

# Infantile Colic, Is There An Association With The Source Of Early Infant Nutrition?

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## Abstract

### Objective

To determine whether breastfeeding has an impact on colic development.

### Design

A prospective cohort study of 436 mother-infant dyads who give birth at term to alive singleton whose birth weight was appropriate for gestational age. Self-administered questionnaires to mothers at 1 and 6 weeks postpartum, requested information on many infant and maternal factors, including source of infant nutrition (exclusively breast fed, complementary fed and exclusively formula fed). Cases of colic were identified by applying modified Wessel criteria or by interpreting the response to the Ames cry score.

### Results

Of 536 mothers, 436 (81%) completed the study. Overall the prevalence of colic at 6 weeks of age was 23.6%. No association was seen between the source of infant nutrition and colic. Higher levels of maternal trait anxiety were associated with increased likelihood of colic after controlling for feeding method, maternal age, and parity although the difference failed to reach significance. No significant difference was seen between employed and unemployed mothers, but some difference was seen between both employed and unemployed mothers in one side and the student mothers in the other side, although the difference was statistically insignificant.

### Conclusion

Breast feeding did not have a protective effect on the development of colic. Although colic was statistically associated with some variables, including pre-existing maternal anxiety and student mothers, some of colic's etiology remains unexplained despite the numerous studies done in this field.

## Introduction

The most frequently accepted definition of colic is that of Wessel of a well thriving infant who has paroxysms of crying lasting for three hours per day for more than three days a week for longer than three weeks [1]. Schmitt suggested the practical definition of "colic as an intermittent unexplained crying during the first three months that reaches a point where parents complain about it and starts before 2 weeks of age and usually stops by 3- 4 months of age, it is not the result of bad parenting, nor caused by too much gas [2]. Colic's onset usually occurs between the 2<sup>nd</sup> and 6<sup>th</sup> weeks of life and its disappearance around 3 months. Its prevalence ranges from 5% to 40%

depending on the definition and methods used. It remains enigmatic, despite its relatively frequent occurrence.

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While apparently innocuous, an infant colic can prompt lasting parental distress [3, 4, 5]. Gastrointestinal causes [6, 7, 8] psychological causes [9, 10, 11] and hormonal factors “increase production of serotonin unopposed by the production of melatonin” [11, 12, 13] have been suggested for the colic’s development. Other investigators found the correlation between infant colic and allergic products of food and advised the use of hypoallergenic formula in the management of some cases of infant colic [14, 15, 16]. An interview of literature, doctors who study colic, have interesting observations: Colicky infants have an increased amount of the hormone motilin in their blood which stimulates gastrointestinal tract to cause cramping [17]. Colicky infants have abnormal contractility of the gallbladder and have increased intestinal permeability [18]. Recently, the composition of intestinal micro flora has been attributed as an independent risk factor for colic. Studies indicate that inadequate amounts of lactobacillus and increased amounts of coli form bacteria like E coli, Klebsiellosis and anaerobes in the intestine may influence gut motor function and gas production which subsequently contributes to the occurrence of colic [19]. Replacing micro flora with probiotic was found to have remarkable improvement and less crying episodes [20]. More recently, the cause of infant colic was uncovered and attributed to Helicobacter pylori infection [21]. The crying of a colicky infant has typical pattern described as loud continuous high pitched cry. It may be accompanied by tensing of the abdominal muscles, flexing the legs, flushing of the face, cold feet, clenching fingers and passing gases. The peak period of crying usually occurs at evening. The attack may terminate when the infant is completely exhausted or with the passage of flatus [22]. Mothers of excessively crying infants experience high levels of depression, and fathers’ depression during the postpartum period is highly correlated with mother depression. Furthermore, depression in fathers during the postpartum period has deleterious effects on infant and child outcomes as the depressed fathers have less optimal interactions with their infants [23].

