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Duration of Breastfeeding and Risk of SIDS: An Individual Participant Data (IPD) Meta-analysis

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Abbreviations: aOR: Adjusted odds ratio; OR: Odds ratio; SES: Socioeconomic status; SIDS: Sudden infant death syndrome

Table of Contents Summary: This study used individual-level data from international studies to assess the associations between duration of any breastfeeding and exclusive breastfeeding and SIDS.

Contributors' Statement:

Dr. Thompson conceptualized and designed the study, carried out the analyses, drafted the initial manuscript and approved the final manuscript as submitted.

Ms. Tanabe, Drs. Moon and Hauck conceptualized and designed the study, participated in the interpretation of the data, critically reviewed and revised the manuscript and approved the final manuscript as submitted.

Drs. Mitchell, McGarvey, Tappin and Blair provided data for the study, reviewed and revised the manuscript and approved the final manuscript as submitted.

Abstract

Context: Sudden infant death syndrome (SIDS) is a leading cause of postneonatal infant mortality. Our prior meta-analyses showed that any breastfeeding is protective against SIDS, and that this effect is stronger with exclusive breastfeeding. It is unclear what duration of breastfeeding is required to confer a protective effect.

Objective: Assess the associations between duration of breastfeeding and SIDS.

Data Sources: Individual-level data from 8 case-control SIDS studies.

Study Selection: All case-control SIDS studies with breastfeeding data.

Data Extraction: Variables for breastfeeding, demographic factors, and other potential confounders were identified. Individual-study and pooled analyses were performed.

Results: 2267 SIDS cases and 6837 control infants were included. In multivariable pooled analysis, breastfeeding for <2 months was not protective (aOR 0.91; 95%CI=0.68-1.22). Any breastfeeding \geq 2 months was protective, with greater protection seen with increased duration (2-4 months: aOR 0.60, 95%CI=0.44-0.82; 4-6 months: aOR 0.40, 95%CI=0.26-0.63; and 6+ months: aOR 0.36; 95%CI=0.22-0.61). Although exclusive breastfeeding for <2 months was not protective (aOR 0.82; 95%CI=0.59-1.14), longer periods were protective: (2-4 months: aOR 0.61, 95%CI=0.42-0.87; and 4-6 months: aOR 0.46, 95%CI=0.29-0.74). In sensitivity analyses only minor changes were seen in the odds ratios, suggesting robustness of the results.

Limitations: The variables collected in each study varied slightly, limiting our ability to include all studies in the analysis controlling for all confounders.

Conclusions and Relevance: Breastfeeding duration of at least 2 months was associated with a protective effect against SIDS, with an approximate halving of risk. Breastfeeding does not need to be exclusive to confer this protection.

Introduction

Breastfeeding has been shown in several studies to be associated with a decreased risk of Sudden Infant Death Syndrome (SIDS).¹⁻³ In a previous meta-analysis, we have shown that breastfeeding is protective against SIDS (adjusted Odds Ratio (aOR)=0.55; 95% CI: 0.44, 0.69 for any breastfeeding), and that this protective effect is stronger with exclusive breastfeeding (OR=0.27; 95% CI: 0.24, 0.31).³

However, it has been difficult to determine what duration of breastfeeding is required to confer a protective effect against SIDS. This may partly be because the incidence of breastfeeding across countries and different cultures varies and because the different studies investigating the association with SIDS use different definitions for any breastfeeding, exclusive breastfeeding and the duration of either practice. Meta-analyses of breastfeeding duration at the study level are difficult to undertake, and so far the effect size and the duration of breastfeeding required to confer this protective effect have not been quantified.

We therefore aimed to use individual-level data from international studies, and with co-operation of the individual authors, to assess the associations between duration of any breastfeeding and exclusive breastfeeding and SIDS.

