.

## Themes:

- Breastfeeding challenges and support
- Milk supply problems
- Concern for infant health





# Delayed Lactogenesis II: Diabetes

- Delayed lactogenesis II in women with diabetes (T1D)
  - Arthur, Smith, Hartmann. (1989). Milk lactose, citrate, and glucose as markers of lactogenesis in normal and diabetic women. *J Pediatr Gastroenterol Nutr*, 9, 488-496.
  - Neubauer, Ferris, Chase, et al. (1993). Delayed lactogenesis in women with insulin-dependent diabetes mellitus. *Am J Clin Nutr*, 58, 54-60.
  - Ostrom, Ferris. (1993). Prolactin concentrations in serum and milk of mothers with and without insulin dependent diabetes mellitus. *Am J Clin Nutr*, 58, 49-53.
  - Hartmann, Cregan. (2001). Lactogenesis and the effects of insulin-dependent diabetes mellitus and prematurity. *J Nutr*, 131, 3016S-3020S.
  - \*Matias, Dewey, Quesenberry, Gunderson. (2014). Maternal prepregnancy obesity and insulin treatment during pregnancy are independently associated with delayed lactogenesis in women with recent gestational diabetes mellitus. *Am J Clin Nutr*, 99, 115-121. [[GDM women- finally!](#)]

# Research Sabbatical in Israel



# Colostrals metabolites as indicators of lactogenesis II in women with GDM

- **Methods:** Part of a larger prospective case-control study of 67 postpartum women in an Israeli hospital, 32 with GDM and 35 without diabetes.
- For the defined first 72-hour time frame, we had 19 women with GDM and 31 without diabetes, to compare colostrals metabolite concentrations.
- Interdisciplinary, international team

# Results: Overall summary

- Significantly, higher proportion of women with GDM perceived they had delayed lactogenesis II
- Composition analysis showed significantly lower specific metabolites in colostrum of women with GDM.
- Even after excluding primipara women to reduce confounding, specific metabolites were still significantly lower.

# Results: Characteristics

**Table 1. Characteristics of women with GDM (n=32) and without GDM (n=35) and their infants.**

<u>Characteristic</u>	<u>non-GDM</u>	<u>GDM</u>	<u>p</u>
Maternal age (years)	30.4	33.0	0.022*
Parity	3.3	3.1	0.704
Cesarean delivery	22.9%	25.0%	0.837
Gestation (weeks)	39.6	39.0	0.041*
Infant birth weight (kg)	3.3	3.4	0.209
Milk coming in was delayed	6.5%	36.8%	0.018*
Neonatal hypoglycemia (BS <45 mg/dL)	0%	36.8%	0.001*
Infant fed formula in hospital	61.3%	89.5	0.050*
Maternal prepregnancy weight (kg)	63.6	74.1	0.018*
Maternal prepregnancy BMI (kg/m <sup>2</sup> )	23.8	27.7	0.011*
Maternal pregnancy weight gain (kg)	12.6	11.0	0.254

# Results: Metabolite Concentrations

Mean differences in metabolites between women without diabetes and women with GDM (31 and 19, respectively).

Metabolite	Non-diabetic	GDM	P
Lactose mM	176.7 $\pm$ 38.9	142.4 $\pm$ 49.8	0.012*
Galactose $\mu$ M	281.5 $\pm$ 251.0	367.7 $\pm$ 311.7	0.307
Glucose-6-phosphate $\mu$ M	4.5 $\pm$ 0.5	4.2 $\pm$ 0.6	0.037*
Glucose $\mu$ M	6.3 $\pm$ 0.8	5.8 $\pm$ 0.6	0.027*
$\beta$ -hydroxybutyrate <sup>1</sup> $\mu$ M	5.6 $\pm$ 0.7	6.2 $\pm$ 0.7	0.053
Lactate $\mu$ M	6.4 $\pm$ 0.6	6.8 $\pm$ 0.2	0.093
Malate $\mu$ M	6.7 $\pm$ 1.1	7.6 $\pm$ 0.7	0.103
Citrate mM	5.2 $\pm$ 1.5	3.3 $\pm$ 1.6	0.005*

# Current Research

- Collected colostrum samples and surveys from 133 postpartum women over the course of the first week postpartum (days 2-7)
- Examine differences in breastfeeding and breast milk changes between GDM and non-GDM women
- Examine correlation between maternal perception of milk “coming in” and biochemical changes indicating lactogenesis II

# Recommendations

## General GDM

- Multidisciplinary team approach
- Include patient and family
- Positively frame approach to care
- Provide access to clear, concise, evidence-based information about GDM, causes, management options, symptoms, and risks
- Present multicultural/multilingual educational materials
- Identify peer support and role models

## Breastfeeding post-GDM

- Educate about the importance of breastfeeding for mother and infant
- Encourage breastfeeding- early and frequent
- If separated, encourage pumping- early and frequent
- Refer to professional lactation assistance
- Encourage extended breastfeeding duration



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Thank you

*Questions?*

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