Isolated ileal perforation in infancy: a lethal initial presentation of Hirschsprung’s disease

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Abstract

A rare case of ileal perforation, as a fatal initial presentation of total colonic aganglionosis (TCA) in infancy is reported. A 10-week-old boy, was brought to the emergency department with symptoms of complicated intestinal obstruction. He looked ill, was lethargic, markedly dehydrated and had a severely distended abdomen. An abdominal X-ray revealed multiple air fluid levels seen in a distended small intestine. During exploratory laparotomy the ileum was massively dilated with distal segment perforation. Ileal perforation repair was performed. A totally collapsed microcolon was identified. Biopsies were taken from the high rectum, sigmoid and hepatic flexure. Appendectomy and ileostomy were performed. All biopsies, as well as the appendix, showed absence of ganglion cells. Despite this procedure the patient progressively deteriorated and later died due to sepsis. Ileal perforation in infants is a rare, but potentially fatal initial presentation of TCA. Early detection is essential to prevent life-threatening complications.

Introduction

Hirschsprung’s disease (HD) is a relatively common developmental disorder characterized by congenital aganglionosis of Auerbach’s myenteric and Meissner’s submucosal neural plexuses of the distal colon, of varying length, resulting in a functional obstruction. The overall incidence of sporadic HD is 1 in 5000 live births,1 with male-to-female ratio of 3:1 to 4:1,2,3 but approaching 1:1 when the whole colon is involved. Total colonic aganglionosis (TCA) is a severe and uncommon form of HD. It occurs in approximately 2-13% of all patients with the disease and involves the entire colon, but may extend proximally into varying lengths of small bowel. Late diagnosis of TCA often occurs, due to milder symptoms; this delay however increases the risk of severe complications, mainly perforation, enterocolitis, sepsis and death, reaching 20 to 50%.1-4

Total colonic HD with an isolated ileal perforation in the neonatal period, as the primary presentation, is very rare, particularly in infants.5,6

We described a case of TCA in a 10 weeks old boy, previously healthy, presenting with abdominal distension due to an intestinal obstruction, complicated by ileal perforation, enterocolitis, sepsis and death. A literature review of this form of HD is also discussed.

Case Report

A 10 weeks old male infant, with no relevant medical history, was brought to the emergency department with severe abdominal distention and lethargy.

Three days prior to presentation, the mother noted fever, poor feeding and vomiting which was mainly post-prandial, non-projectile, non-bilious and non-bloodly. The infant’s condition deteriorated one day prior to presentation, with cessation of bowel movements and subsequent progressive abdominal distention and irritability at first, then lethargy.

The baby was born at term by vaginal delivery, with a weight of 1.860 grams (small for gestational age). No delay in passing meconium at birth nor history of constipation were reported. He was being breast fed and his vaccination schedule was up to date. No family history of Hirschsprung’s disease or congenital disorders was present.

On physical examination, the baby measured 30 cm long (<3 percentile for age and sex) and weighed 3430 grams (<3 percentile for age and sex). He was afebrile (37°C), fair heart rate (124 beats/minute), tachypneic (45 breaths/ minute), hypotension (60-35 mmHg), and maintained an oxygen saturation on room air of 97%. He looked unwell, lethargic (low activity, weak crying, poor sucking), markedly dehydrated (decreased skin turgor, pale mottled skin), and had a markedly distended abdomen which was tense and tender to palpation with reduced bowel sounds (agitation and crying).

Laboratory investigations revealed a high leukocyte count of 20.56×10^9/L neutrophilia (41%), thrombocytosis (994×10^9/L), elevated C-reactive protein (25.5 U/L), electrolyte disturbance (hypokalemia 129 mEq/L), hyperglycemia (196 mg/dL), elevated liver function tests (SGOT 164 U/L and SGPT 232 U/L) and compensated metabolic acidosis on arterial blood gas test (PH 7.324, PaCO2 34.2 and HCO3 -17.2). A plain abdominal X-ray revealed multiple air fluid levels seen in distended loops of small intestine; no pneumoperitoneum was noted (Figure 1).