Methods

We used the same review protocol as that in our previously reported meta-analysis.³ We searched the Ovid Medline database (January 1966 through December 2009) to collect data on breastfeeding and its association with SIDS. The search strategy included published

articles limited to humans with the Medical Subject Headings terms “sudden infant death” and “breast feeding” with key words “sudden infant death syndrome, “SIDS,” “cot death,” and “breastfeeding.” Of the 18 studies included in the meta-analysis individual level data were provided from 8 large case-control studies of SIDS deaths, which comprise all of the published case-control studies with individual-level data about breastfeeding status. In all studies, there were strict definitions and protocols for determining SIDS cases. The cause of death had to be ascertained by local medical examiners, pediatric or forensic pathologists. No studies without individual-level data were included. All data were obtained via direct contact with the original investigators for each case-control study. Data were checked by the original investigators for completeness and consistency before being released for this analysis. The studies include:

The New Zealand Cot Death Study (NZCDS)

The NZCDS was a national case-control study of all SIDS deaths that took place from November 1987 through October 1990. The study successfully recruited and obtained data from 393 cases and 1592 controls, who were randomly selected from all births but with an age distribution to match the age of cases from 1979-1984.⁴ Data were obtained by an interviewer-administered questionnaire and from hospital obstetric records, which included data as to the type of feeding at the time of hospital discharge. Parents were asked whether the infant received any breast milk at any stage of life, in the first 4 weeks and the last 2 days. In addition, parents of infants who received any breast milk were asked at what stage breastfeeding stopped (age in weeks). Coding was available for never started and still breastfeeding.

The Chicago Infant Mortality Study (CIMS)

The CIMS studied all SIDS deaths in Chicago, Illinois, US between November 1993 and April 1996, and included 260 cases and 260 controls, who were matched by maternal ethnicity, age at death and birthweight.⁵ Data on breastfeeding were collected by a standardized interviewer-administered questionnaire. Parents were asked if the child had ever been breastfed, if the child was still being breastfed, and how old the child was when breastfeeding stopped. In addition, data on other methods of feeding and when they started were collected so that duration of exclusive breastfeeding could be calculated.

The German SIDS Study (GeSID)

The GeSID was carried out in 11 of 18 states in the Federal Republic of Germany between November 1998 and October 2001. The study included 333 SIDS cases and 998 controls, who were matched by geographic region, age, gender and reference sleep (i.e., time of sleep was matched to the time of death for the respective case).⁶ Data on breastfeeding were collected by a standardised, interviewer-administered questionnaire. Questions were asked about breastfeeding at 2 weeks of age and each month of age through 12 months (when applicable) and whether this breastfeeding was exclusive.

The Scottish Cot Death Trust Study

The Scottish Cot Death Trust study took place between January 1996 and May 2000. Data were collected on 131 SIDS cases and 278 control infants who were matched by age, season, and obstetric unit.⁷ Data on breastfeeding were collected by a standardised, interviewer-

administered questionnaire. Questions were asked about which types of feeding the infant had, and if not breastfed currently whether they had ever breastfed and when they stopped.

European Concerted Action on SIDS (ECAS)

The ECAS was comprised of case-control studies in 20 regions in Europe between September 1992 and April 1996.⁸ Data for the current analyses were restricted to those centres for which we had not obtained data from elsewhere (Sweden, Norway, Denmark, Netherlands, Austria, Hungary, Ukraine, Spain, Italy, Russia, Slovenia, France, Belgium, Poland and UK (Cambridge)). Data were collected for 382 SIDS cases and 1159 controls. Data on breastfeeding exclusivity and duration were collected by interviewer-administered questionnaires. Questions were asked about how the infant was being fed at the time of death or interview.

Irish Study of Infant Death

The Irish study was part of an ongoing case-control study of infant death in the Republic of Ireland that began collecting data in 1994 and continued until 2010.^{9,10} Controls were matched by date of birth and geographical location. The data included in this analysis comprise 363 cases and 1163 controls for the period 1994-2003. Data on breastfeeding exclusivity and duration were collected during standardized home interviews.

