Review

# Obstruction and pseudo-obstruction in inflammatory bowel disease

K.H. Katsanos, V.E. Tsianos, M. Maliouki, M. Adamidi, I. Vagias, E.V. Tsianos

#### **SUMMARY**

Intestinal obstruction and pseudo-obstruction represents a severe complication and a potential emergency in Crohn's disease and ulcerative colitis. Such intestinal complications are of great importance and may require urgent surgical management. This review discusses the clinical importance, the risk factors and the pathophysiological basis of this phenomenon. Diagnosis and imaging of obstruction together with common and rare causes of obstruction and pseudo-obstruction including toxic megacolon are extensively reviewed. Non-surgical treatment of obstruction in inflammatory bowel disease is reviewed including pharmacologic colonic decompression and infliximab use. Endoscopy in pseudo-obstruction, toxic megacolon, and in stenoses and strictures is reviewed and the setting of the optimal time for surgery in obstructing cases is provided. Strictureplasty or resection in stenoses, post-surgical obstructions as well as available prognostic markers of obstruction and pseudo-obstruction are critically revised.

**Key words:** Obstruction, Crohn's disease, pseudo-obstruction, ulcerative colitis, inflammatory bowel disease

#### 1. Clinical importance of obstruction in IBD

Intestinal obstruction represents a severe complication and a potential emergency in inflammatory bowel disease. In particular, intestinal fibrostenosis in Crohn's disease

1st Department of Internal Medicine & Hepato-Gastroenterology Unit University Hospital of Ioannina, Greece

# Author for correspondence:

Epameinondas V. Tsianos, MD, PhD AGAF, Professor of Internal Medicine, 1st Department of Internal Medicine, Medical School, University of Ioannina, Leoforos Panepistimiou, 451 10 Ioannina, Greece, Tel: 0030-26510-07501,

Fax: 00-30-26510-07016, e-mail: etsianos@cc.uoi.gr

(CD) is a frequent and debilitating complication not only resulting in small bowel obstruction, but eventually in repeated bowel resection and short bowel syndrome.<sup>1</sup>

The prevalence of small bowel stenosis has been reported to range from 20% to 40% and from 7% to 15% in patients with colonic Crohn's disease.<sup>2</sup> By contrast, in patients with ulcerative colitis (UC) intestinal stenosis and obstruction is rather infrequent.<sup>3</sup>

Intestinal pseudo-obstruction usually affects the colon but the small intestine may also be involved, and may present in acute, subacute or chronic form. Acute colonic pseudo-obstruction or Ogilvie's syndrome is the clinical syndrome of acute large bowel dilatation without mechanical obstruction that is an important cause of morbidity and mortality and, therefore, requires urgent evaluation. Physical and radiological findings are identical to those associated with mechanical obstruction of the large bowel. 11-12

Of note, cases of intestinal pseudo-obstruction, <sup>13</sup> or obstruction, <sup>14</sup> proved to be the presenting symptom of Crohn's disease, or a chronic complication in patients with extensive bowel resection. <sup>15</sup> Furthermore, a coexistence of chronic idiopathic intestinal pseudo obstruction and inflammatory bowel disease (IBD) has also been reported. <sup>16</sup>

Particularly in children, such intestinal complications are of great importance and may require urgent surgical management.<sup>17</sup> The presence of intestinal obstruction in children with IBD should be always considered during modification of therapeutic strategies. It is very important to stress that although the Crohn's disease process cannot be eradicated by surgery, the health of a child can be restored to normal for prolonged periods of time.<sup>18</sup>

# 2. Risk factors and pathophysiological basis of obstruction in IBD

Bowel obstruction or pseudo-obstruction in IBD can be the result of mechanical causes or motility disturbanc-

es as it also occurs in non-IBD patients.<sup>19</sup> The syndrome of acute colonic pseudo-obstruction is well delineated but its aetiology remains poorly understood.<sup>20-21</sup>

Small bowel obstruction in Crohn's disease is most frequently due to to strictures, postoperative or inflammatory adhesions, or intestinal neoplasms. Intermittent small bowel obstruction from Crohn's disease may be secondary due to strictures or to chronic adhesive peritonitis. <sup>22-23</sup>

Ileus and colonic pseudo-obstruction are induced and/ or perpetuated by numerous metabolic disorders, drugs that inhibit intestinal motility, severe co-morbidity, and surgery. For example, patients taking anti-psychotic, anti-depressant and anti-parkinsonism drugs seem to be at a higher risk. Exceptional cases of toxic megacolon caused by loperamide administered for Crohn's disease and ileus caused by ferrum tablets with a non-soluble matrix, in the presence of an existing stenosis of the ileum due to Crohn's disease have also been reported. One case of toxic megacolon after sigmoidoscopy for UC has also been reported.

Rarely, the not-congenital (Hirschsprung's disease) chronic megacolon may represent the end-stage of any form of refractory constipation or anal canal stenosis (slow transit constipation or pelvic floor dysfunction) in IBD patients. Chronic intestinal pseudo-obstruction secondary to primary visceral myopathy has never been reported in IBD.

The molecular mechanisms involved in this process of fibrogenesis and subsequent stenosis in the gut have not been identified. However, inflammatory mediators may have an effect on mesenchymal cells in the submucosa and the muscle layers. Transforming growth factor-beta (TGF-beta), has profound anti-inflammatory activity in the mucosa but at the same time it appears to be driving the process of fibrosis in the deeper layers of the gut.<sup>1</sup>

Fibrosis-associated gene expression in the intestine of patients with Crohn's disease and correlation of expression levels with prior medical therapies has been demonstrated,<sup>29</sup> and a positive association was also found between the carriers of at least one CARD15 mutation and the acute intestinal obstruction at diagnosis in Crohn's disease patients.<sup>30</sup>

In transgenic mice, hepatocyte growth factor/scatter factor (HGF/SF) plays an important role in the development of gastrointestinal paresis and chronic intestinal inflammation. Authors suggested that HGF/SF may represent a useful model for the study of molecular mechanisms associated with a subset of IBD and intestinal pseudo-obstruction.<sup>31</sup>

Various pathologic changes in Crohn's disease could be involved in the formation of strictures including obliterative muscularization of the submucosa that may be pathogenetically involved in the formation of strictures either directly by causing a sustained spasm, or indirectly by minimizing the vasoprotective role of the submucosa, impairing repair and enhancing scarring.<sup>32</sup> In addition, histologic alterations of intramural nervous tissue may play a pivotal role<sup>33</sup> and according to a study acute colonic pseudo-obstruction is the result of excessive parasympathetic suppression rather than sympathetic overactivity.<sup>34</sup>

# 3. Diagnosis and imaging of obstruction in IBD

In acute intestinal obstruction, the clinician must characterize the level of emergency of the case and distinguish between acute small bowel and acute colonic obstruction as in non-IBD patients.<sup>35</sup>

Differential diagnosis must be made between, obstructive colitis which is an ulcero-inflammatory and necrotizing condition that occurs in the colon proximal to benign or malignant stenosing lesions and is the result of ischemia or other factors which impair adequate perfusion. The incidence of obstructive colitis among patients with colonic obstruction has been reported to range between 1 and 7%.36 Differential diagnosis must also be made between other inflammatory conditions of the colon mimicking acute obstructive abdomen including epiploic appendagitis, diverticulitis, pseudomembranous colitis, 37 and eosinophilic enteritis.38 In particular cases, other causes of toxic megacolon such as Clostridium difficile and cytomegalovirus infection as well as rare causes of large bowel obstruction such as pelvic actinomycosis also have to be excluded.39

Gastrointestinal contrast studies play an important diagnostic role in partial obstruction. 40 Barium examinations of the large and small bowel seem to be of low value. In the majority of cases computed tomography (CT) is the method of preference for diagnosis and follow up. 42-45 CT is sensitive for diagnosing complete obstruction of the small bowel and for determining the location and cause of obstruction. In addition, in many instances there is a possibility of differential diagnosis between inflammatory and fibrostenotic obstruction based on the presence of edema, thickness of the bowel wall and vascularisation. Although many CT findings in patients with colitis are nonspecific, some features are helpful in suggesting a specific diagnosis46 and radiologists can frequently diagnose a CD-related obstruction in a patient with known CD and differentiate it from an obstruction due to adhesions. 47 A 'beak-like'

luminal narrowing seems the most common CT finding at the obstructed site.<sup>48</sup> CT enterography is a useful technique for bowel imaging and is complementary and often superior to conventional barium studies.<sup>49-50</sup>

Computed tomographic colonography ('virtual colonoscopy') has shown promising results<sup>51-53</sup> and CT colonography has a reasonable sensitivity and specificity for detecting large polyps.<sup>54</sup>

Magnetic enteroclysis is a very valuable method for diagnosing bowel luminal and extraluminal abnormalities. However, it is performed only in specialized centers, with specialized radiologists and usually is not available on an emergency basis.

