

Current Concepts and Controversies in Surgery for IBD

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Surgery improves the quality of life in patients with Crohn's disease (CD) and cures patients with chronic ulcerative colitis (CUC). There are several surgical controversies primarily involving techniques and long-term outcomes. Some debates are long standing; whether to perform a double-stapled ileal pouch-anal anastomosis (IPAA) or a mucosectomy and hand-sewn anastomosis, and whether to divert or not to divert in patients with CUC undergoing an IPAA. Other issues are more recent, such as the effects of age, pregnancy, pouch salvage, and laparoscopic IPAA. In patients with Crohn's disease the anastomosis technique, the management of perianal disease, and the role of laparoscopic surgery are topics of debate. This review shows the current concepts and controversies in the surgical management of patients with CUC or CD.

The term inflammatory bowel disease (IBD) incorporates chronic ulcerative colitis (CUC) at one end of the spectrum and Crohn's disease (CD) at the other. Indeterminate colitis (IC) resides somewhere in-between. The cause of IBD is unknown. The incidence of CUC has been stable over time, whereas the incidence of CD has increased.¹ Between 30% and 40% of patients with CUC will need surgery intervention in the course of their disease, whereas between 70% and 80% of CD patients will require surgery at some point after diagnosis of the disease. The proper role of surgery in patients with CD and UC is controversial. It is a multidisciplinary approach between the patient, surgeon, and gastroenterologist that makes successful treatment of IBD possible. This review shows the current concepts and controversies of surgery for IBD.

CUC

The current controversies surrounding the surgical treatment of CUC center principally around modifications of the ileal J-pouch-anal anastomosis (IPAA) procedure. IPAA has become the standard of care and the most common surgical option offered to patients with UC and familial adenomatous polyposis. Since its introduction,² the technical points of IPAA have been debated. We discuss several of these controversies.

Mucosectomy Versus Double-Staple Technique

The elimination of colonic mucosa is the goal of surgical intervention. This therapeutic aim theoretically eliminates the risk for neoplastic transformation. Since the first description of the technique of double-stapled IPAA,³ the controversy about the risk for dysplasia and residual disease in the remaining anal canal epithelium has been debated. The ease of use and benefits of improved function have made the double-stapled anastomosis the most widely used anastomotic technique. These benefits must be balanced against potential persistence of disease and malignant change in the retained mucosa (Table 1).

Double-stapled IPAA likely preserves the anal transition zone (ATZ) and its nerve endings better than endoanal mucosal resection. The ATZ is that portion of the anal canal between the uninterrupted squamous epithelium of the dentate line below and the uninterrupted columnar epithelium of the rectal epithelium above. The ATZ contains finger-like projections of columnar rectal epithelium and transitional epithelium. We reported in a prospective randomized trial that improved fecal continence at night occurred when the double-stapled technique was used. In this series, 64% of the hand-sewn group experienced occasional or frequent episodes of incontinence compared with 38% of the stapled group.⁴ Higher anal canal resting pressures (49.4 vs. 78.3 mm Hg, $P < 0.05$) and squeeze pressures (144 vs. 195 mm Hg, $P < 0.06$) also were found when the stapled technique was used. These findings have been supported by other randomized, prospective, and retrospective trials.⁵⁻⁸ However, other randomized trials have failed to find a difference in function between the double-stapled technique and mucosectomy.^{9,10}

Abbreviations used in this paper: ATZ, anal transition zone; CUC, chronic ulcerative colitis; IC, indeterminate colitis; IPAA, ileal pouch-anal anastomosis.

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Table 1. Mucosectomy Vs. Double-Stapled Anastomosis

| Findings | Double stapled | Mucosectomy |
|--------------------------------|----------------|-------------|
| Technical ease | Yes | No |
| Preserves ATZ | Yes | No |
| Improved function | Yes | No |
| Decreases septic complications | ? | ? |
| Decreases dysplasia risk | No | Yes |
| Decreases cancer risk | No | No |

Along with function, safety also has been shown to be improved.¹¹ In these series, it has been noted that leaks from a stapled anastomosis had a better prognosis than leaks from a hand-sewn anastomosis (although the reason for the difference is not apparent). Likewise, Ziv et al.¹² found that the double-stapled technique produced fewer septic complications and fewer sepsis-related pouch excisions than did the hand-sewn technique.