Confidential Enquiry into Stillbirth and Deaths in Infancy (CESDI)

CESDI was comprised of 5 regions of England between 1993 and 1996.¹¹ Data were collected for 325 SIDS cases and 1300 controls, who were matched by age and health visitor.

Data were collected for duration of breastfeeding; however, no information on duration of exclusiveness of breastfeeding was collected.

South-West England Infant Sleep Study (SWISS)

The SWISS comprised of 2 regions in the South-West of England between 2003 and 2006.¹²

Data were collected for 80 SIDS cases and 87 controls. Data were collected for duration of breastfeeding; however, no information on duration of exclusiveness of breastfeeding was collected.

Definitions of breastfeeding variables

Duration of any breastfeeding was defined as the length of time that the infant received any human milk, either through breastfeeding or expressed breast milk, either exclusively or in combination with other foods (including infant formula). We defined the duration of any breastfeeding as a continuous variable; we created a categorical variable for duration of any breastfeeding (0-2 months, 2-4 months, 4-6 months, and >6 months).

Duration of exclusive breastfeeding was defined as the length of time that the infant received only human milk, either through breastfeeding or expressed breast milk.¹³ We defined the duration of exclusive breastfeeding as a continuous variable; we created a categorical variable for duration of exclusive breastfeeding (0-2 months, 2-4 months, >4 months). A variable for >6 months was not created due to the very small numbers in this group in most of the studies.

Statistical analysis

Analysis was performed for each study individually, and then data were combined for a pooled analysis. A pooled univariable analysis, using all 8 studies, was conducted, controlling for study. A multivariable model was then fitted using 3 of the studies (NZCDS, CIMS and GeSID) for which all 19 potential confounders were available (Model 1). These confounders had initially been assessed as being available and consistent across these three studies at the inception of the study and have been identified as risk or protective factors for SIDS: sleep position at last sleep (supine, side, prone), maternal smoking during pregnancy (yes/no), bedsharing in the last sleep (infant sleeping with another person on the same surface) (yes/no), roomsharing in the last sleep (infant sleeping in the same room as an adult caregiver but on a separate surface) (yes/no), use of a dummy/pacifier in the last sleep (yes/no), maternal age, prenatal care received (yes/no), marital status (married/not married), parity (primiparous/multiparous), maternal education (university graduate or not), socio-economic status (SES) (low, middle, high), infant age (<13 weeks, 13-19 weeks, 20-26 weeks, >26 weeks), infant sex, admission to a special care baby unit (yes/no), season at death, birthweight (<2500g, 2500g-2999g, 3000g-3499g, ≥3500g), gestational age at birth (28-33 weeks, 34-37 weeks, 38+ weeks), multiple pregnancy (yes/no) and caesarean section (yes/no). Additional models were then fitted to include the other 5 studies at the expense of reducing the number of confounders but increasing sample size. These sequential models did not include the following confounders: caesarean section (CESDI and SWISS included in model, Model 2), SES and season (Irish and ECAS included, Model 3) and finally antenatal care and maternal education level (Scottish included, Model 4).

All analyses were carried out in SAS 9.4 (SAS Institute, Cary, NC, USA). Odds ratios were estimated using the proc logistic procedure, with a strata statement for study in pooled analyses. Survival curves were produced for duration of any breastfeeding for control groups using proc lifetest, with data censored if breastfeeding was still taking place. Statistical significance was defined at the 5% level.

This study was approved by the Institutional Review Board at the University of Virginia. Further, the individual studies were approved by the institutional ethical review boards/ethics committees according to the laws and standards of each country.

Results

There are 8 SIDS case-control studies with individual-level data; all were included (see Figure 1 flow diagram). A total of 2267 SIDS cases and 6837 control infants were included in this analysis. There was great variability in rates of any breastfeeding and exclusive breastfeeding in the studies (Log Rank=1659.6, $p<0.0001$). This is illustrated in Figure 2 which shows survival curves for any breastfeeding for controls from each of the studies. Breastfeeding rates were highest in New Zealand and lowest in the United States, with the European countries having intermediate rates. At six months the rate of any breast feeding ranged from over 50% in NZ and ECAS to less than 10% in several of the studies.