Digestive endoscopy is a mandatory method in the diagnosis of intestinal obstructions. Diagnostic endoscopy in the involvement of the large intestine or enteroscopy in the case of incomplete obstruction of the small intestine is the method indicated in the majority of obstructive intestinal lesions.<sup>55</sup> Fulminant colitis and toxic megacolon represent usual exceptions.

The risks of endoscopy during acute phases include perforation or obstruction worsening. Although the authors are not familiar with CT-guided endoscopy we presume that excessive radiation for both patient and endoscopist is added to the risks of endoscopy performed on emergency. In addition, this method is not always technically feasible.

In study populations, capsule endoscopy is superior to all other modalities for diagnosing small bowel Crohn's disease<sup>56</sup> and for revealing stenoses, which may not be depicted with radiology. The potential for complications when wireless capsule enteroscopy is performed in the presence of intestinal strictures is evident.<sup>57</sup> Retention of the capsule appears to be infrequent. In most cases, capsule retention is asymptomatic and capsule surgical removal is safe and also identifies and treats the underlying small-bowel condition.<sup>58-59</sup>

# 4. Causes of obstruction in IBD

### Toxic megacolon

Toxic megacolon is a serious cause of obstruction in IBD<sup>60</sup> (Table 1) and in a review study from Mount Sinai Hospital toxic megacolon carried a 16% mortality rate.<sup>61</sup> Toxic megacolon may occur in ulcerative colitis but also in Crohn's colitis.<sup>62-64</sup> Megacolon refers to cecal dilatation above the dimension of 12 cm and above 6.5 cm of the sigmoid colon, measured at the pelvic brim. Dilatation of the cecum to greater than 12 cm diameter is a cause for grave concern.<sup>65</sup> Toxic megacolon is a radiology-proved meg-

**Table 1**. Specific causes of obstruction and pseudo-obstruction in IBD

#### Ulcerative colitis

Giant polyposis

Strictures (exclude malignancy)

Toxic megacolon

Volvulus

Cancer (adenocarcinoma)

Ileal pouch-anal anastomosis (IPAA)

Post-operative adhesions

#### Crohn's disease

Strictures

Gallstone ileus

Entreroliths

Endometriosis

Cystic fibrosis

Cancer (adenocarcinoma, lymphoma, carcinoid, GIST)

Capsule retention

Loperamide or labdanum overdosing

Acute steroid withdrawal (post-surgery)

Post-resection stenosis

Temporary 'diverting' loop ileostomy construction

Ileostomy takedown

Continent ileostomy stenosis

Post-operative adhesions

acolon which is also accompanied by signs of systemic toxicity (fever, leucocytosis, tachycardia, tachypnea) and may result from or be complicated with Clostridium difficile infection<sup>66</sup>, cytomegalovirus infection, or common acute gastroenteritis.<sup>67-69</sup>

Adult<sup>70-73</sup> or pediatric<sup>74-75</sup> cases with toxic megacolon reported in ulcerative colitis resulted either in uneventful or complicated<sup>76-78</sup> or fatal outcome.<sup>79</sup> Surgery for toxic megacolon during pregnancy has also been reported.

Surgical therapy of toxic megacolon in ulcerative colitis patients is total colectomy, which actually permits an adequate control of the disease, as well as a satisfactory therapy of the rectal stump, with subsequent recanalization in the majority of cases. In Crohn's disease the frequency of recidive is higher (post-operative recurrence) and it seems more advisable to opt for a radical resection treatment (chiefly right colectomy), but with a conservative purpose as a new operation in the future is quite probable.<sup>80</sup>

#### Acute obstructing malignancy

Adenocarcinoma of the small intestine is a rare complication of Crohn's disease. Diagnosis of early stages of adenocarcinoma of the small intestine is very difficult and thus might be impossible to differentiate from exacerbation or progressive stenosis of preexisting Crohn's disease.

Obstructing adenocarcinoma may be the presenting symptom of small bowel Crohn's disease. 81-84 Carcinoid tumor should be suspected in elderly patients with Crohn's disease presenting with intestinal obstruction and laparotomy should be considered to exclude malignancy. 85 A case of obstructing gastrointestinal stromal tumor in Crohn's has been also reported. In ulcerative colitis obstruction occurs only in advanced cancer cases. 86

#### Gallstone ileus

Gallstone ileus is an uncommon cause of small bowel obstruction, accounting for fewer than 3% of laparotomies for intestinal obstruction. Patients with long-standing Crohn's disease have an increased risk of developing gallstone disease.

So far seven cases with Crohn's disease have presented with gallstone ileus. The correct diagnosed is difficult before surgery. Therapy includes ileal resection and cholecystectomy.<sup>87-92</sup>

# Cystic fibrosis, enteroliths, giant polyposis and endometriosis

A single case of cystic fibrosis, Crohn's colitis and adult meconium ileus equivalent has been reported. 93

Enteroliths associated with Crohn's disease is an extremely rare condition with fewer than 25 cases reported in the literature<sup>22</sup> treated by segmental ileal resection<sup>94</sup> or laparoscopically.<sup>95</sup>

Giant polyposis may present with acute colitic attacks, 96 colonic obstruction, 97-98 toxic megacolon, 99 or colocolonic intussusception. 100

Endometriosis is a rare disease in women of the reproductive age, but should be considered in the differential diagnosis of stenotic Crohn's disease of the terminal ileum.<sup>101</sup>

#### Upper GI obstruction

The presence of gastrointestinal motility abnormalities in children affected by Crohn's disease has been demonstrated by means of electrogastrography. <sup>102</sup> Clinicians should consider impaired gastric emptying when evaluating patients with Crohn's disease and severe symptoms of upper gut dysmotility, which cannot be attributed to active inflammation or organic obstruction of the digestive tract. Symptoms in these patients are refractory to various therapeutic interventions including tube feeding and gastric surgery. <sup>103</sup>

#### Volvulus

Colonic volvulus accounts for 1-7% of cases of large bowel obstruction in the United States and Western Europe and is commoner in tropical and subtropical areas. The sigmoid colon is involved in 65-80% of the cases reported, followed by right colon in 15-30% of cases while transverse colon and splenic flexure volvulus are rare. 104

Intestinal volvulus in patients with inflammatory bowel disease is extremely rare. There are six reports<sup>105-110</sup> of volvulus occurring in Crohn's disease patients; three of which were detected in the sigmoid. <sup>105-107</sup> Only one case of volvulus of the cecum has been reported in ulcerative colitis. <sup>111</sup> Double volvulus of the transverse and the sigmoid colon has been reported only once in patients with inflammatory bowel disease. <sup>112</sup>

High-risk groups among inflammatory bowel disease patients seem to include those with reactivation of the ileal Crohn's disease contributing to the development of volvulus by causing fixation, torsion and dilatation of the distal bowel. Patients diagnosed with dolichocolon should also be included in the high-risk group. Preoperative colonoscopic derotation is beneficial in acute sigmoid volvulus. Correct identification of transverse colon volvulus is necessary in order to reduce high mortality rates. While a conservative treatment of sigmoid volvulus is recommended, it is inadequate in case of transverse colon volvulus.

In the absence of clinical, laboratory or radiological signs of bowel necrosis or perforation, colonoscopic volvulus derotation is recommended in all cases, followed by semi-elective single stage colonic resection. Emergency endoscopic decompression of the sigmoid volvulus is safe and effective as an initial treatment but has a high early recurrence rate.<sup>113</sup>

# 5. Non-surgical treatment of obstruction in IBD

For small bowel obstruction there is an increasing tendency for initial conservative management rather than immediate operative intervention, as a proportion of cases will resolve spontaneously. Hearly postoperative small-bowel obstruction can be safely and effectively managed by nasogastric decompression in the majority of cases. In general, reexploration should be reserved for those patients whose symptoms do not resolve within six days of nasogastric decompression. Hearly

For large bowel obstruction emergent surgery is indicated if there are signs of impending or frank perforation. Scheduled surgery is indicated when non-operative measures fail (Table 2).<sup>117</sup>

## Pharmacologic colonic decompression

General measures include intravenous rehydration, correction of electrolyte abnormalities, discontinuation

**Table 2**. Non-surgical therapy for obstruction and pseudo-obstruction in IBD

#### Pharmacological

Prokinetics (pseudo-obstruction)
Polyethylene glycol solution (after decompression)

#### **Endoscopic**

Ballon dilatation (anastomotic or frank strictures)

Decompression rectal tube (possible steroid infusion via tube) Intrastricture injections via sclerotherapy catheter (Infliximab or triamcinolone)

Stenting

Needle knife cutting

Laser stricture cutting or recanalization

Polypectomy (giant polyps)

Volvulus derotation

#### Toxic megacolon

Infliximab

Cyclosporin?

Tacrolimus?

Endoscopic decompression?

Surgery

of antikinetic drugs, and treatment of other co-morbidities as in non-IBD patients.

