# **Annals of Internal Medicine**

# ORIGINAL RESEARCH

# Lactation and Progression to Type 2 Diabetes Mellitus After Gestational Diabetes Mellitus

## A Prospective Cohort Study

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**Background:** Lactation improves glucose metabolism, but its role in preventing type 2 diabetes mellitus (DM) after gestational diabetes mellitus (GDM) remains uncertain.

**Objective:** To evaluate lactation and the 2-year incidence of DM after GDM pregnancy.

**Design:** Prospective, observational cohort of women with recent GDM. (ClinicalTrials.gov: NCT01967030)

Setting: Integrated health care system.

**Participants:** 1035 women diagnosed with GDM who delivered singletons at 35 weeks' gestation or later and enrolled in the Study of Women, Infant Feeding and Type 2 Diabetes After GDM Pregnancy from 2008 to 2011.

**Measurements:** Three in-person research examinations from 6 to 9 weeks after delivery (baseline) and annual follow-up for 2 years that included 2-hour, 75-g oral glucose tolerance testing; anthropometry; and interviews. Multivariable Weibull regression models evaluated independent associations of lactation measures with incident DM adjusted for potential confounders.

**Results:** Of 1010 women without diabetes at baseline, 959 (95%) were evaluated up to 2 years later; 113 (11.8%) developed incident DM. There were graded inverse associations for lactation intensity at baseline with incident DM and adjusted hazard

ratios of 0.64, 0.54, and 0.46 for mostly formula or mixed/inconsistent, mostly lactation, and exclusive lactation versus exclusive formula feeding, respectively (*P* trend = 0.016). Time-dependent lactation duration showed graded inverse associations with incident DM and adjusted hazard ratios of 0.55, 0.50, and 0.43 for greater than 2 to 5 months, greater than 5 to 10 months, and greater than 10 months, respectively, versus 0 to 2 months (*P* trend = 0.007). Weight change slightly attenuated hazard ratios.

**Limitation:** Randomized design is not feasible or desirable for clinical studies of lactation.

**Conclusion:** Higher lactation intensity and longer duration were independently associated with lower 2-year incidences of DM after GDM pregnancy. Lactation may prevent DM after GDM delivery.

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\* For a list of the Study of Women, Infant Feeding and Type 2 Diabetes After GDM Pregnancy investigators, see the **Appendix** (available at www.annals.org).

Gestational diabetes mellitus (GDM) is a disorder of glucose tolerance affecting 5% to 9% of all U.S. pregnancies (approximately 250 000 annually) (1), with a 7-fold higher risk for type 2 diabetes mellitus (DM) (2). Strategies during the postpartum period for prevention of DM focus on modification of lifestyle behaviors, including dietary intake and physical activity, adapted from the Diabetes Prevention Program to promote weight loss (3, 4). Lactation is a modifiable postpartum behavior that improves glucose and lipid metabolism and increases insulin sensitivity (5-10) and has favorable metabolic effects that persist after weaning (11, 12).

Despite these benefits, evidence that lactation prevents type 2 DM remains inconclusive, particularly among women with GDM (5-10, 13). Two prospective epidemiologic studies, based on self-report of DM and initiation of follow-up many years after childbearing, reported weak to modest risk reductions of 3% to 15% per year of lactation among middle-aged and older U.S. nurses (14) or Chinese women (15). However, these findings are subject to reverse causation (such as

lower perinatal body mass index or greater insulin sensitivity leading to longer lactation) and potential confounding by unmeasured risk factors (such as GDM history, degree of glucose intolerance, perinatal outcomes, and lifestyle behaviors) that may determine lactation initiation and duration.

Among women with GDM, evidence that lactation prevents DM is based on only 2 studies with conflicting findings. A large, retrospective study of U.S. nurses was null (14), whereas a prospective study reported a 46% lower incidence of DM at 15 years or more after delivery with lactation of 3 months or longer (16). Although this study screened for glucose tolerance at regular intervals, statistical power was inadequate for

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# ORIGINAL RESEARCH

Lactation and Incidence of Diabetes After GDM

#### **EDITORS' NOTES**

#### Context

Lactation is a modifiable postpartum behavior that improves glucose and lipid metabolism. Whether lactation prevents type 2 diabetes mellitus (DM) among women with gestational diabetes mellitus (GDM) is unclear.

#### Contribution

Investigators classified baseline infant feeding behaviors, including lactation intensity at baseline and lactation duration during follow-up for 1035 postpartum women with GDM. The women had oral glucose tolerance tests at baseline and annually for 2 years. The primary study outcome was the development of incident DM during follow-up.

#### Caution

Women were followed for only 2 years.

#### **Implication**

Higher intensity and longer duration of lactation were independently associated with decreased incidence of DM among women with GDM.

the 264 women to estimate excess DM incidence earlier than 15 years after delivery or graded associations with lactation (16). This study also lacked measures of perinatal risk factors, lifestyle behaviors, and weight change after delivery. Thus, the entire body of evidence cannot rule out reverse causation and potential confounding from unmeasured determinants of lactation behavior. Also, weight change by 1 year after delivery has not been previously evaluated as a mediator.

To overcome previous limitations and because random assignment of women to either breastfeeding or formula-feeding groups is not desirable or feasible in clinical studies, SWIFT (Study of Women, Infant Feeding and Type 2 Diabetes After GDM Pregnancy) prospectively quantified lactation duration and intensity, assessed risk factors for reverse causation and potential confounders, and conducted standardized glucose tolerance testing 6 to 9 weeks after delivery and thereafter annually for 2 years. We hypothesized that higher intensity and longer duration of lactation prevent progression to DM within 2 years, independent of relevant risk factors and determinants of lactation success.

#### **METHODS**

#### **Study Population**

The SWIFT study design and research method have been previously described (17). In brief, 1035 women diagnosed with GDM by Carpenter-Coustan criteria (18) were enrolled from August 2008 to December 2011. They received prenatal care and delivered at Kaiser Permanente Northern California hospitals and met other eligibility criteria based on telephone screenings during late gestation and after delivery to assess

breastfeeding and formula use. The overall response rate was 55%, and 64% of the eligible respondents enrolled. Participants provided informed consent at the in-person examination at 6 to 9 weeks after delivery (baseline) before collection of blood specimens from a 2-hour, 75-g oral glucose tolerance test (OGTT); completion of surveys; anthropometric and body composition measurements; and annual in-person follow-up examinations for 2 years (October 2008 to June 2014) (17). The Kaiser Permanente Northern California Institutional Review Board approved the study protocol.

#### **Eligibility Criteria**

Eligibility criteria (Supplement, available at www .annals.org) included diagnosis of GDM, live birth at 35 weeks' gestation or later, age 20 to 45 years, no history of diabetes, English- or Spanish-speaking, no serious medical conditions, and classification of infant feeding as intensive lactation or intensive formula feeding. Women recorded daily formula feeding (amount and frequency) from delivery through 6 weeks thereafter to estimate cumulative intake of formula and other supplements (17). Telephone interviews at 4 to 6 weeks after delivery identified women who fell into an intensive lactation (≤6 oz of formula per 24 hours since delivery) or intensive formula feeding (≥14 oz of formula per 24 hours for 3 weeks, no breast milk, or previous breastfeeding and weaned in ≤3 weeks) group. We excluded 83 women who reported mixed feeding (formula supplementation of 7 to 13 oz per 24 hours) or inconsistent feeding (transition to ≥14 oz of formula per 24 hours after 3 weeks of lactation) at the eligibility screening interview within 4 to 6 weeks after delivery (17).

