Proctocolitis in breast fed infants: a contribution to differential diagnosis of haematochezia in early childhood

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Abstract
Dietary protein induced proctocolitis in exclusively breast fed infants is rarely taken into consideration as a cause of rectal bleeding or blood streaked stool in the neonatal period and early infancy. Eleven babies are presented in whom it is believed that bleeding through the rectum was due to proctocolitis as a result of allergy triggered by cows’ milk protein transferred to the infants via the breast milk. Colonoscopy was performed in five infants, revealing benign eosinophilic proctocolitis. Standard treatment was the exclusion of the allergen from the mother’s diet. Resolution of visible rectal bleeding took place within 72 to 96 hours after elimination of the offending protein from the mother’s diet.

Keywords: haematochezia; benign eosinophilic infantile proctocolitis; breast fed infants; cows’ milk allergy

Dietary protein induced eosinophilic proctocolitis in exclusively breast fed infants is rarely taken into consideration as a cause of rectal bleeding in the neonatal period or early infancy. Failure to appreciate this entity may lead to inappropriate diagnostic or therapeutic intervention. The aim of our presentation is to describe clinical signs and symptoms, diagnosis, and treatment of this condition, which is proving to be of increasing importance in neonates and infants.

Methods and results (see table 1)
During a period of 18 months 11 babies were admitted to our department because of rectal bleeding or blood streaked stool. All 11 children (six boys and five girls) were born at term and were exclusively breast fed up to the time of our initial evaluation. The mean age at the onset of symptoms was 7.5 weeks (range 5–10 weeks) and referral for evaluation occurred from the first to the third week after onset. Predominant signs and symptoms were repeated mild rectal bleeding, blood streaked bowel movements, and mucous blood mixed with stool.

Physical examination showed all babies apparently well, thus serious conditions could be ruled out. No anal fissures or perianal dermatitis were found to account for the bleeding. Four children presented with colicky pain and abdominal distension. In only one case was there a possible family history of cows’ milk allergy and a known history of atopy.

Serum haemoglobin was shown to be below the normal range in two infants and investigation of serum albumin in seven infants yielded hypoalbuminaemia in one child. Three children demonstrated peripheral eosinophilia (range 8%–24%). Stool cultures were negative for routine pathogens including ova and parasites.

In five children rectocolonoscopy up to the splenic flexure was performed under general anaesthesia, showing macroscopic evidence of colitis (patchy focal or diffuse mucosal oedema and erythema, friability of the mucosa and small erosions). Involvement of the rectosigmoid was universal, whereas extension into the descending colon was limited. In four infants inflammatory lesions appeared together with nodular lymphoid hyperplasia (“goose pimply appearance”; fig 1). Histological examination of biopsy specimens revealed colitis with hyperaemia and predominance of eosinophils in the epithelium, lamina propria, and submucosa.

In the next six babies with the identical clinical signs and symptoms, invasive examination was not carried out under the assumption of curred from the first to the third week after onset. Predominant signs and symptoms were repeated mild rectal bleeding, blood streaked bowel movements, and mucous blood mixed with stool.

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| Table 1 Clinical data of breast fed children with allergic proctocolitis |
|-------------------------------|------------------|
| Mean age at onset             | 7.5 weeks (5–10 weeks) |
| Previous hospital care due to rectal bleeding | 4/11 |
| Haematochezia or blood streaked stool | 9/11 |
| Gross bleeding through the rectum | 2/11 |
| Diarrhoea (frequent bowel movements) | 11/11 |
| Abdominal pain or distension | 4/11 |
| Anal fissure or perianal dermatitis | 0/11 |
| Anaemia                     | 2/11 |
| Hypoalbuminaemia (<35 g/l)  | 1/7 |
| Leucocytosis (>12 × 10⁹/l)  | 2/11 |
| Peripheral eosinophilia (>6%) | 3/11 |
| Rectocolonoscopy              | 5/11 |
| Eosinophilic colitis         | 5/5 |
| Lymphonodular hyperplasia    | 4/5 |

Figure 1 Endoscopic aspect of lymphonodular hyperplasia of the colon showing multiple 1–4 mm nodules of the colon (“goose pimply appearance”).
Protocolitis due to allergy triggered by cows’ milk protein. All mothers were instructed to exclude cows’ milk from their diet, but none of them replaced breast feeding by formula. Resolution of visible rectal bleeding took place within 72 to 96 hours after elimination of the offending protein. In one case, having presented with gross blood loss, resolution took one week. Reintroduction of cows’ milk in eight children after a period of one year did not result in recurrence of signs and symptoms.