In non-IBD patients specific therapies include prokinetics such as erythromycin, metoclopromide, cisapride, tegaserod, motilin receptor agonists and, diatrizoate meglumine (gastrografin) enema. 118-119 The most effective of all seems to be intravenous neostigmine (2-2.5 mg), which leads to quick decompression in a significant proportion of patients after a single infusion. 65 Administration of polyethylene glycol solution in patients with Ogilvie's syndrome after initial resolution of colonic dilation may increase the sustained response rate but this has not been yet tested in IBD patients. 120 In conservative therapy of toxic megacolon or fulminant colitis, infliximab seems superior to cyclosporin or tacrolimus as described in the next paragraph.

# Infliximab

Tumor necrosis factor, has antifibrotic bioactivity and pharmacologic inhibition of this cytokine carries a theoretical risk of enhanced stricture formation. In addition, a theoretical concern exists that rapid luminal healing in Crohn's disease with infliximab may increase the risk of stenosis, stricture or obstruction. In fact, there have been studies favouring and unfavoring Infliximab use in stenotic or stricturing disease phenotype.

On experimental level infliximab did not ameliorate the degree of fibrosis in alkali burns of the oesophagus in the rat<sup>121</sup> and early obstructive colon after treatment of active refractory Crohn's disease with infliximab has been reported. 122 According to a post-marketing survey exploring adverse event signals with infliximab a signal for bowel obstruction was identified. 123

Among studies favouring Infliximab use there have been those reporting regression of strictures and overall benefit in a subset of patients with subocclusive small bowel stricturing disease<sup>124-127</sup> It has been also suggested that in patients with symptomatic strictures from CD, infliximab should be tested before considering surgery.<sup>128</sup>

Of interest, infliximab prior to abdominal surgery for Crohn's disease is not associated with an increased rate of early postoperative complications. 129-130

Development of strictures, stenosis and obstruction was related to disease duration, severity, ileal location and corticosteroid use. <sup>131</sup> This was confirmed by subsequent studies on the effect of Infliximab on small bowel stenoses, <sup>132</sup> on fibroblasts, <sup>133</sup> and on profibrotic transcriptional response. <sup>29</sup>

Data on the use and efficacy of Infliximab before and after surgery is largely empirical and anecdotal as large controlled trials are lacking. It seems that Infliximab is safe and does not increase postoperative complications. After one week of surgery Infliximab can be administered again with safety. Therapy of toxic megacolon with infliximab has proved to be an important option with a group of UC patients improving and avoiding surgery. 134-136 In the most important pioneer study, forty-five patients with severe to moderately severe ulcerative colitis not responding to conventional treatment were included (24 infliximab and 21 placebo). No patient died. Seven patients in the infliximab group and 14 in the placebo group had a colectomy (P = 017; odds ratio, 4.9; 95% confidence interval, 1.4-17) within 3 months after randomization. No serious side effects occurred. Three patients in the placebo group required operation for septic complications.

Finally, a study combining information from the TREAT and the ACCENT I studies demonstrated that infliximab was not related to such events.

### Endoscopic treatment in pseudo-obstruction

Besides diagnostic importance, colonoscopy enables an effective therapeutic approach, which may immediately follow the diagnostic intervention as it also occurs in non-IBD patients. <sup>137-138</sup> Plain abdominal roentgenogram is a useful diagnostic test. If the cecal diameter is 12 cm or greater, or conservative management is unsuccessful, colonoscopic or operative decompression is needed. Operative colonoscopic endoscopy tends to be increasing by time in referral centers. <sup>139</sup>

In patients failing or having contraindications to prokinetics, colonoscopic decompression is the active intervention of choice. <sup>140</sup> There are no reports on enteroscopic decompression. In pseudo-obstruction, colonoscopy is indicated, <sup>78</sup> with the exception of toxic megacolon. <sup>136</sup>

# Endoscopy in toxic megacolon

We do not advocate endoscopic treatment of toxic megacolon because of the heightened risk of perforation. Of note, there are reports of toxic megacolon treated with colonoscopic decompression<sup>141</sup> and of decompression placement of an indwelling colonic tube through which steroid solution was administered.<sup>142</sup>

In non-IBD patients, average cecal diameter more than 13.3 cm is prognostic for more than one colonoscopies needed for decompression and 13.4 cm is prognostic for failing colonoscopic therapy. 143 Decompressing percutaneous endoscopic cecostomy has been described. 144 In such difficult cases, the use of colonoscopy should be extremely selective after radiology examination, and it should be performed by experts and accompanied generally by tube placement. 145-146

### Endoscopy in stenoses or strictures

Balloon dilatation of strictures with a variety of techniques ranging from divulsion with through-the-scope balloon dilators to laser recanalization<sup>147</sup> palliative treatment of colonic carcinoma by laser procedures or pre-operative endoscopic laser relief of tumor obstruction,<sup>55</sup> self-expanding stents, <sup>118,148</sup> colonoscopic decompression with tube placement, <sup>149-151</sup> reduction of colonic volvulus are the principal indications of therapeutic colonoscopy in the non-surgical treatment of the dilated colon.

Some colonic strictures in Crohn's disease can be dilated using endoscopes of graded caliber or with throughthe-scope balloons, with or without stent placement. Endoscopy is most useful in dilating anastomotic strictures and less useful for long strictures in active inflammatory disease. Strictures in ulcerative colitis are suspicious for neoplasia and, if dilated at all, should be biopsied extensively and followed closely. In obstructed continent ileostomies, endoscopy can be employed effectively to both determine the cause of the obstruction and re-establish patency.<sup>152</sup>

# Endoscopy in re-stricturing

The dilation of benign stenoses of the lower gastrointestinal tract with a balloon catheter is an effective and safe method, which in most cases results in long-term elimination of the obstructive symptoms, so that surgery can be avoided or postponed. 153-154 Repeated dilatations often are required. Treatment with intrastricture triamcinolone injection 155 or intralesional injection of infliximab via the sclerotherapy technique in Crohn's colitis² did not demonstrate remarkable efficacy. Careful patient selection is of paramount importance to ensure favorable long-term outcomes when dilating benign stenoses. 1

# 6. Setting the optimal time for surgery in obstructing IBD

Inflammatory bowel disease can present in numerous ways as an acute surgical emergency. <sup>156-158</sup> Emergency complications of IBD are rare, but may be life threatening and include toxic colitis, hemorrhage, perforation, intraabdominal masses or abscesses with sepsis, and intestinal obstruction. Acute surgical emergencies in patients with inflammatory bowel disease may carry a substantial morbidity, but fortunately today, a low mortality. <sup>159</sup> The successful management of such complications depends on early diagnosis and the judicious and timely use of both medical and surgical therapies. With a multidisciplinary approach, morbidity can be reduced and patients can have a rapid return and improved quality of life. <sup>160</sup>

The most common complications associated with UC are fulminant colitis, toxic megacolon, and bleeding.<sup>161</sup> Some cases can be managed conservatively and may recover completely without any surgical intervention.<sup>162</sup> However surgery should be performed immediately if there is no improvement within 5 days of medical management in case of acute colitis, within 24-72 hours in case of toxic megacolon, intestinal obstruction or severe bleeding, or if the patient deteriorates during this period.<sup>164-165</sup>

When urgent operation is necessary subtotal colectomy with ileostomy and preservation of the rectum is generally indicated (Table 3). 166-167 This will allow subsequent mucosal proctectomy and ileoanal anastomosis.

Subtotal colectomy with ileostomy and mucous fistula of distal sigmoid colon is the operation of choice for patients with fulminant colitis. Restorative proctocolectomy with ileal pouch-anal anastomosis (IPAA) is preferred for patients who undergo scheduled elective surgery for ulcerative colitis. 168

The most common complications associated with CD are abscesses and intestinal obstruction. Although initial treatment includes medical treatment, these complications usually require surgical intervention and conservative intestinal resection. <sup>169</sup> Resection should be minimal but terminated in a macroscopically disease-free area. When urgent colectomy is indicated for Crohn's colitis without

**Table 3**. Surgical therapy in obstruction and pseudo-obstruction in IBD

#### Ulcerative colitis

On emergency [toxic megacolon, fulminat disease] Subtotal colectomy with ileostomy

[preservation of rectum and mucous fistula of distal sigmoid for future ileo-rectal anastomosis]

Scheduled

Restorative proctocolectomy with ileal-pouch anal anastomosis [IPAA]

Permanent ileostomy

Resection preferred to strictureplasty (suspicion of neoplastic stenosis)

#### Crohn's disease

Bowel resection (minimal resections) Strictureplasty (short or long, single or multiple) Multiple bowel bypass (special cases?)

rectal involvement, ileorectal anastomosis can be considered either as a primary or secondary procedure. 166

### 7. Strictureplasty or resection?

# Strictureplasty in Crohn's colitis

In Crohn's disease surgery does not provide cure with radical resection of inflamed bowel. Therefore stricture-plasty has become a useful bowel-preserving surgical technique in the treatment of small-bowel stenosis. In a retrospective study defining the efficacy of strictureplasty and resection in patients with obstructive Crohn's disease of the colon the incidence of postoperative surgical recurrence was 36% in those treated by strictureplasty and 24% in those treated by resection(ns). The authors concluded that strictureplasty in Crohn's colitis is a valuable surgical technique which results in low recurrence rates and in surgical outcome comparable to that in resection without sacrificing functional large bowel length.<sup>170</sup>

# Strictureplasty in Crohn's small bowel disease

Strictureplasty is a good surgical option for stenosing small-bowel Crohn's disease, particularly in patients with multiple obstructions and in those vulnerable to short-bowel syndrome.<sup>171</sup>

In a retrospective study defining the efficacy of strictureplasty and resection in patients with obstructive Crohn's disease of the small bowel postoperative morbidity was 14.8% after strictureplasty and 17% after resection while 50% of the patients treated by strictureplasty and 37% treated by resection developed recurrent disease. The authors concluded that results after strictureplasty are comparable to those after resection in terms of complications, recurrence and quality of life in the treatment of

small bowel strictures in Crohn's disease. In the long run there might be an advantage for stricture plasty because it prevents complications caused by resectional therapy such as short bowel syndrome. <sup>170</sup>

Of interest one multiple small bowel bypass (latero-lateral bowel anastomoses in multiple sites to overcome strictures) case has also been reported.<sup>80</sup> Data on the efficacy of long or short strictureplasty is still conflicting.