#### **Data Collection Procedures**

Women who met study criteria were mailed instructions to follow an unrestricted diet for 3 days preceding the OGTT and to fast at least 10 hours the night before. At each in-person examination, fasting blood specimens were collected between 7:00 and 10:30 a.m. before consumption of 75 g of dextrose (Fisherbrand). Plasma aliquots obtained from fasting and 2-hour postload blood samples were stored at -70 °C until being transported to the University of Washington Northwest Lipid Metabolism and Diabetes Research Laboratories (Seattle, Washington), which performed enzymatic assays for glucose and double-antibody radioimmunoassays for total insulin developed by the Diabetes Endocrinology Research Center Immunoassay Core Laboratory (Seattle, Washington) (5).

### Prospective Quantitative Assessment of Lactation Intensity and Duration

Trained research staff assessed lactation intensity and duration prospectively via feeding diaries, telephone calls, in-person examinations, and monthly mailed questionnaires to collect the exact date and age (months and days) of weaning (17). Lactation intensity at 6 to 9 weeks after delivery (baseline) represents the cumulative amount of formula and breast milk fed since delivery and the intensity for the past 7 days (average

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number of breast milk- and formula-feeding episodes per 24 hours; quantity of formula per feeding; and total number of feedings per day, including water, other liquids, or solids). We classified women into 1 of the 4 following lactation intensity groups at 6 to 9 weeks after delivery: exclusive lactation (no formula, foods, or liquids), mostly lactation (>0 to 6 oz of formula per 24 hours), mostly formula (>17 oz per 24 hours) and mixed (7 to 17 oz of formula per 24 hours) or inconsistent lactation pattern, and exclusive formula feeding (formula only; no breastfeeding or breastfeeding <3 weeks) since birth. Women who transitioned into the mixed or inconsistent group after eligibility screening and before enrollment (n = 101) were classified as mostly formula feeding based on similar DM incidence.

# Classification of Glucose Tolerance and Incident DM

Glucose tolerance at the baseline examination was evaluated by administration of the 2-hour, 75-g OGTT. Women with elevations that indicated a DM diagnosis at baseline confirmed on 2 separate occasions or those without repeated testing were excluded from longitudinal follow-up for incident DM.

Incident DM was identified via this test, which was administered during annual SWIFT follow-up examinations. In addition, electronic medical records from health plans were used to identify incident DM based on clinical laboratory tests (fasting, 2-hour OGTT, or random glucose or glycosylated hemoglobin); International Classification of Diseases, Ninth Revision, Clinical Modification, diagnostic codes confirmed with laboratory results; and receipt of DM medications. For participants receiving outside follow-up care, we also used self-report of DM with health care provider diagnosis. Glucose tolerance categories included normal, glucose intolerant (impaired fasting glucose level of 5.55 to 6.94 mmol/L [100 to 125 mg/dL] or impaired glucose tolerance from the 2-hour, 75-g OGTT postglucose level of 7.77 to 11.04 mmol/L [140 to 199 mg/dL]), or DM (OGTT fasting glucose levels ≥6.99 mmol/L [≥126 mg/dL], 2-hour glucose levels ≥11.10 mmol/L [≥200 mg/dL], random glucose levels ≥11.10 mmol/L [\ge 200 mg/dL], or glycosylated hemoglobin levels ≥6.5%) via the American Diabetes Association diagnostic criteria (18). Elevations after baseline that indicated a DM diagnosis were confirmed by repeated laboratory tests outside of any subsequent pregnancies.

#### Anthropometry and Adiposity Measurements

Research staff used standardized protocol and research-quality calibrated instruments at each examination to measure weight, height, and waist circumference as previously described (17). Body weight was measured in light clothing and at standing height without shoes. Waist circumference was measured midway between the iliac crest and the lowest lateral portion of the rib cage and anteriorly midway between the xiphoid process of the sternum and the umbilicus. The 2 consecutive measurements were followed by a third if these measurements diverged by 1 cm or greater (17). Weight at baseline was subtracted from weight at

1 year after delivery to evaluate changes associated with lactation.

#### **Perinatal Course and Outcomes**

Electronic medical records provided prenatal 3-hour, 100-g OGTT results; GDM treatment type (diet, oral medications, or insulin), prepregnancy or first trimester body weight; delivery weight; and perinatal (GDM diagnosis date, delivery method, and discharge diagnoses) and newborn outcomes (birthweight, length, gestational age, neonatal intensive care unit admission, medical diagnoses, Apgar score, and length of hospital stay).

#### **Lifestyle Behaviors and Depression**

Interviewer- and self-administered, validated questionnaires assessed dietary intake, depression, and physical activity at each examination. PrimeScreen, a brief questionnaire about food frequency (reproducibility of 0.70 for foods and food groups and 0.74 for nutrients), assessed dietary intake of fat, fiber, and several other nutrients (19). A slightly modified version (17) of the Pregnancy Physical Activity Questionnaire, a 32-item, semiquantitative, validated questionnaire (reproducibility measures of 0.78 to 0.93), was used to assess physical activity during the postpartum period (20). The Center for Epidemiologic Studies Depression Scale (21) was used for longitudinal evaluation of depressive symptoms to achieve consistent measurements across the 3 examinations during the 2-year study.

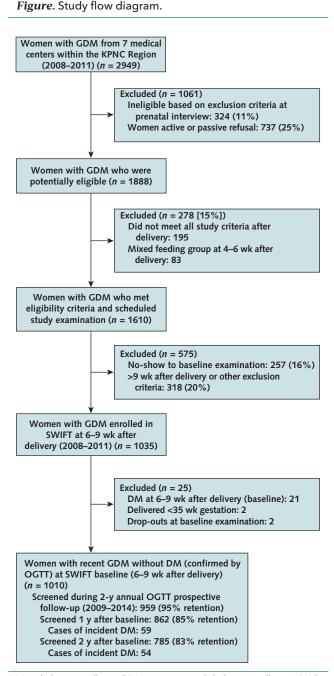
#### Reproductive, Medical, and Sociodemographic Characteristics

Data collection occurred via in-person examination interviews, monthly mailed surveys, and telephone interviews, as well as via health plan electronic medical records to obtain current medical conditions (diagnosis and treatment), medication use (including hormonal contraceptives), subsequent pregnancies and perinatal outcomes, infant health history, family history of diabetes, education, parity, participation in low-income nutrition programs, age, and race/ethnicity.

#### **Analytic Sample and Retention**

Of 1035 enrollees, 21 women with DM who had elevated glucose levels (fasting glucose level ≥6.99 mmol/L [≥126 mg/dL] or 2-hour glucose level ≥11.10 mmol/L [≥200 mg/dL]) from the 2-hour OGTT at 6 to 9 weeks after delivery (baseline) were excluded. We also excluded 2 dropouts at baseline and 2 women who delivered at less than 35 weeks' gestation. After these exclusions, 1010 SWIFT participants without DM at 6 to 9 weeks after delivery were followed and rescreened annually for 2 years to ascertain DM incidence (Figure). We identified 59 DM cases within 1 year (screened 862 of 1010 [85%]) and 54 DM cases from 1 to 2 years (screened 785 of 951 at risk [83%]). Overall, 959 (95%) participants were evaluated for glucose tolerance at least once during the 2-year follow-up (defined as <32 months after baseline): 688 at both years 1 and 2, 174 at year 1 only, and 97 at year 2 only.