Any Breastfeeding

The univariable effects of any breastfeeding stratified by study and the pooled analyses are shown in Table 1. The analysis categorizing duration of any breastfeeding showed that those

who breastfed for less than 2 months incurred a protective effect (OR=0.61; 95% CI: 0.54, 0.69), and those breastfeeding 2-4 months had a greater protective effect (OR=0.26; 95% CI: 0.22, 0.30). Breastfeeding duration beyond 4 months provided further small increases in protection (4-6 months: OR=0.18; 95% CI: 0.14, 0.23; 6+ months: OR=0.13; 95% CI: 0.10, 0.18). The multivariable pooled analysis for the 3 studies with all 19 confounders controlled for found ongoing protective effects of any breastfeeding beyond two months (2-4 months: aOR=0.60; 95% CI: 0.44, 0.82; 4-6 months: aOR= 0.40; 95% CI: 0.26, 0.63, and 6+ months: aOR=0.36; 95% CI: 0.22, 0.61) (Table 2). However, breastfeeding for 0-2 months did not have a statistically significant protective effect (aOR=0.91; 95% CI: 0.68, 1.21). The removal of caesarean section from the model had little effect on the odds ratios; however the removal of SES and season in model 3 saw the protective effects of any breastfeeding become stronger. The further removal of maternal education and antenatal care in model 4 had little additional influence on the aOR, but this result reached statistical significance (aOR 0.83; 95% CI: 0.70, 0.99).

Exclusive breastfeeding

The stratified and pooled analysis for the univariable effects of exclusive breastfeeding is shown in Table 1. The analysis categorizing duration of exclusive breastfeeding showed that those who exclusively breastfed for less than 2 months incurred a protective effect (OR=0.61; 95% CI: 0.53, 0.71), and those breastfeeding 2-4 months had a greater protective effect (OR=0.25; 95% CI: 0.20, 0.30). Exclusive breastfeeding for >4 months provided a further increase in protection (OR=0.16; 95% CI: 0.12, 0.21). As in the multivariable analysis for any breastfeeding which controlled for all potential confounders, those who breastfed

exclusively for less than 2 months did not see any statistically significant protective effect (aOR=0.82; 95% CI: 0.59, 1.14), but those who breastfed for longer than 2 months incurred a protective effect (aOR=0.61; 95% CI: 0.42, 0.97) for 2-4 months, with increasing protection with longer duration (aOR=0.46;95% CI: 0.29, 0.74) for those exclusively breastfeeding >4 months. Similarly, the removal of SES and season from the model made the effect sizes slightly stronger.

Discussion

We conducted a pooled analysis of individual-level data from 8 major international case-control studies with 2259 cases and 6894 controls to assess the association between duration of any breastfeeding and exclusive breastfeeding and SIDS. Although there was some protection seen with breastfeeding for less than 2 months in univariable analysis, after controlling for potential confounders, we found no statistically significant protection against SIDS until infants had breastfed for at least 2 months. After 2 months, the adjusted odds ratio for any breastfeeding was 0.60 (95% CI=0.44, 0.82), while the adjusted odds ratio for exclusive breastfeeding was 0.61 (95% CI=0.42, 0.87). It is thus important that public health messages about SIDS risk reduction emphasize that breastfeeding, if it to be protective, must continue for at least 2 months. This analysis does not demonstrate any advantage to exclusive breastfeeding over partial breastfeeding, which may be reassuring to some parents who cannot or do not wish to exclusively breastfeed their infant.

