

Clinical Practice Guideline for the Surgical Management of Crohn's Disease

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The American Society of Colon and Rectal Surgeons is dedicated to ensuring high-quality patient care by advancing the science, prevention, and management of disorders and diseases of the colon, rectum, and anus. The Clinical Practice Guidelines Committee is composed of Society members who are chosen because they have demonstrated expertise in the specialty of colon and rectal surgery. This Committee was created to lead international efforts in defining quality care for conditions related to the colon, rectum, and anus. This is accompanied by developing Clinical Practice Guidelines based on the best available evidence. These guidelines are inclusive, and not prescriptive. Their purpose is to provide information based on which decisions can be made, rather than dictate a specific form of treatment. These guidelines are intended for the use of all practitioners, health care workers, and patients who desire information about the management of the conditions addressed by the topics covered in these guidelines.

It should be recognized that these guidelines should not be deemed inclusive of all proper methods of care or exclusive of methods of care reasonably directed to obtaining the same results. The ultimate judgment regarding the propriety of any specific procedure must be made by the physician in light of all the circumstances presented by the individual patient.

STATEMENT OF THE PROBLEM

Crohn's disease is a chronic, unremitting, incurable inflammatory disorder that can affect any segment of the intestinal tract, as well as extraintestinal sites. Disease behavior is classified as nonstricturing, nonpenetrating (ie, inflammatory), fibrostenotic, or penetrating, and is

prone to change over the course of the disease. Approximately 19% to 38% of patients have stricturing or penetrating complications at the time of diagnosis, whereas ~61% to 88% of patients will have developed these manifestations after 20 years of disease.¹ Although the etiology of Crohn's disease remains unknown, the disease likely results from a complex interplay between conditioning factors (eg, genetics, triggering events) and effector mechanisms that lead to a dysregulation of both intestinal immune and nonimmune functions.²⁻⁵

Before the introduction of anti-tumor necrosis factor (TNF) medications, population-based cohorts from Canada, Denmark, Norway, Sweden, the United States, and Wales reported that surgical intervention is performed in 27% to 61% of patients with Crohn's disease within 5 years of diagnosis.⁶ Independent risk factors associated with undergoing an initial surgery (versus reoperation) according to similar population-based cohorts include current tobacco usage, disease location, stricturing or penetrating disease behavior, and (although somewhat controversial) the early use of high-dose glucocorticoids or immunomodulators.⁷⁻¹⁰ This clinical practice guideline will focus on the surgical management of patients with Crohn's disease.

METHODOLOGY

This clinical practice guideline is based on the previous American Society of Colon and Rectal Surgeons' *Practice Parameters for the Surgical Management of Crohn's disease* that was published in 2007.¹¹ The management of perianal disease in patients with Crohn's disease is discussed in a separate clinical practice guideline.¹² An organized search of MEDLINE, PubMed, EMBASE, and the Cochrane Database of Collected Reviews was performed from 1978 through February 2015. Key-word combinations included "Crohn's disease"; "colitis"; "ileitis"; "fibrostenotic"; "stricture"; "abscess"; "phlegmon"; "fistula"; "megacolon"; "fulminant";

“complications”; “infliximab”; “steroids”; “TNF”; “tumor necrosis factor”; “immunomodulators”; “Crohn abscess”; “Crohn anastomosis”; “Crohn cancer”; “Crohn colitis”; “Crohn dilation”; “Crohn dysplasia”; “Crohn endoscopy”; “Crohn laparoscopy”; “Crohn laparotomy”; “Crohn operation”; “Crohn resection”; “Crohn stricture”; “Crohn strictureplasty”; and “Crohn surgery,” among others.

Directed searches of the embedded references from the primary articles were also performed in selected circumstances. Although not intended to be exclusionary, the authors primarily focused on English language articles and studies in adults. Recommendations were formulated by the primary authors and reviewed by the entire ASCRS Clinical Practice Guidelines Committee. The final grade of recommendation was performed by using the GRADE system (Table 1).¹³

OPERATIVE INDICATIONS

Failed Medical Therapy

1. **Patients who demonstrate an inadequate response to, develop complications from, or are noncompliant with medical therapy should be considered for surgery. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.**

The goals of medical treatment of Crohn's disease are to achieve remission in patients with active disease and to avoid relapse in those with quiescent disease. Clinical evidence of improvement usually begins in approximately 2 to 4 weeks for the majority of classes of medications used in patients with Crohn's disease. For immunomodulators, a plateau in improvement of clinical symptoms is demonstrated within 12 to 16 weeks.¹⁴ Patients achieving clinical remission should then be considered for maintenance medical therapy.

Surgical intervention is often warranted in patients who are unable to tolerate medical therapy because of side effects,^{15,16} are noncompliant with medical therapy, or do not adequately control their disease activity with medical therapy.¹⁷ One special circumstance involves those patients requiring steroids, where regardless of disease severity, steroids should be transitioned to another medication as soon as possible owing to the deleterious effects of chronic glucocorticoid use. If a patient cannot tolerate medical therapy other than steroids,¹⁸ or if their disease is limited in extent (eg, short-length ileocolic disease), surgery should be strongly considered.

2. **Patients receiving therapy with anti-TNFs, high-dose glucocorticoids, and/or cyclosporine may warrant staged procedures because of concerns about postoperative complications; however, decisions should be individualized based on the patient's risk stratification, overall clinical status, and surgeon judgment. Grade**

of Recommendation: Weak recommendation based on low- or very low-quality evidence, 2C.

The association between antitumor necrosis factor agents and complications in patients with Crohn's disease remains unclear, with studies reporting disparate results.^{19–38} Unfortunately, methodological flaws in the individual studies, ranging from a failure to control for disease activity, being underpowered, or having significant heterogeneity, make it difficult to provide widespread recommendations. The pharmacokinetics of monoclonal antibodies such as infliximab allow for persistent therapeutic concentrations for at least 8 weeks after infusion. Given this controversy, elective surgeries in patients with minimal symptoms may be accordingly delayed as permitted until the end of this period or longer when feasible. However, in both the elective and emergent setting, the decision to perform a staged procedure and/or diversion should involve the patient's overall clinical picture and risk stratification. Factors include nutritional status, intraoperative factors (ie, site of resection, technical issues, associated inflammation), hemodynamic status, and the presence or absence of a surgical delay following failure to respond to medical therapy.^{39–42}

Similarly, the impact of preoperative high-dose glucocorticoid therapy (eg, prednisone >20 mg per day for >6 weeks) on postoperative complications is controversial.^{43–47} Steroids appear to increase the risk for infectious complications, especially when given in combination with anti-TNF agents or cyclosporine.^{18,48,49} This potential risk should be balanced against that of diversion and its inherent complications.⁵⁰ The need for surgery and the timing of operative intervention in patients whose condition initially improves, but never completely normalizes, should be left to the clinical judgment of the treating physician. Continued observation of these patients may exhaust their physiological reserve without benefit, and operative intervention is often recommended despite initial improvement – such as in the case of using cyclosporine to salvage patients with severe colitis. Unfortunately, the literature regarding cyclosporine is principally limited to patients with ulcerative colitis, although this group of patients may be analogous to patients with Crohn's colitis.^{51–53}

Inflammation

1. **Patients with acute colitis who have symptoms or signs of impending or actual perforation should typically undergo surgery. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.**

Acute severe colitis is generally defined as ≥ 6 grossly bloody bowel movements per day plus 1 sign of systemic toxicity,

TABLE 1. The GRADE system-grading recommendations

	Description	Benefit vs Risk and Burdens	Methodological Quality of Supporting Evidence	Implications
1A	Strong recommendation, High-quality evidence	Benefits clearly outweigh risk and burdens or vice versa	RCTs without important limitations or overwhelming evidence from observational studies	Strong recommendation, can apply to most patients in most circumstances without reservation
1B	Strong recommendation, Moderate-quality evidence	Benefits clearly outweigh risk and burdens or vice versa	RCTs with important limitations (inconsistent results, methodological flaws, indirect or imprecise) or exceptionally strong evidence from observational studies	Strong recommendation, can apply to most patients in most circumstances without reservation
1C	Strong recommendation, Low- or very low- quality evidence	Benefits clearly outweigh risk and burdens or vice versa	Observational studies or case series	Strong recommendation but may change when higher-quality evidence becomes available
2A	Weak recommendation, High-quality evidence	Benefits closely balanced with risks and burdens	RCTs without important limitations or overwhelming evidence from observational studies	Weak recommendation, best action may differ depending on circumstances or patients' or societal values
2B	Weak recommendations, Moderate-quality evidence	Benefits closely balanced with risks and burdens	RCTs with important limitations (inconsistent results, methodological flaws, indirect or imprecise) or exceptionally strong evidence from observational studies	Weak recommendation, best action may differ depending on circumstances or patients' or societal values
2C	Weak recommendation, Low- or very low- quality evidence	Uncertainty in the estimates of benefits, risks, and burden; benefits, risk, and burden may be closely balanced	Observational studies or case series	Very weak recommendations, other alternatives may be equally reasonable

GRADE = Grades of Recommendation, Assessment, Development, and Evaluation; RCT = randomized controlled trial.

Adapted from: Guyatt G, Guterman D, Baumann MH, et al. Grading strength of recommendations and quality of evidence in clinical guidelines: report from an American College of Chest Physicians Task Force. *Chest*. 2006;129:174–181.¹³ Used with permission.

which include anemia (<10.5 g/dL), elevated erythrocyte sedimentation rate (>30 mm/h), fever (>37.8°C), and tachycardia (>90 beats per minute).⁵⁴ Alternatively, “fulminant” colitis is characterized by >10 bloody bowel movements per day, daily continuous bleeding, a blood transfusion requirement, an elevated erythrocyte sedimentation rate (>30 mm/h), fever (>37.5°C), tachycardia (>90 beats per minute), abdominal tenderness and distension, and colonic dilation on abdominal radiographs.⁵⁵

Toxic megacolon is defined as a total or segmental nonobstructive dilation (ie, transverse colon >5.5 cm in the midline) of the colon associated with systemic toxicity,^{56,57} and it suggests that the patient is at higher risk for perforation. When severe colitis exists in the setting of persistent or progressing colonic dilation, evolving local peritonitis, and multiple organ failure, impending or actual perforation is of even greater concern and surgery is typically warranted.⁵⁸ Outcomes associated with the medical management of severe or fulminant colitis secondary to Crohn's disease are relatively lacking, but the experience gained from treating patients with severe ulcerative colitis is again generally extrapolated.⁵⁴ Alternative causes of severe colitis (eg, *Clostridium difficile* and cytomegalovirus) should be excluded with stool studies and endoscopic biopsy.⁵⁹ Endoscopy with minimal insufflation may be used with extreme caution to assess for concomitant disease as

well as the endoscopic severity of disease, which is associated with the likelihood for response to medical therapy.⁵⁴ Medical therapy with parenteral glucocorticoids or biological agents may be initiated, whereas narcotics and antidiarrheal agents should generally be avoided to prevent progression to megacolon. Bowel rest and parenteral antibiotics have not proven to reliably improve severe or fulminant colitis, although their use should be dictated by the individual patient's clinical scenario. Medical therapy is judged to have failed in patients when their symptoms are uncontrolled by medications or there is clinical evidence of persistent or progressive disease (eg, persistent tachycardia, hypotension, fever, leukocytosis, or persistent elevation of inflammatory markers such as c-reactive protein). Emerging evidence indicates that therapeutic drug level and anti-TNF antibody assessments are also useful in the determination of medical therapy success or failure.⁶⁰ Patients receiving parenteral glucocorticoids should typically show improvement within 2 to 3 days after initiating therapy.⁶¹ If not, salvage therapy with a biological agent or operative intervention is often required. As seen in ulcerative colitis, a favorable response to a biological agent should normally be demonstrated within 5 to 7 days after receiving the medication.^{62–68} Failure to respond, again, typically indicates a need for operative intervention.