The strongest argument for mucosectomy is the potential risk for cancer developed in the retained anal canal epithelium. At the 10-year follow-up evaluation of the double-stapled IPAA, the incidence of dysplasia in the ATZ is approximately 5%.¹³ Although there have been reports of cancer developing after IPAA, it has occurred after double-stapling^{14,15} and endoanal mucosectomy.^{16–19} Although endoanal mucosectomy theoretically removes all anal canal mucosa to the dentate line, we have shown that residual of rectal mucosa may remain in the denuded muscle cuff in up to 14% of patients and in up to 7% of patients at the anastomosis.²⁰ To decrease potential risks for developing dysplasia or neoplasia in the pouch and ATZ, yearly digital examination and endoscopic biopsy examination of the pouch and ATZ is performed. If dysplasia is found in the ATZ, transanal mucosectomy with ileal pouch advancement is advocated.²¹

In conclusion, stapled IPAA is safe and likely provides improved function compared with a hand-sewn anastomosis. However, in patients with dysplasia of the rectum, endoanal mucosal resection is the procedure of choice.

Role of Temporary Diverting Ileostomy

Among surgeons with the most experience in performing IPAA, the procedure of choice is to construct a temporary ileostomy to divert the fecal stream. Several investigators have, over the years, proposed that a single-staged surgery is no more morbid than a 2-stage procedure.^{22–30} We first looked at these issues several years ago and reported on 37 patients who had a single-stage IPAA. These patients were compared with a case-

matched group of 37 patients who had an IPAA performed with defunctioning ileostomy.³¹ We found no statistical difference in complication rates, or in second surgery rates between the 2 groups. Functional outcomes also were similar in both groups. Others have reported similar long-term functional outcomes in patients undergoing single-staged IPAA.³² Another even more controversial issue is performing an IPAA laparoscopically without diversion. Ky et al.³³ reported on 37 patients who underwent laparoscopic IPAA, 32 of whom had no diverting ileostomy. Results, in terms of morbidity and functional outcomes, were not different between groups.

It seems obvious that eliminating a temporary stoma would decrease the mechanical and functional complications associated with an ileostomy.^{28,34,35} At the Mayo Clinic, we found²⁶ that mechanical complications occurred in 39 of 157 patients treated with diversion (retraction 16%, prolapse 1%, fistula 1%, abscess 1.0%, bowel obstruction related to stoma 6%). Functional complications occurred in 111 patients (peristomal irritation 54%, leakage 8%, high output 4%, incomplete diversion 6%). After stoma closure, bowel obstruction occurred in 15%, peritonitis in 7%, and wound infections in 2%. Although obstructions most often are managed nonsurgically, surgery occasionally is required. Complications such as high ileostomy output lead to dehydration and re-admission in as many as 20% of cases.³⁶

Grobler et al.³⁴ compared 23 patients with IPAA and loop ileostomy with 22 IPAA patients without loop ileostomy. The double-stapled technique was used in all patients, none were on steroids, and all had uncomplicated surgeries. There were no statistically significant differences in postoperative complication rates or function.³⁴ We and others have used similar selection criteria. Other parameters such as general health, comorbid factors, nutritional status, anemia, age, and current use of steroids or other immunosuppressive drugs have been investigated. Furthermore, intraoperative considerations including absence of tension, adequate anastomotic blood supply, lack of intraoperative complications, and satisfactory intraoperative testing of the integrity of the anastomosis are critical. Although not preventing anastomotic complications, a diverting ileostomy likely mitigates their severe complications. At the Mayo Clinic, pelvic sepsis occurs in about 5% of patients undergoing IPAA.³⁷

The data to support our current practice was reported by Garbus et al.³⁶; none of 110 patients who had an ileoanal pouch had clinical evidence of leaks when a diverting ileostomy was used. Three of 36 patients who had IPAA without an ileostomy had leaks that required

fecal diversion. Williamson et al.²² concluded that one-stage restorative proctocolectomy without a diverting ileostomy was associated with an increased risk of death. In 73 patients who developed pelvic sepsis after IPAA, the pouch failure rate (permanent diversion or pouch excision) was 26%, compared with 6% in patients without sepsis.³⁷ Although evacuation frequency was similar, fecal incontinence occurred more frequently in patients with pelvic sepsis.³⁷ Significant lifestyle restrictions also occurred more frequently.³⁷ Aside from the long-term poor functional outcomes, sepsis also may be life threatening.²² It seems reasonable, therefore, to advocate that a diverting ileostomy remain an integral part of IPAA.

