Case Report

Fulminant Clostridium difficile Enteritis after Proctocolectomy and Ileal Pouch-Anal Anastomosis

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Clostridium difficile (C. difficile) infection of the small bowel is very rare. The disease course is more severe than that of C. difficile colitis, and the mortality is high. We present a case of C. difficile enteritis in a patient with ileal pouch-anal anastomosis (IPAA), and review previous case reports in order to better characterize this unusual condition.

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1. INTRODUCTION

C. difficile infection is associated with antibiotic-induced pseudomembranous colitis. This infection is usually thought to be restricted to the colon. Isolated small bowel C. difficile enteritis is rare and can manifest in the absence of a colon.

C. difficile has been shown to colonize small bowel mucosa in about 3% of the population, which then serves as a reservoir for infection [1]. Most carriers are asymptomatic. Altered intestinal anatomy and antibiotic use have been implicated in triggering symptomatic infection. Fecal flora in the small bowel of patients who had undergone a colectomy is altered to resemble that of the colon. Morphological changes (colonic-type metaplasia with partial villous atrophy) which occur in the mucosa of an ileal pouch secondary to altered fecal flow may predispose to infection [2]. These factors may increase small bowel colonization by C. difficile. Alteration of fecal flora by antibiotic can trigger symptomatic infection.

The clinical presentation of C. difficile colitis is typically mild, occasionally progressing to fulminant colitis. The disease course is more fulminant when small bowel is affected, with reported mortality ranging from 60–83% [3]. We report a case of fulminant C. difficile enteritis in a patient with ileal pouch-anal anastomosis (IPAA), and review previous reports of this unusual condition.

2. REPORT OF A CASE

A 42-year-old man underwent proctocolectomy with IPAA and ileostomy for medically refractory ulcerative colitis (UC). The patient returned for ileostomy takedown six months later. His hospital course was complicated by a urinary tract infection, which was treated with ciprofloxacin. The patient was discharged tolerating a regular diet with good bowel function. The patient returned 10 days later complaining of a three-day history of nausea, diarrhea, and abdominal pain. Patient was febrile (38.3°C), tachycardic (138), with elevated white blood cell count (17,000), creatinine (2.5), and platelet count (1450). CT scan of the abdomen showed dilated small bowel with fluid and air to the ileoanal anastomosis. Blood, urine, and stool cultures were sent, and empiric intravenous piperacillin/tazobactam and vancomycin were started. However, the patient became progressively more septic and required vasopressors for blood pressure support. Unexpectedly, C. difficile enzyme immunoassay (EIA) came back positive for toxins A and B, and the patient started on oral vancomycin and metronidazole. Flexible endoscopy was performed and revealed copious amounts of mucus with adherent pseudomembranes throughout the pouch and distal small bowel (Figure 1). Over the next few days, the patient remained in critical condition, but then slowly stabilized. Vasopressors were weaned; WBC and creatinine came down to normal
limits. Within 7 days of admission, patient was restarted on a diet and was ultimately discharged after a 12-day hospitalization. At one year follow up, the patient still occasionally has frequent bowel movements, but his stool cultures have remained negative for *C. difficile* toxins.

### 3. DISCUSSION

PubMed literature search for *C. difficile* enteritis was performed and revealed 26 cases from 1980–2008 (Table 1) [3–22]. There was significant age variability, with a range of 18 to 83 years of age (mean 50.3). Sixteen of the 26 patients had inflammatory bowel disease (IBD), thirteen patients had ulcerative colitis, and three had Crohn’s disease. Ten patients had total colectomies and six underwent IPAA. All but three patients had altered intestinal anatomy. Twenty four patients had recent hospitalization and/or operation as well as recent antibiotic use. Thirty patients were septic and required ICU admission. In all 26 cases, the stool assays were positive for *C. difficile* toxin. Diagnosis of small bowel involvement was made based on biopsy, pathology, or autopsy results. Only seven patients were evaluated endoscopically. Four underwent flexible sigmoidoscopy, and three of those had pseudomembranes. Of the patients with IPAA, only two were examined endoscopically and no pseudomembranes were visualized. One patient had an esophagogastroduodenoscopy (EGD) which demonstrated pseudomembranes in the duodenum. Treatment in all but two patients included metronidazole or vancomycin, or a combination of both. Two patients were resistant to metronidazole. Fourteen of the 26 underwent operative intervention. Mortality rate was 35%.