It is yet unclear why breastfeeding offers protective effects against SIDS. Physiologic, neuropathologic, and genetic studies point to dysfunctional arousal responses as a mechanism

that creates an intrinsic vulnerability in the infant, which predisposes the infant to SIDS,¹⁴ and breastfed infants are more easily aroused from sleep than are formula-fed infants.^{15,16} There are also differences in maternal responses to an infant's behavioral cues, depending on feeding mode, which may impact infant sleep and arousal patterns.^{17,18} Additionally, breastfeeding provides immune benefits and is associated with a lower incidence of viral infections, which are associated with an increased risk of SIDS.¹⁹⁻²¹ Breast milk contains substances that may contribute to myelin development; Kinney and co-authors found that infants who died from SIDS had delayed myelination of the brain compared with control infants.²² Breast milk also contains higher levels of docosahexaenoic acid (DHA) compared with formula, which is an important structural and functional component of the developing infant brain. One study of autopsied brains of SIDS infants found that the frontal lobes of the breastfed infants had higher levels of DHA, compared with those of formula-fed infants, it is unknown if this difference exists in non SIDS infants.²³ Finally, it is possible that breastfeeding is a distal marker of or proxy for complex protective infant care practices that have not yet been measured, though we would expect that such a marker would be related to socio-demographic variables that have been controlled for in these analyses.

It is unclear why exclusive breastfeeding did not offer any additional protection against SIDS than any, i.e., partial breastfeeding. This is a common challenge in studies that have looked at the differential effects of exclusive and partial breastfeeding, because of differing definitions of breastfeeding and confounding factors.^{1,24} The analysis accounted for as many demographic and risk factor variables as were possible, but we acknowledge that the effects reported could be due to residual confounding, though this would be unlikely. It was notable

that inclusion of studies that did not have data on SES increased the protective effect further from the null, thus seemingly showing the importance of SES as a confounder in relation to breastfeeding. Given that lower SES is a risk factor for SIDS, it is possible that the protective effect of SES may in part be explained by increased breastfeeding rates. However, Model 3, which did not have data on SES also did not have data on season. While SES is associated with breastfeeding it is unlikely there is a relationship between season and breastfeeding, thus we believe that these changes in estimates are likely to be associated with SES.

Other limitations of this study are related to issues with combining data in the individual case-control studies. These case-control studies were all conducted in a rigorous manner and are the basis for most of the current infant safe sleep guidelines in developed countries.²⁵⁻²⁷ However, as noted above, the variables collected in the course of each study varied slightly, limiting our ability to include all studies in the analysis controlling for all confounders. However, the results of the univariable analysis using just the 3 countries included in the completely controlled multivariable model (model 1) did not differ greatly from the univariable analysis with all 8 studies, so it is unlikely that including the additional studies would have changed the results of the analysis in any meaningful way.

Given these findings, there should be ongoing concerted efforts to increase rates of breastfeeding initiation and maintenance. Among the control infants in five of the eight countries in this analysis, the proportion of infants who were breastfeeding at 2 months of age was <50%, and <30% at 4 months of age. In more recent years, national breastfeeding rates have increased; 2007 OECD data show that the proportions of infants who were ever

breastfed in the countries included in our study were 42% in Ireland, 75% in the US, 77% in the UK, 85% in New Zealand, and 89% in the European Union.²⁸ The World Health Organization's 2025 targets for breastfeeding are to have >50% of infants exclusively breastfeeding for at least 6 months.¹³ Further increases in breastfeeding rates will result in lower infant mortality as a whole,^{24,29} and decreases in SIDS rates,³ specifically.

In conclusion, breastfeeding duration of a minimum of 2 months appears to be necessary to confer a significant protective effect against SIDS, with an almost halving of the risk. The protective benefits of breastfeeding increase as the duration increases. However, exclusive breastfeeding does not confer additional benefits over partial breastfeeding with regards to SIDS risk reduction. Therefore, mothers should be encouraged to breastfeed for at least 2 months (and preferably longer). Even if mothers are unable to exclusively breastfeed, they can feel reassured that any breastfeeding provides protection against SIDS to their infants. Further study is still needed to better understand the mechanisms by which breastfeeding offers protection.