Stricture

1. Endoscopic dilation may be considered for patients with symptomatic small-bowel or anastomotic strictures that are not amenable to medical therapy. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.

Strictures complicating Crohn's disease can arise anywhere in the intestinal tract, and they are conventionally classified as either inflammatory or fibrostenotic in nature. However, these categories are not mutually exclusive: intestinal fibrosis is a dynamic process, and inflammation and fibrosis can readily coexist within the same stricture or region.⁶⁹ Computed tomography enterography and magnetic resonance enterography can diagnose small-bowel strictures with a high level of accuracy, and may be considered apart or in addition to standard CT.^{70,71} Both CT enterography and MR enterography have a relatively high sensitivity and specificity for identifying disease activity that helps discriminate between inflammatory and fibrotic strictures.⁷² However, MRI is not associated with radiation exposure like CT imaging, and may be more appropriate in younger patients who potentially face an increased lifetime cumulative radiation exposure. At present, neither modality is recommended for the routine assessment of ileocolic or colonic strictures unless the stricture or more proximal areas of concern are inaccessible by endoscopy.

Medical therapy remains the first line of treatment, especially for inflammatory strictures^{14,18}; however, endoscopic dilation of the affected area can also be utilized for strictures that are readily accessible, especially anastomotic strictures.⁷³ The most commonly utilized approach is a "through-the-scope" method using 10- to 20-mm balloons, although a standardized methodology of sequential dilations has not been established.⁷⁴ The typical indication is a limited, short-segment (<5 cm) stricture in the absence of associated penetrating disease (eg, abscess, fistula).

The most commonly reported complications following dilation are hemorrhage, perforation, and sepsis. Although the average technical success rate is nearly 90%,⁷⁵ the 5-year clinical recurrence rate is ~36%.⁷⁶ Dilatation of strictures <4 cm in length in the setting of Crohn's disease has been associated with the best long-term outcome.⁷⁷ Patients usually require at least 2 dilations to achieve patency over a 5-year period.⁷⁴ Experience is limited with other reported techniques including expandable metal⁷⁸ or biodegradable stent placement,⁷⁹ as well as the off-label use of intralesional glucocorticoid⁸⁰ or infliximab injection,⁸¹ prohibiting a generalized recommendation until more experience is available.

2. Surgery is indicated for patients with symptomatic small-bowel or anastomotic strictures that are not amenable to medical therapy and/or dilation. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.

Surgery (eg, resection or strictureplasty) is warranted for small-bowel and anastomotic strictures when medical and/or endoscopic treatment fails to adequately improve the patient's symptoms or when there is concern about concomitant malignancy (see below).⁸² Strictureplasty may be more appropriate for nonphlegmonous strictures in selected patients with diffuse involvement of the small bowel, existing or impending short-bowel syndrome, or rapidly recurring disease. The type of strictureplasty performed is largely based on the length of the stricture and the pliability of the affected bowel.^{83,84} Complications after strictureplasty have been reported in 4% to 15% of patients in large series, and include obstruction, bleeding, sepsis, perforation, and death.^{84,85} Older age,⁸⁶ preoperative anemia,⁸⁷ hypoalbuminemia,⁸⁸ weight loss,⁸⁶ emergency surgery,⁸⁹ and the presence of an intra-abdominal abscess with peritoneal contamination⁹⁰⁻⁹² are all reported risk factors for postoperative morbidity. Conversely, preoperative glucocorticoid use, number, site and length of strictureplasties, and synchronous bowel resection have not been associated with a higher rate of complications.⁸⁵ Meta-regression analysis has reported 5-year recurrence rates for strictureplasty in jejunoileal and ileocolonic locations between 25% and 30%, including a ~3% site-specific recurrence, with the majority of recurrences developing in intestinal segments remote from the previous strictureplasty sites.⁸⁵ Although patients undergoing strictureplasty alone are at risk for recurrence,⁹³ this is, in general, comparable to resection.⁹⁴ Factors such as malnutrition, the presence of phlegmon/perforation/fistula at the site, multiple strictures within a small segment, and suspicion of a malignancy are all relative contraindications to strictureplasty.⁹⁴ The procedure can also be safely performed for disease involving the neo-terminal ileum associated with an ileocolostomy when short-bowel syndrome is a concern.^{95,96} However, the 5-year cumulative surgical recurrence rate in this location is ~45% and 24% for strictureplasty and resection of the ileocolostomy.⁹²

3. Patients with strictures of the colon that cannot be adequately surveyed endoscopically should be considered for resection. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.

Because of the transmural inflammation associated with Crohn's disease, it is not uncommon to have luminal narrowing within the colon, especially with repeated flares. Obstructive symptoms from colonic strictures occur in up to 17% of patients.⁹⁷ Even in the absence of symptoms, colonic strictures harbor occult carcinoma in approximately 7% of instances.⁹⁷ It is often difficult to differentiate malignant from benign strictures on a strictly clinical basis. Short strictures, as opposed to long strictures, appear to be more commonly associated with cancer, as is advanced age and longer duration of disease (3.3% with less than 20 years of Crohn's disease vs 11% with disease >20 years).⁹⁷

However, any colorectal stricture should be extensively assessed with multiple endoscopic biopsies to ensure the absence of malignancy. Cytological investigation of the large bowel (although not widely utilized) has been reported to aid in the diagnosis of colorectal carcinomas in this setting. Brushing with cytology appears to be an easy, highly specific, but relatively insensitive method to help in the diagnosis of colorectal cancer, and can be used as a complement to biopsy.⁹⁸ If the stricture cannot be adequately surveyed to exclude a concomitant carcinoma, resection of the affected large intestine should be considered. In this situation, appropriate resection following standard oncological principles should be performed. Although colonic strictureplasty has been reported for segmental disease,⁹⁹ it is generally discouraged owing to a lack of proven benefit, as well as concerns about concomitant carcinoma being left in situ.

Penetrating Disease

1. Patients with a free perforation should undergo surgery. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.

Patients with Crohn's disease can present with free perforation of the small or large bowel, and perforation is the indication for surgery in ~1% to 16% of instances.¹⁰⁰ Immediate resection of the perforated segment is preferred over simple suture closure¹⁰¹ because of a relatively high failure rate and concomitant increased mortality associated with the latter procedure.¹⁰² After resection of a small-bowel perforation, an anastomosis is typically possible. A diverting or an end stoma in the setting of a small-bowel perforation may be performed depending on the presentation and operative findings, and is often reserved for patients with severe hemodynamic instability, significantly edematous bowel, or technical challenges in constructing the anastomosis. In addition, a stoma following a small-bowel perforation may be needed in patients with other coexisting risk factors such as severe malnutrition or significant abdominal contamination. Benefits of stoma creation to divert a low-risk small-bowel anastomosis should always be balanced against the risk of additional surgeries that will subsequently be needed to close it.

In patients presenting with a free colonic perforation, a primary anastomosis with fecal diversion or an end stoma may be considered, especially in a patient presenting with feculent peritonitis, associated hemodynamic instability, moderate to severe protein-calorie malnutrition, chronic steroid or anti-TNF use, and treatment delay.¹⁰³ Additional factors such as the extent of colonic resection, location of the anastomosis, baseline continence, and the presence or absence of proctitis or perianal disease also need to be taken into account when considering the need for diversion.¹⁰⁴ Carefully selected patients (ie, hemodynamically stable, healthy bowel, technically sound anasto-

mosis, minimal comorbidities) presenting with purulent peritonitis but without widespread feculent contamination may be suitable for a primary anastomosis. Finally, any unstable patient or those with severe peritoneal contamination from perforation at any site may be left in temporary discontinuity and brought back to the operating room for a washout and/or definitive procedure following appropriate resuscitation.

2. Patients with enteroparietal, interloop, intramesenteric, or retroperitoneal abscesses may be managed by antibiotics with or without percutaneous drainage. Surgical drainage with or without resection should be considered when this is not successful. Grade of Recommendation: Weak recommendation based on moderate-quality evidence, 2B.

Intra-abdominal abscesses in patients with Crohn's disease typically result from a perforation or penetrating ulcers that are contained by surrounding structures, which may include adjacent segments of bowel. Small (<3 cm) abscesses can generally be sterilized with parenteral antibiotics alone.¹⁰⁵ Antibiotics and percutaneous drainage (PD) of a larger abscess usually resolve symptoms of sepsis, infrequently results in an enterocutaneous fistula, and occasionally obviates the need for future surgery.¹⁰⁶⁻¹¹¹ Comparative analyses of PD and initial surgery for spontaneous abdominal abscess in Crohn's disease indicate that successful PD, defined as abscess resolution and avoidance of subsequent surgery, occurs in the range of 23% to 78%.¹¹²⁻¹¹⁶ Factors associated with PD failure include concomitant steroid use, colonic disease, and abscesses that are large, multiloculated, or multifocal.^{115,117,118} In comparison with initial surgery, PD as a bridge to surgical resection results in a decrease in overall complications, diverting stomas, and overall cost, with similar rates of postoperative enterocutaneous fistula and anastomotic leak.^{115,116} Initial PD combined with intravenous antibiotics, steroids, total parenteral nutrition, and same admission bowel resection has been shown to result in the ability to perform a primary anastomosis (84%), with low risk of recurrent abscess and the need for subsequent surgery.¹¹⁹ The role of anti-TNF therapy in the setting of inflammatory mass (ie, phlegmon) with a concomitant abscess may be safe if the regimen includes initial intravenous antibiotics, followed by anti-TNF therapy, and continuation of antibiotics for 3 months total duration. With 2-year follow-up, it has been shown that most patients remain on anti-TNF therapy and do not require surgery.¹²⁰ Following successful resolution of an abscess with PD, consideration should be given to performing a drain study to exclude a concomitant fistula before drain removal, although this practice has been largely anecdotal with no large-scale trials to evaluate its utility.

Although centers comparing surgical versus medical management of these abscesses report similar

outcomes,^{113,121,122} efforts should be made to avoid surgery as the first line of treatment. When medical management is not successful, operative intervention with resection should be considered instead of operative drainage alone, the latter of which has been associated with higher rates of enterocutaneous fistula and reoperation.¹²³ Overall, surgeons must make every effort to avoid overaggressive resection, because unnecessary removal of bowel can lead to immediate or future development of short-bowel syndrome.¹⁰⁸

3. Patients with enteric fistulas and symptoms or signs of localized or systemic sepsis that persist despite appropriate medical therapy should be considered for surgery. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.