Age

IPAA generally is performed in young patients: the mean age at surgery is approximately 30–35 years in almost all of the large reported series. Recently, IPAA has been performed increasingly in older patients. Bauer et al.³⁸ reported on 66 IPAA patients who were greater than 50 years of age and compared them with 253 patients less than 50 years old. Overall morbidity, mortality, and functional outcomes were similar.³⁸ Interestingly, however, a recent large trial reported by Delaney et al. found that nighttime seepage, incontinence, and overall quality of life actually were decreased in older patients.³⁹ We, too, have found the incidence of nocturnal evacuation and incontinence to be increased in older patients,^{40–42} although satisfaction remains high (age >65 years, 89% satisfied; age <65 years, 98% satisfied).^{39,40} It appears that in the properly motivated patient, age may not be a contraindication to pouch surgery.

Indeterminate Colitis

Even with the entire colonic specimen in hand, surgical pathologists are unable to differentiate between CD and CUC in 10%–15% of patients. When no clear distinction can be made between CUC and CD, patients have IC. There is much confusion in the literature regarding IC. In reports of patients with IC from the Mayo Clinic, we confirmed the diagnosis of IC in those patients who had no evidence of CD preoperatively (specifically, no evidence of perianal pathology), but who on review intraoperatively had one more confounding factor (skip areas, deep ulcerations, and so forth). Thus, a patient is diagnosed with IC only when a preoperative diagnosis of CUC is confounded by histopathology suggestive of CD intraoperatively. When IC is diagnosed, IPAA is performed.

Although failure rates in patients with IC (6.5% to 19.0%)^{43–45} are much better than in patients with CD (15% to 43%), the results still are not as good as those with CUC (1.4% to 8.0%).⁴⁶ Unfortunately, not only is failure of the pouch more frequent, but anorectal septic complications^{47,48} are increased as well with IC. We found the long-term function to be noted as good in up to 73% of patients.^{49,50} Although some of these IC patients go on to develop CD, it still is reasonable given the acceptable results to precede with IPAA in a patient with IC features and no ileal or perianal disease.

Pregnancy

Fecundity describes the biological ability to conceive and is based on the time periods of unprotected intercourse and expressed as fecundability (the probability of becoming pregnant per month with unprotected intercourse).⁵¹ Women with UC are known to have fecundity equal to that of the general population.^{52–54} After IPAA, parturition is normal and IPAA function appears to be unaffected.⁵⁵ Among 544 women who had IPAA and were under the age of 40, there were 142 pregnancies.⁵⁵ Of these, more than 50% had vaginal deliveries and many had multiple births after IPAA.

Two Dutch reports determined fecundity among patients after IPAA. The first study⁵⁶ assessed 237 women 12 months after stoma closure until the time of the study, or age 50 years. Birth rates were compared with the general population. The investigators found a slight decrease in the rate of fertility before IPAA at 87% of predicted ($P < 0.05$), but a significant decrease in fertility after IPAA of 49% of predicted ($P > 0.001$).^{12,27} The second trial followed-up 290 women after IPAA and compared them with 661 women in a reference population.⁵¹ The investigators found equivalent fecundity before IPAA, but after IPAA the fecundity decreased by 80%. This reduction in fecundity persisted over the 60-month follow-up period. The most likely cause of decreased fecundity is tubal occlusion from adhesive disease. These same problems have been reported in other series. Whether a laparoscopic approach might ameliorate this decreased fecundity after IPAA is unknown.