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Figure legends

Figure 1: PRISMA flow diagram for pooled analyses using individual-level data

Figure 2: Kaplan Meier survival curves for proportion of controls still Breastfeeding, stratified by study

Figure 2

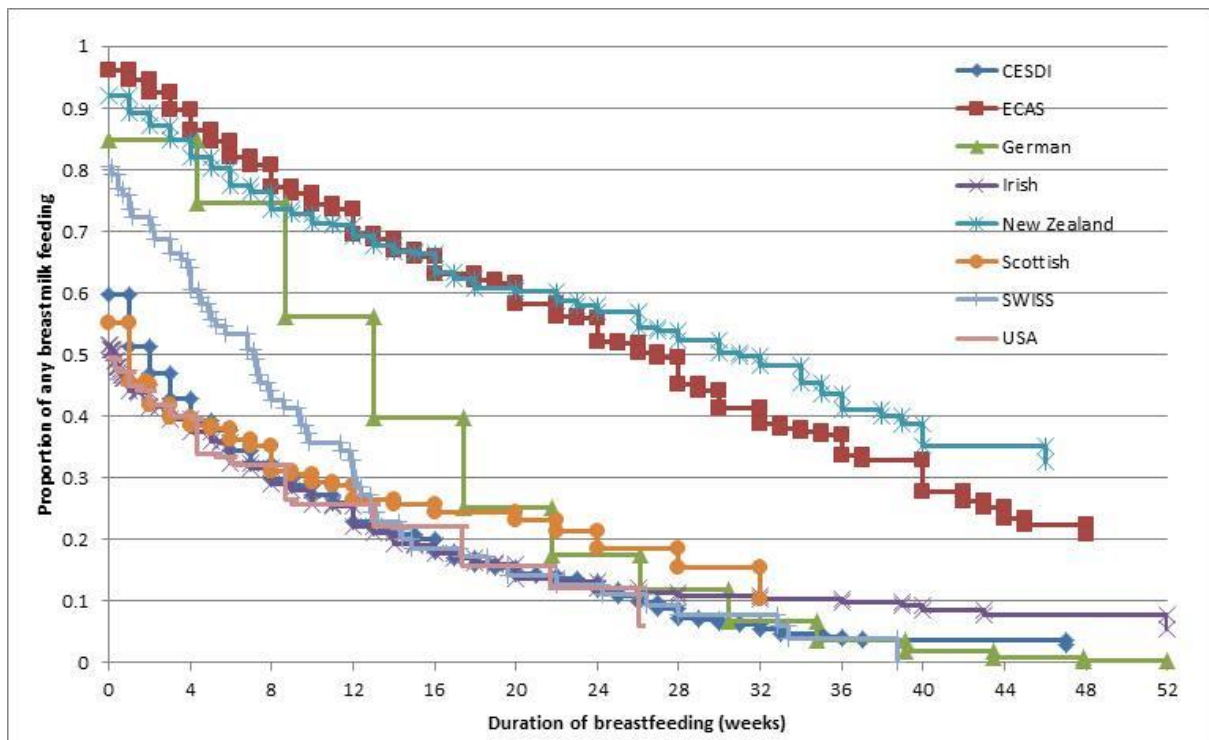


Table 1: Stratified and pooled univariable odds ratio (95% CI) of SIDS for duration of any and exclusive breastfeeding

