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Beyond birth outcomes: Interpregnancy interval and injuryrelated infant mortality

Marie E. Thoma¹, Lauren M. Rossen², Dane A. De Silva¹, Margaret Warner³, Alan E. Simon⁴, Susan Moskosky⁵, Katherine A. Ahrens^{5,6}

¹Department of Family Science, School of Public Health, University of Maryland, College Park, MD

²National Center for Health Statistics, Centers for Disease Control and Prevention, Division of Vital Statistics, Reproductive Health Statistics Branch, Hyattsville, MD

³National Center for Health Statistics, Centers for Disease Control and Prevention, Division of Vital Statistics, Mortality Statistics Branch, Hyattsville, MD

⁴Environmental Influences on Child Health Outcomes (ECHO) Project Office, Office of the Director, National Institutes of Health, Rockville, MD

⁵Office of Population Affairs, Office of the Assistant Secretary for Health, U.S. Department of Health and Human Services, Rockville, MD

⁶Muskie School of Public Service, University of Southern Maine, Portland, ME

Abstract

Background: Several studies have examined the association between IPI and birth outcomes, but few have explored the association between interpregnancy interval (IPI) and postnatal outcomes.

Objective: We examined the association between IPI and injury-related infant mortality, a leading cause of postneonatal mortality.

Methods: We used 2011-2015 US period-linked birth-infant death vital statistics data to generate a multiyear birth cohort of non-first-born singleton births (N = 9 782 029). IPI was defined as the number of months between a live birth and the start of the pregnancy leading to the next live birth.

Correspondence Marie Thoma, University of Maryland, School of Public Health, 4200 Valley Dr., College Park, MD 20742. mthoma@umd.edu.

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DISCLAIMER

The findings and conclusions in this article are those of the authors and do not necessarily represent the official position of the National Center for Health Statistics, Centers for Disease Control and Prevention, ECHO Program Office, Office of the Director, National Institutes of Health, or the Office of Population Affairs.

All authors approved the final manuscript as submitted and agreed to be accountable for all aspects of the work. The corresponding author attests that all listed authors meet authorship criteria and that no others meeting the criteria have been omitted.

SUPPORTING INFORMATION

Additional supporting information may be found online in the Supporting Information section at the end of the article.

Causes of death in the first year of life were identified using ICD-10 codes. Hazard ratios (HR) for IPI categories were estimated using Cox proportional hazards models adjusted for birth order, county poverty level, and maternal characteristics (marital status, race/ethnicity, education, age at previous birth).

Results: After adjustment, overall infant mortality (48.1 per 10 000 births) was higher for short and long IPIs compared with IPI 18-23 months (reference): <6, aHR 1.61, 95% CI 1.54, 1.68; 6-11, aHR 1.22, 95% CI 1.17, 1.26; and 60+ months, aHR 1.12, 95% CI 1.08, 1.16. In comparison, the risk of injury-related infant mortality (4.4 per 10 000 births) decreased with longer IPIs: <6, aHR 1.77, 95% CI 1.55, 2.01; 6-11, aHR 1.41, 95% CI 1.25, 1.59; 12-17, aHR 1.25, 95% CI 1.10, 1.41; 24-59, aHR 0.78, 95% CI 0.69, 0.87; and 60+ months, aHR 0.55, 95% CI 0.48, 0.62.

Conclusion: Unlike overall infant mortality, injury-related infant mortality decreased with IPI length. While injury-related deaths are rare, these patterns suggest that the timing between births may be a marker of risk for fatal infant injuries. The first year postpartum may be an ideal time for the delivery of evidence-based injury prevention programmes as well as family planning services.

Keywords

birth spacing; infant	mortality; injury; in	iterpregnancy in	iterval; parity	

1 | BACKGROUND

Considerable progress has been made to reduce infant mortality in the United States over the past few decades. In addition to increased survival of preterm births, this decline has been partly attributed to successful public health programmes aimed at reducing infant mortality caused by unsafe sleep environments. At the same time, infant mortality due to sudden infant death syndrome (SIDS) and unintentional injury persist as two of the ten leading causes of infant mortality in the United States, highlighting the need for additional intervention strategies.

Examining cause-specific mortality can provide greater insight into factors that may be amenable to public health interventions. In particular, external (ie non-pathological) causes of death represent a heterogeneous grouping of potentially preventable injury-related deaths. Higher birth order and plurality have been associated with an increased risk of external causes of infant mortality, consistent with the hypothesis that divided parental time and resources combined with increased childcare responsibilities may play a role in infant injury risk.^{6,7} A related factor, closely spaced births, may also be associated with external causes of infant death, but has not been examined for the United States, in part because national data have not been available since the early 1990s.

Closely spaced births, often operationalised using interpregnancy interval (IPI; the time between a live birth and start of a subsequent pregnancy), have been associated with increased risks of adverse birth outcomes and infant mortality. 8–10 The most common explanation for these associations is the 'maternal depletion hypothesis', which posits that adverse birth outcomes associated with short IPI are the result of depleted maternal

nutritional stores due to insufficient recuperation time between pregnancies. ¹¹ However, recent analysis using matched designs has cast doubt on the causal effect of short IPI on adverse outcomes, arguing that previous associations have been confounded by factors such as socio-economic status (SES). ^{12–14} A recent series of articles on birth spacing published in this journal discussed the strengths and limitations of these approaches and called for an examination of additional outcomes, beyond the perinatal period, to guide evidence-based recommendations. ^{15,16} Limited data exist on the association between short IPI and external causes of infant death, which may operate through causal mechanisms outside of maternal nutritional depletion, such as divided parental time and resources described above. Investigating this association could further inform public health guidance on birth spacing. ¹⁷

Therefore, our objective was to examine the relationship between IPI and injury-related infant mortality using recently available birth-infant death data from the United States. We use enhanced information on SES to provide better control for this potential confounding factor. In addition, we contrast our findings with those for the association between IPI and other measures of infant mortality to expand upon existing research.