Revisional Pouch Surgery

IPAA fails in about 8% of patients, and some patients are candidates for pouch revision. We looked at outcomes of pouch revision and found that two thirds of patients had good clinical outcomes (Table 2).^{56–64} When pelvic sepsis was the cause of pouch failure, the long-term outcomes were less optimistic. Others have reported as well that pouch failure caused by pelvic sepsis

Table 2. Reoperations for Failure After IPAA

| Study | Year | No. patients | Salvage rate, % | Good result, % |
|---|------|--------------|-----------------|----------------|
| Galandiuk et al. ⁵⁷ | 1990 | 114 | 80 | 70 |
| Poggioli et al. ⁶⁰ | 1993 | 6 | 83 | 66 |
| Sagar et al. ⁵⁶ | 1996 | 23 | 74 | 48 |
| Herbst et al. ⁶¹ | 1996 | 16 | 81 | 80 |
| Ogunbiyi et al. ⁶⁶ | 1997 | 32 | 50 | 50 |
| Fazio et al. ⁶³ | 1998 | 35 | 86 | 49 |
| Cohen et al. ⁶⁴ | 1998 | 24 | 75 | 62 |
| Fonkalsrud and Bustorff-Silva ⁶² | 1998 | 164 | 97 | 93 |
| Saltzberg et al. ⁵⁹ | 1999 | 29 | 60 | NS |
| Zmora et al. ⁵⁸ | 2000 | 25 | 84 | 64 |

NS, not specified.

resulted in poor long-term outcomes.^{65,66} However, with increasing experience, salvage rates have climbed from 50% to 90% (Table 1).

Laparoscopic IPAA

The use of laparoscopic surgery for diseases of the colon and rectum began in the early 1990s. The need for advanced skills is obvious given that surgeries on the colon involve all 4 quadrants and procedures are performed on organs that move. Likewise, the unforgiving and friable nature of the disease adds to the complexity of the procedure.

Initially it appeared that laparoscopic approaches to treat CUC added little or no benefit (Table 3)⁶⁷⁻⁷⁰ compared with standard surgeries. Indeed, the earliest reports detailed increased morbidity and no benefits.⁷¹⁻⁷³ More recently, reports that laparoscopic IPAA is not only feasible, but may confer benefits over open IPAA, have appeared.^{68-70,74,75} Benefits include improved cosmesis, reduced pain, earlier return of bowel function, decreased morbidity, earlier discharge from the hospital, improved nutrition,⁷⁶ preservation of immune response,^{77,78} and decreased long-term complications.^{74,79-82} Functional outcomes and quality of life after laparoscopic IPAA appear to be no different from open IPAA.⁷⁰

Potential longer-term benefits include fewer adhesions and less hernia formation,⁸³⁻⁸⁵ improved fertility, and better quality of life.^{70,72} The consequences of the adhesions typically associated with open IPAA include: chronic pain, small bowel obstruction, ileus, and infertility.⁸⁶ The rate of small bowel obstruction after IPAA is about 20%, with half of patients requiring surgery. Moreover, the cumulative risk for small bowel obstruction at 10 years after IPAA is 31%.⁸⁷

Although the actual technique of laparoscopic IPAA varies among institutions, at the Mayo Clinic we use a simplified 4-port technique, with extracorporeal mesen-

teric vascular ligation and J-pouch construction, through a 4-5-cm periumbilical incision. Our early experience with laparoscopic IPAA has been reported.⁷⁵ Seven patients undergoing laparoscopic IPAA were compared with 7 case-matched controls undergoing conventional open IPAA. We found significant reductions in intravenous narcotic use, time to resumption of regular diet (2 vs. 7 days, $P = 0.010$), and length of hospital stay (4 vs. 9 days, $P = 0.012$). Complication rates were the same in both groups. Surgical times were significantly longer with the laparoscopic approach (median = 340 vs. 237 minutes, $P = 0.013$), however, with experience, surgical times have diminished, with most procedures now requiring approximately 3 hours.

Marcello et al.⁶⁸ published their case-matched comparative study of open vs. laparoscopic IPAA and found that laparoscopic IPAA not only had the same complication rate as open IPAA (20% vs. 25%), but patients had a faster return of bowel function and shorter hospital stay as well. Similarly, Hasegawa et al.⁶⁹ reported that laparoscopic IPAA is both feasible and safe in their study of 18 patients. Dunker et al.^{88,89} found cosmetic results and improved body image, as well as improved global physical function in patients after laparoscopic IPAA. In summary, laparoscopic IPAA may confer measurable benefits to patients requiring IPAA for CUC and the technique continues to evolve.