Fistulas originating in diseased bowel and secondarily involving other intra-abdominal organs or the skin are not usually associated with localized or systemic sepsis. If sepsis is present, the patient should be treated with appropriate resuscitation and parenteral antibiotics, and imaging studies should be performed to identify a source. If a concomitant abscess is identified,¹²⁴ it should be drained when feasible. Regardless of whether an abscess is present or not, persistent sepsis usually warrants excision of the diseased bowel.¹²⁵ Management of the affected organs is based on differentiating diseased bowel from other loops of bowel or organs that are secondarily involved and may or may not require resection. In general, diseased bowel is resected, whereas noninflamed bowel can be primarily closed,^{126–128} and other internal organs (eg, bladder, vagina) are primarily closed or left to heal by secondary intention.^{129,130} Surgery may not be required in the absence of symptoms, because the presence of an internal fistula does not necessarily mandate surgery, especially in the absence of malabsorption, diarrhea, or recurrent infections (ie, enterovesical fistulas).¹³¹

Hemorrhage

1. Stable patients with significant GI hemorrhage may be evaluated and treated by endoscopic and/or interventional radiological techniques. Unstable patients should typically undergo operative exploration. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.

Acute lower GI hemorrhage is an unusual complication of Crohn's disease, occurring in 0.9% to 6% of patients.^{132–137} Significant lower GI bleeding in patients with Crohn's disease is more often secondary to severe inflammation and, unlike other sources of massive lower GI bleeding (ie, diverticula, arteriovenous malformation), rarely necessitates the diagnostic evaluation required in a bleed of unknown origin. Patients with more significant bleeding should undergo concomitant resuscitation along with appropriate

diagnostic measures. Stable patients can undergo CT imaging with prominent contrast arterial phase, endoscopy, and/or mesenteric arteriography.¹³⁸ Although not specific to Crohn's disease, CT angiography has reported sensitivity rates for acute GI bleeding of up to 89% and specificity rates of up to 92%.^{139,140} In addition, the latter 2 modalities also offer the benefit of allowing concomitant therapeutic intervention. The likelihood of identifying a precise bleeding source is somewhat limited, because spontaneous cessation of bleeding occurs in nearly one-half of all patients with lower GI bleeding.^{133,137} However, patients remain at risk for recurrent bleeding in nearly 40% of cases.¹³³ More specific to patients with Crohn's disease, the probability of rebleeding in those successfully managed with non-operative measures may be reduced with anti-TNF agent therapy.¹³³ Therefore, treating the underlying inflammation with the broad range of Crohn's disease medications will often help minimize the risk of significant recurrent lower GI bleeding.

In general, surgical treatment is recommended in patients with life-threatening bleeding, persistent hemodynamic instability, or recurrent significant GI bleeding. If the source of bleeding has been localized by the use of preoperative or intraoperative modalities, a targeted resection with adherence to accepted surgical tenets is recommended. In the case of Crohn's colitis, a total colectomy is recommended for patients who have persistent hemodynamic instability or persistent, serious bleeding that cannot be localized to 1 segment of the colon. The decision regarding creating an anastomosis versus an end or a diverting ileostomy should again depend on factors such as the patient's degree of immunosuppression, age, hemodynamic stability, continence, and the presence of active Crohn's disease in the rectum or perianal area.

Growth Retardation

1. Prepubertal patients with significant growth retardation despite appropriate medical therapy should be considered for surgery. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.

Crohn's disease manifests itself during childhood or adolescence in up to 20% of patients,¹⁴¹ and the potential for linear growth impairment as a complication of chronic intestinal inflammation is unique to this population. Growth retardation exists in ~10% to 40% of children with Crohn's disease, may present at the time of diagnosis or occur after diagnosis, and potentially causes compromised height at maturity.^{142–144} The treatment of young patients may include medical, nutritional, and surgical therapy that aims to decrease mucosal inflammation and alleviate symptoms, all while improving growth rates and promoting normal pubertal development. Resection of localized disease maximizes "catch-up" growth, and is warranted early in the disease process in otherwise

treatment-resistant, prepubertal children because they usually will achieve catch-up growth within the ensuing 6 months.¹⁴⁴ However, early surgical intervention for growth retardation is only indicated in cases with clearly localized disease because of the risk for recrudescence with more diffuse involvement, as well as owing to the association between malnourishment and further developmental difficulties with extensive resections.

Neoplasia

1. Patients with long-standing Crohn's disease of the ileocolic region or colon should have endoscopic surveillance of the large bowel. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.

The incidence of colorectal carcinoma in a patient with Crohn's disease is 0.5/1000 person-years duration, which is a 2- to 3-fold increase in risk compared with the incidence in an age-matched general population.¹⁴⁵ The impact of age at disease onset, disease duration, extent of disease, primary sclerosing cholangitis, and a positive family history of sporadic colorectal cancer are debated, but may be associated with an increased risk of colorectal cancer.^{146–150} Furthermore, postinflammatory pseudopolyps are thought to increase the risk of colorectal cancer, and inflammation is believed to be a risk factor for progression to colorectal neoplasia.¹⁵¹

All patients, regardless of the extent of disease at initial diagnosis, should undergo a screening colonoscopy no later than 8 years after the onset of symptoms with multiple biopsy specimens obtained throughout the entire colon to assess the true microscopic extent of inflammation.^{151,152} It has also been recommended that patients with more than one-third of the colon and rectum affected by Crohn's disease should be surveyed beginning within 1 to 2 years of the initial screening colonoscopy,^{151–153} and continued at 1- to 3-year intervals, despite a lack of high-quality evidence to support this recommendation.^{151,152} Surveillance colonoscopy should be ideally performed when the colonic disease is in remission. Although no prospective studies have determined the optimal number of biopsy specimens that should be acquired to reliably detect dysplasia, a minimum of 33 specimens has been recommended in patients with pancolitis.^{151,152,154} Chromoendoscopy with targeted biopsies is an alternative to random biopsies for endoscopists who have expertise with this technique, because limited reports have shown a higher sensitivity for detecting dysplasia by chromoendoscopy than with traditional white light endoscopy.¹⁵⁵ Conversely, high-definition colonoscopy has been reported to improve targeted detection of dysplastic lesions during surveillance colonoscopy.¹⁵⁶

Patients with concomitant primary sclerosing cholangitis (PSC) should undergo a screening colonoscopy when PSC is initially diagnosed, followed by annual surveillance

colonoscopy with biopsies because of the reported increased risk of malignancy.^{151,152,157,158} The presence of concomitant PSC may also be a consideration for prophylactic total proctocolectomy, although the relative risk of malignancy for PSC patients and Crohn's is somewhat controversial, and appears to be less than that of PSC in patients with ulcerative colitis.^{159–161} Of note, when malignancy does occur in the setting of PSC and Crohn's disease, it has a predilection for occurring proximal to the splenic flexure.¹⁵⁹ In addition, patients with a history of colorectal cancer in a first-degree relative, ongoing active endoscopic or histological inflammation, or anatomic abnormalities such as multiple inflammatory pseudopolyps or a stricture may also benefit from more frequent surveillance examinations or prophylactic bowel resection.

2. Total proctocolectomy should be considered for patients with carcinoma, a nonadenoma-like dysplasia-associated lesion or mass (DALM), high-grade dysplasia, or multifocal low-grade dysplasia of the colon or rectum. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.

Dysplasia is regarded to be the best indicator of colorectal cancer risk in Crohn's disease, although the diagnosis of dysplasia in mucosal biopsy specimens is associated with a high level of interobserver variability, especially when diagnosing indeterminate and low-grade dysplasia.^{162,163} Therefore, pathologists with expertise in GI diseases should normally confirm all cases diagnosed as indefinite, low-grade or high-grade dysplasia before a management plan is implemented, when possible.¹⁶⁴

Unfortunately high-grade evidence detailing the natural history of dysplasia is still lacking, which ultimately affects surveillance and treatment recommendations. Although dysplasia is present in many patients with adenocarcinoma,¹⁶⁵ colorectal cancer can also be the index neoplastic lesion^{146,148} or develop in the setting of low-grade dysplasia without intervening high-grade dysplasia.¹⁴⁶ Furthermore, indefinite dysplasia and unifocal low-grade dysplasia are associated with a higher incidence of synchronous and metachronous advanced neoplasms,^{166,167} although the magnitude and significance of that risk are still unknown.

A non-adenoma dysplasia-associated lesion or mass (DALM) is a raised, endoscopically visible lesion that harbors dysplastic mucosa and does not demonstrate the endoscopic feature of a sporadic adenoma. Nonadenomatous DALMs are an indication for resection because the incidence of associated cancer approaches 40%.¹⁶⁶ Conversely, sporadic adenomas can develop in a background of Crohn's disease, and they can be safely managed by polypectomy and continued surveillance, especially if flat dysplasia is absent both from the mucosa immediately surrounding the polyp, as well as elsewhere in the colon.^{148,151,168}

The extent of resection is influenced by the site of neoplasm and recognition that multifocal dysplasia is ultimately found in more than one-third of specimens from patients undergoing colectomy for low- or high-grade dysplasia detected during diagnostic or surveillance colonoscopy.¹⁶⁶ Accordingly, resection of all large bowel (ie, total proctocolectomy) has been advocated over segmental resection.^{166,167} Further support for such an approach stems from observing that ~14% to 40% of the patients who undergo segmental resection for cancer of the large intestine develop metachronous colorectal cancers.^{166,167}

3. Suspicious lesions (ie, mass, ulcer) identified in patients with Crohn's disease should typically be biopsied, especially when considering a small-bowel strictureplasty. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.

Suspicious lesions anywhere along the GI tract in patients with Crohn's disease warrant further evaluation. This requires a diagnosis confirmed by pathology when possible. The incidence of small-bowel carcinoma in a patient with Crohn's disease is 0.3/1000 person-years duration, and this represents an 18.75-fold increase compared with an age-matched standard population.¹⁴⁵ Although uncommon, a few cases of adenocarcinoma arising at or near a previous strictureplasty site have been reported.¹⁶⁹⁻¹⁷² Biopsy of any suspicious mucosal lesions or suspicious ulcerations at the time of strictureplasty is accordingly suggested. Failure to do so may result in leaving malignant or premalignant lesions in situ that would otherwise warrant resection.

SITE-SPECIFIC OPERATIONS

Terminal Ileum, Ileocolon, and Upper GI Tract

1. Patients who require surgery for disease of the jejunum, proximal ileum, terminal ileum, or ileocolon without existing or impending short-bowel syndrome should usually undergo resection of the affected bowel. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.

Resection continues to be the most commonly performed operation for symptomatic inflammatory or penetrating disease affecting the small bowel and proximal colon, especially when existing or impending short-bowel syndrome is unlikely and medical management fails. Limited (2-cm) macroscopic disease-free resection margins are adequate, because this conserves bowel length and is not associated with an increased risk of disease recurrence.¹⁷³ Certain factors are associated with an increased risk for infectious complications following surgery. In addition to steroids and immunomodulators, other associated factors include a concomitant abscess,^{46,129} or fistula,¹²⁹ malnutrition (>10% weight loss within past 3 months),⁴⁶ anemia

(<10 g/dL),¹⁷⁴ low serum albumin (<3.0 mg/dL),¹²⁹ penetrating disease,¹⁷⁵ prolonged operative times (>180 minutes),¹⁷⁵ and urgent surgery.¹⁷⁶ When one or more of these factors is present, surgery should be postponed, when possible, until the patient can be safely optimized. Alternatively, proximal fecal diversion may be more frequently indicated, although resolution of distal inflammation following diversion is more variable in the setting of IBD.