2 | METHODS

2.1 | Data source

We used the 2011-2015 period-linked birth-infant death vital statistics files. ¹⁸ The period-linked numerator files consist of all deaths within 1 year of birth occurring in a given calendar year, retrospectively linked to their corresponding birth certificate data from the same or prior year. The period-linked denominator files consist of all births occurring in a given calendar year. These files are used by the National Center for Health Statistics to tabulate infant mortality rates by various maternal and infant characteristics on the birth certificate. We linked deaths in the numerator files to their corresponding birth records in the denominator files using unique identifiers to create a single multiyear birth cohort file, which we used for multiple regression analyses. Maternal identifiers were not available to link multiple or repeat births to the same woman. Ethical approval was not required as the deidentified data are publically available through a data use agreement with the National Center for Health Statistics.

The analytic sample included non-first-born singleton births to residents in states using the 2003 revised US birth certificate during 2011-2015. Given the small number of infant deaths due to external causes per year (~1400),⁴ we pooled across years 2011-2015 to ensure stable estimates. We restricted analyses to resident births among states using the 2003 revised US birth certificate because information to determine IPI had not been available since 1993.¹⁹ This information was reinstated with the 2003 revision; however, states adopted the 2003 revised US birth certificate at different times. Newly revised items on the birth certificate were first released in the 2011 period-linked national data files.²⁰ By 2011, 36 states had adopted the revision and 48 states had adopted by 2015 (representing 88% of all US non-first-born, singleton resident births during 2011-2015 period). Detailed information on birth certificate revisions and its implications for IPI analyses are described elsewhere.^{19,21}

2.2 | Measures

The linked birth-infant death data provide information on maternal and infant characteristics at birth from birth certificates and information on age and cause of death from death certificates. We also merged additional data from the 2011-2015 American Community Survey on the percentage of families living below the federal poverty threshold in the mother's county of residence.²²

2.2.1 | **Birth certificate measures**—The exposure variable was IPI, defined as the number of months between a live birth and the start of the pregnancy leading to the next live birth. This information is obtained from three data items collected on the birth certificate —'Date of current birth', 'Date of last live birth', and 'obstetric estimate of gestational age' —which have high agreement with medical records and other nationally representative estimates of IPI.^{21,23,24} The data file includes a recoded variable, live birth interval (in months), which is calculated by subtracting the date of last live birth from the date of current birth. We calculated IPI by subtracting the obstetric estimate of gestational age (in weeks converted to months) from the recoded live birth interval. We categorised IPI as <6, 6-11, 12-17, 18-23, 24-59, and 60+ months, consistent with other studies.^{9,11,25} Plural births could not be included in the calculation of IPI because information on 'date of last live birth' for multiples does not necessarily capture information on the preceding pregnancy. IPI was calculated using information on the birth certificate and not by linking births within mothers, which is currently not possible with national birth certificate data.

Descriptive information from the birth certificate on maternal sociodemographic and health characteristics included: maternal age at the current birth, race/ethnicity, level of education, marital status, principal source of payment at delivery, use of the Special Supplemental Nutrition Program for Women, Infants, and Children (WIC) during pregnancy, prepregnancy body mass index (BMI), and pre-pregnancy smoking. Maternal age at prior birth was derived by subtracting IPI from her current age and grouped into 5-year age categories. Infant characteristics included birth order, gestational age (using the obstetric estimate), birthweight, and sex. Maternal race/ ethnicity was classified consistent with 1997 Office of Management and Budget standards. Infant sex refers to the sex (male/female) assigned on the birth certificate at the time of the birth.

2.2.2 | **Death certificate measures**—Information on the age at death (days) and cause of death was obtained from the death certificate. We categorised infant deaths by age into either neonatal (<28 days) or postneonatal (28-365 days) infant death and subcategories of early neonatal (<7), late neonatal (7-27), early postneonatal (28-90), early middle postneonatal (91-182), late middle postneonatal (183-273), and late postneonatal (274-365).

Underlying cause of death was coded using the International Classification of Disease, tenth revision (ICD-10), and used to determine external causes of infant deaths (ie injury-related) from birth to 365 days: accidents (unintentional injuries; V01-X59); assault (homicide; *U01, X85-Y09); complications of medical and surgical care (Y40-Y84); and other external causes (Y10-Y36). Accidental suffocation and strangulation in bed (W75) is the most common external cause of death code among infants^{6,7} and one of three codes used to

classify sudden unexpected infant death (SUID) (W75, R95, and R99). Due to diagnostic shifts in classification of sleep-related deaths over time from R95 to W75, 3,27,28 we examined the following alternative groupings of injury-related ICD-10 codes: external causes excluding W75, external causes plus two additional causes of death used to classify SUID (SIDS [R95] and other ill-defined and unspecified causes of mortality [R99]), and SUID alone (W75, R95, and R99). Unintentional injury and homicide were examined separately in sensitivity analyses.

2.3 | Statistical analysis

Tabulations of births and deaths were based on period-linked data to be consistent with national reporting of infant mortality rates by maternal and infant characteristics, which rely on period-linked data. For deaths, we tabulated infant deaths overall, by age at the time of death, and cause of death categories across IPI categories; rates were calculated using births as the denominator. We used the statistical weights provided in the period-linked data²⁰ to account for unlinked infant deaths (<1%), to ensure that counts of deaths were consistent with national reports.

Using the multiyear birth cohort file, we fit Cox proportional hazard models to estimate hazard rate ratios (HR) by age at time of death and cause of death categories. We selected a reference group of 18-23 months to be consistent with other studies showing this interval has the lowest risk of adverse birth outcomes. ^{12,29,30} Time at risk was based on age in days. For surviving infants, births occurring during 2011-2014 were censored after 365 days of life. For births occurring in 2015, data on deaths occurring in 2016 were not available at the time of analysis; thus, time at risk for these infants was censored at the age of the infant (in days) as of 31 December 2015. Infants who died from causes that were not the outcome under evaluation were censored at their age of death. For neonatal deaths, infants who died after 27 days were censored at 28 days. Graphs of Schoenfeld residuals showed uniform distributions, indicating the Cox model proportionality assumption was not violated. Additionally, cumulative and instantaneous hazard of external cause of death curves by IPI categories were generated using the Kaplan-Meier survival function estimator. In situations such as ours, where the risk of outcome was rare, this estimator provides a reasonable approximation of risk even in the presence of competing risks. ^{6,7,31}