Laparoscopy in CD

Minimally invasive techniques are ideally suited for patients with CD. Both randomized and nonrandomized studies have shown that morbidity and mortality after laparoscopic resection are the same as after open resection (Table 4).^{86,90-104} Duepre et al.⁹⁵ showed that patients undergoing laparoscopic ileocolic resection for CD had a shorter time to resumption of diet, time to bowel function, and length of hospital stay.²³ Moreover, Milsom et al.⁹⁰ found that recovery of pulmonary function returned earlier in the laparoscopic group, but time to first bowel movement and hospital stay were not significantly different.⁹⁰ Finally, Bergamaschi et al.⁹³

Table 3. Laparoscopic IPAA

| Study | Year | No. patients | Conclusion |
|-------------------------------|------|-----------------|------------|
| Wexner et al. ⁷³ | 1992 | 5 | No benefit |
| Peters et al. ⁶⁷ | 1992 | 2 | — |
| Schmitt et al. ⁷² | 1994 | 22 | No benefit |
| Santoro et al. ⁸⁰ | 1999 | 5 | Benefit |
| Dozois et al. ⁷⁵ | 1999 | 7 (7 matched) | Benefit |
| Marcello et al. ⁶⁸ | 2000 | 20 (20 matched) | Benefit |
| Dunker et al. ⁷⁰ | 2001 | 16 | Benefit |
| Hasegawa et al. ⁶⁹ | 2002 | 18 | Feasible |

Table 4. Laparoscopic Surgery for CD

| Study | Year | No. patients | Obstruction rates: open vs. laparoscopic | Hernia rates: open vs. laparoscopic | Morbidity | Length of stay (days) |
|-----------------------------------|------|-------------------------|---|--|-----------|--------------------------|
| Milsom et al. ¹⁰¹ | 1993 | 9 | | | 0 | 7 |
| Liu et al. ⁹⁹ | 1995 | 10 | | | | 7 |
| Ludwig et al. ¹⁰⁰ | 1996 | 31 | | | 2 | 6 |
| Reissman et al. ¹⁰² | 1996 | 51 | | | 14 | 5.1 |
| Reissman et al. ¹⁰³ | 1996 | 72 total/49 CD | | | 18 total | 6.5 |
| Bauer et al. ⁹² | 1996 | 25 | | | 0 | 6.5 |
| Wu et al. ¹⁰⁴ | 1997 | 46 | | | 7 | 4.5 |
| Hildebrandt et al. ⁹⁸ | 1998 | 222 | | | 15 | .5-8 |
| Canin-Endres et al. ⁹⁴ | 1999 | 88 | | | 8 | 4.2 |
| Hamel et al. ⁹⁷ | 2001 | 130 | | | 18 | 8.8 |
| Milsom (PRT) et al. ⁹⁰ | 2001 | 31 laparoscopic/29 open | | | 6 | 5 |
| Duepre et al. ⁹⁵ | 2002 | 21 | | | 14.3 | 3 |
| Bergamaschi et al. ⁹³ | 2003 | 39 | 35.4% vs. 11.1% | | | 5.6 |
| Duepre et al. ⁹⁶ | 2003 | 211 total/123 IBD | 24.3% vs. 8.1% | 14% vs. 3.3% | | NR |

PRT, prospective randomized control trial; NR, not reported.

compared 39 laparoscopic ileocelectomies with 53 open ileocelectomies and found a shorter hospital stay in the laparoscopic group. Moreover, they found that the rate of small bowel obstruction was 35% and 11% in favor of the laparoscopic group.

Anastomotic Techniques

Whether the actual surgical techniques impact on the rates of recurrence and need a second surgery is unknown. Eventually about 80% of patients with CD will require surgery and 30% will require a second surgery within 5 years. Patients with the ileocolic variant of CD have the highest risk for recurrence; fully 42% of patients will require a second surgery by 15 years.^{105,106} This rate of recurrence is not affected by achieving clear margins. Because disease can recur at the site of anastomosis, it seems reasonable to question whether the type of anastomosis constructed influenced the rate of recurrence of disease.

Several retrospective studies have found that a stapled anastomosis achieved longer intervals of time between recurrences. Moreover, the trials by Hashemi et al.,¹⁰⁷ Yamamoto et al.,¹⁰⁸ and Munoz et al.¹⁰⁹ have documented that stapled anastomosis is superior to hand-sewn techniques in this regard. However, these studies were not randomized, and will need to be addressed by a randomized trial comparing stapled vs. hand-sewn anastomosis.