Any breastfeeding (months)					
	New Zealand	Germany	USA	Scottish	Pooled
Never	1.00	1.00	1.00	1.00	
>0 to 2	0.79 (0.55, 1.13)	0.70 (0.48, 1.02)	0.43 (0.28, 0.65)	0.65 (0.41, 1.05)	
>2 to 4	0.49 (0.33, 0.71)	0.18 (0.13, 0.25)	0.07 (0.03, 0.21)	0.16 (0.06, 0.47)	
>4 to 6	0.32 (0.19, 0.53)	0.10 (0.06, 0.16)	0.06 (0.01, 0.51)	0.20 (0.04, 0.90)	
>6 months	0.19 (0.09, 0.37)	0.15 (0.10, 0.25)	undefined	undefined	
	ECAS	CESDI	SWISS	Irish	
Never	1.00	1.00	1.00	1.00	1.00
>0 to 2	0.56 (0.36, 0.88)	0.61 (0.46, 0.81)	0.79 (0.37, 1.67)	0.68 (0.51, 0.92)	0.61 (0.54, 0.69)
>2 to 4	0.21 (0.13, 0.33)	0.42 (0.28, 0.64)	0.42 (0.16, 1.12)	0.16 (0.09, 0.29)	0.26 (0.22, 0.30)
>4 to 6	0.20 (0.12, 0.33)	0.39 (0.20, 0.78)	0.15 (0.02, 1.40)	0.04 (0.01, 0.26)	0.18 (0.14, 0.23)
>6 months	0.07 (0.03, 0.14)	0.25 (0.09, 0.70)	0.11 (0.01, 0.95)	0.22 (0.07, 0.72)	0.13 (0.10, 0.18)
Exclusive breastfeeding					
	New Zealand	Germany	USA	Scottish	Pooled
Never	1.00	1.00	1.00	1.00	
>0 to 2	0.88 (0.58, 1.34)	0.40 (0.28, 0.56)	0.46 (0.28, 0.76)	0.63 (0.39, 1.03)	
>2 to 4	0.41 (0.27, 0.64)	0.22 (0.15, 0.33)	0.18 (0.02, 1.51)	0.21 (0.06, 0.73)	
4-6	0.32 (0.10, 1.03)	0.13 (0.09, 0.20)	undefined	undefined	
	ECAS	CESDI	SWISS	Irish	
Never	1.00	Not available	Not available	1.00	1.00
>0 to 2	0.98 (0.73, 1.32)			0.40 (0.28, 0.58)	0.61 (0.53, 0.71)
>2 to 4	0.29 (0.21, 0.41)			0.12 (0.05, 0.30)	0.25 (0.20, 0.30)
4-6 months	0.19 (0.12, 0.31)			0.48 (0.16, 1.41)	0.16 (0.12, 0.21)

Table 2: Stratified and pooled multivariable odds ratios (95% CI) of SIDS for duration of any breastfeeding

	New Zealand	Germany	USA	Scottish
Never	1.00	1.00	1.00	1.00
>0 to 2	0.86 (0.53, 1.40)	0.89 (0.49, 1.63)	0.69 (0.36, 1.31)	0.69 (0.31, 1.52)
>2 to 4	0.67 (0.40, 1.11)	0.51 (0.29, 0.88)	0.16 (0.04, 0.71)	0.38 (0.09, 1.54)
>4 to 6	0.39 (0.19, 0.80)	0.37 (0.18, 0.74)	0.16 (0.01, 1.72)	0.20 (0.03, 1.57)
>6 months	0.44 (0.17, 1.13)	0.30 (0.15, 0.63)	undefined	undefined
	ECAS	CESDI	SWISS	Irish
Never	1.00	1.00	1.00	1.00
>0 to 2	0.79 (0.25, 2.51)	0.96 (0.65, 1.40)	0.12 (0.01, 2.18)	1.13 (0.59, 2.17)
>2 to 4	0.82 (0.25, 2.73)	0.78 (0.45, 1.34)	0.02 (<0.001, 0.93)	0.19 (0.07, 0.51)
>4 to 6	0.94 (0.23, 3.94)	0.64 (0.24, 1.75)	<0.001 (<0.001, 3.50)	0.08 (0.01, 0.86)
>6 months	0.06 (0.00, 0.94)	0.26 (0.05, 1.25)	0.001 (<0.001, 1.71)	0.45 (0.06, 3.09)
	Pooled Model 1^a	Pooled Model 2^b	Pooled Model 3^c	Pooled Model 4^d
	n=3386	n=5008	n=6121	n=7842
Never	1.00	1.00	1.00	1.00
>0 to 2	0.91 (0.68, 1.21)	0.90 (0.72, 1.12)	0.83 (0.69, 1.01)	0.83 (0.70, 0.99)
>2 to 4	0.60 (0.44, 0.82)	0.62 (0.48, 0.80)	0.52 (0.41, 0.65)	0.46 (0.37, 0.56)
>4 to 6	0.40 (0.26, 0.63)	0.42 (0.29, 0.61)	0.38 (0.27, 0.54)	0.40 (0.30, 0.53)
>6 months	0.36 (0.22, 0.61)	0.34 (0.22, 0.54)	0.33 (0.21, 0.50)	0.25 (0.17, 0.37)