2. Select patients with symptomatic disease of the stomach or duodenum should be considered for endoscopic dilation, bypass, or strictureplasty of the affected area. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.

Gastroduodenal disease occurs in 0.5% to 4% of all patients with Crohn's disease. Obstructive symptoms arising from single, short, moderately thick strictures of the duodenum can be successfully treated with endoscopic dilation as an initial procedure.¹⁷⁷⁻¹⁷⁹ Symptomatic Crohn's disease of the stomach or duodenum requiring operative treatment is often best managed by bypass (ie, gastrojejunostomy, duodenojejunostomy) or strictureplasty, rather than resection. Proton pump inhibitor therapy has supplanted the need for concomitant vagotomy in patients at risk of sustaining marginal or duodenal ulceration following bypass.¹⁸⁰ Because the operation has become more simplified, a laparoscopic approach may be considered because of its decreased risk for operative morbidity and equivalent recurrence rates.¹⁸⁰ Nonperforated, nonphlegmonous stenotic lesions in this region can alternatively be managed by strictureplasty.^{92,180-183} Finally, solitary strictures of the second and third portions of the duodenum are often best suited for strictureplasty.¹⁸³

Colon

1. The procedure of choice for emergency surgery in Crohn's colitis is a total abdominal colectomy with end ileostomy. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.

Subtotal or total colectomy with construction of an end ileostomy and Hartmann closure of the distal bowel or creation of a mucous fistula is a safe procedure in patients with severe or fulminant Crohn's colitis who require surgery.¹⁸⁴ This approach removes most of the inflamed intestine with a relatively simple operation that avoids a pelvic dissection and the risks of an anastomosis in this setting. Extrafascial placement of the closed rectosigmoid stump may be associated with fewer pelvic septic complications than an intraperitoneal position, although it may be technically difficult to achieve.¹⁸⁵⁻¹⁸⁸ Abdominal drains as well as transanal drainage of the distal stump may further decrease the risk of pelvic sepsis.¹⁸⁹

2. **Patients with colonic disease and rectal sparing that require elective surgery may undergo segmental colectomy for single-segment disease or total colectomy for more extensive disease. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.**

Outside the aforementioned concerns regarding malignancy, symptomatic disease of the colon with rectal sparing can be managed by removal of only the diseased segment or by total abdominal colectomy. The operations are comparable with respect to operative complications, need for a permanent stoma, and risk of recurrence; however, segmental resection is associated with surgical recurrence that occurs a median of 4.4 years earlier.¹⁹⁰ Total colectomy with ileoproctostomy is preferable if 2 or more colonic segments are affected, because this subgroup has a higher recurrence rate when segmental resections are performed.¹⁹⁰

3. **Patients who require surgery for disease of the rectum typically undergo total proctocolectomy, or proctectomy with creation of a stoma. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.**

Refractory proctitis generally warrants total proctocolectomy with a permanent ileostomy owing to a frequent association with colonic involvement; however, proctectomy alone with creation of a colostomy can be considered if the colon is spared.^{191,192} When proctectomy is warranted, the entire rectum should be resected because carcinoma has been described in patients with even a short Hartmann remnant.¹⁹³ Furthermore, an intersphincteric dissection and primary closure of the perineal wound is preferred, because it is associated with fewer wound complications and less morbidity compared with a traditional abdominoperineal resection.¹⁹⁴

4. **Total proctocolectomy with IPAA is not generally recommended for patients with known Crohn's disease. Grade of Recommendation: Strong recommendation based on low- or very low-quality evidence, 1C.**

Two centers have suggested that a restorative proctocolectomy with an IPAA can be considered in Crohn's disease in highly selected patients if the small bowel is unaffected and there is no history of perianal disease.^{195,196} In appropriately selected patients, the 5-year pouch survival rate is at least 85%.^{195,196} However, this must be balanced against the high risk of further complications, pouch loss, poor function, and the need for further immunosuppressive therapy. Therefore, although there are advocates for restorative proctocolectomy in specific cohorts,¹⁹⁷ this should be approached with extreme caution and, at the present time, is not widely recommended.^{198–200}

TECHNICAL CONSIDERATIONS

1. **Patients who require resection of the affected bowel should be offered a laparoscopic approach where appropriate experience and expertise are available. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.**

A laparoscopic approach is preferred to an open operation where appropriate surgeon expertise is available. For small-bowel and ileocolic Crohn's disease, laparoscopy has been associated with earlier return of bowel function, shorter length of stay, and fewer postoperative complications.^{201–203} Long-term outcomes, including clinical recurrence, do not appear to be compromised with a laparoscopic approach.^{204–206} Furthermore, a laparoscopic approach can also be safely used when managing complex (eg, penetrating, recurrent) disease, when appropriate surgical experience and expertise is available.^{207–211} Single-port laparoscopy may be also considered by those facile with this approach, but its advantages over multiport laparoscopy have not been consistently described.^{212–216}

When performing a total abdominal colectomy for Crohn's colitis in the elective setting, laparoscopy is associated with decreased perioperative morbidity and improved short-term outcomes (ie, length of stay, return of bowel function) in comparison with an open approach.^{209,217,218} The risk of wound infection and intra-abdominal abscess are significantly lower with a laparoscopic total abdominal colectomy compared with open surgery in the patient with nontoxic Crohn's disease.²¹⁹ Even in an emergent setting, a laparoscopic approach for severe acute Crohn's colitis that failed adequate medical management is associated with a shorter length of stay²²⁰ and fewer minor morbidities.²²¹

2. **Patients who are candidates for primary anastomosis following resection of diseased intestine can have the anastomosis constructed as deemed most appropriate by the surgeon. Grade of Recommendation: Strong recommendation based on moderate-quality evidence, 1B.**

Despite earlier reports, the anastomosis following resection in the setting of Crohn's disease can be configured in any manner (eg, end-to-end, side-to-side, end-to-side) without increasing the risk for endoscopic, clinical, or operative recurrence.²²² A side-to-side (ie, functional end-to-end) anastomosis constructed with staples or sutures does not reduce the likelihood for anastomotic leak compared with other configurations and techniques.^{223,224} However, it may reduce the overall rate of postoperative complications.²²² Although a stapled side-to-side anastomosis has been shown to reduce reoperation rates,²²² the use of the mechanical stapler may be contraindicated when the bowel wall is thickened as seen with bowel proximal to a chronic obstruction.

APPENDIX A

Contributing Members of the ASCRS Clinical Practice Guideline Committee

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REFERENCES

- Louis E. Epidemiology of the transition from early to late Crohn's disease. *Dig Dis*. 2012;30:376–379.
- Jostins L, Ripke S, Weersma RK, et al. Host-microbe interactions have shaped the genetic architecture of inflammatory bowel disease. *Nature*. 2012;491:119–124.
- Rutgeerts P, Goboos K, Peeters M, et al. Effect of fecal diversion on recurrence of Crohn's disease in the neoterminal ileum. *Lancet*. 1991;338:771–774.
- Swidsinski A, Loening-Baucke V, Herber A. Mucosal flora in Crohn's disease and ulcerative colitis - an overview. *J Physiol Pharmacol*. 2009;60 (suppl 6):61–71.
- Gersemann M, Wehkamp J, Stange EF. Innate immune dysfunction in inflammatory bowel disease. *J Intern Med*. 2012;271:421–428.
- Bouguen G, Peyrin-Biroulet L. Surgery for adult Crohn's disease: what is the actual risk? *Gut*. 2011;60:1178–1181.
- Bernell O, Lapidus A, Hellers G. Risk factors for surgery and postoperative recurrence in Crohn's disease. *Ann Surg*. 2000;231:38–45.
- Solberg IC, Vatn MH, Hoie O, et al; IBSEN Study Group. Clinical course in Crohn's disease: results of a Norwegian population-based ten-year follow-up study. *Clin Gastroenterol Hepatol*. 2007;5:1430–1438.
- Ramadas AV, Gunesh S, Thomas GA, Williams GT, Hawthorne AB. Natural history of Crohn's disease in a population-based cohort from Cardiff (1986-2003): a study of changes in medical treatment and surgical resection rates. *Gut*. 2010;59:1200–1206.
- Peyrin-Biroulet L, Harmsen WS, Tremaine WJ, Zinsmeister AR, Sandborn WJ, Loftus EV Jr. Surgery in a population-based cohort of Crohn's disease from Olmsted County, Minnesota (1970-2004). *Am J Gastroenterol*. 2012;107:1693–1701.
- Strong SA, Koltun WA, Hyman NH, Buie WD; Standards Practice Task Force of The American Society of Colon and Rectal Surgeons. Practice parameters for the surgical management of Crohn's disease. *Dis Colon Rectum*. 2007;50:1735–1746.
- Steele SR, Kumar R, Feingold DL, Rafferty JL, Buie WD; Standards Practice Task Force of the American Society of Colon and Rectal Surgeons. Practice parameters for the management of perianal abscess and fistula-in-ano. *Dis Colon Rectum*. 2011;54:1465–1474.
- Guyatt G, Gutterman D, Baumann MH, et al. Grading strength of recommendations and quality of evidence in clinical guidelines: report from an american college of chest physicians task force. *Chest*. 2006;129:174–181.
- Lichtenstein GR, Hanauer SB, Sandborn WJ; Practice Parameters Committee of American College of Gastroenterology. Management of Crohn's disease in adults. *Am J Gastroenterol*. 2009;104:465–483.
- Azzopardi N, Ellul P. Risk factors for osteoporosis in Crohn's disease: infliximab, corticosteroids, body mass index, and age of onset. *Inflamm Bowel Dis*. 2013;19:1173–1178.
- Hudesman D, Lichtiger S, Sands B. Risk of extraintestinal solid cancer with anti-TNF therapy in adults with inflammatory bowel disease: review of the literature. *Inflamm Bowel Dis*. 2013;19:644–649.
- Dignass A, Van Assche G, Lindsay JO, et al; European Crohn's and Colitis Organisation (ECCO). The second European evidence-based Consensus on the diagnosis and management of Crohn's disease: current management. *J Crohns Colitis*. 2010;4:28–62.
- Yang ZP, Hong L, Wu Q, Wu KC, Fan DM. Preoperative infliximab use and postoperative complications in Crohn's disease: a systematic review and meta-analysis. *Int J Surg*. 2014;12:224–230.
- Syed A, Cross RK, Flasar MH. Anti-tumor necrosis factor therapy is associated with infections after abdominal surgery in Crohn's disease patients. *Am J Gastroenterol*. 2013;108:583–593.
- Appau KA, Fazio VW, Shen B, et al. Use of infliximab within 3 months of ileocolonic resection is associated with adverse postoperative outcomes in Crohn's patients. *J Gastrointest Surg*. 2008;12:1738–1744.
- El-Hussuna A, Theede K, Olaison G. Increased risk of post-operative complications in patients with Crohn's disease treated with anti-tumour necrosis factor α agents - a systematic review. *Dan Med J*. 2014;61:A4975.
- Uchino M, Ikeuchi H, Matsuoka H, et al. Risk factors for surgical site infection and association with infliximab administration during surgery for Crohn's disease. *Dis Colon Rectum*. 2013;56:1156–1165.
- Myrelid P, Marti-Gallostra M, Ashraf S, et al. Complications in surgery for Crohn's disease after preoperative antitumour necrosis factor therapy. *Br J Surg*. 2014;101:539–545.
- Krane MK, Allaix ME, Zoccali M, et al. Preoperative infliximab therapy does not increase morbidity and mortality after laparoscopic resection for inflammatory bowel disease. *Dis Colon Rectum*. 2013;56:449–457.
- Rosenfeld G, Qian H, Bressler B. The risks of post-operative complications following pre-operative infliximab therapy for Crohn's disease in patients undergoing abdominal surgery: a systematic review and meta-analysis. *J Crohns Colitis*. 2013;7:868–877.
- Nørgård BM, Nielsen J, Qvist N, Gradel KO, de Muckadell OB, Kjeldsen J. Pre-operative use of anti-TNF- α agents and the risk of post-operative complications in patients with Crohn's disease—a nationwide cohort study. *Aliment Pharmacol Ther*. 2013;37:214–224.
- El-Hussuna A, Andersen J, Bisgaard T, et al. Biologic treatment or immunomodulation is not associated with postoperative anastomotic complications in abdominal surgery for Crohn's disease. *Scand J Gastroenterol*. 2012;47:662–668.
- Kasperek MS, Bruckmeier A, Beigel F, et al. Infliximab does not affect postoperative complication rates in Crohn's patients undergoing abdominal surgery. *Inflamm Bowel Dis*. 2012;18:1207–1213.
- Regueiro M, El-Hachem S, Kip KE, et al. Postoperative infliximab is not associated with an increase in adverse events in Crohn's disease. *Dig Dis Sci*. 2011;56:3610–3615.