The baby was treated for clinical sepsis, which involved immediate fluid resuscitation and initiation of broad spectrum parenteral antibiotics. An initial diagnosis of complicated intestinal obstruction was made and urgent laparotomy was performed. During surgery, the ileum was massively dilated with identification of a distal segment perforation (10 inches from ileocelecal valve). Primary repair of the ileal perforation, enterocolitis, sepsis and death.
Case Report

The initial presentation of HD is often as a functional obstruction at or shortly after birth (failure to pass meconium within 24-48h of birth, abdominal distension, bilious emesis, and enterocolitis). Paradoxically, the perforation was situated in aganglionic bowel. The mechanism of perforation in HD appears to be directly related to increased intraluminal pressure from distal obstruction, rather than any inflammatory process, subsequent bacterial overgrowth or intestinal wall ischemia.1,24

In this case enterocolitis was associated with perforation in TCA. This association is infrequently reported in infants less than 1 year old,7-26 and in the majority of cases, absence of enterocolitis was inferred from the case descriptions.

Theoretically, every HD patient is at risk for developing associated enterocolitis. Studies have shown that several factors contribute to this risk, such as family history, long segment disease and previous episodes of enterocolitis.1-6,27 Our little patient had no personal or familial anamnestic data of enterocolitis, but pathology results confirmed the long segment character of the disease. Additionally, as in our report, untreated HD patients are at high risk of associated enterocolitis. Pathologically, enterocolitis is defined as an inflammation of the lining of either the colon or the small intestine. As the disease progresses, the lining erodes, and the inside of the intestine becomes filled with pus.27 If the process is left to progress untreated, patients become very ill and perforation of the intestine may occur. Enterocolitis is

Discussion

A case of ileal perforation associated with enterocolitis, as a fatal initial presentation of TCA in an infant older than 2 months is reported.

Hirschsprung’s disease is a relatively common developmental disorder in neonates characterized by aganglionosis in the distal colon, resulting in a functional obstruction. Clinical classification of HD has evolved with further understanding of the disease, from ultra-short, short and long segment HD, to colonic and TCA with or without involvement of the small bowel.1,4

Total colonic aganglionosis represents a relatively uncommon form of HD and is defined as aganglionosis extending from the anus to at least the ileocecal valve but involving no more than 50 cm of small bowel proximal to the ileocecal valve. Even thought, in this case, there was no related family history of HD, nor any and associated anomalies found, it had been reported that the familial incidence of TCA is higher (12.4% to 33%) than that in patients with classical HD. A number of developmental conditions, such as chromosomal and congenital hypoventilation syndrome as well as ileal atresia and tumors of neural origin have been also associated with TCA.1-6

Compared to the classic cases of HD, TCA presents more diagnostic problems and is usually associated with higher morbidity and mortality rates.1-3

The initial presentation of HD is often

Figure 1. Multiple air fluid levels seen in distended small intestine, but no pneumoperitoneum.
usually accompanied by abdominal distension in 99% of cases, explosive diarrhea in 82%, vomiting in 61%, fever in 40%, lethargy in 32%, rectal bleeding in 6%, shock in 6%. Our baby presented with acute onset abdominal distension and lethargy. Fever and vomiting were reported by the mother several days earlier, responsive to anti-pyretics which were administered rectally, but no actual documentation of fever exists before and during hospitalization. Enterocolitis in patients with HD may be considered a risk factor for the fulminant, rapidly progressive and lethal form, with subsequent death, especially in the context of a late presentation.1-4, 8,10

Total colonic aganglionosis presents other diagnostic problems. X-ray studies may show enlarged loops of bowel, and a contrast enema may show a question mark-shaped colon owing to the rounded edges of the large intestines. However, there are no specific pathological findings on barium enema and X-ray studies are diagnostic in only 20% to 30% of all patients with TCA.3,4 The diagnosis of TCA must be entertained if clinical symptoms of intestinal obstruction persist in the absence of any other known causes despite a radiologically normal-looking colon.3 The definitive diagnosis of TCA is obtained by performing laparotomy and intraoperative seromuscular biopsies of the rectum, colon, and ileum. In our case, during the laparotomy, the transitional zone was very well identified at the ileocecal valve; the entire colon was collapsed and the distal ileum was very dilated and perforated around 10 inches proximal to the valve. This strongly suggested a diagnosis of TCA, which was confirmed by the biopsy results; because of the absence of an ileal biopsy, the ileal involvement remains unknown.