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DM = diabetes mellitus; GDM = gestational diabetes mellitus; KPNC = Kaiser Permanente Northern California; OGTT = oral glucose tolerance test; SWIFT = Study of Women, Infant Feeding and Type 2 Diabetes After GDM Pregnancy.

#### **Statistical Methods**

We compared characteristics of women and newborns by incident DM status during follow-up using *t* tests and chi-square tests for continuous and categorical variables, respectively. Prenatal, postpartum, and newborn characteristics were compared across lactation intensity groups using 1-way analysis of variance for continuous variables, chi-square tests for categorical variables, and Wilcoxon rank-sum tests for medians. We used the Fisher exact test for cell sizes of less than

5. The Weibull regression model (a flexible, parametric survival model) was used for point and interval estimation of hazard ratios and 95% CIs of DM associated with lactation intensity and duration and controlled for confounding variables (22). Analyses accounted for standard right censoring due to not having DM at the last examination during follow-up and for interval censoring due to DM screening at annual study examinations or routine clinical examinations. With interval censoring, the exact failure time for a woman without DM at an examination and diagnosed at the next examination is unknown. In addition to interval censoring with unequal screening intervals, regression analyses accommodated time-varying lactation duration in which women were appropriately categorized at each point in time during follow-up according to current lactation duration status, moving into the next highest-duration category when, and if, the duration category boundary was reached. On the basis of a priori decisions (17), selected groups of baseline, perinatal, and newborn covariates were included in regression models in succession to control for potential confounding or reverse causation due to perinatal risk factors and lifestyle behaviors. Weight change was added to the final model (all covariates included) to assess its role in the mediation of the lactation-DM association. We used SAS, version 9.3 (SAS Institute), for all analyses and PARM\_ICE macro, version 3.2 (Free Software Foundation) (22), for regression analyses. A more detailed description of the modeling approach is provided in the Supplement. Tests of trends across increasing lactation intensity and duration categories were calculated by treating the ordinal categories as continuous variables in the regression models (coded as 1 through 4 across the 4 categories). Statistical significance was defined as P values (all 2-sided) less than 0.05.

#### **Role of the Funding Source**

This study was funded by the National Institute of Child Health and Human Development. The funding source had no role in the scientific conduct of this study, including the design, method, data analysis, interpretation of the results, or dissemination of findings.

#### RESULTS

Of 1010 eligible participants without DM at baseline, median follow-up was 1.8 years (range, 0.2 to 2.6 years). During the 2-year follow-up, 959 women were assessed, and 113 (11.8%) developed incident DM: 59 of 862 (6.8%) at 1 year and 54 of 782 (6.9%) at 2 years. The overall incidence rate of DM was 5.64 cases per 1000 person-months (95% CI, 4.60 to 6.68) and ranged from 3.95 cases per 1000 person-months (CI, 2.07 to 5.83) for exclusive lactation to 8.79 cases per 1000 person-months (CI, 5.47 to 12.11) for exclusive formula use at 6 to 9 weeks after delivery (Table 1) in a graded manner (*P* trend = 0.004). Of 113 DM cases, 77 were classified by 2-hour OGTTs; 16 by fasting or random glucose or glycosylated hemoglobin levels based on the American Diabetes Association diagnostic criteria

Table 1. Crude Incidence Rates (95% CIs) of Type 2 DM Within 2 y of Follow-up Among Women With GDM Pregnancy, by Lactation Intensity Groups at 6 to 9 wk After Delivery\*

Lactation Intensity Group	Sample, n	Participants With Incident DM, n	Person-Months	Incidence Rate (95% CI)†
Exclusive formula	153	27	3071.2	8.79 (5.47-12.11)
Mostly formula and mixed or inconsistent lactation	214	29	4480.5	6.47 (4.11-8.83)
Mostly lactation	387	40	8191.2	4.88 (3.37-6.39)
Exclusive lactation	205	17	4299.6	3.95 (2.07-5.83)
Total	959	113	20 042.5	5.64 (4.60-6.68)

(18) (Methods section); 11 by International Classification of Diseases, Ninth Revision, Clinical Modification, diagnostic codes and treatment with DM medications; 1 by self-report based on the health care provider's diagnosis; 1 by a 2-hour postload glucose level of 11.04 mmol/L (199 mg/dL); and 7 by fasting glucose levels of 6.88 to 69.94 mmol/L (124 to 125 mg/dL).

At baseline, women who later developed DM were heavier, of Hispanic or Asian race/ethnicity, more likely to receive insulin or oral medications to treat GDM, and more likely to deliver by cesarean section (Table 2). They also had slightly higher dietary consumption of animal fat, and physical activity scores (Table 2), as well as adverse newborn outcomes (large for gestational age and heavier birthweight), (Appendix Table 1, available at www.annals.org). Lactation intensity at 6 to 9 weeks after delivery (Appendix Tables 2 and 3, available at www.annals.org) was inversely associated with prepregnancy body mass index and neonatal intensive care unit admission and hospital stay but not with GDM treatment, gestational weight gain, severity of prenatal glucose intolerance (that is, the sum of 3-hour OGTT Z score), or history of polycystic ovarian syndrome. Lactation intensity at 6 to 9 weeks after delivery was directly associated with lactation duration (Appendix Table 4, available at www.annals.org)-the exclusive and mostly lactation groups breastfed for longer periods (P < 0.001).

In multivariable regression models (Table 3), higher lactation intensity was associated with lower adjusted rates of incident DM in a monotonic gradient adjusted for age and covariates, including maternal and perinatal risk factors, newborn outcomes, and lifestyle behaviors (all P trend < 0.025). Longer lactation duration (>2 to 5 months, >5 to 10 months, and >10 months compared with 0 to 2 months) was also independently associated with lower rates of incident DM (Table 4) in a monotonic gradient (all P trend < 0.01). Addition of newborn outcomes and lifestyle covariates to models adjusted for age and maternal and perinatal risk factors had minimal effect on observed associations. In a sensitivity analysis, exclusion of 7 DM cases classified by a fasting glucose level of 6.88 to 69.94 mmol/L (124 to 125 mg/dL) did not change our findings. Addition of weight change from delivery to 1 year thereafter, a potential mediator, modestly attenuated the associations between lactation intensity and duration with incident DM in models adjusted for age, maternal and perinatal risk factors, newborn outcomes, and lifestyle behaviors (Tables 3 and 4). However, protective associations remained statistically significant (P trend < 0.05). The results of additional exploratory analyses are presented in the Supplement.

#### DISCUSSION

Our prospective assessment of both intensity and duration of lactation and annual evaluation of type 2 DM after GDM pregnancy provides robust evidence for the role of lactation in preventing DM among high-risk young women. We saw 36% to 57% relative reductions in the 2-year DM incidence with higher intensity of lactation at 6 to 9 weeks after delivery as well as for longer periods (>2 months through >10 months), independent of obesity, gestational glucose tolerance, and perinatal outcomes that can delay lactogenesis and thereby shorten lactation duration (23). Lifestyle may also affect lactation behavior but did not confound our findings, possibly due to high breastfeeding rates among racial and ethnic subgroups, particularly Hispanic and Asian women, in whom lifestyle behaviors may not influence infant feeding choices (that is, higher breastfeeding intensity and duration may not correlate with higher physical activity levels). Our findings also show that postpartum weight change only slightly mediated the lactation-DM association, a finding that had not been evaluated in previous studies.