Colorectal strictures are not uncommon in ulcerative colitis and approximately a quarter of them are malignant. According to a retrospective single centre study three principal features distinguished the malignant from the benign strictures: appearance late in the course of ulcerative colitis, location proximal to the splenic flexure and symptomatic large bowel obstruction. Moreover, cancer associated with strictures tended to be more advanced than that which does not produce strictures.<sup>172</sup>

To conclude, in short fibrous stenosis of the small bowel or ileocecal anastomosis without acute inflammation strictureplasty should be performed. In all other cases, especially in colonic strictures and patients with long term disease with their increased risk of malignancy, resection is the treatment of choice. For the same reason resection is the treatment of choice in insisting colonic stenoses of patients with ulcerative colitis.<sup>173</sup>

# 8. Post-surgical obstruction in IBD

Early postoperative small-bowel obstruction in general population was observed in 9.5 percent of cases. 115 No independent risk factors predisposing to early postoperative small-bowel obstruction were identified. The predisposing factors for postoperative ileus have not been clearly identified.115 Resident intestinal muscularis macrophages initially involved in inflammatory responses may be of importance.<sup>174</sup> Susceptibility to postoperative ileus following abdominal surgery increases with advancing age. There is both an age-dependent increase in the proinflammatory mediator expression and an age-dependent decrease in anti-inflammatory mediator expressions following minor insult to the bowel. Such imbalances between pro- and anti-inflammatory mechanisms may form the basis for increased susceptibility to ileus and for the increased severity and duration of ileus observed in the elderly. 175

Early post-surgery small intestinal obstruction is not an uncommon complication following excisional operation for Crohn's disease. <sup>176</sup> Late restricturing after ileocolonic resection for Crohn's disease is not infrequent <sup>155</sup> and the need for intestinal repeat resection for recurrence of obstructing Crohn's disease is considerable. <sup>177</sup> Loop

ileostomy construction and takedown is associated with risk of obstruction. However, no differences exist between technique used for closure or the baseline pathology of the patient.<sup>178</sup>

Small bowel obstruction is common following pouch surgery for ulcerative colitis and is the dominating cause of hospitalisation postoperatively. In such a study about 25% of patients developed small bowel obstruction and half of them needed surgery. The use of a diverting loopileostomy was related to an increased risk of surgery for small bowel obstruction.<sup>179</sup> The obstruction may occur either before or after closure of the temporary ileostomy and according to a study patients who had a temporary Brooke ileostomy were more likely to develop obstruction than those who had a loop ileostomy. Also, patients who had had previous operations were at greater risk of obstruction than those who had not.<sup>180</sup>

Acute ileus due to steroid withdrawal caused symptoms of intestinal obstruction. Since ileus from acute steroid withdrawal occurred four times as frequently as mechanical small-bowel obstruction, prompt recognition and treatment may reduce postoperative morbidity in IBD patients.<sup>181</sup>

# 9. Prognosis of obstruction in IBD

Acute colonic pseudo-obstruction remains a serious medical disorder, carrying a considerable mortality rate of 15% while assessment of predictors of response is important. 182-184

For toxic megacolon, dismal prognostic factors affecting mortality include very young age<sup>185</sup> or age over 40 years old,<sup>61</sup> female sex,<sup>186</sup> increased cecal diameter,<sup>6-9</sup> and especially the occurrence of small bowel distension,<sup>187</sup> any type of colonic perforation<sup>188</sup> and multiple organ dysfunction syndrome.<sup>189</sup>

#### Emergency of obstruction

- -Yes (acute abdomen)
- -No (chronic or 'relapsing' symptoms)

# Site of obstruction

- -upper GI
- -small bowel
- -large bowel
- -post-surgery

# Reason of obstruction

-mechanical obstruction -pseudo-obstruction

#### Time allocated for each therapy

- -pharmacological
- -endoscopic
- -surgery

-In short-term always exclude need for urgent operation

-In long-term always exclude cancer

Figure. The ESRT scheme (Emergency, Site, Reason, Time) in the management of obstruction symptoms in inflammatory bowel disease.

Prognosis of the operated obstruction is related to underlying cause and type of disease. In UC patients, operation usually results in symptom relief. In Crohn's disease, relapses may occur and the best strategy is to postpone next intervention when possible. In most cases a short-bowel syndrome is the consequence of extensive bowel resection. 190

To conclude, the most important factors influencing outcome are physician experience, careful patient follow up, the choice of the appropriate therapy, the timing for surgery and the quality of the procedures performed. We suggest the ESRT (Emergency, Site, Reason, Time) scheme [Figure] for assessment and management of obstruction and pseudo-obstruction cases. Following this scheme we can increase our confidence and avoid unnessecary manipulations during management of patients with inflammatory bowel disease presenting with symptoms of obstruction. In addition, this scheme provides time limits for each therapy and can be used as an algorithm of collaboration among surgeons, radiologists and gastroenterologists in order to provide optimal care and efficient solutions to patients with bowel obstruction.

#### REFERENCES

- Van Assche G, Geboes K, Rutgeerts P. Medical therapy for Crohn's disease strictures. Inflamm Bowel Dis 2004; 10:55-60
- Swaminath A, Lichtinger S. Dilation of colonic strictures by intralesional injection of infliximab in patients with Crohn's colitis. Inflamm Bowel Dis 2008; 14:213-216
- Miyagawa A, Yasuoka T, Sakabe H, Matsuda H, Nakajima S, Aoyama H, Kushima R, Fujiyama Y, Bamba T. A case of ulcerative colitis with ileus due to extensive stricture of the colon.Nippon Shokakibyo Gakkai Zasshi. 2000; 97:1267-1271.
- 4. Delgado-Aros S, Camilleri M. Clinical management of acute colonic pseudo-obstruction in patients: a systematic review of the literature. Gastroenterol Hepatol 2003; 26:646-655
- Delgado-Aros S, Camilleri M. Pseudo-obstruction in the critically ill. Best Pract Res Clin Gastroenterol 2003; 17:427-444
- Saunders MD, Cappell MS. Endoscopic management of acute colonic pseudo-obstruction. Endoscopy 2005; 37:760-763
- Saunders MD, Kimmey MB. Colonic pseudo-obstruction: the dilated colon in the ICU. Semin Gastrointest Dis 2003; 14;20-27
- Saunders MD, Kimmey MB. Systematic review: acute colonic pseudo-obstruction. Aliment Pharmacol Ther 2005; 22:917-925
- Saunders MD. Acute colonic pseudo-obstruction. Best Pract Res Clin Gastroenterol 2007; 21:671-687
- 10. Spira IA, Rodrigues R, Wolf WI. Pseudo-obstruction of the