Strictureplasty plays a prominent role in the surgical management of small bowel CD. Isolated strictures under 10 cm in length often are considered for strictureplasty. Our original experience with 35 patients in whom 71 strictureplasties were performed and who were followed-up for over 3 years found no significant increase in postoperative morbidity and symptomatic recurrence

rates of 20%.¹¹⁰ Two more recent reports of more than 1400 strictureplasties followed-up for more than 7 years found second surgery rates of between 34% to 44% and symptomatic relief in more than 95% of patients.^{111,112} Strictureplasty is indeed valuable in the surgical treatment of CD.

Perianal Disease

The primary treatment approach for patients with CD with perineal involvement is combined medical and surgical management.¹¹³ Either one alone is not efficacious. Surgically, a noncutting seton is used most commonly and its effectiveness in managing complex fistulas is well documented.¹¹⁴⁻¹¹⁸ A seton is a nonabsorbable suture (or vessel loop) that is placed through the fistula tract. Passing it through the cutaneous opening of the fistula and out of the associated anal canal opening allows the 2 ends to be tied loosely together. Although sometimes these draining setons are uncomfortable for the patient, drainage is maintained, thus decreasing the risk for recurrent abscess while aggressive medical therapy is instituted.

Among 27 patients with fistulizing CD, Scott and Northover¹¹⁷ reported that 85% of patients treated with noncutting setons experienced fistula closure. Although other investigators have reported good initial results by using such setons,^{114,119} the recurrence rate approaches 40% after removing the seton. This high recurrence rate appears to lend legitimacy to our use of concomitant antibiotics, azathioprine, or 6-mercaptopurine and infliximab.^{120,121} Recent experience at the Mayo Clinic showed that infliximab and surgery has led to the resolution of perianal fistulas in 68% of patients.¹²² We also found that the addition of seton placement with infliximab reduced the rate of recurrent abscess. Topstad et

al.¹²³ found that 67% of patients had a complete response to combination therapy and 19% a partial response. In a comparative study by Regueiro and Mardini,¹²⁴ perianal fistulas were treated with infliximab alone vs. combination infliximab plus seton placement. They found that initial response was improved with seton placement (100% vs. 82.6%), lower recurrence rates (44% vs. 79%), and longer time to recurrence (13.5 vs. 3.6 months). At the Mayo Clinic, medical therapy is instituted together with placement of setons. Once the inflammation subsides, the seton is downsized as the fistula fibroses and narrows in caliber, or it is removed.

Summary

Controversies in IBD surgery are many and varied. There will no doubt be continued improvement in our understanding of both the techniques and outcomes of surgery for IBD. The most interesting and promising developments are the adaptation of minimally invasive techniques for CD and CUC patients and the recognition that medical and surgical management are required to manage perianal CD efficaciously.