To our knowledge, to date SWIFT was the largest and most ethnically diverse prospective cohort of women with GDM to conduct glucose tolerance testing annually from the early postpartum period, and it is the only study that prospectively quantified lactation intensity and duration and controlled for several perinatal and newborn potential confounders. Two previous studies of lactation and DM incidence in women with a history of GDM reported conflicting findings and did not account for perinatal outcomes, subsequent pregnancies, or postpartum lifestyle behaviors (14, 16). These same limitations characterize all other studies that enrolled women with unknown GDM history and initiated follow-up for most women many years after childbearing had ended (14, 15, 24-26). Thus, findings may result from reverse causation and potential confounding due to unmeasured risk factors for suboptimal lactation. The only previous prospective study of

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DM = diabetes mellitus; GDM = gestational diabetes mellitus.

\* 959 female participants in SWIFT (Study of Women, Infant Feeding, and Type 2 Diabetes After GDM Pregnancy). P value for trend = 0.004.
† Incident DM cases per 1000 person-months.

Characteristic	Incident DM ( $n = 113$ )	No DM $(n = 846)$	P Valu
Sociodemographic			
Mean age at baseline (SD), y	33.9 (5.4)	33.3 (4.7)	0.197
Mean educational level, n (%)	, ,	, ,	0.002
High school or less	41 (36.3)	189 (22.3)	
Some college	33 (29.2)	237 (28.0)	
College graduate	39 (34.5)	420 (49.7)	
Race/ethnicity, n (%)†			0.00
Non-Hispanic white	17 (15.0)	208 (24.6)	
Non-Hispanic black	13 (11.5)	59 (7.0)	
Hispanic	46 (40.7)	246 (29.0)	
Asian	36 (31.9)	316 (37.4)	
Other	1 (0.9)	17 (2.0)	
Parity, n (%)	V/	( /	0.56
Primiparous	38 (33.6)	308 (36.4)	
Multiparous	75 (66.4)	538 (63.6)	
Polycystic ovarian syndrome, n (%)	18 (15.9)	44 (5.2)	<0.00
WIC recipient, n (%)‡	42 (37.2)	204 (24.1)	0.00
Family history of DM, n (%)	69 (61.1)	404 (47.8)	0.00
Talling history of Divi, II (70)	07 (01.1)	404 (47.0)	0.00
Prenatal	00.470.0	00.077.5	
Mean prepregnancy BMI (SD), kg/m²	33.4 (8.3)	29.0 (6.9)	<0.00
Prepregnancy weight status, n (%)	45 (40.0)	07//00//	<0.00
BMI <25 kg/m <sup>2</sup>	15 (13.3)	276 (32.6)	
BMI 25 to <30 kg/m <sup>2</sup>	30 (26.6)	257 (30.4)	
BMI ≥30 kg/m <sup>2</sup>	68 (60.1)	313 (37.0)	
Mean gestational weight gain (SD), kg Mean 3-h, 100-g OGTT results (SD)	8.1 (8.4)	10.6 (6.5)	<0.00
Fasting glucose level			< 0.00
mmol/L	5.6 (0.7)	5.0 (0.6)	<0.00
mg/dL	100.3 (13.5)	90.9 (11.5)	
1-h glucose level	100.3 (13.3)	70.7 (11.3)	< 0.00
mmol/L	11 0 /1 E)	11.0 (1.2)	<0.00
	11.8 (1.5)		
mg/dL	212.1 (26.5)	197.7 (21.9)	-0.00
2-h glucose level	10.3 (1.0)	0.7.(4.5)	<0.00
mmol/L	10.3 (1.8)	9.7 (1.5)	
mg/dL	186.4 (32.5)	175.1 (26.5)	0.00
3-h glucose level	7.5 (0.4)	7.0 (4.0)	0.00
mmol/L	7.5 (2.1)	7.0 (1.8)	
mg/dL	135.1 (38.2)	125.5 (32.3)	
Mean sum of 3-h, 100-g OGTT Z scores (SD)§	1.9 (3.1)	-0.2 (2.4)	<0.00
Mean gestational age at GDM diagnosis (SD), wk	22.3 (8.5)	25.8 (6.7)	< 0.00
Treatment of GDM, n (%)			< 0.00
Diet only	46 (40.7)	609 (72.0)	
Oral hypoglycemic agents	52 (46.0)	218 (25.7)	
Insulin	15 (13.3)	19 (2.3)	
Hospital labor and delivery			
Mean length of gestation (SD), wk	39.1 (1.2)	39.1 (1.1)	0.80
Mean length of stay (SD), d	2.9 (1.0)	2.5 (1.1)	0.00
Cesarean section, n (%)	47 (41.6)	258 (30.5)	0.01
t to 0 mile office delivery /hossiline			
6 to 9 wk after delivery (baseline)  Mean BMI (SD), kg/m²	22 4 (7 4)	20 6 (6 4)	<0.00
	33.4 (7.4)	29.6 (6.4)	
Mean waist circumference (SD), cm	97.0 (14.1)	89.0 (13.5)	<0.00
Mean postdelivery weight change (SD), kg	-8.2 (4.0)	-9.2 (3.6)	0.00
Mean HOMA-IR index score (SD)	8.5 (5.7)	5.1 (3.4)	<0.00
Median HOMA-IR index score (IQR)	7.4 (4.5–10.5)	4.1 (2.8-6.3)	<0.00
Mean 2-h, 75-g OGTT results (SD)			
Fasting glucose level			<0.00
mmol/L	5.7 (0.6)	5.2 (0.4)	
mg/dL	102.4 (10.9)	93.6 (8.1)	
2-h postload glucose level			< 0.00
mmol/L	7.4 (1.7)	6.1 (1.5)	
mg/dL	133.4 (30.4)	109.5 (26.5)	
Glucose tolerance, n (%)			< 0.00
Normal	33 (29.2)	595 (70.3)	
IFG level only	34 (30.1)	142 (16.8)	
IGT only	18 (15.9)	77 (9.1)	
IFG level and IGT	28 (24.8)	32 (3.8)	

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Table 2-Continued

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Characteristic	Incident DM ( $n = 113$ )	No DM (n = 846)	P Value
Depression, n (%)			0.68
None	91 (85.1)	710 (86.5)	
Moderate or severe	16 (14.9)	111 (13.5)	
Median lifestyle behaviors (IQR)			
Total PA, metabolic equivalent h/wk	46.5 (36.0-71.7)	42.6 (31.5-55.4)	< 0.001
Moderate to vigorous PA, metabolic equivalent h/wk	24.4 (16.6-40.7)	21.4 (14.0-30.6)	0.005
Dietary glycemic index	220.5 (150.6-280.3)	228.1 (172.0-302.1)	0.24
Dietary fiber, g/100 kcal	1.0 (0.8-1.2)	1.0 (0.8-1.2)	0.82
Dietary animal fat, percentage of kcal	26.8 (22.2-31.7)	24.9 (19.7-30.8)	0.023
Contraception, n (%)†			0.95
Progesterone only	3 (2.7)	23 (2.7)	
Combination oral pills	14 (12.4)	104 (12.3)	
Intrauterine device	12 (10.6)	79 (9.3)	
None or barrier	84 (74.3)	640 (75.7)	
Subsequent birth (1 birth during 2-y follow-up), n (%)	8 (7.1)	106 (12.5)	0.093

BMI = body mass index; DM = diabetes mellitus; GDM = gestational diabetes mellitus; HOMA-IR = Homeostatic Metabolic Assessment of Insulin Resistance; IFG = impaired fasting glucose; IGT = impaired glucose tolerance; IQR = interquartile range; OGTT = oral glucose tolerance test; PA = physical activity; WIC = Special Supplemental Nutrition Program for Women, Infants, and Children.