## Materials and Method

Approval of this study was received from the administration of Armed Forces Hospital in Najran, Saudi Arabia. The study was a prospective cohort study of mother-infant dyads. A self-administered questionnaire to mothers at one and six weeks postpartum including, maternal health behaviors, demographic variables,

biological factors and the current source of infant nutrition (exclusive breastfed, complementary fed and exclusively formula-fed). Infants who received any quantity of non-human milk along with breast milk were defined as complementary fed. Standardized **measures** that assessed maternal anxiety, postnatal depression and social support [24, 25] were incorporated into the first questionnaire because these constructs have been suspected to play a role in colic’s development. The psychometric properties of the state-trait anxiety inventory are well established. So, this scale was used in this study. The state-trait anxiety inventory is composed of 2 scales that measure 2 distinct anxiety concepts: state anxiety and trait anxiety. The trait scale asks participants how they generally feed and is indicative of anxiety proneness [26, 27]. The state anxiety asks participants how they currently feel and sensitivity indicates participant’s levels of transitory anxiety. Possible scores range from 20-80 on each scale [28, 29]. Cases of colic were identified as those infants whose cry/ fuss behaviors fulfilled modified Wessel criteria. For this study, a colicky infant was defined as one who was otherwise healthy and well fed, but who had paroxysms of irritability, fussing, or crying lasting for three or more hours in any one day and occurring on three or more days in any week. Mothers were also asked to complete a short questionnaire based instrument, the Ames cry score, which is composed of three questions, each with four response categories that are scored from 0-3. It asks about the frequency, average and maximum duration of an infant’s cries during the past week. Overall scores, calculated by summing the scores of individual items, range from 0-9, with a score of 3 or greater indicating colic. The overall prevalence of colic at 6 weeks of age was calculated among exclusively breast fed, exclusively formula fed and complementary fed infants. A questionnaire was mailed to mothers approximately one week after they were discharged from the hospital. Another questionnaire was mailed to the mothers at six weeks and four months after delivery. For all analyses, the exposure variable was the source of infant nutrition at one week postpartum, while the outcome was the colic defined at 6 weeks postpartum. This approach addressed one of the shortcomings of earlier works because it permitted an examination of the potential temporal relationship between the source of infant nutrition and colic’s development.

## Results

Eligible cases were approached between November 2013 and October 2014. About 68% of

participants were breast feeding their infants at 6 weeks postpartum. Only 81% of cases completed the study at 6 weeks postpartum. Overall the prevalence of colic was 23.6% when examined separately. The prevalence of colic based on the interpretation derived from the Ames Cry Score was 22%. The mean amount of crying/fussing recorded during an infant sixth weeks of life was more than 10 hours (SD= 6.1 hrs.) while this is 1.5 hr./day. The prevalence of colic among exclusively breast fed, exclusively formula fed and complementary fed infants was 24%,21% and 25% respectively (Table 2). There was

no statistically significant relationship between the source of early infant nutrition, measured at one week postpartum and the development of colic at 6 weeks postpartum. In the 436 mother-infant dyads participating in the study at 4 months, 17 infants (3.9%) had colic at 4 months of age, compared with 103 infants (23.6%) who had colic at six weeks of age. The mothers of infants who had colic at six weeks of age did not have significant differences in rates of anxiety and depression, compared with the mothers of infants who did not have colic.

**Table 1.** Characteristics of the study sample

| Variable                                       | Participants<br>n= 436 (81.3%) | Nonparticipants<br>n= 100 (18.7%) | P Value |
|--|--------------------------------|-----------------------------------|---------|
| <b>Mode of delivery</b>                        |                                |                                   |         |
| • Vaginal                                      | 367(80.7)                      | 88 (19.3)                         | 0.47    |
| • Cesarean section                             | 69 (85.2)                      | 12 (14.8)                         |         |
| <b>Maternal age (Yr.)</b>                      |                                |                                   |         |
| • <20  | 12 (75.0)                      | 4(25.0)                           | 0.000   |
| • 20-35  | 370 (81.7)                     | 83(18.3)                          |         |
| • >35  | 54 (80.6)                      | 13(19.4)                          |         |
| <b>Parity</b>                                  |                                |                                   |         |
| • Primipara                                    | 242 (81.2)                     | 56(18.8)                          | 0.49    |
| • Multiparous                                  | 194 (81.5)                     | 44(18.5)                          |         |
| <b>Maternal education</b>                      |                                |                                   |         |
| • < High school                                | 76 (79.2)                      | 20(20.8)                          | 0.000   |
| • High school                                  | 221(81.0)                      | 52(19.0)                          |         |
| • >High school                                 | 139 (83.2)                     | 28(16.8)                          |         |
| <b>Employment status 2 mo. Before delivery</b> |                                |                                   |         |
| • Unemployed (Housewife)                       | 288(82.1)                      | 63(17.9)                          | 0.000   |
| • Student                                      | 36(78.3)                       | 10(21.7)                          |         |
| • Employed                                     | 122(76.7)                      | 37(23.3)                          |         |

-Data are given as number (percentage) of mothers. Percentage are based on row totals.