Confounders were selected based on the literature and whether the variable was associated (and preceded) the interpregnancy interval and outcome. 6,7,16,25,32 Our final models were adjusted for: maternal age at prior birth, race/ethnicity, and birth order; and, at the time of the current birth, maternal marital status, education, and county poverty level. Pre-pregnancy BMI, pre-pregnancy smoking, source of payment at delivery, and WIC use during pregnancy were not included in final models, since they were measured at the time of the current birth and may have represented a factor along the causal pathway. However, we considered this latter group of variables as proxy indicators of socio-economic and health status and additionally adjusted for these characteristics in sensitivity analysis to assess consistency of findings. Consistent with previous studies, 33 we further adjusted for preterm birth in sensitivity analyses to assess potential attenuation of associations due to mediation by gestational age. Additionally, we ran adjusted models with an interaction term between IPI

categories and birth year (2011-2015) and IPI categories and indicators of SES (eg maternal education, marital status) to assess variation in risk of external cause of death over time and SES indicators, respectively. The significance of the interaction terms was assessed using a likelihood ratio test.

Analyses were performed in 2018-2019 using Stata/MP and Stata/SE, version 14.

2.4 | Missing data

In our analytic sample, information on IPI was missing for 7.5% of births. All other covariates had missing values of less than 5%. We conducted sensitivity analyses using multiple imputation. Our imputation models used chained equations with predictive mean matching, generating 50 imputations, and included all covariates from the final models described above, plus the survival time and log of the survival. We collapsed continuous variables into categories (eg grouped age at death in days to weeks) and applied frequency weights for computational efficiency. We compared findings from the main complete case analysis (with continuous variables), frequency-weighted complete case analysis (with collapsed variables), and frequency-weighted multiple imputation analysis (with collapsed variables).

3 | RESULTS

During 2011-2015, IPI information was available for 9 782 029 non-first-born singleton births to residents who gave birth in states using the 2003 revised birth certificate. Of these births, the percentage that followed an IPI < 6, 6-11, 12-17, 18-23, 24-59, and 60+ months were 4.8%, 10.8%, 13.5%, 11.7%, 37.5%, and 21.7%, respectively. In general, compared with the other IPI categories, women having the shortest (<12 months) and longest (60+ months) IPI had similar covariate profiles (Table 1), which included lower SES and higher pre-pregnancy BMI. Births among the shortest and longest IPI groups were also more likely to be preterm and low birthweight. Being a fourth-or-higher-born infant was more common among the shorter interval groups; no differences in infant sex were found.

Among our analytic sample, there were 47 011 infant deaths, which corresponded to an infant mortality rate of 48.1 per 10 000 births (Table 2). Rates of neonatal and postneonatal deaths were 28.4 and 19.6 per 10 000 births, respectively. The infant death rate was highest for IPI < 6 months (87.0) and decreased to 56.9, 44.9, and 41.2 per 10 000 births for IPI of 6-11, 12-17, and 18-23 months, respectively, before increasing slightly to 41.7 for IPI of 24-29 months and more appreciably to 51.6 for IPI of 60+ months (Figure S1). Neonatal mortality rates, which accounted for approximately 60% of all infant deaths, demonstrated a similar J-shaped pattern with the highest rate per 10 000 births found among the shortest IPI (45.9) and longest IPI (35.3). Postneonatal mortality rates decreased with increasing IPI length—from 41.1 (IPI < 6 months) to 16.3 per 10 000 births (IPI of 60+ months).

The rates of infant deaths due to external causes and SUID were 4.4 and 9.8 per 10 000 births, respectively (Table 2). In contrast to the overall and neonatal death rates, external cause and alternative groupings demonstrated declines with increasing IPI length, similar to postneonatal mortality (Figures S1 and S2). The cumulative and instantaneous hazard

functions for infant mortality due to external causes by IPI category, respectively, show the steepest increases in risk were during the first 30-150 days of life (Figures 1 and 2).

After adjustment, the HRs for all-cause and neonatal infant mortality were highest for IPI < 6, 6-11, and 60+ months (all-cause aHR 1.61, 1.22, and 1.12; neonatal aHR 1.60, 1.13, and 1.40, respectively), whereas the aHRs for postneonatal mortality decreased from 1.61 (<6 months) to 0.77 (60+ months) relative to IPI of 18-23 months (Table 3). Similarly, infant mortality due to all external causes and alternative groupings of these cause of death codes showed diminishing risk with increasing length of IPI. In particular, the risk of infant mortality due to external causes was 77% higher among women with IPI < 6 months and reduced by 45% among women with IPI of 60+ months compared to 18-23 months. Finally, when we examined specific external causes in sensitivity analyses, the aHRs for infant deaths due to unintentional injuries decreased across length of IPI from 1.73 (95% CI 1.43, 2.09) for IPI < 6 months to 0.54 (95% CI: 0.45, 0.66) for 60+ months relative to IPI of 18-23 months; the corresponding aHRs for homicide decreased from 1.83 (95% CI 1.22, 2.76) to 0.51 (95% CI: 0.33, 0.77) (data not shown in tables).

Further adjustment for proxy variables of SES (health insurance, WIC use) and prepregnancy health (smoking, BMI) had minimal impact on the hazard ratio estimates (Table S1). Adjustment for preterm birth attenuated the HR estimates. This attenuation was considerably greater for overall infant mortality and neonatal mortality than for postneonatal, external causes, or SUID deaths. For external causes of death, there were no significant interactions between IPI and study year or SES indicators (all *P*-values >.12). Imputation of missing data resulted in attenuation of HRs for IPI of less than 6 or 6-11 months, but inferences remained similar (Table S2). After imputation, the percentage in each IPI category changed to 5.4, 11.1, 13.7, 12.0, 36.4, and 21.4 for less than 6 months through 60+ months, respectively.