Our patient was similar to the previously reported cases of *C. difficile* enteritis in that he had a history of IBD, recent surgery, and antibiotic use. He required ICU admission secondary to sepsis, but he did not require operative intervention. Unlike any of the previously reported cases, our patient’s pouch endoscopy revealed pseudomembranes, facilitating timely intervention and his ultimate recovery.

*C. difficile* enteritis appears to have a fulminant course, with high risk of sepsis, need for operation, and mortality. It is unclear why the disease course is more severe than in colitis. Increased small bowel permeability is one potential explanation. Delay in diagnosis and treatment may play a role as well.

The clinical presentation can be similar for both enteritis and colitis. Symptoms include diarrhea, dehydration, and increased ileostomy output. Unlike colitis, enteritis more commonly presents with systemic manifestations such as fever, hypotension, leukocytosis and thrombocytosis [3], and occasionally with peritonitis or bowel perforation [7, 17].

*C. difficile* enteritis may be difficult to differentiate from other inflammatory processes, and requires high degree of suspicion to make the diagnosis. *C. difficile* has also been implicated as a cause of chronic pouchitis in patients with IPAA [16], and should be suspected in this setting. Given
Table 1: Previously reported cases of small bowel *C. difficile*.

<table>
<thead>
<tr>
<th>Author</th>
<th>Age</th>
<th>IBD</th>
<th>Intestinal operation</th>
<th>Recent hospitalization/operation</th>
<th>Recent Abx</th>
<th>ICU/Sepsis</th>
<th>OR</th>
<th>Endoscopy</th>
<th>Treatment</th>
<th>Death</th>
<th>Notes</th>
</tr>
</thead>
<tbody>
<tr>
<td>1 LaMont and Trnka [4] 1980</td>
<td>23</td>
<td>Crohn's</td>
<td>Partial colectomy</td>
<td>No</td>
<td>No</td>
<td>No</td>
<td>No</td>
<td>EGD— pseudomembranes in duodenum</td>
<td>Vancomycin</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>2 Shortland et al. [5] 1983</td>
<td>70</td>
<td>No</td>
<td>Ileal conduit</td>
<td>Yes</td>
<td>—</td>
<td>No</td>
<td></td>
<td>Sigmodioscopy— pseudomembranes</td>
<td>Vancomycin</td>
<td>Yes</td>
<td></td>
</tr>
<tr>
<td>3 Testore et al. [6] 1984</td>
<td>69</td>
<td>No</td>
<td>APR</td>
<td>Yes</td>
<td>Yes</td>
<td>No</td>
<td>No</td>
<td></td>
<td>—</td>
<td>Yes</td>
<td></td>
</tr>
<tr>
<td>4 Miller et al. [7] 1989</td>
<td>18</td>
<td>No</td>
<td>—</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Flexible sigmoidoscopy— inflammation; no pseudomembranes</td>
<td>Streptomycin</td>
<td>No</td>
<td>2 jejunal perforations</td>
</tr>
<tr>
<td>5 Kuntz et al. [8] 1993</td>
<td>53</td>
<td>UC</td>
<td>TAC</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>—</td>
<td></td>
<td>Vancomycin, flagyl</td>
<td>Yes</td>
<td>Intramural gas on CT</td>
</tr>
<tr>
<td>6 Tsutaoka et al. [9] 1994</td>
<td>66</td>
<td>No</td>
<td>Rt hemicolectomy</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td></td>
<td>Vancomycin, flagyl</td>
<td>Yes</td>
<td></td>
</tr>
<tr>
<td>7 Yee et al. [10] 1996</td>
<td>71</td>
<td>No</td>
<td>TAC</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>—</td>
<td></td>
<td>Flagyl</td>
<td>Yes</td>
<td></td>
</tr>
<tr>
<td>8 Kravovich et al. [11] 1997</td>
<td>65</td>
<td>No</td>
<td>Jejunileal bypass</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Flexible sigmoidoscopy— pseudomembranes</td>
<td>Vancomycin, flagyl</td>
<td>Yes</td>
<td></td>
</tr>
<tr>
<td>9 Vesoulis et al. [12] 2000</td>
<td>56</td>
<td>Crohn's</td>
<td>TPC</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td></td>
<td>Flagyl</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>10 Freiler et al. [13] 2001</td>
<td>26</td>
<td>UC</td>
<td>TAC</td>
<td>Yes</td>
<td>Yes</td>
<td>No</td>
<td>No</td>
<td></td>