References

- Gasche C, Scholmerich J, Brynskov J, D'Haens G, Hanauer SB, Irvine EJ, Jewell DP, Rachmilewitz D, Sachar DB, Sandborn WJ, Sutherland LR. A simple classification of Crohn's disease: report of the Working Party for the World Congresses of Gastroenterology, Vienna 1998. *Inflamm Bowel Dis* 2000;6:8-15.
- Parks AG, Nicholls RJ. Proctocolectomy without ileostomy for ulcerative colitis. *Br Med J* 1978;2:85-88.
- Knight CD, Griffen FD. An improved technique for low anterior resection of the rectum using the EEA stapler. *Surgery* 1980;88:710-714.
- Reilly WT, Pemberton JH, Wolff BG, Nivatvongs S, Devine RM, Litchy WJ, McIntyre PB. Randomized prospective trial comparing ileal pouch-anal anastomosis performed by excising the anal mucosa to ileal pouch-anal anastomosis performed by preserving the anal mucosa. *Ann Surg* 1997;225:666-676.
- Deen KI, Williams JG, Grant EA, Billingham C, Keighley MR. Randomized trial to determine the optimum level of pouch-anal anastomosis in stapled restorative proctocolectomy. *Dis Colon Rectum* 1995;38:133-138.
- Gemlo BT, Belmonte C, Wiltz O, Madoff RD. Functional assessment of ileal pouch-anal anastomotic techniques. *Am J Surg* 1995;169:137-141.
- Hallgren TA, Fasth SB, Oresland TO, Hulten LA. Ileal pouch anal function after endoanal mucosectomy and handsewn ileoanal anastomosis compared with stapled anastomosis without mucosectomy. *Eur J Surg* 1995;161:915-921.
- McIntyre PB, Pemberton JH, Beart RW Jr, Devine RM, Nivatvongs S. Double-stapled vs. handsewn ileal pouch-anal anastomosis in patients with chronic ulcerative colitis. *Dis Colon Rectum* 1994;37:430-433.
- Choen S, Tsunoda A, Nicholls RJ. Prospective randomized trial comparing anal function after hand sewn ileoanal anastomosis with mucosectomy versus stapled ileoanal anastomosis without mucosectomy in restorative proctocolectomy. *Br J Surg* 1991;78:430-434.
- Luukkonen P, Jarvinen H. Stapled vs hand-sutured ileoanal anastomosis in restorative proctocolectomy. A prospective, randomized study. *Arch Surg* 1993;128:437-440.
- MacRae HM, McLeod RS, Cohen Z, O'Connor BI, Ton EN. Risk factors for pelvic pouch failure. *Dis Colon Rectum* 1997;40:257-262.
- Ziv Y, Fazio VW, Church JM, Lavery IC, King TM, Ambrosetti P. Stapled ileal pouch anal anastomoses are safer than handsewn anastomoses in patients with ulcerative colitis. *Am J Surg* 1996;171:320-323.
- Remzi FH, Fazio VW, Delaney CP, Preen M, Ormsby A, Bast J, O'Riordain MG, Strong SA, Church JM, Petras RE, Gramlich T, Lavery IC. Dysplasia of the anal transitional zone after ileal pouch-anal anastomosis: results of prospective evaluation after a minimum of ten years. *Dis Colon Rectum* 2003;46:6-13.
- Baratsis S, Hadjimiditriou F, Christodoulou M, Lariou K. Adenocarcinoma in the anal canal after ileal pouch-anal anastomosis for ulcerative colitis using a double stapling technique: report of a case. *Dis Colon Rectum* 2002;45:687-691.
- Sequens R. Cancer in the anal canal (transitional zone) after restorative proctocolectomy with stapled ileal pouch-anal anastomosis. *Int J Colorectal Dis* 1997;12:254-255.
- Lauret S, Ugolini F, D'Errico A, Rago S, Poggioli G. Adenocarcinoma below ileoanal anastomosis for ulcerative colitis: report of a case and review of the literature. *Dis Colon Rectum* 2002;45:418-421.
- Puthu D, Rajan N, Rao R, Rao L, Venugopal P. Carcinoma of the rectal pouch following restorative proctocolectomy. Report of a case. *Dis Colon Rectum* 1992;35:257-260.
- Rodriguez-Sanjuan JC, Polavieja MG, Naranjo A, Castillo J. Adenocarcinoma in an ileal pouch for ulcerative colitis. *Dis Colon Rectum* 1995;38:779-780.
- Stern H, Walfisch S, Mullen B, McLeod R, Cohen Z. Cancer in an ileoanal reservoir: a new late complication? *Gut* 1990;31:473-475.
- O'Connell PR, Pemberton JH, Weiland LH, Beart RW Jr, Dozois RR, Wolff BG, Telander RL. Does rectal mucosa regenerate after ileoanal anastomosis? *Dis Colon Rectum* 1987;30:1-5.
- Fazio VW, Tjandra JJ. Transanal mucosectomy. Ileal pouch advancement for anorectal dysplasia or inflammation after restorative proctocolectomy. *Dis Colon Rectum* 1994;37:1008-1011.
- Williamson ME, Lewis WG, Sagar PM, Holdsworth PJ, Johnston D. One-stage restorative proctocolectomy without temporary ileostomy for ulcerative colitis: a note of caution. *Dis Colon Rectum* 1997;40:1019-1022.
- Sugerman HJ, Sugerman EL, Meador JG, Newsome HH Jr, Kellum JM Jr, DeMaria EJ. Ileal pouch anal anastomosis without ileal diversion. *Ann Surg* 2000;232:530-541.
- Mowschenson PM, Critchlow JF, Rosenberg SJ, Peppercorn MA. Factors favoring continence, the avoidance of a diverting ileostomy and small intestinal conservation in the ileoanal pouch operation. *Surg Gynecol Obstet* 1993;177:17-26.
- Mowschenson PM, Critchlow JF. Outcome of early surgical complications following ileoanal pouch operation without diverting ileostomy. *Am J Surg* 1995;169:143-145.
- Metcalfe AM, Dozois RR, Kelly KA, Wolff BG. Ileal pouch-anal anastomosis without temporary, diverting ileostomy. *Dis Colon Rectum* 1986;29:33-35.
- Matikainen M, Santavirta J, Hiltunen KM. Ileoanal anastomosis without covering ileostomy. *Dis Colon Rectum* 1990;33:384-388.
- Jarvinen HJ, Luukkonen P. Comparison of restorative proctocolectomy with and without covering ileostomy in ulcerative colitis. *Br J Surg* 1991;78:199-201.
- Hainsworth PJ, Bartolo DC. Selective omission of loop ileostomy