4 | COMMENT

4.1 | Principal findings

To our knowledge, this is the first manuscript to examine the relationship between IPI and injury-related causes of infant mortality in the United States.³⁴ We found a decreasing risk of infant death due to external causes, postneonatal mortality, and each alternative grouping of external cause of death as the time between pregnancies increased. This is in contrast to the observed J-shaped relationship between IPI and overall and neonatal infant mortality found in our study and previous studies.⁹ In the United States, the neonatal mortality rate is almost twofold higher than postneonatal mortality, encompassing a larger proportion of overall infant deaths.³⁵ Leading causes of neonatal mortality are more likely due to factors related to the pregnancy or birth (eg low birthweight or maternal complications during pregnancy) compared to leading causes of postneonatal mortality (eg SIDS and unintentional injury).³⁵ Accordingly, our findings suggest a different relationship between IPI and injury-related mortality as compared with overall and neonatal infant mortality, highlighting that research on alternative outcomes beyond the perinatal or neonatal period could contribute to additional information when considering birth spacing guidelines.

Previous research using 1991 US birth certificate data showed an elevated association between short IPI (defined as <19 months) only and infant mortality. Similarly, more recent state-based studies using 2003 revised birth certificate data also found similar associations between only short IPI (defined as either <18 months or <12 months) and infant mortality. On IPI and increased risk of overall infant mortality. This discrepancy may be explained by our larger sample size or differences between our cohort and a given state or time period.

4.2 | Strengths of the study

We expand on previous studies using cross-sectional vital statistics data by examining the impact of residual confounding from SES. Estimates were attenuated after initial adjustment for SES and other confounders in our models, but there was minimal change when county-level poverty was added to the model or after adjustment for additional SES proxy variables (Table S1). Moreover, the underlying risk profiles for both short (<6 months) and long (60+months) IPI were similar, which would not explain why we would observe a continued decline in risk of injury-related mortality. The consistency of our findings with the proposed mechanism and prior studies on birth order and plurality and injury-related infant mortality underscore the need for further investigation on this relationship and other postneonatal infant health outcomes. Additional data sources are needed to further explore the association between short IPI and non-fatal injuries, including hospitalisations due to injury.

4.3 | Limitations of the data

While this study is unique in exploring injury-related infant mortality, it has limitations. In the absence of maternally linked records, we could not conduct a within-mother analysis for comparison. Recent studies, which rely on a sibling comparison design, have questioned the causal association between IPI and birth outcomes, suggesting that these associations may be attributable to unmeasured confounding, primarily due to SES. 12,13,37 However, it is unclear whether these new studies represent a better analytic approach or introduce both bias and lack of generalisability. 38 For example, estimates from these studies are based on women with 3 or more births who are discordant on both exposure and outcome status. Furthermore, these study designs have the potential to introduce greater bias in cases where unmeasured confounders, such as SES, vary between pregnancies within women—a plausible scenario in the context of increasing family size and short IPI. 39–41

4.4 | Interpretation

While mechanisms for the relationship between IPI and infant mortality are likely multifactorial, one proposed mechanism postulates that closely spaced births may be at higher risk of death due to sibling competition for resources and parental care and attention. ¹¹ A prior study also found a higher likelihood of maternal depression and neglectful parenting behaviours towards children born after closely spaced births. ⁴² In addition, epidemiologic studies have found a relationship between short IPI and postneonatal mortality, which may be served as a proxy for injury-related infant mortality. ^{11,30} Our study supports and expands these findings by specifically examining external causes of death. In combination with our other recent studies that have observed increased risk of injury-related death for plural and higher parity births, ^{6,7} our findings are consistent with the hypothesis

that parental time and resources may be more divided among children who are closely spaced (ie short IPI, plural births) or who have older siblings. In line with our findings, a study from New Zealand found that children living in families with high parenting demand were at higher risk of injury-related death and SUID.⁴³

4.5 | Conclusions

In sum, short birth spacing may be a marker for increased risk of mortality due to injury in infants. While injury-related infant mortality is rare in the United States, relative associations with IPI were consistent across multiple analyses. These findings have implications for understanding potential mechanisms for the relationship between IPI and infant mortality beyond the maternal depletion hypothesis and for targeting prevention strategies accordingly—as both short IPI and injury-related deaths are preventable. The first year postpartum is an important time for the delivery of evidence-based interventions to ensure optimal birth spacing and prevent injury.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Synopsis

Study question

Is interpregnancy interval (IPI) associated with injury-related infant mortality?

What is already known

IPI has been associated with increased risk of adverse birth outcomes and infant mortality, but little is known on its association with other postnatal outcomes, including external (injury-related) causes of infant death.

What this study adds

Unlike the J-shaped association with overall infant mortality, there is a decreasing risk of infant death due to injury-related causes with increasing IPI. This finding underscores the need to look beyond the perinatal or neonatal period in order to better inform birth spacing guidelines.

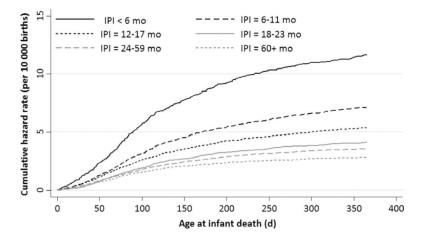


FIGURE 1. Cumulative hazard function of external cause of infant mortality by interpregnancy interval categories

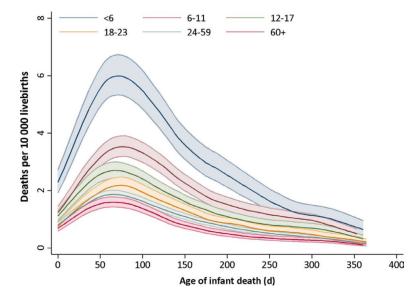


FIGURE 2. Instantaneous hazard function of external cause of infant mortality by interpregnancy interval categories. Footnote to Figure 2. The darker line indicates the point estimate for each IPI category. The shaded area in the same colour around each line corresponds to the 95% confidence interval estimates

TABLE 1

Per cent distribution of selected maternal and infant characteristics by interpregnancy interval among singleton, second-born, or higher births: 2003 revised US birth certificate data linked to infant death data, 2011-2015