- colon. Am J Gastroenterol 1976; 65:397-408
- De Giorgio R, Barbara G, Stanghellini V, Tonini M, Vasina V, Cola B, Corinaldesi R, Biagi G, De Ponti F. The pharmacologic treatment of acute colonic pseudo-obstruction. Aliment Pharmacol Ther 2001; 15:1717-1727
- Carethers JM, McDonnell WM, Owyang C, Scheiman JM.Massive secretory diarrhea and pseudo-obstruction as the initial presentation of Crohn's disease. J Clin Gastroenterol 1996; 23:55-59
- Miyajima S, Kuwano T.3 cases of Crohn's disease presenting as ileus. Geka Chiryo 1968; 19:353-356.
- Myrhoj T, Ladefoged K, Jarnum S. Chronic intestinal pseudo-obstruction in patients with extensive bowel resection for Crohn's disease. Scand J Gastroenterol 1988; 23:380-384
- Ogilvie WH. Large-intestine colic due to sympathetic deprivation. Br Med J 1948; 2:671-673
- Ohlsson B, Fork FT, Veress B, Toth E. Coexistent chronic idiopathic intestinal pseudoobstruction and inflammatory bowel disease. Gut 2005; 54:729-730
- Stawarski A, Iwańczak B, Krzesiek E, Iwańczak F. Intestinal complications and extraintestinal manifestations in children with inflammatory bowel disease. Pol Merkur Lekarski 2006; 20:22-25
- Telander RL, Schmeling DJ.Current surgical management of Crohn's disease in childhood.Semin Pediatr Surg 1994; 3:19-27.
- 19. Farmer KC, Phillips RK. True and false large bowel obstruction. Baillieres Clin Gastroenterol 1991; 5:563-585
- Dorudi S, Berry AR, Kettlewell MG. Acute colonic pseudoobstruction. Br J Surg 1992; 79:99-103
- Eisen GM, Baron TH, Dominitz JA, et al. Acute colonic pseudo-obstruction. Gastrointest Endosc 2002; 56:789-792
- Bruni R, Chirco L, Lemeni AR, Petrocca S. Intermittent small bowel obstruction by jejunal enterocolitis in a patient with a Crohn's disease stricture. Chir Ital 2002; 54:903-905
- Sorrentino D, Terrosu G, Vadalů S, Avellini C. Fibrotic strictures and anti-TNF-alpha therapy in Crohn's disease. Digestion 2007; 75:22-4.
- Batke M, Cappell MS. Adynamic ileus and acute colonic pseudo-obstruction. Med Clin North Am 2008; 92:649-670
- Hitosugi M, Kitamura O, Shigeta A, Takatsu A, Yoshino Y, Ohtsuki M. Analysis of sudden death caused by intestinal obstruction. Nihon Hoigaku Zasshi 1997; 51:423-429
- Cuadrado-Gomez LM, Arranz-Caso A, Albarran-Hernandez F, Alvarez de Mon M. [Toxic megacolon caused by loperamide as initial form of Crohn disease]. Rev Clin Esp 1994; 194:201-202
- Witteman BJ, Weterman IT, Griffioen G, Lamers CB. Intestinal obstruction caused by non-absorbable tablets and Budd-Chiari syndrome in a patient with Crohn's disease. Ned Tijdschr Geneeskd. 1991; 135:766-769
- Reddy KR, Thomas E. Toxic megacolon after proctosigmoidoscopy in ulcerative colitis. South Med J 1983; 76:1072-1073
- Burke JP, Ferrante M, Dejaegher K, Watson RW, Docherty NG, De Hertogh G, Vermeire S, Rutgeerts P, D'Hoore A,

Penninckx F, Geboes K, Van Assche G, O'Connell PR. Transcriptomic analysis of intestinal fibrosis-associated gene expression in response to medical therapy in Crohn's disease. Inflamm Bowel Dis 2008; 14:1197-1204

- Guagnozzi D, Cossu A, Viscido A, Corleto V, Annese V, Latiano A, Delle Fave G, Caprilli R. Acute intestinal obstruction and NOD2/CARD15 mutations among Italian Crohn's disease patients. Eur Rev Med Pharmacol Sci 2004; 8:179-185
- Takayama H, Takagi H, Larochelle WJ, Kapur RP, Merlino G. Ulcerative proctitis, rectal prolapse and intestinal pseudo-obstruction in transgenic mice overexpressing hepatocyte growth factor/scatter factor. Lab Invest 2001; 81:297-305
- 32. Koukoulis G, Ke Y, Henley JD, Cummings OW. Obliterative muscularization of the small bowel submucosa in Crohn disease: a possible mechanism of small bowel obstruction. Arch Pathol Lab Med 2001; 125:1331-1334
- Oehmichen M, Reifferscheid P. Intramural ganglion cell degeneration in inflammatory bowel disease. Digestion 1977; 15:482-496
- Trevisani GT, Hyman, NH, Church JM. Neostigmine: safe and effective treatment for acute colonic pseudo-obstruction. Dis Colon Rectum 2000; 43:599-603
- 35. Burke M. Acute intestinal obstruction: diagnosis and management. Hosp Med 2002; 63:104-107
- Gratama S, Smedts F, Whitehead R. obstructive colitis: an analysis of 50 cases and a review of the literature. Pathology 1995; 27:324-329
- Horton KM, Corl FM, Fisherman EK. CT evaluation of the colon: inflammatory disease. Radiographics 2000; 20:399-418
- Uenishi T, Sakata C, Tanaka S, Yamamoto T, Shuto T, Hiro-hashi K, Kubo S, et al. Eosinophilic enteritis presenting as acute intestinal obstruction: a case report and review of the literature. Dig Surg 2003; 20:326-329
- 39. Yeguez JF, Marinez SA, Sands LR, Hellinger MD. Pelvic actinomycosis presenting as malignant large bowel obstruction: a case report and a review of the literature. Am Surg 2000; 66:85-90
- Kerber GW, Frank PH. Carcinoma of the small intestine and colon as a complication of Crohn disease: radiologic manifestations. Radiology 1984; 150:639-645
- Balthazar EJ. George W Holmes lecture. CT of small bowel obstruction. AJR 1994; 162:255-261
- 42. Frager D. Instestinal obstruction: role of CT Gastroenterol Clin North Am 2002; 31:777-799
- Jacobs JE, Birnbaum BA. CT of inflammatory disease of the colon. Semin Ultrasound CT MR 1995; 16:91-101
- Megibow AJ. Bowel obstruction. Evaluation with CT. Radiol Clin North Am 1994; 32:861-870
- Romano S, Lombardo P, Cinque T, Tortora G, Romano L. Acute colonic disease: how to image in emergency. Eur J Radiol 2007; 61:424-432
- 46.Philpotts LE, Heiken JP, Westcott MA, Gore RM. Colitis: use of CT findings in differential diagnosis. Radiology 1994; 190:445-449
- Zissin R, Hertz M, Paran H, Bernheim J, Shapiro-Feiberg M, Gayer G. Small bowel obstruction secondary to Crohn

- disease: CT findings. Abdom Imaging 2004; 29:320-325
- 48. Ha HK, Park CH, Kim SK et al. CT analysis of intestinal obstruction due to adhesions: early detection of strangulation. J Comput Assist Tomogr 1993; 17:386-389
- 49. Raptopoulos V, Schwartz RK, Minicholas MJ, Morsono J, Pearlman J, Joffe N. Multiplanar helical CT enterography in patients with Crohn's disease. AJR 1997; 169:1545-1550
- Roberts-Thomson IC, Tucker GR, Hewett PJ, Cheung P, Sebben RA, Khoo EE, et al. Single-center study comparing computed tomography colonography with conventional colonoscopy. World J Gastroenterol 2008; 14:469-473
- Florie J, van Gelder RE, Stoker J. Colonography by computed tomography. Eur J Gastroenterol Hepatol 2005; 17:809-813
- 52. Frager D, Medwid SW, Baer JW, et al. CT of small bowel obstruction:value in establishing the diagnosis and determining the degree and cause. AJR 1994; 162:37-41
- 53. Farrell RJ, Morrin MM, McGee JB. Virtual colonoscopy: a gastroenterologist's perspective. Dig Dis 1999;17:185-193
- Rosman AS, Korsten MA. Meta-analysis comparing CT colonography, air contrast barium enema and colonoscopy. Am J Med 2007; 120:203-210
- Dvte P, Lata J, Novotnú I.Intestinal obstruction and perforation--the role of the gastroenterologist. Dig Dis. 2003; 21:63-67.
- 56. Triester SL, Leighton JA, Leontiadis GI, Gurudu SR, Fleischer DE, Hara AK, Heigh RI, Shiff AD, Sharma VK.A meta-analysis of the yield of capsule endoscopy compared to other diagnostic modalities in patients with non-stricturing small bowel Crohn's disease. Am J Gastroenterol. 2006; 101:954-964.
- Kastin DA, Buchman AL, Barrett T, Halverson A, Wallin A. Strictures from Crohn's disease diagnosed by video capsule endoscopy. J Clin Gastroenterol 2004; 38:346-349
- 58. Li F, Gurudu SR, De Petris G, Sharma VK, Shiff AD, Heigh RI, Fleischer DE, Post J, Erickson P, Leighton JA. Retention of the capsule endoscope: a single-center experience of 1000 capsule endoscopy procedures. Gastrointest Endosc 2008;68:174-180
- Magdeburg R, Riester T, Hummel F, Lohr M, Post S, Sturm J. Ileus secondary to wireless capsule enteroscopy. Int J Colorectal Dis 2006; 21:610-613
- Benchimol EI, Turner D, Mann EH, Thomas KE, Gomes T, McLernon RA, Griffiths AM. Toxic megacolon in children with inflammatory bowel disease: clinical and radiographic characteristics. Am J Gastroenterol 2008; 103:1524-1531
- Greenstein AJ, Sachar DB, Gibas A, Schrag D, Heimann T, Janowitz HD, Aufses AH Jr. Outcome of toxic dilatation in ulcerative and Crohn's colitis. J Clin Gastroenterol 1985; 7:137-143
- Fazio VW. Toxic megacolon in ulcerative colitis and Crohn's colitis. Clin Gastroenterol 1980; 9:389-407
- 63. Fukata M, Sugisaka H, Kijima H, Sanjo A, Takekuma Y, Inoue F, Nagayama K, Tanaka T, Hada T, Takao Y, Fukunaga M. A case of ulcerative colitis complicated by nontoxic megacolon. Nippon Shokakibyo Gakkai Zasshi 2000; 97:920-924
- 64. Funayama Y, Sasaki I, Naito H, Fukushima K, Shibata C,