\* Data were missing for depression score (n = 31), HOMA-IR index score (n = 5), and lifestyle behaviors (n = 5).

‡ Eligibility based on income ≤185% of the federal poverty level.

women with GDM (n = 264) had additional limitations, including inadequate power to assess graded associations and excess risks due to formula feeding only, or to evaluate excess DM rates within 2 to 5 years after delivery.

Modest weight loss is highly effective for preventing type 2 DM. In the Diabetes Prevention Program, participants in the lifestyle intervention group had an average 3- to 4-kg weight loss over 2 to 3 years that was associated with a 58% reduction in DM incidence (3, 27). Lactation increases energy expenditure (28) and mobilizes adipose stores in the femoral region and possibly visceral fat (29, 30). However, it has only slight

effects on body composition and may have a minimal or no effect on postpartum weight loss, although evidence is mixed (9, 31-34). In our study, greater weight loss (1.0 to 1.3 kg) at 1 year after delivery for the exclusive or mostly lactation groups versus formula groups slightly mediated the association between lactation and lower DM incidence, although findings remained statistically significant. Previous studies examining the association have not evaluated postpartum weight change. Our findings indicate that the strong protective associations between lactation measures and progression to type 2 DM may involve mechanisms other than weight loss.

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Table 3. Lactation Intensity Groups at 6 to 9 wk After Delivery and Adjusted HRs of Incident DM (95% CIs) Within 2-y Follow-up Among Women With GDM Pregnancy\*

Weibull Regression Proportional Hazards Model	Adjusted HR (95% CI) of Incident DM Within 2 y of Follow-up, by Lactation Intensity Groups at 6 to 9 wk After Delivery				
	Exclusive Formula (n = 153)	Mostly Formula† and Mixed or Inconsistent Lactation (n = 214)	Mostly Lactation‡ (n = 387)	Exclusive Lactation (n = 205)	
Age-adjusted	1.00 (Reference)	0.72 (0.43-1.23)	0.54 (0.33-0.89)	0.43 (0.23-0.82)	0.003
Maternal and perinatal risk factors§	1.00 (Reference)	0.64 (0.37-1.12)	0.54 (0.32-0.92)	0.46 (0.24-0.88)	0.016
Maternal and perinatal risk factors and newborn outcomes§¶	1.00 (Reference)	0.65 (0.37-1.13)	0.53 (0.31-0.91)	0.47 (0.25-0.91)	0.017
Maternal and perinatal risk factors, newborn outcomes, and postpartum lifestyle behaviors§**	1.00 (Reference)	0.66 (0.38-1.14)	0.56 (0.32-0.95)	0.48 (0.25-0.92)	0.023
Add potential mediator: weight change from delivery to 1 y††	1.00 (Reference)	0.72 (0.41-1.28)	0.67 (0.38-1.16)	0.47 (0.24-0.93)	0.048

<sup>†</sup> Fisher exact text for cell sizes <5.

<sup>§</sup> Sum of the 4 Z scores for the prenatal 100-g OGTT glucose values (fasting and 1, 2, and 3 h).

DM = diabetes mellitus; GDM = gestational diabetes mellitus; HR = hazard ratio.

\* 959 participants in SWIFT (Study of Women, Infant Feeding, and Type 2 Diabetes After GDM Pregnancy) and 113 incident cases of DM within 32 mo after baseline. Covariate data are complete except for those for lifestyle behaviors, which were missing for 5 women, and 1-y weight change, which were missing for 35 women.

<sup>†&</sup>gt;17 oz/24 h.

<sup>‡0-6</sup> oz/24 h

<sup>§</sup> Addition of this selected group of covariates in succession to the age-adjusted model.

| Race/ethnicity: education: prepregnancy body received.

Race/ethnicity; education; prepregnancy body mass index; GDM treatment; sum of prenatal 3-h, 100-g oral glucose tolerance test Z score;

<sup>¶</sup> kaceretimicity, education, prepregnancy body mass madex, GDM treatment, sum of prenatar 3-11, 1700-g oral glacose tolerance test 2 score, gestational age at GDM diagnosis; and subsequent birth (1 vs. 0) during 2-y follow-up.

¶ Large for gestational age vs. not large for gestational age (reference), hospital stay >3 d, and neonatal intensive care unit admission.

\*\* Total physical activity (metabolic equivalent h/wk), glycemic index, and percentage of kcal from animal fat (greater than the median percentage).

†† Added to full model adjusted for age, maternal and perinatal risk factors, newborn outcomes, and postpartum lifestyle behaviors.

Table 4. Lactation Duration Categories and Adjusted HRs of Incident DM (95% Cls) Within 2-y Follow-up Among Women With **GDM Pregnancy\*** 

Weibull Regression Proportional Hazards Model	Adjusted HR (95% CI) of Incident DM Within 2 y of Follow-up, by Time-Dependent Lactation Duration				
	0-2 mo (n = 189)	>2-5 mo (n = 190)	>5-10 mo (n = 208)	>10 mo (n = 372)	
Age-adjusted	1.00 (Reference)	0.64 (0.36-1.13)	0.44 (0.22-0.88)	0.38 (0.20-0.71)	0.001
Maternal and perinatal risk factors†‡	1.00 (Reference)	0.55 (0.30-1.01)	0.51 (0.25-1.01)	0.44 (0.23-0.84)	0.009
Maternal and perinatal risk factors and newborn outcomes†§	1.00 (Reference)	0.55 (0.31-1.01)	0.50 (0.25-0.99)	0.43 (0.23-0.82)	0.007
Maternal and perinatal risk factors, newborn outcomes, and postpartum lifestyle behaviors†	1.00 (Reference)	0.54 (0.30-0.99)	0.55 (0.28-1.09)	0.43 (0.22-0.82)	0.010
Add potential mediator: weight change from delivery to 1 y¶	1.00 (Reference)	0.48 (0.25-0.90)	0.65 (0.33-1.24)	0.47 (0.24-0.91)	0.037

† Addition of this selected group of covariates in succession to the age-adjusted model.

Added to full model adjusted for age, maternal and perinatal risk factors, newborn outcomes, and postpartum lifestyle behaviors.