		Interpregnan % of total	Interpregnancy Interval (months) % of total	.hs)			
	Total	<6 (4.8%)	6-11 (10.8%)	12-17 (13.5%)	18-23 (11.7%)	24-59 (37.5%)	60+ (21.7%)
Maternal and infant characteristics	N = 9.782029	N = 471 165	N = 1 056 915	N = 1 319 210	N = 1 148 398	N = 3 667 239	N = 2 119 102
Maternal characteristics at birth							
County of residence at birth (% below poverty threshold) a	ald) ^a						
Less than 10%	34.9	28.0	34.0	37.7	38.3	35.7	31.9
10%-19%	59.7	64.8	60.3	57.6	57.1	59.0	62.1
20%-29%	5.2	8.9	5.4	4.5	4.4	5.1	5.8
30% or greater	0.3	0.4	0.2	0.2	0.2	0.3	0.3
County of residence at birth (% poverty level, mean)	12.0	12.8	12.1	11.7	11.6	11.9	12.3
Maternal age at prior birth (years)							
Under 20	13.0	15.7	9.2	9.4	8.3	12.8	19.4
20-24	31.9	37.0	29.9	26.0	25.6	30.7	40.8
25-29	30.7	26.3	30.2	32.5	32.8	31.5	28.0
30-34	18.9	15.2	21.8	24.1	25.2	19.8	10.3
35-39	5.0	5.2	7.8	7.3	7.4	8.4	1.4
40-54	0.5	9.0	1.0	8.0	8.0	0.4	0.1
Maternal age at prior birth (years, mean)	26.1	25.7	26.9	27.4	27.3	26.2	24.1
Maternal race/ethnicity							
Non-Hispanic white	52.2	45.2	54.8	8.09	61.0	52.9	41.2
Non-Hispanic black	14.5	21.6	15.9	12.0	11.5	13.5	17.0
Hispanic	25.9	26.6	22.4	20.0	20.2	25.8	34.4
$Other^{b}$	7.4	9.9	7.0	7.2	7.4	7.7	7.5
Marital status							
Not married	36.6	53.8	37.9	29.2	28.3	34.6	44.6
Married	63.5	46.2	62.1	70.8	71.8	65.5	55.4