- Ohtani N, Hiwatashi N, Masuda T, Matsuno S. A case of severe ulcerative colitis complicated by toxic megacolon. Nippon Shokakibyo Gakkai Zasshi 1998; 95:66-69
- Bharucha AE, Phillips SF. Megacolon: acute, toxic and chronic. Curr Treat Options Gastroenterol 1999; 2:517-523
- 66. Garcia-Osogobio S, Takahashi T, Gamboa-Dominguez A, Medina H, Arch J, Mass W, Sierra-Madero J, Uscanga L. Toxic pseudomembranous colitis in a patient with ulcerative colitis. Inflamm Bowel Dis 2000; 6:188-190
- Knapp AB, Mirsky FJ, Dillon EH, Korelitz BI. Successful infliximab therapy for a duodenal stricture caused by Crohn's disease. Inflamm Bowel Dis. 2005; 11:1123-1125.
- Kotanagi H, Fukuoka T, Shibata Y, Yoshioka T, Aizawa O, Saito Y, Koyama K, Otaka M, Chiba M, Saito M. A case of toxic megacolon in ulcerative colitis associated with cytomegalovirus infection. J Gastroenterol 1994; 29:501-505
- Sanchez Yubero S, Lopez Gil A, Perez Rojo JA, Kessler Saiz P. Toxic megacolon presenting as Crohn's disease. An Med Interna 1999; 16;580-582
- 70. Heise W, Kersten O, Kassner KM, Birkenmeyer G, Grosse G, Niedobitek F. Z Gastroenterol 1997; 35:481-490
- 71. Karjoo M, McCarthy B. Toxic megacolon of ulcerative colitis in infancy. Pediatrics 1976; 57:962-966
- Kisloff B, Adkins JC. Toxic megacolon developing in a patient with long-standing distal ulcerative colitis. Am J Gastroenterol 11981; 75:451-453
- Laurent S, Reenaers C, Detroz B, Detry O, Delvenne P, Belaiche J, Meurisse M. A patient who survived total colonic ulcerative colitis surinfected by cytomegalovirus complicated by toxic megacolon and disseminated intravascular coagulation. Acta Gastroenterol Belg 2005; 68:276-279
- Bauer W, Tepper R, Katz S. Megacolon as a presenting finding of acute pancreatitis associated with chronic ulcerative colitis:unusual presentation of an unusual association. Am J Gastroenterol 1995; 90:1013-1015.
- Lindahl JA, Williams FH, Newman SL. Small bowel obstruction in chronic granulomatous disease. J Pediatr Gastroenterol Nutr 1984; 3:637-640
- Molina-Infante J, Sanz-Garcia C, Catalina-Rodriguez MV, Nogales-Rincon O, Matilla-Pena A, Nunez-Martinez O, Clemente-Ricote G. Massive abdominal vein thrombosis with acute liver failure and toxic megacolon as onset of ulcerative colitis. Gastroenterol Hepatol 2005; 28:551-554
- 77. Arai H, Hanai H, Furuta T, Sato Y, Yamada M, Kaneko E, Baba S, Sugimura H. A patient who survived total colonic type ulcerative colitis complicated by toxic megacolon, disseminated intravascular coagulation, methicillin-resistant staphylococcus aureus infection and bilateral femoral phlebothrombosis. J Gastroenterol 1999; 34:395-399
- Melange M, Van Gossum A, Houben JJ, de Ronde T, Vanheuverzwyn R, Adler M. Acute dilatation of the colon. Acta Gastroenterol Belg 1991; 54:233-236
- Cooksey G, Gunn A, Wotherspoon WC. Surgery for acute ulcerative colitis and toxic megacolon during pregnancy. Br J Surg 1985; 72:547
- 80. Cunsolo A, Bragaglia RB, Arena N, Mastromatteo MG, Vecchi R, Gozzetti G. Toxic megacolon complicating ulcerative colitis and Crohn's disease. Int Surg 1985; 70:339-343

- 81. Christodoulou D, Skopelitou AS, Katsanos KH, Katsios C, Agnantis N, Price A, Kappas A, Tsianos EV. Small bowel adenocarcinoma presenting as a first manifestation of Crohn's disease: report of a case, and a literature review. Eur J Gastroenterol Hepatol. 2002; 14:805-810
- Eitan B, Zisman I, Qarawany M, Grishkan A, Mazor A. Crohn's disease of small bowel manifested by obstructive carcinoma. Harefuah 1993; 125:209-211
- Lingohr P, Knoefel WT, Kleimann E, Rheinwalt KP.Laparoscopic coincidental finding in a case of incomplete ileus: adenocarcinoma of the small intestine as first manifestation of Crohn's disease. Zentralbl Chir 2007; 132:564-568
- 84. Perzin KH, Peterson M, Castiglione CL, Fenoglio CM, Wolff M. Intramucosal carcinoma of the small intestine arising in regional enteritis (Crohn's disease). Report of a case studied for carcinoembryonic antigen and review of the literature. Cancer 1984; 54:151-162
- Cioffi U, De Simone M, Ferrero S, Ciulla MM, Lemos A, Avesani EC. Synchronous adenocarcinoma and carcinoid tumor of the terminal ileum in a Crohn's disease patient. BMC Cancer 2005; 5:157
- 86. Mijandrusić Sincić B, Kovac D, Jasić M, Grbas H, Uravić M, Depolo A. Crohn's disease and a gastrointestinal stromal tumor in an 81-year-old man--a rare coincidence. Zentralbl Chir 2005;130:597-599
- 87. Senofsky GM, Stabile BE. Gallstone ileus associated with Crohn's disease. Surgery 1990; 108:114-117
- Almogy G, Bauer JJ, Venturero M, Presen DH.Gallstone ileus and Crohn's disease without biliary-enteric fistula: report of a unique case. Mt Sinai J Med 2000; 67:159-162
- 89. Basili G, Lorenzetti L, Celona G, Biondi G, Preziuso E, Angrisano C, Goletti O, Belcari C, Venturini G.Gallstone ileus in patient with Crohn's disease: report of a clinical observation. Surg Endosc 2006; 20:703-704
- Highman L, Jagelman DG. Gallstone ileus complicating terminal ileal Crohn's disease. Br J Surg 1981; 68:201-202
- Scott EM, Freeman AH. Prominet omental and mesenteric vasculature in inflammatory bowel disease shown by computed tomography. Eur J Radiol 1996; 22:104-106
- 92. La Meir M, Van Molhem Y. Recurrence of gallstone ileus with Crohn's disease. Acta Chir Belg 2001; 101:35-37
- Ojeda VJ, Levitt S, Ryan G, Laurence BH. Cystic fibrosis, Crohn's colitis, and adult meconium ileus equivalent. Dis Colon Rectum. 1986; 29:567-571
- Jones MW, Koper B, Weatherhead WF. Crohn's disease with enterolith treated laparoscopically. JSLS 2005; 9:339-341
- Johnson CD, Ahlquist DA. Computed tomography colonography (virtual colonoscopy): a new method for colorectal screening. Gut 1999; 44:301-305
- Adelson JW, deChadarevian JP, Azouz EM, Guttman FM. Giant inflammatory polyposis causing partial obstruction and pain in 'healed' ulcerative colitis in an adolescent. J Pediatr Gastroenterol Nutr 1988; 7:135-140
- 97. Yada S, Matsumoto T, Kudo T, Hirahashi M, Yao T, Mibu R, Iida M. Colonic obstruction due to giant inflammatory polyposis in a patient with ulcerative colitis. J Gastroenter-ol 2005; 40:536-539

 Balazs M. Giant inflammatory polyps associated with idiopathic inflammatory bowel disease. An ultrastructural study of five cases. Dis Colon Rectum 1990; 33:773-777