^a Model 1 controlled for sleep position at last sleep, maternal smoking during pregnancy, bedsharing in the last sleep (infant sleeping with another person on the same surface), roomsharing in the last sleep (infant sleeping in the same room as an adult caregiver but on a separate surface), dummy/pacifier in the last sleep, maternal age, prenatal care, marital status, parity, maternal education, socio-economic status, infant age, infant sex, admission to a special care baby unit, season at death, birthweight, gestational age, multiple pregnancy and caesarean section.

^b Model 2 controlled for variables in model 1 except for caesarean Section to include CESDI and SWISS studies.

^c Model 3 controlled for variables in model 2 except season and socio-economic status to include ECAS and Irish studies.

^d Model 4 controlled for variables in model 3 except for antenatal care and maternal education to include Scottish study.

Table 3: Stratified and pooled multivariable odds ratios (95% CI) of SIDS for duration of exclusive breastfeeding

	New Zealand	Germany	USA	Scottish
Never	1.00	1.00	1.00	1.00
>0 to 2	1.02 (0.56, 1.84)	0.70 (0.41, 1.19)	0.81 (0.39, 1.69)	0.61 (0.26, 1.44)
>2 to 4	0.47 (0.27, 0.83)	0.51 (0.29, 0.90)	0.61 (0.04, 8.83)	0.63 (0.12, 3.23)
4-6 months	0.56 (0.15, 2.07)	0.31 (0.17, 0.58)	undefined	Undefined
	ECAS	CESDI	SWISS	Irish
Never	1.00	Not available	Not available	1.00
>0 to 2	1.27 (0.46, 3.48)			0.68 (0.33, 1.42)
>2 to 4	0.48 (0.15, 1.53)			0.09 (0.02, 0.53)
4-6 months	0.60 (0.14, 2.54)			3.14 (0.56, 17.55)
	Pooled Model 1^a	Pooled Model 2^b	Pooled Model 3^c	Pooled Model 4^d
	n=3397		n=4319	n=6006
Never	1.00		1.00	1.00
>0 to 2	0.82 (0.59, 1.14)		0.75 (0.58, 0.98)	0.82 (0.67, 1.01)
>2 to 4	0.61 (0.42, 0.87)		0.44 (0.32, 0.60)	0.40 (0.31, 0.51)
4-6 months	0.46 (0.29, 0.74)		0.47 (0.31, 0.71)	0.37 (0.26, 0.52)

^a Model 1 controlled for sleep position at last sleep, maternal smoking during pregnancy, bedsharing in the last sleep (infant sleeping with another person on the same surface), roomsharing in the last sleep (infant sleeping in the same room as an adult caregiver but on a separate surface), dummy/pacifier in the last sleep, maternal age, prenatal care, marital status, parity, maternal education, socio-economic status, infant age, infant sex, admission to a special care baby unit, season at death, birthweight, gestational age, multiple pregnancy and caesarean section.

^b Model 2 excluded CESDI and SWISS studies, which had no data on exclusive breastfeeding. Not available as these two studies had no data on exclusive breastfeeding

^c Model 3 controlled for variables in model 2 except for season and socio-economic status to include ECAS and Irish studies.

^d Model 4 controlled for variables in model 3 except for antenatal care and maternal education to include Scottish study.