Potential mechanisms to explain the lower incidence of DM with higher intensity and duration of lactation include preservation of pancreatic  $\beta$  cells (35), less inflammation, and improved endothelial function; however, biochemical evidence is sparse. The hormone prolactin is known to increase pancreatic  $\beta$ -cell mass and function during human pregnancy, but effects on  $\beta$ cells during the peripartum and postpartum periods have not been delineated. In mice, a higher rate of maternal  $\beta$ -cell proliferation was reported in lactating groups than in nonlactating groups (36). In postpartum women with recent GDM, lactation enhances  $\beta$ -cell compensation for insulin resistance, resulting in better insulin sensitivity, glucose effectiveness, and first-phase insulin response to glucose according to the Bergman Minimal Model (9). In SWIFT, we previously reported an inverse association for lactation intensity and fasting blood lipids, glucose, and insulin resistance as well as the prevalence of prediabetes at study baseline independent of body mass index, race, or other risk factors

Prolactin also regulates adipogenesis to suppress lipid storage as well as release of the inflammatory cytokine, interleukin-6 (37), and adiponectin from adipocytes (38). In SWIFT, lower fasting plasma adiponectin and leptin levels were associated with higher lactation intensity at baseline. As a risk factor for type 2 DM, this finding is consistent with prolactin's and possibly leptin's suppression of adiponectin release (37, 39) but conflicts with lower circulating adiponectin levels as a risk factor for type 2 DM (40). In other crosssectional studies that measured biomarkers once at 3 years after delivery, there was no association between lactation duration and nonfasting levels of blood glucose, lipids, adiponectin, leptin, or inflammatory markers (41, 42). Longitudinal changes in biomarkers from early postpartum to postweaning periods have not been directly linked with lactation and subsequent type 2 DM. These measurements are required to determine

the biological plausibility for lactation's protection against type 2 DM after GDM delivery.

Limitations of the SWIFT study include the lack of longitudinal biomarker measures as mediators of lactation and progression to DM and the inability to evaluate the associations beyond 2 years of follow-up. However, SWIFT has several strengths, including recruitment of the largest and most racially or ethnically diverse cohort of women with standardized criteria for GDM diagnosis and treatment within a large integrated health care system; detailed prospective and quantitative assessment of lactation intensity and duration; annual screening for incident DM from the early postpartum period and annually thereafter for 2 years via the 2-hour, 75-g OGTT; research-quality measurements of body anthropometry; and high retention rates (17). Other strengths include evaluation of GDM severity, maternal risk factors, newborn outcomes, and postpartum lifestyle behaviors that can lead to delayed lactogenesis or shorter duration of lactation (23). By design, SWIFT minimized the reverse causation and residual confounding that lessen the validity of all previous studies. The SWIFT cohort was very diverse: 75% were Hispanic, Asian, or black, and 25% had household incomes that fell at or below 185% of the U.S. federal poverty guidelines. This contrasts with almost all previous studies, which predominantly included women of European ancestry, except a study of older Chinese women, and heightens the relevance of our findings for Hispanic, Asian, and black women at highest risk for GDM.

Women with a history of GDM are faced with an extremely high risk for type 2 DM; up to 50% diagnosed within 5 years after delivery (2, 43). In our study, both higher lactation intensity and duration showed strong, graded protective associations with DM incidence independent of risk factors (sociodemographic characteristics, prenatal metabolic status and course, perinatal outcomes, and lifestyle behaviors) that were

DM = diabetes mellitus; GDM = gestational diabetes mellitus; HR = hazard ratio.
\* 959 participants in SWIFT (Study of Women, Infant Feeding, and Type 2 Diabetes After GDM Pregnancy) and 113 incident cases of DM within 32 mo after baseline. Covariate data are complete except for those for lifestyle behaviors, which were missing for 5 women, and 1-y weight change, which were missing for 35 women.

<sup>‡</sup>Race/ethnicity; education; prepregnancy body mass index; GDM treatment; sum of prenatal 3-h, 100-g oral glucose tolerance test Z score; gestational age at GDM diagnosis; and subsequent birth (1 vs. 0) during 2-y follow-up.

§ Large for gestational age vs. not large for gestational age (reference), hospital stay >3 d, and neonatal intensive care unit admission.

Total physical activity (metabolic equivalent h/wk), glycemic index, and percentage of kilocalories from animal fat (greater than the median

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not explained by weight loss. Thus, early DM strategies for postpartum women may need to refocus efforts to support optimal lactation intensity and duration, with later implementation of dietary and physical activity modifications after the period of intensive lactation. The coordinated and sequential timing of postpartum interventions among GDM women may involve several biological pathways and thereby maximize DM risk reduction.

The American Academy of Pediatrics recommends exclusive breastfeeding for approximately 6 months, followed by continued breastfeeding with complementary foods for 1 year or longer (44). International professional bodies have recommended breastfeeding for women with GDM but have acknowledged that evidence was insufficient to conclude that it conferred longer-term metabolic benefits for women (45). Currently, only 43% of U.S. women report exclusively breastfeeding at 3 months, and by 6 months, only 51% are breastfeeding at all (46). Breastfeeding promotion may be a practical, low-cost intervention during the postpartum period to prevent diabetes in high-risk women, with the potential for benefits that are complementary to lifestyle interventions targeting weight loss. Modification of lactation behaviors to increase intensity and duration should be considered a high priority for pregnant and postpartum women with GDM because of their lasting metabolic benefits. Greater allocation of health care resources to promote and support exclusive and extended breastfeeding may benefit high-risk women by reducing their risk for midlife progression to DM.

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Appendix Table 1. Characteristics of Infants Born to Women With GDM, by Incident DM Status Within 2-y Follow-up After **GDM Pregnancy\*** 

Characteristic	Incident DM ( $n = 113$ )	No DM $(n = 846)$	P Value
Mean birthweight (SD), g	3546 (576.9)	3390 (484.8)	0.002
Mean length (SD), cm	51.0 (2.5)	50.5 (2.4)	0.026
Mean weight for length Z score (SD)	-0.2 (1.5)	-0.3 (1.4)	0.52
Median weight for length Z score (IQR)	-0.3 (-1.1 to 0.7)	-0.3 (-1.2 to 0.5)	0.47
Sex, n (%)			0.145
Female	46 (40.7)	406 (48.0)	
Male	67 (59.3)	440 (52.0)	
Gestational age, n (%)			0.38
35 to <37 wk (preterm)	7 (6.2)	37 (4.4)	
≥37 wk (term)	106 (93.8)	809 (95.6)	
Birthweight, n (%)†			0.21
1500-2499 g	3 (2.7)	23 (2.7)	
2500-2999 g	17 (15.0)	159 (18.8)	
3000-3999 g	74 (65.5)	577 (68.2)	
≥4000 g	19 (16.8)	87 (10.3)	
Size for gestational age, n (%)†			0.013
Small	1 (0.9)	18 (2.1)	
Appropriate	75 (66.4)	655 (77.4)	
Large	37 (32.7)	173 (20.5)	
Apgar score at 5 min, n (%)†			0.26
6 or lower	0 (0.0)	9 (1.1)	
7 or higher	110 (97.3)	826 (97.6)	
Missing	3 (2.7)	11 (1.3)	
Newborn nursery, n (%)			0.47
NICU	6 (5.3)	39 (4.6)	
Brief/intermediate level	10 (8.9)	51 (6.0)	
Well care	97 (85.8)	756 (89.4)	
Hospital stay, n (%)			0.149
≥3 d	28 (24.8)	161 (19.0)	
<3 d	85 (75.2)	685 (81.0)	

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DM = diabetes mellitus; GDM = gestational diabetes mellitus; IQR = interquartile range; NICU = neonatal intensive care unit.

\* Chi-square test for categorical variables; t test or analysis of variance for continuous variables. Kruskal-Wallis test to compare medians.

† Fisher exact test for cell sizes <5. Two-sided P values.