30. Canedo J, Lee SH, Pinto R, Murad-Regadas S, Rosen L, Wexner SD. Surgical resection in Crohn's disease: is immunosuppressive medication associated with higher postoperative infection rates? *Colorectal Dis.* 2011;13:1294–1298.
31. Nasir BS, Dozois EJ, Cima RR, et al. Perioperative anti-tumor necrosis factor therapy does not increase the rate of early postoperative complications in Crohn's disease. *J Gastrointest Surg.* 2010;14:1859–1865.
32. 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.
33. Colombel JF, Loftus EV Jr, Tremaine WJ, et al. 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.
34. Marchal L, D'Haens G, Van Assche G, et al. The risk of postoperative complications associated with infliximab therapy for Crohn's disease: a controlled cohort study. *Aliment Pharmacol Ther.* 2004;19:749–754.
35. Kopylov U, Ben-Horin S, Zmora O, Eliakim R, Katz LH. Anti-tumor necrosis factor and postoperative complications in Crohn's disease: systematic review and meta-analysis. *Inflamm Bowel Dis.* 2012;18:2404–2413.
36. Narula N, Charleton D, Marshall JK. Meta-analysis: peri-operative anti-TNF α treatment and post-operative complications in patients with inflammatory bowel disease. *Aliment Pharmacol Ther.* 2013;37:1057–1064.
37. Billioud V, Ford AC, Tedesco ED, Colombel JF, Roblin X, Peyrin-Biroulet L. Preoperative use of anti-TNF therapy and postoperative complications in inflammatory bowel diseases: a meta-analysis. *J Crohns Colitis.* 2013;7:853–867.
38. Randall J, Singh B, Warren BF, Travis SP, Mortensen NJ, George BD. Delayed surgery for acute severe colitis is associated with increased risk of postoperative complications. *Br J Surg.* 2010;97:404–409.
39. Ahmed Ali U, Martin ST, Rao AD, Kiran RP. Impact of preoperative immunosuppressive agents on postoperative outcomes in Crohn's disease. *Dis Colon Rectum.* 2014;57:663–674.
40. El-Hussuna A, Krag A, Olaison G, Bendtsen F, Gluud LL. The effect of anti-tumor necrosis factor alpha agents on postoperative anastomotic complications in Crohn's disease: a systematic review. *Dis Colon Rectum.* 2013;56:1423–1433.
41. Lau C, Dubinsky M, Melmed G, et al. The impact of preoperative serum anti-TNF α therapy levels on early postoperative outcomes in inflammatory bowel disease surgery. *Ann Surg.* 2015;261:487–496.
42. Post S, Betzler M, von Ditfurth B, Schürmann G, Küppers P, Herfarth C. Risks of intestinal anastomoses in Crohn's disease. *Ann Surg.* 1991;213:37–42.
43. Yamamoto T, Allan RN, Keighley MR. Risk factors for intra-abdominal sepsis after surgery in Crohn's disease. *Dis Colon Rectum.* 2000;43:1141–1145.
44. Aberra FN, Lewis JD, Hass D, Rombeau JL, Osborne B, Lichtenstein GR. Corticosteroids and immunomodulators: postoperative infectious complication risk in inflammatory bowel disease patients. *Gastroenterology.* 2003;125:320–327.
45. Alves A, Panis Y, Bouhnik Y, Pocard M, Vicaut E, Valleur P. Risk factors for intra-abdominal septic complications after a first ileocecal resection for Crohn's disease: a multivariate analysis in 161 consecutive patients. *Dis Colon Rectum.* 2007;50:331–336.
46. Tzivanakis A, Singh JC, Guy RJ, Travis SP, Mortensen NJ, George BD. Influence of risk factors on the safety of ileocolic anastomosis in Crohn's disease surgery. *Dis Colon Rectum.* 2012;55:558–562.
47. Panaccione R, Rutgeerts P, Sandborn WJ, Feagan B, Schreiber S, Ghosh S. Review article: treatment algorithms to maximize remission and minimize corticosteroid dependence in patients with inflammatory bowel disease. *Aliment Pharmacol Ther.* 2008;28:674–688.
48. Burger D, Travis S. Conventional medical management of inflammatory bowel disease. *Gastroenterology.* 2011;140:1827–1837.e2.
49. Serradori T, Germain A, Scherrer ML, et al. The effect of immune therapy on surgical site infection following Crohn's Disease resection. *Br J Surg.* 2013;100:1089–1093.
50. Treem WR, Cohen J, Davis PM, Justinich CJ, Hyams JS. Cyclosporine for the treatment of fulminant ulcerative colitis in children: immediate response, long-term results, and impact on surgery. *Dis Colon Rectum.* 1995;38:474–479.
51. Nelson R, Liao C, Fichera A, Rubin DT, Pekow J. Rescue therapy with cyclosporine or infliximab is not associated with an increased risk for postoperative complications in patients hospitalized for severe steroid-refractory ulcerative colitis. *Inflamm Bowel Dis.* 2014;20:14–20.
52. Lazarev M, Present DH, Lichtiger S, et al. The effect of intravenous cyclosporine on rates of colonic surgery in hospitalized patients with severe Crohn's colitis. *J Clin Gastroenterol.* 2012;46:764–767.
53. Brown SR, Haboubi N, Hampton J, George B, Travis SP; ACPGBI. The management of acute severe colitis: ACPGBI position statement. *Colorectal Dis.* 2008;10(suppl 3):8–29.
54. Hanauer SB. Inflammatory bowel disease. *N Engl J Med.* 1996;334:841–848.
55. Jones JH, Chapman M. Definition of megacolon in colitis. *Gut.* 1969;10:562–564.
56. Sheth SG, LaMont JT. Toxic megacolon. *Lancet.* 1998;351:509–513.
57. Autenrieth DM, Baumgart DC. Toxic megacolon. *Inflamm Bowel Dis.* 2012;18:584–591.
58. Criscuoli V, Casà A, Orlando A, et al. Severe acute colitis associated with CMV: a prevalence study. *Dig Liver Dis.* 2004;36:818–820.
59. Lewis JD. The utility of biomarkers in the diagnosis and therapy of inflammatory bowel disease. *Gastroenterology.* 2011;140:1817–1826.e2.
60. Greenstein AJ, Sachar DB, Gibas A, et al. Outcome of toxic dilatation in ulcerative and Crohn's colitis. *J Clin Gastroenterol.* 1985;7:137–143.
61. Sands BE, Tremaine WJ, Sandborn WJ, et al. Infliximab in the treatment of severe, steroid-refractory ulcerative colitis: a pilot study. *Inflamm Bowel Dis.* 2001;7:83–88.
62. Lichtiger S, Present DH, Kornbluth A, et al. Cyclosporine in severe ulcerative colitis refractory to steroid therapy. *N Engl J Med.* 1994;330:1841–1845.