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Maternal and infant characteristics Total cd.(48%) col (110,8%) Lot (111,68%)			Interpregnan % of total	Interpregnancy Interval (months) % of total	ths)			
N=9782029 N=471166 N=1105010 N=1319210 N=1148398 N=3667229 182 270 197 158 148 170 259 340 272 229 224 255 212 211 204 189 190 213 440 179 327 424 438 362 440 48 403 438 450 450 445 484 403 389 450 450 446 49 403 389 450 450 448 52 51 48 47 44 45 52 51 48 47 44 462 628 481 482 44 47 453 37.2 519 60.7 520 55.7 588 37.2 27.6 223 44.4 47 48.1 38.6 23.1 24.6 52.1 44.		Total	<6 (4.8%)	6-11 (10.8%)	12-17 (13.5%)	18-23 (11.7%)	24-59 (37.5%)	60+ (21.7%)
FIGH 182 27.0 19.7 15.8 14.8 17.0 17.0 ED 25.9 34.0 27.2 22.9 22.4 25.5 21.3 21.2 21.1 20.4 18.9 19.0 21.3 21.3 21.3 21.4 20.4 18.9 19.0 21.3 21.3 21.3 21.4 20.4 21.3 22.4 25.5 21.3 20.4 21.3 21.4 20.4 21.3 20.4 21.3 20.4 21.3 20.4 21.3 20.4 21.3 20.4 21.3 20.4 21.3 20.4 21.3 20.4 21.3 20.4 21.4 20.4 21.4 20.4 21.4 20.4 21.4 20.4 21.4 20.4 21.4 21.4 21.4 21.4 21.4 21.4 21.4 21	Maternal and infant characteristics	N = 9.782 029	N = 471 165	N = 1 056 915	N = 1 319 210	N = 1 148 398	N = 3 667 239	N = 2 119 102
FIGE 18.2 27.0 19.7 15.8 14.8 17.0 17.0 18.9 18.9 17.0 17.0 17.2 17.2 17.2 17.2 17.2 17.2 17.2 17.2	Maternal educational attainment							
ED 259 340 272 229 224 255 247	No high school diploma or GED	18.2	27.0	19.7	15.8	14.8	17.0	20.8
r. 34.7 17.9 20.4 18.9 19.0 21.3 r. 44.0 25.7 40.4 40.5 38.9 26.0 44.0 25.7 40.4 40.3 38.9 45.0 44.0 42.8 40.3 38.9 45.0 44.1 4.8 5.2 5.1 4.8 4.7 4.7 44.9 44.2 5.2 5.1 4.8 5.2 5.2 4.7 4.4 45.2 5.2 5.1 4.8 5.2 5.2 4.7 4.4 48.1 37.6 48.1 30.3 38.0 44.4 58.6 31.7 31.7 26.2 23.1 23.1 58.7 38.8 43.1 31.7 26.2 23.1 23.1 58.8 43.1 84.3 88.6 91.1 91.1 89.0 52.9 44.4 50.2 54.5 55.0 57.6 52.9 44.4 50.2 54.5 55.0 57.6 52.9 54.6 55.0 57.6 57.6 57.0 57.6 52.9 54.6 52.0 57.0 57.6 57.0 57.6 52.9 54.6 57.0 57.0 57.0 57.0 57.6 52.9 54.6 57.0 57.0 57.0 57.0 57.0	High school diploma or GED	25.9	34.0	27.2	22.9	22.4	25.5	28.2
rr 440	Some college	21.2	21.1	20.4	18.9	19.0	21.3	24.2
17. 440 25.7 40.4 49.7 51.7 46.0 46.0 49.1 49.1 49.1 49.1 49.1 46.2 45.0 45.0 49.4 40.3 88.9 45.0 45.0 48.1 48.8 5.2 4.7 4.4 4.4 4.8 48.2 4.7 4.4 4.7 4.4 4.8 48.1 89.3 88.0 44.4 4.8 48.2 48.1 89.3 88.0 44.4 48.1 89.2 88.1 88.6 89.1 11.0 89.0 11.0 88.0 88	Bachelor's degree or higher	34.7	17.9	32.7	42.5	43.8	36.2	26.8
440 25.7 40.4 49.7 51.7 46.0 465 64.3 49.4 40.3 38.9 45.0 4.7 4.8 5.2 5.2 4.7 4.4 4.8 5.2 5.1 4.7 4.4 4.8 5.2 5.1 4.7 4.4 46.2 62.8 4.81 3.3 4.7 4.7 45.1 62.8 4.81 3.3 4.4 4.7 45.1 37.6 4.81 3.8 4.8 4.4 4.4 45.1 37.6 4.8 4.8 4.8 4.8 4.8 4.8 1.1 8.6 31.7 27.6 25.3 24.9 25.1	Source of payment for delivery							
bx 465 643 49,4 403 389 450 450 450 447 448 452 452 477 489 452 477 489 452 477 489 452 478 481 481 482 477 489 447 478 481 481 482 481 482 481 482 481 481 481 481 481 481 481 481 481 481	Private	44.0	25.7	40.4	49.7	51.7	46.0	38.7
lex 4.7 4.8 5.2 5.1 4.8 4.7 4.4 4.8 5.2 5.1 4.8 4.7 4.4 4.8 5.2 5.1 4.8 4.7 4.7 4.1 4.8 5.2 5.1 4.8 4.7 4.7 4.2 5.2 4.8 1 39.3 38.0 4.44 4.3 3.2 3.1 3.6 6.7 6.2 6.3 3.8 5. 3.1 3.6 3.6 3.8 4.8 4.8 4.8 4.8 4.8 4.8 4.8 4.8 4.8 4	Medicaid	46.5	64.3	49.4	40.3	38.9	45.0	51.8
lex 46.2 62.8 48.1 39.3 38.0 44.4 44.4 45.2 53.8 37.2 51.9 60.7 62.0 55.7 44.4 44.4 43.1 3.6 3.6 3.6 3.8 3.7 55.7 51.9 60.7 62.0 55.7 51.9 60.7 62.0 55.7 51.0 52.0 52.0 52.1 52.1 52.1 52.1 52.1 52.1 52.1 52.1	Self-pay	4.7	4.8	5.2	5.2	4.7	4.4	8.8
lex 3.2 3.1 3.2 3.1 3.2 3.1 3.2 3.1 3.2 3.1 3.2 3.1 3.2 3.2	Other	4.8	5.2	5.1	8.8	4.7	4.7	8.8
lex	WIC use during pregnancy							
lex 3.2 3.1 3.2 3.1 3.6 43.1 3.6 43.8 43.8 43.8 43.8 43.8 43.1 43.1 26.9 31.7 26.3 31.7 26.2 31.7 26.2 27.1 26.2 31.7 26.2 27.1 26.2 31.7 26.2 27.1 26.2 27.1 26.2 27.1 26.2 27.1 26.2 27.1 27.2 2	Yes	46.2	62.8	48.1	39.3	38.0	44.4	53.5
lex 3.2 3.1 3.6 43.1 43.8 43.8 43.8 48.2 48.1 56.8 26.8 27.6 27.6 27.6 26.3 27.7 26.3 31.7 31.7 26.2 27.1 26.2 27.1 26.2 27.1 26.2 27.1 26.2 27.1 26.2 27.1 27.1 27.2 27.2 27.2 27.3 27.3 27.3 27.4 27.3 27.4 27.5 27.5 27.7 27.9 27.9 27.9 27.9 27.9 27.0	No	53.8	37.2	51.9	2.09	62.0	55.7	46.5
3.2 3.1 3.6 3.6 3.8 3.7 3.7 43.8 43.2 44.1 5.0 5.0 5.3 3.8 3.7 48.1 5.8 5.8 5.8 5.8 5.8 5.8 5.8 5.8 5.8 5.8	Pre-pregnancy body mass index							
56.8 27.6 26.3 24.9 48.1 48.1 48.1 56.8 24.9 24.1 25.1 26.9 31.7 31.7 26.2 25.1 24.1 25.1 25.1 25.1 25.1 25.1 25.1 25.1 25	Underweight	3.2	3.1	3.6	3.6	3.8	3.7	2.3
26.8 27.6 27.6 26.3 24.9 25.1 26.9 31.7 31.7 26.2 23.1 23.1 88.7 84.3 88.6 91.1 91.1 89.0 11.3 15.7 11.4 8.9 8.9 11.0 52.9 44.4 50.2 54.6 55.9 54.6 27.6 27.7 25.9 24.6 25.0 27.6 19.6 27.9 23.9 20.9 19.1 17.8	Normal	43.1	37.6	43.8	43.8	48.2	48.1	37.0
sc 31.7 31.7 26.2 23.1 23.1 88.7 84.3 88.6 91.1 91.1 89.0 111.3 15.7 11.4 8.9 8.9 11.0 52.9 44.4 50.2 54.5 55.9 54.6 27.6 27.7 25.9 24.6 25.0 27.6 19.6 27.9 23.9 20.9 19.1 17.8 7.6 11.1 7.8 6.4 6.2 7.0	Overweight	26.8	27.6	27.6	26.3	24.9	25.1	29.1
88.7 84.3 88.6 91.1 91.1 89.0 11.0 11.3 15.7 11.4 8.9 8.9 8.9 11.0 11.0 11.0 11.0 11.0 11.0 11.0 11	Obese	26.9	31.7	31.7	26.2	23.1	23.1	31.6
88.7 84.3 88.6 91.1 99.0 11.3 15.7 11.4 8.9 8.9 11.0 52.9 44.4 50.2 54.5 55.9 54.6 27.6 27.7 25.9 24.6 25.0 27.6 19.6 27.9 23.9 20.9 19.1 17.8 7.6 11.1 7.8 6.4 6.2 7.0	Pre-pregnancy smoking status $^{\mathcal{C}}$							
11.3 15.7 11.4 8.9 8.9 11.0 11.0 11.4 8.9 8.9 11.0 11.0 11.1 11.4 8.9 8.9 11.0 11.1 11.4 8.9 8.9 11.0 11.0 11.1 11.4 8.9 8.9 11.0 11.0 11.1 11.8 6.4 6.2 7.0	No	88.7	84.3	9.88	91.1	91.1	0.68	86.3
52.9 44.4 50.2 54.5 55.9 54.6 27.6 27.7 25.9 24.6 25.0 27.6 19.6 27.9 23.9 20.9 19.1 17.8 7.6 11.1 7.8 6.4 6.2 7.0	Yes	11.3	15.7	11.4	8.9	8.9	11.0	13.7
orm 52.9 44.4 50.2 54.5 55.9 54.6 m 27.6 27.7 25.9 24.6 25.0 27.6 om or higher 19.6 27.9 23.9 20.9 19.1 17.8 week at birth 7.6 11.1 7.8 6.4 6.2 7.0	Infant characteristics at birth							
52.9 44.4 50.2 54.5 55.9 54.6 54.6 27.6 27.7 25.9 24.6 25.0 27.6 19.6 27.9 23.9 20.9 19.1 17.8 7.6 11.1 7.8 6.4 6.2 7.0	Birth order							
27.6 27.7 25.9 24.6 25.0 27.6 19.6 27.9 23.9 20.9 19.1 17.8 7.6 11.1 7.8 6.4 6.2 7.0	Second-born	52.9	44.4	50.2	54.5	55.9	54.6	50.4
19.6 27.9 23.9 20.9 19.1 17.8 1 7.6 11.1 7.8 6.4 6.2 7.0	Third-born	27.6	Z7.7	25.9	24.6	25.0	27.6	31.5
7.6 11.1 7.8 6.4 6.2 7.0	Fourth-born or higher	19.6	27.9	23.9	20.9	19.1	17.8	18.1
7.6 11.1 7.8 6.4 6.2 7.0	Gestational week at birth							
	<37 (preterm birth)	7.6	11.1	7.8	6.4	6.2	7.0	9.3