Appendix Table 2. Prenatal and Baseline Characteristics of Women With GDM, by Lactation Intensity Groups at 6 to 9 wk After Delivery (Study Baseline)\*

Characteristic	Exclusive Formula (n = 153)	Mostly Formula, Mixed/Inconsistent (n = 214)	Mostly Lactation (n = 387)	Exclusive Lactation (n = 205)
Sociodemographic				
Mean age at baseline (SD), y	33.0 (4.9)	33.4 (5.0)	33.4 (4.8)	33.5 (4.4)
Education, <i>n</i> (%)†	,	(2.2)	( ) - ( )	,
High school or less	60 (39.2)	56 (26.2)	79 (20.4)	35 (17.1)
Some college	55 (36.0)	55 (25.7)	106 (27.4)	54 (26.3)
College graduate	38 (24.8)	103 (48.1)	202 (52.2)	116 (56.6)
Race/ethnicity, n (%)†‡	30 (24.0)	100 (40.1)	202 (32.2)	110 (30.0)
Non-Hispanic white	34 (22.2)	43 (20.1)	79 (20.4)	69 (33.6)
Non-Hispanic black	22 (14.4)	19 (8.9)	20 (5.2)	11 (5.4)
Hispanic	48 (31.4)	68 (31.8)	118 (30.5)	58 (28.3)
Asian	43 (28.1)	80 (37.3)	166 (42.9)	63 (30.7)
Other	6 (3.9)			
	6 (3.7)	4 (1.9)	4 (1.0)	4 (2.0)
Parity, <i>n</i> (%)	40 (33 0)	02 (42 E)	122 (24.1)	70 (25.4)
Primiparous	49 (32.0)	93 (43.5)	132 (34.1)	72 (35.1)
Multiparous	104 (68.0)	121 (56.5)	255 (65.9)	133 (64.9)
Polycystic ovarian syndrome, n (%)	10 (6.5)	11 (5.1)	31 (8.0)	10 (4.9)
WIC recipient, n (%)†	59 (38.6)	55 (25.7)	87 (22.5)	45 (22.0)
Family history of diabetes, n (%)	70 (45.8)	103 (48.1)	204 (52.7)	96 (46.8)
Prenatal PMI(CD) ( / 2)	24.0.42.23	20.0 (7.0)	00.277	00.244.63
Mean prepregnancy BMI (SD), kg/m²†	31.2 (8.3)	30.0 (7.8)	29.3 (6.7)	28.3 (6.1)
Prepregnancy weight status, n (%)§				
BMI <25 kg/m <sup>2</sup>	38 (24.8)	66 (30.8)	114 (29.5)	73 (35.6)
BMI 25 of <30 kg/m <sup>2</sup>	37 (24.2)	60 (28.0)	125 (32.3)	65 (31.7)
BMI ≥30 kg/m²	78 (51.0)	88 (41.2)	148 (38.2)	67 (32.7)
Mean gestational weight gain (SD), kg	10.8 (7.9)	10.2 (6.5)	10.2 (6.5)	10.3 (6.9)
Mean 3-h, 100-g OGTT (SD), mg/dL				
Fasting glucose level	92.2 (11.4)	93.4 (11.1)	92.0 (12.6)	90.4 (12.8)
1-h glucose level	199.0 (22.1)	201.3 (23.0)	198.6 (23.2)	199.1 (23.1)
2-h glucose level	177.1 (25.0)	176.3 (28.8)	176.0 (29.2)	176.7 (24.7)
3-h glucose level	126.3 (34.7)	128.5 (31.1)	127.8 (33.3)	122.7 (33.7)
Mean sum of 3-h OGTT Z scores (SD)	0.0 (2.3)	0.3 (2.6)	0.0 (2.8)	-0.2 (2.6)
Median sum of 3-h OGTT Z scores (IQR)	-0.3 (-1.6 to 1.3)	-0.3 (-1.5 to 1.4)	-0.6 (-1.8 to 1.3)	-0.7 (-1.9 to 0.7
Mean gestational age at GDM diagnosis (SD), $wk$ Treatment of GDM, $n$ (%)‡	24.9 (7.8)	24.9 (7.2)	25.4 (6.9)	26.4 (6.5)
Diet modification only	108 (70.6)	141 (65.9)	263 (68.0)	143 (69.8)
Oral hypoglycemic agents	43 (28.1)	65 (30.4)	105 (27.1)	57 (27.8)
Insulin	2 (1.3)	8 (3.7)	19 (4.9)	5 (2.4)
Hospital labor and delivery				
Mean length of gestation (SD), wk	39.0 (1.1)	39.3 (1.1)	38.9 (1.2)	39.2 (1.0)
Mean length of stay (SD), $d^{\dagger}$	2.6 (1.1)	2.8 (1.3)	2.5 (1.0)	2.3 (0.9)
Cesarean section, n (%)§	61 (39.9)	74 (34.6)	113 (29.2)	
Cesarean section, II (70/8	01 (37.7)	74 (34.0)	113 (29.2)	57 (27.8)
6 to 9 wk after delivery (baseline)				
Mean BMI (SD), kg/m <sup>2</sup> †	32.0 (7.7)	30.6 (7.1)	29.8 (6.4)	28.5 (5.2)
Mean waist circumference (SD), cm†	93.6 (14.9)	91.5 (15.8)	89.1 (13.1)	86.9 (11.0)
Mean weight change postdelivery (SD), kg	-8.9 (4.0)	-8.7 (3.7)	-9.1 (3.6)	-9.5 (3.4)
Mean HOMA-IR index score (SD)†	7.0 (4.1)	6.5 (5.1)	4.9 (3.2)	4.3 (2.8)
Median HOMA-IR index score (IQR)†	5.9 (4.0-8.7)	5.1 (3.1-8.5)	4.1 (2.7-6.3)	3.5 (2.6-5.1)
Mean 2-h, 75-g OGTT results (SD)				
Fasting glucose level, mg/dL†	97.8 (9.1)	97.3 (10.2)	93.2 (8.0)	92.2 (7.9)
2-h postload glucose level, mg/dL†	110.6 (25.8)	117.4 (27.6)	113.0 (28.5)	106.9 (28.4)
Mean 2-h, 75-g OGTT results (SD)				
Fasting glucose level, mmol/L†	5.4 (0.5)	5.4 (0.6)	5.1 (0.4)	5.1 (0.4)
2-h postload glucose level, mmol/L†	6.1 (1.4)	6.5 (1.5)	6.2 (1.6)	5.9 (1.6)
Glucose tolerance, n (%)†	, ,	, ,	, -,	, ,
Normal	89 (58.2)	119 (55.6)	265 (68.5)	155 (75.6)
IFG level only	43 (28.1)	53 (24.8)	53 (13.7)	27 (13.2)
IGT only	9 (5.9)	20 (9.3)	50 (12.9)	16 (7.8)
IFG level and IGT	12 (7.8)	22 (10.3)	19 (4.9)	7 (3.4)
Depression, n (%)	12 (7.0)	22 (10.5)	17 (4.7)	7 (3.4)
None	126 (86.3)	174 (84.5)	325 (86.0)	176 (88.9)
Moderate or severe				
iviouerate or severe	20 (13.7)	32 (15.5)	53 (14.0)	22 (11.1)