- Hurlstone DP. Large-bowel obstruction secondary to localized rectal giant pseudopolyposis complicating ulcerative colitis: first reported case. Endoscopy 2002; 34:1025
- 100. Maldonado TS, Firoozi B, Stone D, Hiotis K. Colocolonic intussusception of a giant pseudopolyp in a patient with ulcerative colitis: a case report and review of the literature. Inflamm Bowel Dis 2004; 10:41-44
- 101. Kφrber J, Grammel S, Lobeck H, Weidemann H.Stenosis of the terminal ileum. Endometriosis as the differential diagnosis of Crohn's disease.Dtsch Med Wochenschr 1997; 122:926-929
- 102. Bracci F, Iacobelli BD, Papadatou B, Ferretti F, Lucchetti MC, Cianchi D, Francalanci P, Ponticelli A. Role of electrogastrography in detecting motility disorders in children affected by chronic intestinal pseudo-obstruction and Crohn's disease. Eur J Pediatr Surg 2003; 13:31-34
- 103. Kristinsson JO, Hopman WP, Oyen WJ, Drenth JP. Gastroparesis in patients with inactive Crohn's disease: a case series. BMC Gastroenterol 2007; 7:11
- 104. Jones IT, Fazio VW. Colonic volvulus. Etiology and management. Dig Dis 1989; 7:203-209
- 105. Kinjo F, Sunagawa T, Hokama A et al. Crohn's disease and volvulus. Gastrointest Endosc 2003; 58:758-759
- 106. Lashner BA, Anastaplo SM, Kirsner JB. Sigmoid volvulus as a complication of ileal Crohn's disease. J Clin Gastroenterol 1989: 11:82-84
- 107. Payne-James JJ. Sigmoid volvulus in association with Crohn's disease of the colon. Br J Surg 1986; 73:1005
- 108. Fink W, Cohen A, Greenstein AJ. Small bowel volvulus in association with recurrent Crohn's disease. Am J Gastroenterol 1980; 74:337-341
- 109. Javett SL. Volvulus of the ileum in Crohn's disease. S Afr Med J 1972; 46:967-968
- 110. Diamond MT. Regional ileitis with volvulus of the terminal ileum Gastroenterology 1958; 34:126-129
- 111. Masters H. Volvulus and gangrene of the cecum in a patient with ulcerative colitis. Calif Med 1961; 94:36-37
- 112. Katsanos KH, Ignatiadou E, Markouizos G, Doukas M, Siafakas M, Fatouros M, Tsianos E. V. Non-toxic megacolon due to transverse and sigmoid colon volvulus in a patient with ulcerative colitis. Journal of Crohn's and Colitis 2009; 3:38-41
- 113. Renzulli P, Maurer CA, Netzer P, Buchler MW. Preoperative colonoscopic derotation is beneficial in acute colonic volvulus. Dig Surg 2002; 19:223-229
- 114. Burkill G, Bell J, Healy J. Small bowel obstruction: the role of computed tomography in this diagnosis and management with reference to other imaging modalities. Eur Radiol 2001; 11:1405-1422
- 115. Ellozy SH, Harris MT, Bauer JJ, Gorfine SR, Kreel I. Early postoperative small-bowel obstruction: aprospective evaluation in 242 consecutive abdominal operations. Dis Colon Rectum 2002; 45:1214-1217
- 116. Frager D, Medwid SW, Baer JW, Mollinelli B, Friedman M. CT of small bowel obstruction: value in establishing the di-

- agnosis and determining the degree and cause. AJR 1994; 162:37-41
- 117. Nivatvongs S, Vermeulen FD, Fang DT. Colonoscopic decompression of acute pseudo-obstruction of the colon. Ann Surg 1982; 196:598-600
- 118. Eaker EY. Update on acute colonic pseudo-obstruction. Curr Gastroenterol Rep 2001; 3:433-436
- 119. MacColl C, MacCannell KL, Baylis B, et al. Treatment of acute colonic pseudo-obstruction (Ogilvie's syndrome) with cisapride. Gastroenterology 1990; 98:773-776
- 120. Sgouros SN, Vlachogiannakos J, Vassiliadis K, Bergele C, Stefanidis G, Nastos H, Avgerinos A, Mantides A. Effect of polyethylene glycol electrolyte balanced solution on patients with acute colonic-pseudo-obstruction after resolution of colonic dilation: a prospective, randomized, placebo controlled trial. Gut 2006; 55:638-642
- 121. Yukselen V, Vardar E, Yukselen O, Oruc N, Yenisey C, Karaoglu A, Ozutemiz O. Infliximab in experimental alkali burns of the oesophagus in the rat. Acta Gastroenterol Belg 2008; 71:21-26.
- 122. Belaiche J, Louis E.Early obstructive colon after treatment of active refractory Crohn's disease with infliximab. Gastroenterol Clin Biol 2003; 27:1045-1047.
- 123. Hansen RA, Gartlehner G, Powell GE, Sandler RS. Serious adverse events with infliximab: analysis of spontaneously reported adverse events. Clin Gastroenterol Hepatol 2007; 5:729-735.
- 124. Bouguen G, Trouilloud I, Siproudhis L, Oussalah A, Bigard MA, Bretagne JF, Peyrin-Biroulet L.Long-term outcome of non-fistulizing (ulcers, stricture) perianal Crohn's disease in patients treated with infliximab. Aliment Pharmacol Ther 2009; 30:749-756.
- 125. Louis E, Boverie J, Dewit O, Baert F, De Vos M, D'Haens G. Belgian IBD Research Group. Treatment of small bowel subocclusive Crohn's disease with infliximab: an open pilot study. Acta Gastroenterol Belg 2007; 70:15-19
- 126. Pelletier AL, Kalisazan B, Wienckiewicz J, Bouarioua N, Soult JC. Infliximab treatment for symptomatic Crohn's disease strictures. Aliment Pharmacol Ther 2009; 29:279-285
- 127. Sorrentino D. Factors associated with the development of intestinal strictures or obstructions in patients with Crohn's disease. Am J Gastroenterol 2006; 101:2892-2893.
- 128. Sorrentino D, Avellini C, Beltrami CA, Pasqual E, Zearo E.Selective effect of infliximab on the inflammatory component of a colonic stricture in Crohn's disease. Int J Colorectal Dis 2006; 21:276-281.
- 129. Colombel JF, Loftus EV Jr, Tremaine WJ, Pemberton JH, Wolff BG, Young-Fadok T, Harmsen WS, Schleck CD, Sandborn WJ. Early postoperative complications are not increased in patients with Crohn's disease treated perioperatively with infliximab or immunosuppressive therapy. Am J Gastroenterol. 2004; 99:878-883
- 130. Kunitake H, Hodin R, Shellito PC, Sands BE, Korzenik J, Bordeianou L. Perioperative treatment with infliximab in patients with Crohn's disease and ulcerative colitis is not associated with an increased rate of postoperative complications. J Gastrointest Surg 2008; 12:1730-1736
- 131. Lichtenstein GR, Olson G, Travers S, Diamont RH, Chen

- DM, Prichard ML, Feagan B, Cohen RD, et al. Factors associated with the development of intestinal strictures or obstructions in patients with Crohn's disease. Am J Gastroenterol 2006; 101:1030-1038
- 132. Pallotta N, Barberani F, Abdulkadir Hassan N Guagnozzi D, Vincoli G, Corazziari E. Effect of Infliximab on small bowel stenoses in patients with Crohn's disease. World J Gastroenterol 2008; 14:1885-1890
- 133. Di Sabatino A, Ciccocioppo R, Benazzato L, Sturniolo GC, Corazza GR. Infliximab downregulates basic fibroblast growth factor and vascular endothelial growth factor in Crohn's disease patients. Aliment Pharmacol Ther 2004; 19:111019-111024
- 134. Järnerot G, Hertervig E, Friis-Liby I, Blomquist L, Karlén P, Grännö C, et al. Infliximab as rescue therapy in severe to moderately severe ulcerative colitis: a randomized, placebocontrolled study. Gastroenterology 2005; 128:1805-1811.
- 135. Castro Fernandez M, Garcia Romero D, Sanchez Munoz D, Grande L, Larrraona JL. Severe ulcerative colitis with toxic megacolon resolved with infliximab therapy. Rev Esp Enferm Dig 2007; 99:426-427
- 136. Sriram PV, Reddy KS, Rao GV, Santosh D, Reddy DN. Infliximab in the treatment of ulcerative colitis with toxic megacolon. Indian J Gastroenterol 2004; 23:22-23
- 137. Blomberg B.Endoscopic treatment modalities in inflammatory bowel disease. Endoscopy 1992; 24:578-581.
- 138. Vanek VW, Al-Salti M. Acute pseudo-obstruction of the colon (Ogilvie's syndrome). An analysis of 400 cases. Dis Colon Rectum 1986; 29:203-210
- 139. Van Gossum A, Bourgeois F, Gay F, Lievens P, Adler M, Cremer M. Operative colonoscopic endoscopy. Acta Gastroenterol Belg 1992; 55:314-326
- 140. Nedin D, Kostadinova R, Deleva V, Popova D, Timev I, Tsenova V. Surgical tactic in a severe form of Crohn's disease complicated by ileus and cachexia. Khirurgiia 2002; 58:52-53
- 141. Hoashi T, Tsuda S, Yao T, Ichimaru T, Nakahara T, Okamoto T, Futami, K, Arima S, Iwashita A. A case of ulcerative colitis with toxic megacolon, successfully treated with colonoscopic decompression. Nippon Shokakibyo Gakkai Zasshi 1991; 88:91-95
- 142. Banez AV, Yamanishi F, Crans CA. Endoscopic colonic decompression of toxic megacolon, placement of colonic tube and steroid colonclysis. Am J Gastroenterol 1987; 82:692-694
- 143. Jetmore AB, Timmcke AE, Garthright Jr BJ, et al. Ogilvie's syndrome: coloscopic decompression and analysis of predisposing factors. Dis Colon Rectum 1992; 35:1135-1142
- 144. Ramage JI, Baron TH. Percutaneous endoscopic cecostomy: a case series. Gastrointest Endosc 2003; 57:752-755
- 145. Rex DK. Colonoscopy and acute colonic pseudo-obstruction. Gastrointest Endosc Clin North Am 1997; 7:499-508
- 146. Freeman AH. CT and bowel disease. Br J Radiol 2001;74:4-14
- 147. Forde KA. Therapeutic colonoscopy. World J Surg 1992; 16:1048-1053
- 148. Baron TH. Expandable metal stents for the treatment of cancerous obstruction of the gastrointestinal tract. N Engl J Med