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		Interpregnan % of total	Interpregnancy Interval (months) % of total	ths)			
	Total	<6 (4.8%)	6-11 (10.8%)	12-17 (13.5%)	18-23 (11.7%)	24-59 (37.5%)	60+ (21.7%)
Maternal and infant characteristics	N = 9.782029	N = 471 165	N = 1 056 915	N = 1 319 210	N = 1 148 398	N = 3 667 239	N = 2 119 102
37-38 (early term)	26.3	31.9	26.9	24.9	24.6	25.8	27.4
39-40 (term)	61.2	53.4	60.3	63.0	63.8	62.5	58.8
41 (late term)	4.6	3.2	4.6	5.2	5.1	4.5	4.3
42 or over (post-term birth)	0.3	0.4	0.5	0.4	0.4	0.3	0.3
Birthweight (grams) d							
<1500	6.0	1.3	8.0	9.0	9.0	8.0	1.3
1500–2499	4.6	6.9	4.6	3.7	3.6	4.2	5.9
2500 or over	94.5	91.8	94.7	95.7	95.8	95.0	92.8
Sex							
Female	48.9	48.9	48.9	48.9	48.8	48.8	48.9
Male	51.2	51.1	51.1	51.1	51.2	51.2	51.1

(1.1%), source of payment for delivery (1.0%), Special Supplemental Nutrition Program for Women, Infants, and Children (WIC) use during pregnancy (2.0%), pre-pregnancy body mass index (3.6%), pre-Note: Unweighted counts of average live births per year were used. GED stands for General Education Diploma. Missing values were <4% of the following characteristics: maternal educational attainment pregnancy smoking status (3.8%), and imputed birthweight (<0.1%).

bIncludes Asian or Pacific Islander, American Indian or Alaska Native, Other, and Hispanic origin unknown or not stated. of residence at the time of birth.

^aGenerated from restricted-use data files merged by county Federal Information Processing Standard Publication 6-4 (FIPS) county codes with families below the poverty threshold in the mother's county

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 $^{^{\}mathcal{C}}_{\text{In the three months prior to pregnancy.}}$

dMissing birthweight values are imputed from the previous record with the same period of gestation, race, sex, and plurality.

TABLE 2

Rate of infant death by all-cause, neonatal, postneonatal, and external cause categories among singleton, second-born, or higher births: 2003 revised US birth certificate data linked to infant death data, 2011-2015

			Interp	regnancy	Interva	Interpregnancy Interval (IPI) in months	months							
	Total deaths	eaths	ş		6-11		12–17		18–23		24–59		+09	
Number of births (2011–2015)	9 782 029	59	47 11 65	55	10 56 915	15	1 319 210	10	1 148 398	869	3 667 239	6	2 119 102	12
Cause of death	п	Rate	п	Rate	u	Rate	u	Rate	u	Rate	u	Rate	u	Rate ^a
All causes	47 011	48.1	4098	87.0	6012	56.9	5927	44.9	4737	41.2	15 299	41.7	10 938	51.6
Age at death (days)														
Total neonatal (0-27)	27 802	28.4	2161	45.9	3087	29.2	3214	24.4	2667	23.2	9192	25.1	7482	35.3
Early neonatal (under 7)	21 643	22.1	1678	35.6	2393	22.6	2460	18.6	2082	18.1	7131	19.4	2900	27.8
Late neonatal (7–27)	6159	6.3	483	10.3	694	9.9	754	5.7	585	5.1	2061	5.6	1582	7.5
Total postneonatal (28–365)	19 210	19.6	1937	41.1	2925	27.7	2714	20.6	2071	18.0	6107	16.7	3456	16.3
Early postneonatal (28–90)	8775	0.6	840	17.8	1259	11.9	1205	9.1	965	8.4	2831	7.7	1674	7.9
Early middle postneonatal (91–182)	6422	9.9	694	14.7	1011	9.6	925	7.0	929	5.9	2028	5.5	1088	5.1
Late middle postneonatal (183–273)	2643	2.7	276	5.9	433	4.1	372	2.8	286	2.5	820	2.2	455	2.1
Late postneonatal (274–365)	1370	1.4	127	2.7	222	2.1	212	1.6	143	1.2	427	1.2	239	1.1
External causes and SUID (ICD-10 codes)														
All external causes b	4344	4.4	531	11.3	752	7.1	269	5.3	477	4.2	1305	3.6	581	2.7
Accidents/unintentional injuries	3435	3.5	399	8.5	591	5.6	548	4.2	383	3.3	1049	2.9	466	2.2
Assault/homicide	634	9.0	96	2.0	113	1:1	100	8.0	69	9.0	172	0.5	83	0.4
Complications of medical/surgical care $^{\mathcal{C}}$	29	0.0	1	ı	1	1	ı	1	1	ŀ	10	0.0	I	1
Other external causes (undetermined intent)	246	0.3	35	0.7	43	0.4	45	0.3	23	0.2	74	0.2	25	0.1
External causes excluding W75 ^d	1966	2.0	264	5.6	347	3.3	316	2.4	210	1.8	577	1.6	252	1.2
External causes and $\mathrm{SUID}^{oldsymbol{e}}$	11 573	11.8	1399	29.7	1957	18.5	1744	13.2	1255	10.9	3616	6.6	1602	7.6
$\mathrm{SUID}\mathrm{only}^f$	2096	8.6	1134	24.1	1610	15.2	1429	10.8	1045	9.1	3039	8.3	1350	6.4

Note: Weighted counts of infant deaths are presented. SUID, sudden unexpected infant death. '--' suppressed due to fewer than 10 infant deaths.