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Appendix Table 2–Continued	·	·	·	
Characteristic	Exclusive Formula (n = 153)	Mostly Formula, Mixed/Inconsistent (n = 214)	Mostly Lactation (n = 387)	Exclusive Lactation (n = 205)
Median lifestyle behaviors (IQR)				
Total PA, metabolic equivalent h/wk§	46.5 (35.4-61.1)	41.8 (32.2-59.4)	42.8 (30.4-54.6)	42.9 (31.3-54.1)
Moderate to vigorous PA, metabolic equivalent h/wk	24.3 (16.2-35.6)	22.0 (14.3-33.5)	21.5 (13.7-30.3)	20.5 (13.5-29.7)
Glycemic index†	186.5 (136.4-375.2)	223.6 (166.5-280.3)	232.5 (177.7-313.1)	248.9 (186.4-337.5)
Dietary fiber, g/100 kcal†	0.9 (0.7-1.1)	1.0 (0.8-1.2)	1.0 (0.8-1.3)	1.1 (0.8-1.3)
Dietary animal fat, percentage of kcal†	28.4 (18.8-33.5)	25.8 (19.4-31.3)	24.4 (18.9-30.3)	24.2 (19.5-29.4)
Contraception methods, n (%)‡				
Progesterone only	8 (5.2)	4 (1.9)	10 (2.6)	4 (2.0)
Combination oral pills	24 (15.7)	20 (9.4)	46 (11.9)	28 (13.7)
Intrauterine device	17 (11.1)	18 (8.3)	41 (10.6)	15 (7.3)
None/barrier	104 (68.0)	172 (80.4)	290 (74.9)	158 (77.0)
Subsequent birth (1 birth during 2-y follow-up), n (%)	16 (10.5)	32 (15.0)	44 (11.4)	22 (10.7)

BMI = body mass index; GDM = gestational diabetes mellitus; HOMA-IR = Homeostatic Metabolic Assessment of Insulin Resistance; IFG = impaired fasting glucose; IGT = impaired glucose tolerance; IQR = interquartile range; OGTT = oral glucose tolerance test; PA = physical activity; sum of 3-h OGTT Z scores = sum of the 4 Z scores for the prenatal 100-g OGTT glucose values (fasting and 1, 2, and 3 h); WIC = Special Supplemental Nutrition Program for Women, Infants, and Children (eligibility based on income  $\leq$ 185% of the federal poverty level).

\* Chi-square test for categorical variables; t test or analysis of variance for continuous variables. Kruskal-Wallis test to compare medians. Data were missing for depression score (n = 31), HOMA-IR index score (n = 5), and lifestyle behaviors (n = 5).

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<sup>‡</sup> Fisher exact test for cell sizes <5. Two-sided P values.

<sup>§</sup> P < 0.050. || P < 0.010.

Appendix Table 3. Characteristics of Infants Born to Women With GDM, by Lactation Intensity Groups at 6 to 9 wk After Delivery\*

Characteristic	Exclusive Formula (n = 153)	Mostly Formula, Mixed/Inconsistent (n = 214)	Mostly Lactation $(n = 387)$	Exclusive Lactation (n = 205)
Mean birthweight (SD), g†	3435 (478.9)	3426 (495.7)	3355 (523.5)	3470 (460.6)
Mean length (SD), cm†	50.5 (2.5)	51.0 (2.4)	50.3 (2.5)	50.7 (2.4)
Mean weight for length Z score (SD)†	-0.2 (1.5)	-0.5 (1.3)	-0.3 (1.4)	-0.2 (1.3)
Median weight for length Z score (IQR)†	-0.2 (-1.1 to 0.6)	-0.5 (-1.4 to 0.2)	-0.4 (-1.2 to 0.6)	-0.2 (-1.1 to 0.8)
Sex, n (%)†				
Female Male	76 (49.7) 77 (50.3)	81 (37.8) 133 (62.2)	192 (49.6) 195 (50.4)	103 (50.2) 102 (49.8)
Gestational age, n (%)‡§				
35 to <37 wk (preterm)	6 (3.9)	5 (2.3)	29 (7.5)	4 (2.0)
≥37 wk (term)	147 (96.1)	209 (97.7)	358 (92.5)	201 (98.0)
Birthweight, n (%)†‡				
1500-2499 g	3 (2.0)	2 (0.9)	18 (4.7)	3 (1.5)
2500-2999 g	26 (17.0)	38 (17.8)	85 (22.0)	27 (13.2)
3000-3999 g	108 (70.5)	150 (70.1)	241 (62.2)	152 (74.1)
≥4000 g	16 (10.5)	24 (11.2)	43 (11.1)	23 (11.2)
Birth size for gestational age, n (%)‡				
Small	0 (0.0)	7 (3.3)	9 (2.3)	3 (1.5)
Appropriate	119 (77.8)	161 (75.2)	293 (75.7)	157 (76.6)
Large	34 (22.2)	46 (21.5)	85 (22.0)	45 (22.0)
Apgar score at 5 min, n (%)‡				
≤6	1 (0.7)	4 (1.9)	4 (1.0)	0 (0.0)
≥7	149 (97.3)	209 (97.6)	377 (97.4)	201 (98.0)
Missing	3 (2.0)	1 (0.5)	6 (1.6)	4 (2.0)
Newborn nursery, n (%)‡§				
NICU	10 (6.5)	11 (5.1)	22 (5.7)	2 (1.0)
Brief/intermediate level	10 (6.5)	21 (9.8)	26 (6.7)	4 (2.0)
Well care	133 (86.9)	182 (85.1)	339 (87.6)	199 (97.1)
Hospital stay, n (%)†				
≥3 d	37 (24.2)	48 (22.4)	77 (19.9)	27 (13.2)
<3 d	116 (75.8)	166 (77.6)	310 (80.1)	178 (86.8)

GDM = gestational diabetes mellitus; IQR = interquartile range; NICU = neonatal intensive care unit.

Appendix Table 4. Distribution of 959 SWIFT Participants Among Lactation Intensity Groups at 6 to 9 wk After Delivery and Lactation Duration Groups at the End of Follow-up\*

Lactation Intensity Groups at 6 to 9 wk After Delivery (Baseline)	Lactation Duration Groups					
o to 7 wk Arter Denvery (Basenne)	0-2 mo	>2-5 mo	>5-10 mo	>10 mo	Total	
Exclusive lactation	2 (0.2)	20 (2.1)	45 (4.7)	138 (14.4)	205 (21.4)	
Mostly lactation (≤6 oz of formula per day)	5 (0.5)	73 (7.6)	116 (12.1)	193 (20.1)	387 (40.3)	
Mostly formula feeding (>17 oz of formula per day) and mixed/inconsistent lactation	29 (3.0)	97 (10.1)	47 (4.9)	41 (4.3)	214 (22.3)	
Exclusive formula feeding	153 (16.0)	0 (0)	0 (0)	0 (0)	153 (16.0)	
Overall total	189 (19.7)	190 (19.8)	208 (21.7)	372 (38.8)	959 (100)	

SWIFT = Study of Women, Infant Feeding, and Type 2 Diabetes After GDM Pregnancy. \* P < 0.001. Values are numbers (percentages).

<sup>\*</sup> Study baseline. Chi-square test was used for categorical variables; F test was used for continuous variables. Kruskal-Wallis test for comparison of medians. Two-sided P values.

<sup>+</sup>P < 0.050.

<sup>‡</sup> Fisher exact test for cell sizes <5. § *P* < 0.010.