**Table 2.** Factors associated with the development of colic at 6 weeks' age

| Factor                             | Colic             |                   | P value |
|------------------------------------|-------------------|-------------------|---------|
|                                    | Yes               | No                |         |
| <b>Source of infant nutrition.</b> |                   |                   |         |
| -Breast milk only                  | 30(24.0)          | 95(76.0)          | 0.000   |
| -Formula only                      | 29(21.2)          | 108(78.8)         |         |
| -Breast milk and powder            | 44(25.3)          | 130(74.7)         |         |
| <b>Sex</b>                         |                   |                   |         |
| -Male                              | 51(25.8)          | 147(74.2)         | 0.000   |
| -Female                            | 52(21.8)          | 186(78.2)         |         |
| <b>Maternal employment</b>         |                   |                   |         |
| -Unemployment                      | 31(19.4)          | 129(80.6)         | 0.001   |
| -Student                           | 22(27.2)          | 59(72.8)          |         |
| -Employment                        | 44(22.6)          | 151(77.4)         |         |
| <b>Marital status</b>              |                   |                   |         |
| -Single                            | 6(26.1)           | 17(73.9)          | 0.04    |
| -Partner                           | 90(21.8)          | 323(78.2)         |         |
| State anxiety, Mean(SD) (SEM)      | 34.87(9.5) (0.95) | 32.12(9.8) (0.53) | 0.014   |
| Trait anxiety, Mean(SD) (SEM)      | 36.72(8.2) (0.82) | 34.56(8.3) (0.45) | 0.026   |

-Data are presented as number(percentage) and percentages are based on row totals. The factors not significantly associated with colic at 6 weeks of age are: infant sex, mode of delivery, family history of atopy, maternal education level, and maternal smoking or caffeine consumption during pregnancy and postnatal depression.

-Other sociodemographic variables such as alcohol consumption, family income, etc. are not included due to the conservative culture and the approximate social level of cases in this community.

Table 3. Factors associated with colic remission at 4 months

| Variable                                    | Colic persisted<br>no=17(16.5%) | Colic remitted<br>no=86(83.5%) | P value |
|---|---------------------------------|--------------------------------|---------|
| <b>Infant sex</b>                           |                                 |                                |         |
| -Male                                       | 14(82.4)                        | 40(46.5)                       | 0.003   |
| -Female                                     | 3(17.6)                         | 46(43.5)                       |         |
| <b>Employment in pregnancy</b>              |                                 |                                |         |
| -Unemployment                               | 6(35.3)                         | 36(41.9)                       | 0.01    |
| -Student                                    | 1(5.9)                          | 6(7.0)                         |         |
| -Employment                                 | 10(58.8)                        | 44(51.2)                       |         |
| <b>Marital status</b>                       |                                 |                                |         |
| -Single                                     | 2(11.8)                         | 4(4.7)                         | 0.000   |
| -Partner                                    | 15(88.2)                        | 82(95.3)                       |         |
| <b>Parity</b>                               |                                 |                                |         |
| -Primipara                                  | 11(64.7)                        | 38(44.2)                       | 0.004   |
| -Multipara                                  | 6(35.3)                         | 48(55.8)                       |         |
| <b>Source of infant nutrition</b>           |                                 |                                |         |
| -Breast fed only                            | 7(41.2)                         | 27(31.4)                       | 0.002   |
| -Formula only                               | 5(29.4)                         | 30(34.9)                       |         |
| -Both breast and formula                    | 5(29.4)                         | 29(33.7)                       |         |
| Trait anxiety <sup>§</sup> change* Mean(SD) | -1.80(8.95)                     | -2.60(7.58)                    | 0.34    |
| Postnatal depression" change* Mean(SD)      | -1.12(4.78)                     | -1.82(4.45)                    | 0.12    |

-Data are presented as the number of dyads, restricted to the 103 infants with colic at 6 weeks of age. Other factors not found to be significantly associated with persistence of colic at 4 months included, maternal age, marital status, parity, mode of delivery, history of atopy, maternal education, postnatal depression and source of infant nutrition [30].

<sup>§</sup>Measured by means of the State-Trait Anxiety Inventory [31]

"Measured by means of the Edinburgh Postnatal Depression Scale [32]

\*Change in the score is measured as the difference between the score at 4 months and the score at 1 week postpartum.