63. D'Haens G, Lemmens L, Geboes K, et al. Intravenous cyclosporine versus intravenous corticosteroids as single therapy for severe attacks of ulcerative colitis. *Gastroenterology*. 2001;120:1323–1329.
64. D'Haens GR, Panaccione R, Higgins PD, et al. The London Position Statement of the World Congress of Gastroenterology on Biological Therapy for IBD with the European Crohn's and Colitis Organization: when to start, when to stop, which drug to choose, and how to predict response? *Am J Gastroenterol*. 2011;106:199–212; quiz 213.
65. Sjöberg M, Magnuson A, Björk J, et al; Swedish Organization for the Study of Inflammatory Bowel Disease (SOIBD). Influximab as rescue therapy in hospitalised patients with steroid-refractory acute ulcerative colitis: a long-term follow-up of 211 Swedish patients. *Aliment Pharmacol Ther*. 2013;38:377–387.
66. Gibson DJ, Heetun ZS, Redmond CE, et al. An accelerated infliximab induction regimen reduces the need for early colectomy in patients with acute severe ulcerative colitis. *Clin Gastroenterol Hepatol*. 2015;13:330–335.
67. Protic M, Seibold F, Schoepfer A, et al. The effectiveness and safety of rescue treatments in 108 patients with steroid-refractory ulcerative colitis with sequential rescue therapies in a subgroup of patients. *J Crohns Colitis*. 2014;8:1427–1437.
68. Rieder F, Fiocchi C. Intestinal fibrosis in IBD—a dynamic, multifactorial process. *Nat Rev Gastroenterol Hepatol*. 2009;6:228–235.
69. Allen BC, Leyendecker JR. MR enterography for assessment and management of small bowel Crohn disease. *Radiol Clin North Am*. 2014;52:799–810.
70. Seastedt KP, Trencheva K, Michelassi F, et al. Accuracy of CT enterography and magnetic resonance enterography imaging to detect lesions preoperatively in patients undergoing surgery for Crohn's disease. *Dis Colon Rectum*. 2014;57:1364–1370.
71. Panés J, Bouzas R, Chaparro M, et al. Systematic review: the use of ultrasonography, computed tomography and magnetic resonance imaging for the diagnosis, assessment of activity and abdominal complications of Crohn's disease. *Aliment Pharmacol Ther*. 2011;34:125–145.
72. Hagel AF, Hahn A, Dauth W, et al. Outcome and complications of endoscopic balloon dilatations in various types of ileocaecal and colonic stenosis in patients with Crohn's disease. *Surg Endosc*. 2014;28:2966–2972.
73. Vrabie R, Irwin GL, Friedel D. Endoscopic management of inflammatory bowel disease strictures. *World J Gastrointest Endosc*. 2012;4:500–505.
74. Wibmer AG, Kroesen AJ, Gröne J, Buhr HJ, Ritz JP. Comparison of strictureplasty and endoscopic balloon dilatation for stricturing Crohn's disease—review of the literature. *Int J Colorectal Dis*. 2010;25:1149–1157.
75. Gustavsson A, Magnuson A, Blomberg B, Andersson M, Halfvarson J, Tysk C. Endoscopic dilation is an efficacious and safe treatment of intestinal strictures in Crohn's disease. *Aliment Pharmacol Ther*. 2012;36:151–158.
76. Hassan C, Zullo A, De Francesco V, et al. Systematic review: endoscopic dilatation in Crohn's disease. *Aliment Pharmacol Ther*. 2007;26:1457–1464.
77. Attar A, Maunoury V, Vahedi K, et al; GETAID. Safety and efficacy of extractible self-expandable metal stents in the treatment of Crohn's disease intestinal strictures: a prospective pilot study. *Inflamm Bowel Dis*. 2012;18:1849–1854.
78. Rejchrt S, Kopacova M, Brozik J, Bures J. Biodegradable stents for the treatment of benign stenoses of the small and large intestines. *Endoscopy*. 2011;43:911–917.
79. 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.
80. Swaminath A, Lichtiger S. Dilation of colonic strictures by intralesional injection of infliximab in patients with Crohn's colitis. *Inflamm Bowel Dis*. 2008;14:213–216.
81. Bellolio F, Cohen Z, MacRae HM, et al. Strictureplasty in selected Crohn's disease patients results in acceptable long-term outcome. *Dis Colon Rectum*. 2012;55:864–869.
82. Ambe R, Campbell L, Cagir B. A comprehensive review of strictureplasty techniques in Crohn's disease: types, indications, comparisons, and safety. *J Gastrointest Surg*. 2012;16:209–217.
83. Campbell L, Ambe R, Weaver J, Marcus SM, Cagir B. Comparison of conventional and nonconventional strictureplasties in Crohn's disease: a systematic review and meta-analysis. *Dis Colon Rectum*. 2012;55:714–726.
84. Yamamoto T, Fazio VW, Tekkis PP. Safety and efficacy of strictureplasty for Crohn's disease: a systematic review and meta-analysis. *Dis Colon Rectum*. 2007;50:1968–1986.
85. Dietz DW, Laureti S, Strong SA, et al. Safety and longterm efficacy of strictureplasty in 314 patients with obstructing small bowel Crohn's disease. *J Am Coll Surg*. 2001;192:330–337.
86. Sampietro GM, Cristaldi M, Porretta T, Montecamozzo G, Danelli P, Taschieri AM. Early perioperative results and surgical recurrence after strictureplasty and miniresection for complicated Crohn's disease. *Dig Surg*. 2000;17:261–267.
87. Fazio VW, Tjandra JJ, Lavery IC, Church JM, Milsom JW, Oakley JR. Long-term follow-up of strictureplasty in Crohn's disease. *Dis Colon Rectum*. 1993;36:355–361.
88. Hurst RD, Michelassi F. Strictureplasty for Crohn's disease: techniques and long-term results. *World J Surg*. 1998;22:359–363.
89. Yamamoto T, Allan RN, Keighley MR. Strategy for surgical management of ileocolonic anastomotic recurrence in Crohn's disease. *World J Surg*. 1999;23:1055–1060.
90. Yamamoto T, Bain IM, Connolly AB, Allan RN, Keighley MR. Outcome of strictureplasty for duodenal Crohn's disease. *Br J Surg*. 1999;86:259–262.
91. Yamamoto T, Keighley MR. Factors affecting the incidence of postoperative septic complications and recurrence after strictureplasty for jejunoileal Crohn's disease. *Am J Surg*. 1999;178:240–245.
92. Reese GE, Purkayastha S, Tilney HS, von Roon A, Yamamoto T, Tekkis PP. Strictureplasty vs resection in small bowel Crohn's disease: an evaluation of short-term outcomes and recurrence. *Colorectal Dis*. 2007;9:686–694.
93. Ozuner G, Fazio VW, Lavery IC, Milsom JW, Strong SA. Reoperative rates for Crohn's disease following strictureplasty: long-term analysis. *Dis Colon Rectum*. 1996;39:1199–1203.
94. Hesham W, Kann BR. Strictureplasty. *Clin Colon Rectal Surg*. 2013;26:80–83.
95. Poggioli G, Stocchi L, Laureti S, et al. Conservative surgical management of terminal ileitis: side-to-side enterocolic anastomosis. *Dis Colon Rectum*. 1997;40:234–237.
96. Sampietro GM, Cristaldi M, Maconi G, et al. A prospective, longitudinal study of nonconventional strictureplasty in Crohn's disease. *J Am Coll Surg*. 2004;199:8–20.

97. Yamazaki Y, Ribeiro MB, Sachar DB, Aufses AH, Greenstein AJ. Malignant strictures in Crohn's disease. *Am J Gastroenterol*. 1991;86:882–885.
98. Kontzoglou K, Moulakakis KG, Alexiou D, et al. The role of liquid-based cytology in the investigation of colorectal lesions: a cytohistopathological correlation and evaluation of diagnostic accuracy. *Langenbecks Arch Surg*. 2007;392:189–195.
99. Broering DC, Eisenberger CF, Koch A, et al. Strictureplasty for large bowel stenosis in Crohn's disease: quality of life after surgical therapy. *Int J Colorectal Dis*. 2001;16:81–87.
100. Werbin N, Haddad R, Greenberg R, Karin E, Skornick Y. Free perforation in Crohn's disease. *Isr Med Assoc J*. 2003;5:175–177.
101. Veroux M, Angriman I, Ruffolo C, et al. A rare surgical complication of Crohn's disease: free peritoneal perforation. *Minerva Chir*. 2003;58:351–354.
102. Greenstein AJ, Sachar DB, Mann D, Lachman P, Heimann T, Aufses AH Jr. Spontaneous free perforation and perforated abscess in 30 patients with Crohn's disease. *Ann Surg*. 1987;205:72–76.
103. Hyman NH, Cataldo P, Osler T. Urgent subtotal colectomy for severe inflammatory bowel disease. *Dis Colon Rectum*. 2005;48:70–73.
104. Hyman N, Manchester TL, Osler T, Burns B, Cataldo PA. Anastomotic leaks after intestinal anastomosis: it's later than you think. *Ann Surg*. 2007;245:254–258.
105. Feagins LA, Holubar SD, Kane SV, Spechler SJ. Current strategies in the management of intra-abdominal abscesses in Crohn's disease. *Clin Gastroenterol Hepatol*. 2011;9:842–850.
106. Casola G, vanSonnenberg E, Neff CC, Saba RM, Withers C, Emarine CW. Abscesses in Crohn disease: percutaneous drainage. *Radiology*. 1987;163:19–22.
107. Sahai A, Bélair M, Gianfelice D, Coté S, Gratton J, Lahaie R. Percutaneous drainage of intra-abdominal abscesses in Crohn's disease: short and long-term outcome. *Am J Gastroenterol*. 1997;92:275–278.
108. Jawhari A, Kamm MA, Ong C, Forbes A, Bartram CI, Hawley PR. Intra-abdominal and pelvic abscess in Crohn's disease: results of noninvasive and surgical management. *Br J Surg*. 1998;85:367–371.
109. Lambiase RE, Cronan JJ, Dorfman GS, Paoletta LP, Haas RA. Percutaneous drainage of abscesses in patients with Crohn disease. *AJR Am J Roentgenol*. 1988;150:1043–1045.
110. Gervais DA, Hahn PF, O'Neill MJ, Mueller PR. Percutaneous abscess drainage in Crohn disease: technical success and short- and long-term outcomes during 14 years. *Radiology*. 2002;222:645–651.
111. Golfieri R, Cappelli A, Giampalma E, et al. CT-guided percutaneous pelvic abscess drainage in Crohn's disease. *Tech Coloproctol*. 2006;10:99–105.
112. Lobatón T, Guardiola J, Rodríguez-Moranta F, et al. Comparison of the long-term outcome of two therapeutic strategies for the management of abdominal abscess complicating Crohn's disease: percutaneous drainage or immediate surgical treatment. *Colorectal Dis*. 2013;15:1267–1272.
113. Gutierrez A, Lee H, Sands BE. Outcome of surgical versus percutaneous drainage of abdominal and pelvic abscesses in Crohn's disease. *Am J Gastroenterol*. 2006;101:2283–2289.
114. Bermejo F, Garrido E, Chaparro M, et al. Efficacy of different therapeutic options for spontaneous abdominal abscesses in Crohn's disease: are antibiotics enough? *Inflamm Bowel Dis*. 2012;18:1509–1514.
115. da Luz Moreira A, Stocchi L, Tan E, Tekkis PP, Fazio VW. Outcomes of Crohn's disease presenting with abdominopelvic abscess. *Dis Colon Rectum*. 2009;52:906–912.
116. He X, Lin X, Lian L, et al. Preoperative percutaneous drainage of spontaneous intra-abdominal abscess in patients with Crohn's disease: a meta-analysis (published online ahead of print September 11, 2014). *J Clin Gastroenterol*. doi: 10.1097/MCG.0000000000000219
117. Kumar RR, Kim JT, Haukoos JS, et al. Factors affecting the successful management of intra-abdominal abscesses with antibiotics and the need for percutaneous drainage. *Dis Colon Rectum*. 2006;49:183–189.
118. Lee H, Kim YH, Kim JH, et al. Nonsurgical treatment of abdominal or pelvic abscess in consecutive patients with Crohn's disease. *Dig Liver Dis*. 2006;38:659–664.
119. Poritz LS, Koltun WA. Percutaneous drainage and ileocelectomy for spontaneous intraabdominal abscess in Crohn's disease. *J Gastrointest Surg*. 2007;11:204–208.
120. Cullen G, Vaughn B, Ahmed A, et al. Abdominal phlegmons in Crohn's disease: outcomes following antitumor necrosis factor therapy. *Inflamm Bowel Dis*. 2012;18:691–696.
121. Garcia JC, Persky SE, Bonis PA, Topazian M. Abscesses in Crohn's disease: outcome of medical versus surgical treatment. *J Clin Gastroenterol*. 2001;32:409–412.
122. Nguyen DL, Sandborn WJ, Loftus EV Jr, et al. Similar outcomes of surgical and medical treatment of intra-abdominal abscesses in patients with Crohn's disease. *Clin Gastroenterol Hepatol*. 2012;10:400–404.
123. Ayuk P, Williams N, Scott NA, Nicholson DA, Irving MH. Management of intra-abdominal abscesses in Crohn's disease. *Ann R Coll Surg Engl*. 1996;78:5–10.
124. Maconi G, Sampietro GM, Parente F, et al. Contrast radiology, computed tomography and ultrasonography in detecting internal fistulas and intra-abdominal abscesses in Crohn's disease: a prospective comparative study. *Am J Gastroenterol*. 2003;98:1545–1555.
125. Poritz LS, Gagliano GA, McLeod RS, MacRae H, Cohen Z. Surgical management of entero and colcutaneous fistulae in Crohn's disease: 17 years' experience. *Int J Colorectal Dis*. 2004;19:481–485.
126. Saint-Marc O, Vaillant JC, Frileux P, Ballardur P, Tiret E, Parc R. Surgical management of ileosigmoid fistulas in Crohn's disease: role of preoperative colonoscopy. *Dis Colon Rectum*. 1995;38:1084–1087.
127. Young-Fadok TM, Wolff BG, Meagher A, Benn PL, Dozois RR. Surgical management of ileosigmoid fistulas in Crohn's disease. *Dis Colon Rectum*. 1997;40:558–561.
128. Melton GB, Stocchi L, Wick EC, Appau KA, Fazio VW. Contemporary surgical management for ileosigmoid fistulas in Crohn's disease. *J Gastrointest Surg*. 2009;13:839–845.
129. Yamamoto T, Keighley MR. Enterovesical fistulas complicating Crohn's disease: clinicopathological features and management. *Int J Colorectal Dis*. 2000;15:211–215.