- 2001; 344:1681-1687
- 149. Geller A, Petersen BT, Gostout CJ. Endoscopic decompression for acute colonic pseudo-obstruction. Gastrointest Endosc 1996; 44:144-150
- 150. Gay SSB, Shaffer HA Jr, Futterer SSF, Atchison PM, Patel SJ. Gastrointestinal case of the day. Toxic megacolon with underlying ulcerative colitis. Am J Roetgenol 1996;167:240-242
- 151. Brolin RE. The role of gastrointestinal tube decompression in the treatment of mechanical intestinal obstruction. Am Surg 1983; 49:131
- 152. Goldstein ES, Rubin PH. Endoscopic Therapy for inflammatory bowel disease. Curr Treat Options Gastroenterol 2003; 6:237-243
- 153. Shimada Y, Iai T, Okamoto H, Suda T, Hatekeyama K, Honma T, Ajioka Y. Toxic megacolon associated with cytomegalovirus infection in ulcerative colitis. J Gastroenterol 2003; 38:1107-1108
- 154. Solt J, Hertelendy A, Szilαgyi K. Long-term results of balloon catheter dilation of lower gastrointestinal tract stenoses. Dis Colon Rectum 2004; 47:1499-1505.
- 155. East JE, Brooker JC, Rutter MD, Saunders BP.A pilot study of intrastricture steroid versus placebo injection after balloon dilatation of Crohn's strictures. Clin Gastroenterol Hepatol 2007; 5:1065-1069.
- 156. Jalan KN, Sircus W, Card WI et al. An experience in ulcerative colitis. Toxic dilatation in 55 cases. Gastroenterology 1969; 57:68-82
- 157. Stoianov G, Aleksandrova A, Vasilev V. Crohn's disease in the cecal area. Apropos of a case with a course like ileus. Khirurgiia 1984; 37:143-144
- 158. Berg DF, Bahadursingh AM, Kaminski DL, Longo WE. Acute surgical emergencies in inflammatory bowel disease. Am J Surg 2002;184:45-51.
- 159. Cheung O, Regueiro MD. Inflammatory bowel disease emergencies. Gastroenterol Clin North Am 2003; 32:1269-1288
- 160. Bitton A, Peppercorn MA. Emergencies in inflammatory bowel disease. Crit Care Clin 1995; 11:513-529
- 161. D'Addazzio G, Scordamaglia R, Tedeschi U, Barra M, Arpe F, Beatini L, Meola V. Emergency surgery in inflammatory bowel disease. Experience in 26 cases. Minerva Chir 1998; 53:1001-1007
- 162. Igarashi C, Hori T, Yoshida M, Yokozawa M, Fujita S, Yonezawa K, Tosaka M, Yoshida Y. Acute fulminant ulcerative colitis with toxic megacolon. Acta Paediatr Jpn 1997; 39:237-240
- 163. Linssen A, Tytgat GN. Fulminant onset of Crohn's disease of the colon. An observation of six cases. Dig Dis Sci 1982; 27:731-736
- 164. Ginanneschi U, Fabiani P, Rizzi M, Fanti G. Emergency surgical treatment of ulcerative rectocolitis and Crohn's disease of the colon. Ann Ital Chir 1996; 67:193-196
- 165. Goebell H. Acute manifestations in Crohn disease and ulcerative colitis. Conservative treatment of acute situations. Langenbecks Arch Chir 1988; Suppl2:77-81
- 166. Zenilman ME, Becker JM. Emergencies in inflammatory bowel disease. Gastroenterol Clin North Am 11988; 17:387-408

167. Metcalf AM. Elective and emergent operative management of ulcerative colitis. Surg Clin North Am 2007; 87:633-641

- 168. Ziv Y, Fazio VW, Church JM, Milsom JW, Schroeder TK. Safety of urgent restorative proctocolectomy with ileal pouch-anal anastomosis for fulminant colitis. Dis Colon Rectum 1995; 38:345-349
- 169. Eigler FW, Lange R, Luetkens S. Surgical treatment of acute conditions of Crohn disease. Langebecks Arch Chir 1988; Suppl 2:83-87
- 170. Broering DC, Eisenberger CF, Koch A, Bloechle C, Knoefel WT, Izbicki JR. Quality of life after surgical therapy of small bowel stenosis in Crohn's disease. Dig Surg 2001; 18:124-130
- 171. Ozuner G, Fazio VW, Lavery IC, Church JM, Hull TL. How safe is stricture plasty in the management of Crohn's disease? Am J Surg 1996;171:57-60
- 172. Gumaste V, Sachar DB, Greenstein AJ. Benign and malignant colorectal strictures in ulcerative colitis. Gut 1992; 33:938-941
- 173. Makowiec F, Starlinger M. Gastroduodenal involvement and circumscribed intestinal stricture in Crohn's disease. Zentraldl Chir 1998; 123:3338-3343
- 174. Wehner S, Behrendt FF, Lyutenski BN, Lysson M, Bauer AJ, Hirner A, Kalff JC. Inhibition of macrophage function prevents intestinal inflammation and postoperative ileus in rodents. Gut 2007; 56:176-185
- 175. Moore BA, Albers KM, Davis BM, Grandis JR, Tögel S, Bauer AJ. Altered inflammatory gene expression underlies increased susceptibility to murine postoperative ileus with advancing age. Am J Physiol Gastrointest Liver Physiol 2007; 292:G1650-1659.
- 176. Hughes ES, McDermot FT, Masterton JP.Intestinal obstruction following operation for inflammatory disease of the bowel. Dis Colon Rectum 1979; 22:469-471
- 177. Landsend E, Johnson E, Johannessen HO, Carlsen E. Longterm outcome after intestinal resection for Crohn's disease. Scand J Gastroenterol 2006; 41:1204-1208
- 178. Garcia-Botello SA, Garcia-Armengol J, Garcia-Granero E, Espi A, Juan C, Lopez-Mozos F, Lledo S. A prospective au-

- dit of the complications of loop ileostomy construction and takedown. Dig Surg 2004; 21:440-446
- 179. Aberg H, Pahlman L, Karlbom U. Small-bowel obstruction after restorative proctocolectomy in patients with ulcerative colitis. Int J Colorectal Dis 2007; 22:637-642
- 180. Francois Y, Dozois RR, Kelly KA, Beart RW Jr, Wolf BG, Pemberton JH, Ilstrup DM. Small intestinal obstruction complicating ileal pouch-anal anastomosis. Ann Surg 1989; 209:46-50
- 181. Stelzner M, Phillips JD, Fonkalsrud EW. Acute ileus from steroid withdrawal simulating intestinal obstruction after surgery for ulcerative colitis. Arch Surg 1990; 125:914-917
- 182. Fazel A, Verne GN. New solutions to an old problem: acute colonic pseudo-obstruction. J Clin Gastroenterol 2005; 39:17-20
- 183. Pliego Perez AR, Cardenas Salinas LJ, Rodriguez Ballesteros R. Rectal catheter decompression in colonic pseudo-obstruction (Ogilvie syndrome). Report of a case and review of the literature. Ginecol Obstet Mex 2000; 68:39-41
- 184. Loftus CG, Harewood GC, Baron TH. Assessment of predictors of response to neostgmine for acute colonic pseudoobstruction. Am J Gastroenterol 2002; 97:3118-3122
- 185. Carman R, Iannuccilli EA, Thayer WR. Toxic megacolon in inflammatory bowel disease: a new perspective. R I Med J 1978; 61:342-346
- 186. Page MJ, Poritz LS, Kunselman SJ, Koltun WA. Factors affecting surgical risk in elderly patients with inflammatory bowel disease. J Gastrointest Surg 2002; 6:606-613
- 187. Latella G, Vernia P, Viscido A, Frieri G, Cadau G, COcco A, Cossu A, Tomei E, Caprilli R. GI distension in severe ulcerative colitis. Am J Gastroenterol 2002; 97:11169-11175
- 188. Mogan GR, Sachar DB, Bauer J, Salky B, Janowitz HD. Toxic megacolon in ulcerative colitis complicated by pneumo-mediastinum: report of two cases. Gastroenterology 1980; 79:559-562
- 189. Caprilli R, Vernia P, Colaneri O, Frieri G. Risk factors in toxic megacolon. Dig Dis Sci 1980; 25:817-822
- 190. Forssmann K, Singer MV.Therapy of Crohn disease in internal medicine:short bowel syndrome and toxic megalon. Praxis 1994; 87:1652-1656