### Factors Associated with the Development of Colic at 6 Weeks of Age

If a mother was employed or attended school full-time before being delivered of an infant, her infant was significantly more likely to exhibit colic. Infants born to mothers who were married or had a common-law partner were less likely to exhibit colic, relative to infants born to single mothers. Elevated levels of postnatal depression also seemed associative with colic's development, but this association failed to reach statistical significance. At 4 months of age, colic persisted in 16.5% of cases. Additionally, the trait anxiety score and the postnatal depression score changed at 4 months of age, compared to the basal scores at one week postpartum. Because pacifier use was expected to affect breastfeeding and infant cry/fuss behavior, we considered it as a potential confounder.

However, the evidence that pacifier use does not markedly affect the daily duration of cry/fuss behavior [30] prompted our decision not to include pacifier use in the multivariate model.

### Comment

Despite using prospective methods, an objective case definition, a population-based relatively large sample and multivariate techniques, our results failed to document any relationship between the source of early infant nutrition and colic's development. Unlike other studies, potential etiologic agents including the source of infant nutrition and levels of maternal distress were assessed before the development of colic, allowing for the exploration of temporal relationships. Additionally, the study provides evidence that most cases of infant colic

remit by 4 months with little lasting impact on the levels of maternal anxiety or depression. Admittedly, because the participants in our study were, on average, well educated, our findings may not generalize to populations that differ in their socioeconomic background and/or access to health care. Nevertheless, the internal validity of our study is high because of the inclusive nature of the study population, the representative sample, and the high response rates. This internal validity lends support to our conclusion that, in most cases, recommendations to alter an infant's diet in the hope of ameliorating colic are not logically established. Moreover, we examined the potential for an episode of colic to adversely affect maternal mental health after controlling for earlier pre-colic levels of anxiety and depression. Elevated levels of maternal anxiety have received considerable attention as a contributing factor in colic's development [33-35]. Elevated levels of maternal trait anxiety, measured at 1 week postpartum, predicted colicky behavior at 6 weeks; however, this finding failed to reach statistical significance. Our results suggest that about 1 in 4 infants experienced colic at 6 weeks of age. The collection of cry/fuss data at this point of time was expected to capture most cases of colic because the sixth week of life represents the peak of infant crying [36]. Our results document no any relationship between the source of early infant nutrition and colic's development. The absence of a protective effect of breastfeeding on colic's development is firmly considered across time, geography, and study design [37-41]. The assumption that infantile colic is developmental, psycho-biological problem (i.e. developmental sleep disorder) suggested by some investigators [42] is not an evidence-based theory, and their study designed to test the hypothesis that kangaroo care is more effective than conventional care on infantile colic. Moreover, their work was conducted on preterm infants. In this model, colic is viewed as a delay or disturbance in the biorhythmic organization of the infant and colicky infant becomes overly stimulated and overly tired and is less able to initiate a downward shift in the state from awake and crying to a sleep state. This recurrent cycle is theorized to affect the overall interact total synchrony of the mother-infant dyad.

Although continued uncertainty about colic's etiology may be viewed in a negative light, our findings should provide comfort to parents as we provide further evidence that parents are not responsible for their infants' colic. Of course, much work remains to be done to improve our understanding of the underlying processes responsible

for colic and to minimize their potential effects on families. Colic's relatively frequent occurrence and potential sequelae command our continued attention. Thus, the author's novel finding of "Helicobacter pylori might be the causative agent of infant colic" is a clearly relevant work of attention. Further research needs to determine an optimal manner for identifying infants who are at an elevated risk of state regulatory functional disorders.

### **What Does This Study Add?**

Infant colic is a relatively common condition, characterized by excessive and inconsolable crying as formerly estimated. Some have suggested that food intolerance is to blame but this has limited confidence. Because numerous intervention strategies are based on dietary incompatibility theories, there is a need to determine whether this focus is justified. By using prospective methods in a community sample of infants, this study addressed many of the methodological issues of previous works. In doing so, the study provided evidence that about 1 in 4 infants exhibited colicky behavior at 6 weeks of age, suggested that breastfeeding was not protective in colic's development, and although colic was statistically associated with several factors, including preexisting maternal anxiety, the clinical importance of this finding is limited. So, while the origins of colic remain a mystery, its clinical salience cannot be questioned. This stimulates recalling other related functional disorders that are commonly occurring (i.e. recurrent abdominal pain of childhood and irritable bowel syndrome of adolescence). This underlies the importance of continued research particularly in the field of molecular genetics approach. The future studies should be centered and directed to the genetic basis of these interrelated functional disorders.

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