Ultimate treatment for TCA is surgical. Despite recent advances in surgical techniques and meticulous pre- and post-operative management, surgical treatment for TCA remains challenging. The affected patients often require multiple procedures; long-term results are suboptimal and complications are common.1-4 In this report, definitive surgery was not performed as the condition of the baby was critical and his small bowel severely inflamed. A protective ileostomy only was created with definitive surgery planned once there had been improvement of the general status of the infant.

### Table 1. Literature survey of patients with Hirschsprung’s disease presenting with ileal or other location of perforation as initial presentation

| Reference | N. of patients | Age       | Location of perforation | Extent of aganglionic segment |
|------------|----------------|-----------|-------------------------|------------------------------|
| **Ileal perforation** | | | | |
| Bowden et al.7 | 1 | 18 days | Terminal ileum | Total colonic aganglionosis |
| Keefer8 | 1 | 13 days | Terminal ileum | Total colonic aganglionosis |
| Patel et al.5 | 1 | 2 days | Terminal ileum | Total colonic aganglionosis |
| Swenson et al.3 | 1 | Newborn | Ileum | Total colonic aganglionosis |
| This report | 1 | 2 months and 2 weeks | Terminal ileum | Total colonic aganglionosis |
| **Other location of perforation** | | | | |
| Arliss et al.10 | 1 | 7 days | Appendix | Rectosigmoid |
| Asch et al.11 | 2 | 8 weeks | Appendix | Total colonic aganglionosis |
| Bowden et al.7 | 1 | 5 days | Cecum | Total colonic aganglionosis |
| Donald et al.12 | 2 | 2 days; 4 days | Cecum | Total colonic aganglionosis |
| Forshall et al.13 | 1 | 4 days | Cecum | Sigmoid; transverse colon |
| Frech14 | 4 | 1 month | Colon | Total colonic aganglionosis |
| Gastrin et al.15 | 1 | 11 days | Appendix | Disatll half colon |
| Grobl16 | 3 | 1-2 days | Cecum (1); splenic flexure (2) | Distal (2); not recorded (1) |
| Hiatt17 | 2 | Newborns | Sigmoid colon | Rectosigmoid |
| Leonidas et al.18 | 1 | 2 days | Cecal perforation with entero Colitis | Rectosigmoid |
| Martin et al.19 | 3 | 1 day; 15 days; 33 days | Appendix | Transverse colon; splenic flexure (2) |
| Newman et al.6 | 3 | 1 day; 2 days; 4 months | Cecum (2); appendix | Distal (sigmoid); total colonic aganglionosis (2) |
| Ojyäi et al.20 | 2 | 1 day | Cecum | Distal |
| Sane et al.31 | 2 | 1 month | Proximal colon (2) | Total colonic aganglionosis |
| Sarioglu et al.22 | 2 | 40 days; 27 days | Appendix | Total colonic aganglionosis; descending colon |
| Sing et al.23 | 14 | Newborns | Transverse-sigmoid colon or splenic flexure (14, but not individually defined) | Not defined |
| Soltero-Harrington et al.34 | 1 | 6 weeks | Ascending colon | Total colonic aganglionosis |
| Soper et al.25 | 1 | 1 day | Cecum | Sigmoid |
| Swenson et al.3 | 15 | Newborns | Colon (14); appendix (1) | Descending colon (2); transverse colon (4); cecum (1); sigmoid (5); terminal ileum (2); total colonic aganglionosis (1) |
| Wylie16 | 1 | 7 days | Appendix | Not recorded |
Conclusions

Ileal perforation in infants is a rare but potentially lethal initial presentation of TCA. TCA must be a major differential consideration in all neonates or young infants with persistent symptoms referable to the gastrointestinal tract even when no delay in passing meconium at birth was reported. It is challenging, but imperative to establish early diagnosis and proper management in order to prevent unexpected progression of the disease to life threatening complications.

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