130. Gruner JS, Sehon JK, Johnson LW. Diagnosis and management of enterovesical fistulas in patients with Crohn's disease. *Am Surg*. 2002;68:714–719.
131. Levy C, Tremaine WJ. Management of internal fistulas in Crohn's disease. *Inflamm Bowel Dis*. 2002;8:106–111.
132. Papi C, Gili L, Tarquini M, Antonelli G, Capurso L. Infliximab for severe recurrent Crohn's disease presenting with massive gastrointestinal hemorrhage. *J Clin Gastroenterol*. 2003;36:238–241.
133. Kim KJ, Han BJ, Yang SK, et al. Risk factors and outcome of acute severe lower gastrointestinal bleeding in Crohn's disease. *Dig Liver Dis*. 2012;44:723–728.
134. Robert JR, Sachar DB, Greenstein AJ. Severe gastrointestinal hemorrhage in Crohn's disease. *Ann Surg*. 1991;213:207–211.
135. Cirocco WC, Reilly JC, Rusin LC. Life-threatening hemorrhage and exsanguination from Crohn's disease: report of four cases. *Dis Colon Rectum*. 1995;38:85–95.
136. Driver CP, Anderson DN, Keenan RA. Massive intestinal bleeding in association with Crohn's disease. *J R Coll Surg Edinb*. 1996;41:152–154.
137. Belaiche J, Louis E, D'Haens G, et al. Acute lower gastrointestinal bleeding in Crohn's disease: characteristics of a unique series of 34 patients. Belgian IBD Research Group. *Am J Gastroenterol*. 1999;94:2177–2181.
138. Daperno M, Sostegni R, Rocca R. Lower gastrointestinal bleeding in Crohn's disease: how (un-)common is it and how to tackle it? *Dig Liver Dis*. 2012;44:721–722.
139. García-Blázquez V, Vicente-Bártulos A, Olavarria-Delgado A, Plana MN, van der Winden D, Zamora J; EBM-Connect Collaboration. Accuracy of CT angiography in the diagnosis of acute gastrointestinal bleeding: systematic review and meta-analysis. *Eur Radiol*. 2013;23:1181–1190.
140. Wu LM, Xu JR, Yin Y, Qu XH. Usefulness of CT angiography in diagnosing acute gastrointestinal bleeding: a meta-analysis. *World J Gastroenterol*. 2010;16:3957–3963.
141. Auvin S, Molinié F, Gower-Rousseau C, et al. Incidence, clinical presentation and location at diagnosis of pediatric inflammatory bowel disease: a prospective population-based study in northern France (1988–1999). *J Pediatr Gastroenterol Nutr*. 2005;41:49–55.
142. Motil KJ, Grand RJ, Davis-Kraft L, Ferlic LL, Smith EO. Growth failure in children with inflammatory bowel disease: a prospective study. *Gastroenterology*. 1993;105:681–691.
143. Sawczenko A, Sandhu BK. Presenting features of inflammatory bowel disease in Great Britain and Ireland. *Arch Dis Child*. 2003;88:995–1000.
144. Heuschkel R, Salvestrini C, Beattie RM, Hildebrand H, Walters T, Griffiths A. Guidelines for the management of growth failure in childhood inflammatory bowel disease. *Inflamm Bowel Dis*. 2008;14:839–849.
145. Laukoetter MG, Mennigen R, Hannig CM, et al. Intestinal cancer risk in Crohn's disease: a meta-analysis. *J Gastrointest Surg*. 2011;15:576–583.
146. Maykel JA, Hagerman G, Mellgren AF, et al. Crohn's colitis: the incidence of dysplasia and adenocarcinoma in surgical patients. *Dis Colon Rectum*. 2006;49:950–957.
147. von Roon AC, Reese G, Teare J, Constantinides V, Darzi AW, Tekkis PP. The risk of cancer in patients with Crohn's disease. *Dis Colon Rectum*. 2007;50:839–855.
148. Friedman S, Rubin PH, Bodian C, Harpaz N, Present DH. Screening and surveillance colonoscopy in chronic Crohn's colitis: results of a surveillance program spanning 25 years. *Clin Gastroenterol Hepatol*. 2008;6:993–8; quiz 953.
149. Basseri RJ, Basseri B, Vassilaki ME, et al. Colorectal cancer screening and surveillance in Crohn's colitis. *J Crohns Colitis*. 2012;6:824–829.
150. Sebastian S, Hernández V, Myrelid P, et al. Colorectal cancer in inflammatory bowel disease: results of the 3rd ECCO pathogenesis scientific workshop (I). *J Crohns Colitis*. 2014;8:5–18.
151. Farraye FA, Odze RD, Eaden J, Itzkowitz SH. AGA technical review on the diagnosis and management of colorectal neoplasia in inflammatory bowel disease. *Gastroenterology*. 2010;138:746–774.
152. Cairns SR, Scholefield JH, Steele RJ, et al; British Society of Gastroenterology; Association of Coloproctology for Great Britain and Ireland. Guidelines for colorectal cancer screening and surveillance in moderate and high risk groups (update from 2002). *Gut*. 2010;59:666–689.
153. Levin B, Lieberman DA, McFarland B, et al; American Cancer Society Colorectal Cancer Advisory Group; US Multi-Society Task Force; American College of Radiology Colon Cancer Committee. Screening and surveillance for the early detection of colorectal cancer and adenomatous polyps, 2008: a joint guideline from the American Cancer Society, the US Multi-Society Task Force on Colorectal Cancer, and the American College of Radiology. *Gastroenterology*. 2008;134:1570–1595.
154. Van Assche G, Dignass A, Panes J, et al; European Crohn's and Colitis Organisation (ECCO). The second European evidence-based Consensus on the diagnosis and management of Crohn's disease: definitions and diagnosis. *J Crohns Colitis*. 2010;4:7–27.
155. Subramanian V, Mannath J, Ragunath K, Hawkey CJ. Meta-analysis: the diagnostic yield of chromoendoscopy for detecting dysplasia in patients with colonic inflammatory bowel disease. *Aliment Pharmacol Ther*. 2011;33:304–312.
156. Subramanian V, Ramappa V, Telakis E, et al. Comparison of high definition with standard white light endoscopy for detection of dysplastic lesions during surveillance colonoscopy in patients with colonic inflammatory bowel disease. *Inflamm Bowel Dis*. 2013;19:350–355.
157. Annese V, Daperno M, Rutter MD, et al; European Crohn's and Colitis Organisation. European evidence based consensus for endoscopy in inflammatory bowel disease. *J Crohns Colitis*. 2013;7:982–1018.
158. Aalykke C, Jensen MD, Fallingborg J, et al. Colonoscopy surveillance for dysplasia and colorectal cancer in patients with inflammatory bowel disease. *Dan Med J*. 2015;62:B4995.
159. Navaneethan U, Rai T, Venkatesh PG, Kiran RP. Primary sclerosing cholangitis and the risk of colon neoplasia in patients with Crohn's colitis (published online ahead of print February 26, 2015). *Gastroenterol Rep (Oxf)*. doi: 10.1093/gastro/gov007
160. Navaneethan U, Venkatesh PG, Jegadeesan R, et al. Comparison of outcomes for patients with primary sclerosing cholangitis associated with ulcerative colitis and Crohn's disease (published online ahead of print October 29, 2014). *Gastroenterol Rep (Oxf)*. doi: 10.1093/gastro/gou074