 $^a\mathrm{Per}$ 10 000 live births.

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b ICD-10 codes: U01, V01-Y84. Accidents/unintentional injuries (V01-X59); assault/homicide (U01, X85-Y09); complications of medical/surgical care (Y40-Y84); other external causes (Y10-Y36) of

The underlying cause of death will underestimate the number of deaths in which this complication is a factor, unless the cause of or reason for the medical/surgical care is not reported. undetermined intent.

^dICD-10 codes: U01, V01-W74, W76-Y84.

^eICD-10 codes: U01, V01-Y84, R95, R99.

fICD-10 codes: R95, R99, W75.

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TABLE 3

Hazard ratios of interpregnancy interval and infant death among singleton, second-born, or higher births by ICD-10 grouping: 2003 revised US birth certificate data linked to infant death data, 2011-2015

	Interpregnancy Interval (months)	nterval (months)				
	9 >	6–11	12–17	18–23	24–59	+09
Cause of death	HR (95% CI)	HR (95% CI)	HR (95% CI)	HR (95% CI)	HR (95% CI)	HR (95% CI)
All causes						
Unadjusted	2.12 (2.03, 2.21)	1.38 (1.32, 1.43)	1.09 (1.05, 1.13)	1.00 (Reference)	1.02 (0.98, 1.05)	1.27 (1.23, 1.32)
Adjusted ^a	1.61 (1.54, 1.68)	1.22 (1.17, 1.26)	1.06 (1.02, 1.11)	1.00 (Reference)	0.96 (0.93, 1.00)	1.12 (1.08, 1.16)
Neonatal (0–27 days)						
Unadjusted	1.98 (1.87, 2.09)	1.26 (1.19, 1.33)	1.05 (1.00, 1.11)	1.00 (Reference)	1.08 (1.03, 1.13)	1.52 (1.46, 1.59)
Adjusted ^a	1.60 (1.51, 1.69)	1.13 (1.07, 1.19)	1.03 (0.98, 1.08)	1.00 (Reference)	1.04 (1.00, 1.09)	1.40 (1.34, 1.47)
Postneonatal (28-365 days)						
Unadjusted	2.32 (2.17, 2.47)	1.54 (1.45, 1.63)	1.14 (1.08, 1.21)	1.00 (Reference)	0.93 (0.89, 0.98)	0.92 (0.87, 0.98)
Adjusted ^a	1.61 (1.51, 1.72)	1.32 (1.24, 1.40)	1.11 (1.05, 1.18)	1.00 (Reference)	0.86 (0.81, 0.90)	0.77 (0.73, 0.82)
All external causes b						
Unadjusted	2.83 (2.49, 3.22)	1.72 (1.53, 1.94)	1.31 (1.16, 1.48)	1.00 (Reference)	0.87 (0.78, 0.97)	0.70 (0.62, 0.79)
Adjusted ^a	1.77 (1.55, 2.01)	1.41 (1.25, 1.59)	1.25 (1.10, 1.41)	1.00 (Reference)	0.78 (0.69, 0.87)	0.55 (0.48, 0.62)
External causes excluding W75						
Unadjusted	3.29 (2.71, 3.99)	1.81 (1.50, 2.18)	1.42 (1.18, 1.71)	1.42 (1.18, 1.71) 1.00 (Reference)	0.90 (0.76, 1.07)	0.70 (0.58, 0.86)
Adjusted ^a	2.11 (1.73, 2.56)	1.53 (1.27, 1.84)	1.36 (1.13, 1.64)	1.00 (Reference)	0.80 (0.67, 0.95)	0.54 (0.44, 0.67)
External causes and SUID^d						
Unadjusted	2.73 (2.52, 2.95)	1.69 (1.57, 1.82)	1.21 (1.13, 1.31)	1.00 (Reference)	0.91 (0.85, 1.97)	0.71 (0.66, 0.77)
Adjusted ^a	1.73 (1.59, 1.87)	1.40 (1.30, 1.51)	1.17 (1.08, 1.26)	1.00 (Reference)	0.81 (0.76, 0.86)	0.56 (0.52, 0.60)
$\mathrm{SUID}\mathrm{only}^{\boldsymbol{e}}$						
Unadjusted	2.63 (2.41, 2.87)	1.67 (1.54, 1.81)	1.18 (1.08, 1.28)	1.00 (Reference)	0.91 (0.84, 0.98)	0.71 (0.66, 0.77)
Adjusted ^a	1.66 (1.52, 1.81)	1.38 (1.27, 1.49)	1.13 (1.04, 1.23)	1.00 (Reference)	0.81 (0.75, 0.87)	0.56 (0.52, 0.61)

Abbreviations: HR, hazard ratio; SUID, sudden unexpected infant death.

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Models were adjusted for maternal age at prior birth (under 20, 20-24, 25-29, 30-34, 35-39, 40+), birth order (2, 3, and 4+), maternal race/ethnicity (non-Hispanic black, non-Hispanic white, Hispanic, and other race or unknown ethnicity), maternal education (no high school diploma/no General Education Diploma (GED), high school diploma/GED, some college, bachelor's degree, or higher), marital status (currently married or not), and county poverty level (continuous).

 b ICD-10 codes: U01, V01-Y84.

 $^{\mathcal{C}}_{\text{ICD-10 codes: U01, V01-W74, W76-Y84.}}$

 $^{d}_{\rm ICD-10\;codes:\;U01,\;VO1-Y84,\;R95,\;R99.}$

^eICD-10 codes: R95, R99, W75.