<td>Flagyl</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>11 Jacobs et al. [14] 2001</td>
<td>83</td>
<td>No</td>
<td>—</td>
<td>Yes</td>
<td>—</td>
<td>Yes</td>
<td>Yes</td>
<td></td>
<td>—</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>12 Tjandra et al. [15] 2001</td>
<td>60</td>
<td>No</td>
<td>Sigmoid colectomy</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Flexible sigmoidoscopy— pseudomembranes</td>
<td>Vancomycin, flagyl</td>
<td>Yes</td>
<td></td>
</tr>
<tr>
<td>13 Mann et al. [16] 2003</td>
<td>35</td>
<td>UC</td>
<td>IPAA</td>
<td>No</td>
<td>No</td>
<td>No</td>
<td>No</td>
<td></td>
<td>Vancomycin; resistant to flagyl</td>
<td>No</td>
<td>Chronic pouchitis</td>
</tr>
<tr>
<td>14 Hayetian et al. [17] 2006</td>
<td>80</td>
<td>No</td>
<td>LAR</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td></td>
<td>Flagyl</td>
<td>Yes</td>
<td>Ileal perforation</td>
</tr>
<tr>
<td>15 Hayetian et al. [17] 2006</td>
<td>83</td>
<td>No</td>
<td>None</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td></td>
<td>Vancomycin, flagyl</td>
<td>No</td>
<td>Ileal perforation</td>
</tr>
<tr>
<td>16 Kim et al. [18] 2007</td>
<td>65</td>
<td>Crohn's</td>
<td>TPC</td>
<td>Yes</td>
<td>Yes</td>
<td>No</td>
<td>No</td>
<td></td>
<td>Flagyl</td>
<td>Yes</td>
<td></td>
</tr>
<tr>
<td>17 Lundeen et al. [3](6 patients) 2007</td>
<td>35.3</td>
<td>UC (6 patients)</td>
<td>3 IPAA 3 TAC</td>
<td>6/6</td>
<td>1/6</td>
<td>1/6</td>
<td>—</td>
<td></td>
<td>Vancomycin, flagyl</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>18 Wood et al. [19] 2008</td>
<td>48</td>
<td>UC</td>
<td>IPAA</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Yes</td>
<td>Flexible endoscopy—normal pouch</td>
<td>Flagyl</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>19 Follmar et al. [20] 2008</td>
<td>49</td>
<td>UC</td>
<td>IPAA</td>
<td>Yes</td>
<td>No</td>
<td>Yes</td>
<td>—</td>
<td></td>
<td>Vancomycin, resistant to flagyl</td>
<td>No</td>
<td>Or-mesh removal</td>
</tr>
<tr>
<td>20 Fleming et al. [21] 2008</td>
<td>54</td>
<td>UC</td>
<td>TAC</td>
<td>Yes</td>
<td>?</td>
<td>No</td>
<td>—</td>
<td></td>
<td>Flagyl, vancomycin, rifampin</td>
<td>No</td>
<td></td>
</tr>
<tr>
<td>21 Yafi et al. [22] 2008</td>
<td>21</td>
<td>UC</td>
<td>TAC</td>
<td>Yes</td>
<td>?</td>
<td>Yes</td>
<td>—</td>
<td></td>
<td>Vancomycin</td>
<td>No</td>
<td>Pelvic abscess</td>
</tr>
<tr>
<td>Total</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td>16/26</td>
<td>23/26</td>
<td>24/26</td>
<td>24/26</td>
<td></td>
<td>13/26</td>
<td>7/26</td>
</tr>
</tbody>
</table>
the higher risk that IBD patients may have for developing *C. difficile* enteritis, it is important to be able to differentiate it from an exacerbation of IBD. Diagnosis is made by identifying *C. difficile* toxin A or B in the stool. Similarly, endoscopy should be utilized in patients with suspected small bowel involvement even with history of prior colectomy. This may facilitate differentiation between Crohn’s enteritis, pouchitis, and *C. difficile* enteritis.

As with our patient, most cases will respond to treatment with metronidazole or vancomycin. However, more virulent and resistant strains have been reported [23]. Some patients will need emergent surgical resection of any perforated or gangrenous bowel if they fail to respond to medical treatment.

*C. difficile* enteritis is emerging with increased frequency and can have devastating results. Patients with IBD and prior colectomy are at increased risk. Prompt identification of the organism via stool culture and endoscopy may result in more favorable outcomes.

**NOMENCLATURE**

APR: Abdominoperineal resection  
TAC: Total abdominal colectomy  
TPC: Total proctocolectomy  
IPAA: Ileal pouch-anal anastomosis

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**REFERENCES**


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