161. Halliday JS, Djordjevic J, Lust M, et al. A unique clinical phenotype of primary sclerosing cholangitis associated with Crohn's disease. *J Crohns Colitis*. 2012;6:174–181.
162. Eaden J, Abrams K, McKay H, Denley H, Mayberry J. Inter-observer variation between general and specialist gastrointestinal pathologists when grading dysplasia in ulcerative colitis. *J Pathol*. 2001;194:152–157.
163. Odze RD, Goldblum J, Noffsinger A, Alsaigh N, Rybicki LA, Fogt F. Interobserver variability in the diagnosis of ulcerative colitis-associated dysplasia by telepathology. *Mod Pathol*. 2002;15:379–386.
164. Itzkowitz SH, Present DH; Crohn's and Colitis Foundation of America Colon Cancer in IBD Study Group. Consensus conference: colorectal cancer screening and surveillance in inflammatory bowel disease. *Inflamm Bowel Dis*. 2005;11:314–321.
165. Kiran RP, Khoury W, Church JM, Lavery IC, Fazio VW, Remzi FH. Colorectal cancer complicating inflammatory bowel disease: similarities and differences between Crohn's and ulcerative colitis based on three decades of experience. *Ann Surg*. 2010;252:330–335.
166. Kiran RP, Nisar PJ, Goldblum JR, et al. Dysplasia associated with Crohn's colitis: segmental colectomy or more extended resection? *Ann Surg*. 2012;256:221–226.
167. Maser EA, Sachar DB, Kruse D, Harpaz N, Ullman T, Bauer JJ. High rates of metachronous colon cancer or dysplasia after segmental resection or subtotal colectomy in Crohn's colitis. *Inflamm Bowel Dis*. 2013;19:1827–1832.
168. Rubin PH, Friedman S, Harpaz N, et al. Colonoscopic polypectomy in chronic colitis: conservative management after endoscopic resection of dysplastic polyps. *Gastroenterology*. 1999;117:1295–1300.
169. Marchetti F, Fazio VW, Ozuner G. Adenocarcinoma arising from a strictureplasty site in Crohn's disease: report of a case. *Dis Colon Rectum*. 1996;39:1315–1321.
170. Jaskowiak NT, Michelassi F. Adenocarcinoma at a strictureplasty site in Crohn's disease: report of a case. *Dis Colon Rectum*. 2001;44:284–287.
171. Partridge SK, Hodin RA. Small bowel adenocarcinoma at a strictureplasty site in a patient with Crohn's disease: report of a case. *Dis Colon Rectum*. 2004;47:778–781.
172. Tonelli F, Bargellini T, Leo F, Nesi G. Duodenal adenocarcinoma arising at the strictureplasty site in a patient with Crohn's disease: report of a case. *Int J Colorectal Dis*. 2009;24:475–477.
173. Fazio VW, Marchetti F, Church J, et al. Effect of resection margins on the recurrence of Crohn's disease in the small bowel: a randomized controlled trial. *Ann Surg*. 1996;224:563–573.
174. Bruewer M, Utech M, Rijcken EJ, et al. Preoperative steroid administration: effect on morbidity among patients undergoing intestinal bowel resection for Crohn's disease. *World J Surg*. 2003;27:1306–1310.
175. Kanazawa A, Yamana T, Okamoto K, Sahara R. Risk factors for postoperative intra-abdominal septic complications after bowel resection in patients with Crohn's disease. *Dis Colon Rectum*. 2012;55:957–962.
176. Simi M, Leardi S, Minervini S, Pietroletti R, Schietroma M, Speranza V. Early complications after surgery for Crohn's disease. *Neth J Surg*. 1990;42:105–109.
177. Mottet C, Vader JB, Felley C, et al; EPACT II Study Group. Appropriate management of special situations in Crohn's disease (upper gastro-intestinal; extra-intestinal manifestations; drug safety during pregnancy and breastfeeding): Results of a multidisciplinary international expert panel-EPACT II. *J Crohns Colitis*. 2009;3:257–263.
178. Kochhar R, Kochhar S. Endoscopic balloon dilation for benign gastric outlet obstruction in adults. *World J Gastrointest Endosc*. 2010;2:29–35.
179. Kelly SM, Hunter JO. Endoscopic balloon dilatation of duodenal strictures in Crohn's disease. *Postgrad Med J*. 1995;71:623–624.
180. Shapiro M, Greenstein AJ, Byrn J, et al. Surgical management and outcomes of patients with duodenal Crohn's disease. *J Am Coll Surg*. 2008;207:36–42.
181. Worsey MJ, Hull T, Ryland L, Fazio V. Strictureplasty is an effective option in the operative management of duodenal Crohn's disease. *Dis Colon Rectum*. 1999;42:596–600.
182. Takesue Y, Yokoyama T, Akagi S, et al. Strictureplasty for short duodenal stenosis in Crohn's disease. *J Gastroenterol*. 2000;35:929–932.
183. Tonelli F, Alemanno G, Bellucci F, Focardi A, Sturiale A, Giudici F. Symptomatic duodenal Crohn's disease: is strictureplasty the right choice? *J Crohns Colitis*. 2013;7:791–796.
184. Teeuwen PH, Stommel MW, Bremers AJ, van der Wilt GJ, de Jong DJ, Bleichrodt RP. Colectomy in patients with acute colitis: a systematic review. *J Gastrointest Surg*. 2009;13:676–686.
185. Carter FM, McLeod RS, Cohen Z. Subtotal colectomy for ulcerative colitis: complications related to the rectal remnant. *Dis Colon Rectum*. 1991;34:1005–1009.
186. Ng RL, Davies AH, Grace RH, Mortensen NJ. Subcutaneous rectal stump closure after emergency subtotal colectomy. *Br J Surg*. 1992;79:701–703.
187. Trickett JP, Tilney HS, Gudgeon AM, Mellor SG, Edwards DP. Management of the rectal stump after emergency sub-total colectomy: which surgical option is associated with the lowest morbidity? *Colorectal Dis*. 2005;7:519–522.
188. McKee RE, Keenan RA, Munro A. Colectomy for acute colitis: is it safe to close the rectal stump? *Int J Colorectal Dis*. 1995;10:222–224.
189. Karch LA, Bauer JJ, Gorfine SR, Gelernt IM. Subtotal colectomy with Hartmann's pouch for inflammatory bowel disease. *Dis Colon Rectum*. 1995;38:635–639.
190. Tekkis PP, Purkayastha S, Lanitis S, et al. A comparison of segmental vs subtotal/total colectomy for colonic Crohn's disease: a meta-analysis. *Colorectal Dis*. 2006;8:82–90.
191. Ritchie JK, Lockhart-Mummery HE. Non-restorative surgery in the treatment of Crohn's disease of the large bowel. *Gut*. 1973;14:263–269.
192. Yamamoto T, Watanabe T. Surgery for luminal Crohn's disease. *World J Gastroenterol*. 2014;20:78–90.
193. Cirincione E, Gorfine SR, Bauer JJ. Is Hartmann's procedure safe in Crohn's disease? Report of three cases. *Dis Colon Rectum*. 2000;43:544–547.
194. Zeitels JR, Fiddian-Green RG, Dent TL. Intersphincteric proctectomy. *Surgery*. 1984;96:617–623.
195. Regimbeau JM, Panis Y, Pocard M, et al. Long-term results of ileal pouch-anal anastomosis for colorectal Crohn's disease. *Dis Colon Rectum*. 2001;44:769–778.
196. Melton GB, Fazio VW, Kiran RP, et al. Long-term outcomes with ileal pouch-anal anastomosis and Crohn's disease: pouch

- retention and implications of delayed diagnosis. *Ann Surg.* 2008;248:608–616.
197. Joyce MR, Fazio VW. Can ileal pouch anal anastomosis be used in Crohn's disease? *Adv Surg.* 2009;43:111–137.
 198. Brown CJ, Maclean AR, Cohen Z, Macrae HM, O'Connor BI, McLeod RS. Crohn's disease and indeterminate colitis and the ileal pouch-anal anastomosis: outcomes and patterns of failure. *Dis Colon Rectum.* 2005;48:1542–1549.
 199. Braveman JM, Schoetz DJ Jr, Marcello PW, et al. The fate of the ileal pouch in patients developing Crohn's disease. *Dis Colon Rectum.* 2004;47:1613–1619.
 200. Martínez-Ramos D, Gibert-Gerez J, Escrig-Sos J, Alcalde-Sánchez M, Salvador-Sanchis JL. Ileal pouch-anal anastomosis for Crohn's disease. Current status [in Spanish]. *Cir Esp.* 2009;85:69–75.
 201. Tilney HS, Constantinides VA, Heriot AG, et al. Comparison of laparoscopic and open ileocecal resection for Crohn's disease: a metaanalysis. *Surg Endosc.* 2006;20:1036–1044.
 202. Tan JJ, Tjandra JJ. Laparoscopic surgery for Crohn's disease: a meta-analysis. *Dis Colon Rectum.* 2007;50:576–585.
 203. Lee Y, Fleming FJ, Deeb AP, Gunzler D, Messing S, Monson JR. A laparoscopic approach reduces short-term complications and length of stay following ileocolic resection in Crohn's disease: an analysis of outcomes from the NSQIP database. *Colorectal Dis.* 2012;14:572–577.
 204. Stocchi L, Milsom JW, Fazio VW. Long-term outcomes of laparoscopic versus open ileocolic resection for Crohn's disease: follow-up of a prospective randomized trial. *Surgery.* 2008;144:622–627.
 205. Patel SV, Patel SV, Ramagopalan SV, Ott MC. Laparoscopic surgery for Crohn's disease: a meta-analysis of perioperative complications and long term outcomes compared with open surgery. *BMC Surg.* 2013;13:14.
 206. Bellinger J, Munoz-Bongrand N, Pariente B, et al. Endoscopic and clinical recurrences after laparoscopic or open ileocolic resection in Crohn's disease. *J Laparoendosc Adv Surg Tech A.* 2014;24:617–622.
 207. Goyer P, Alves A, Bretagnol F, Bouhnik Y, Valleur P, Panis Y. Impact of complex Crohn's disease on the outcome of laparoscopic ileocecal resection: a comparative clinical study in 124 patients. *Dis Colon Rectum.* 2009;52:205–210.
 208. Holubar SD, Dozois EJ, Privitera A, et al. Laparoscopic surgery for recurrent ileocolic Crohn's disease. *Inflamm Bowel Dis.* 2010;16:1382–1386.
 209. Holubar SD, Dozois EJ, Privitera A, Pemberton JH, Cima RR, Larson DW. Minimally invasive colectomy for Crohn's colitis: a single institution experience. *Inflamm Bowel Dis.* 2010;16:1940–1946.
 210. Beyer-Berjot L, Mancini J, Bege T, et al. Laparoscopic approach is feasible in Crohn's complex enterovisceral fistulas: a case-match review. *Dis Colon Rectum.* 2013;56:191–197.
 211. Tavernier M, Lebreton G, Alves A. Laparoscopic surgery for complex Crohn's disease. *J Visc Surg.* 2013;150:389–393.
 212. Maeda K, Noda E, Nagahara H, et al. A comparative study of single-incision versus conventional multiport laparoscopic ileocecal resection for Crohn's disease with strictures. *Asian J Endosc Surg.* 2012;5:118–122.
 213. Rijcken E, Mennigen R, Argyris I, Senninger N, Bruewer M. Single-incision laparoscopic surgery for ileocolic resection in Crohn's disease. *Dis Colon Rectum.* 2012;55:140–146.
 214. Stewart DB, Messaris E. Early experience with single-site laparoscopic surgery for complicated ileocolic Crohn's disease at a tertiary-referral center. *Surg Endosc.* 2012;26:777–782.
 215. Gardenbroek TJ, Verlaan T, Tanis PJ, et al. Single-port versus multiport laparoscopic ileocecal resection for Crohn's disease. *J Crohns Colitis.* 2013;7:e443–e448.
 216. Moftah M, Nazour F, Cunningham M, Cahill RA. Single port laparoscopic surgery for patients with complex and recurrent Crohn's disease. *J Crohns Colitis.* 2014;8:1055–1061.
 217. da Luz Moreira A, Stocchi L, Remzi FH, Geisler D, Hammel J, Fazio VW. Laparoscopic surgery for patients with Crohn's colitis: a case-matched study. *J Gastrointest Surg.* 2007;11:1529–1533.
 218. Umanskiy K, Malhotra G, Chase A, Rubin MA, Hurst RD, Fichera A. Laparoscopic colectomy for Crohn's colitis: a large prospective comparative study. *J Gastrointest Surg.* 2010;14:658–663.
 219. Bartels SA, Gardenbroek TJ, Ubbink DT, Buskens CJ, Tanis PJ, Bemelman WA. Systematic review and meta-analysis of laparoscopic versus open colectomy with end ileostomy for non-toxic colitis. *Br J Surg.* 2013;100:726–733.
 220. Marceau C, Alves A, Ouaiissi M, Bouhnik Y, Valleur P, Panis Y. Laparoscopic subtotal colectomy for acute or severe colitis complicating inflammatory bowel disease: a case-matched study in 88 patients. *Surgery.* 2007;141:640–644.
 221. Messenger DE, Mihailovic D, MacRae HM, O'Connor BI, Victor JC, McLeod RS. Subtotal colectomy in severe ulcerative and Crohn's colitis: what benefit does the laparoscopic approach confer? *Dis Colon Rectum.* 2014;57:1349–1357.
 222. Guo Z, Li Y, Zhu W, Gong J, Li N, Li J. Comparing outcomes between side-to-side anastomosis and other anastomotic configurations after intestinal resection for patients with Crohn's disease: a meta-analysis. *World J Surg.* 2013;37:893–901.
 223. McLeod RS, Wolff BG, Ross S, Parkes R, McKenzie M; Investigators of the CAST Trial. Recurrence of Crohn's disease after ileocolic resection is not affected by anastomotic type: results of a multicenter, randomized, controlled trial. *Dis Colon Rectum.* 2009;52:919–927.
 224. Zurbuchen U, Kroesen AJ, Knebel P, et al; German Advanced Surgical Treatment Study Group. Complications after end-to-end vs. side-to-side anastomosis in ileocecal Crohn's disease—early postoperative results from a randomized controlled multi-center trial (ISRCTN-45665492). *Langenbecks Arch Surg.* 2013;398:467–474.