**Discussion**

Rectal bleeding and haematochezia, particularly in neonates and infants, give understandable reason for serious concern and parental anxiety leading to urgent investigation and hospital care. A useable management algorithm should provide the practitioner with a better appraisal of unexplained rectal bleeding in infants. It is important to determine when a case of rectal bleeding should be aggressively evaluated (for example, volvulus, necrotising enterocolitis, intussusception). If the episode of the bleeding has been mild or self limiting, it is reasonable to defer extensive investigation (fig 2).

Although allergic proctocolitis due to cows’ milk or soy protein is a significant cause of rectal bleeding in infants, dietary protein induced proctocolitis in apparently exclusively breast fed infants has been less emphasised. A concept of a classical type IV hypersensitivity reaction at the level of the colonic mucosa is disputed, although the exact responsible mechanisms remain unknown. Proteins in the maternal diet have been recovered in breast milk and may act as an allergen.

A presumptive diagnosis of proctocolitis can be established on the ground of a precise case history. Onset occurs in the first four months of life (range 1 week to 5 months), and apart from the bleeding the exclusively breast fed infants are apparently well. Mild-to-moderate diarrhoea or grossly blood streaked stool with mucus are the consistent features. No anal fissures or perianal dermatitis are found to account for the bleeding. Signs and symptoms of malabsorption, abdominal distension, severe diarrhoea, vomiting or impaired growth, which would indicate food allergy, are absent. Features of type I hypersensitivity such as anaphylaxis and rash are lacking. Infectious origin can be excluded by stool tests including ova and parasites.

Peripheral eosinophilia, raised serum IgE concentration, a positive family history of allergy, and positive skin sensitivity tests may be present, but are not pathognomonic.

In ignorance of the true nature of the disease, colonoscopy will be performed in many infants. Endoscopy demonstrates patchy focal or diffuse mucosal oedema and erythema of various distribution and severity, mucosal friability, haemorrhagic suffusion, and small erosions. The sigmoid colon and rectum are predominantly involved, whereas extension of the disease into the descending colon is less common. The colon proximal to the splenic flexure is always excluded. The same limitation

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**Learning points**

- A useable management algorithm should provide the practitioner with a better appraisal of unexplained rectal bleeding in the neonatal period and early infancy.
- Dietary protein induced eosinophilic proctocolitis in apparently exclusively breast fed infants has been less emphasised.
- In the case of strong evidence for allergic colitis we suggest deferring further invasive investigation.
- Strict exclusion of cows’ milk products is recommended and will result in resolution of haematochezia.

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**Figure 2** Algorithm in diagnosis and treatment of haematochezia and blood streaked stool in early infancy.
of the inflammation to the sigmoid and descending colon has been reported in both cows’ milk protein and soy protein induced enteritis.\textsuperscript{7,9} Up to 40% of infants additionally demonstrate lymphoid nodular hyperplasia with multiple, distinct and raised 1–4 mm nodules uniformly distributed over the colonic mucosa (“goose pimply appearance”; fig 1).\textsuperscript{1,11}

Histological examination shows the uniform feature of distinct eosinophilic and plasma cell infiltration of the epithelium, lamina propria, and submucosa.\textsuperscript{7,11} The regular structure of the mucosa is preserved, but all specimens showed a striking hyperaemia.\textsuperscript{1,7}

In the case of proved or suggested allergic proctocolitis we recommend strict exclusion of cows’ milk products from the mother’s diet. It is important to emphasise that the presence of allergic colitis caused by cows’ milk proteins should not raise concern about breast feeding. Resolution of the visible rectal bleeding takes place within 72 to 96 hours after elimination of the offending protein.\textsuperscript{6,7} Resolution has also been observed when a casein-hydrolysate based formula is substituted for cows’ milk. In singular cases, soy protein, corn, egg, and chocolate may cause a similar reaction as the sole allergen or coincidentally with cows’ milk.\textsuperscript{7}

Most infants experience a very benign course of the disease, and in some series up to 20% of infants had gradual spontaneous resolution of the bleeding without changes in the mother’s diet. This fact suggests that the condition might be self limiting.\textsuperscript{6,7,10} Up until now no long term risk of development of an allergic disease or chronic intestinal inflammatory disease has been reported and all children tolerated introduction of the offending protein by 1 year of age.\textsuperscript{7}

The condition appears to be common enough that practitioners should take it into account as a distinct clinical syndrome in apparently well neonates and infants with bleeding through the rectum. In the case of strong evidence for allergic colitis we suggest deferring further invasive investigation. Further evaluation is recommended only if use of the algorithm of diagnosis and treatment (fig 2) does not result in resolution of rectal bleeding.
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