- in restorative proctocolectomy. *Int J Colorectal Dis* 1998;13:119–123.
30. Gorfine SR, Gelernt IM, Bauer JJ, Harris MT, KreeI I. Restorative proctocolectomy without diverting ileostomy. *Dis Colon Rectum* 1995;38:188–194.
 31. Galandiuk S, Wolff BG, Dozois RR, Beart RW Jr. Ileal pouch-anal anastomosis without ileostomy. *Dis Colon Rectum* 1991;34:870–873.
 32. Tjandra JJ, Fazio VW, Milsom JW, Lavery IC, Oakley JR, Fabre JM. Omission of temporary diversion in restorative proctocolectomy—is it safe? *Dis Colon Rectum* 1993;36:1007–1014.
 33. Ky AJ, Sonoda T, Milsom JW. One-stage laparoscopic restorative proctocolectomy: an alternative to the conventional approach? *Dis Colon Rectum* 2002;45:207–210.
 34. Grobler SP, Hosie KB, Keighley MR. Randomized trial of loop ileostomy in restorative proctocolectomy. *Br J Surg* 1992;79:903–906.
 35. Hosie KB, Grobler SP, Keighley MR. Temporary loop ileostomy following restorative proctocolectomy. *Br J Surg* 1992;79:33–34.
 36. Garbus JE, Potenti F, Wexner SD. Current controversies in pouch surgery. *South Med J* 2003;96:32–36.
 37. Farouk R, Dozois RR, Pemberton JH, Larson D. Incidence and subsequent impact of pelvic abscess after ileal pouch-anal anastomosis for chronic ulcerative colitis. *Dis Colon Rectum* 1998;41:1239–1243.
 38. Bauer JJ, Gorfine SR, Gelernt IM, Harris MT, KreeI I. Restorative proctocolectomy in patients older than fifty years. *Dis Colon Rectum* 1997;40:562–565.
 39. Delaney CP, Fazio VW, Remzi FH, Hammel J, Church JM, Hull TL, Senagore AJ, Strong SA, Lavery IC. Prospective, age-related analysis of surgical results, functional outcome, and quality of life after ileal pouch-anal anastomosis. *Ann Surg* 2003;238:221–228.
 40. Delaney CP, Dadvand B, Remzi FH, Church JM, Fazio VW. Functional outcome, quality of life, and complications after ileal pouch-anal anastomosis in selected septuagenarians. *Dis Colon Rectum* 2002;45:890–894.
 41. Farouk R, Pemberton JH, Wolff BG, Dozois RR, Browning S, Larson D. Functional outcomes after ileal pouch-anal anastomosis for chronic ulcerative colitis. *Ann Surg* 2000;231:919–926.
 42. Karlbom U, Raab Y, Ejerblad S, Graf W, Thorn M, Pahlman L. Factors influencing the functional outcome of restorative proctocolectomy in ulcerative colitis. *Br J Surg* 2000;87:1401–1408.
 43. Breen EM, Schoetz DJ Jr, Marcello PW, Roberts PL, Collier JA, Murray JJ, Rusin LC. Functional results after perineal complications of ileal pouch-anal anastomosis. *Dis Colon Rectum* 1998;41:691–695.
 44. Marcello PW, Schoetz DJ Jr, Roberts PL, Murray JJ, Collier JA, Rusin LC, Veidenheimer MC. Evolutionary changes in the pathologic diagnosis after the ileoanal pouch procedure. *Dis Colon Rectum* 1997;40:263–269.
 45. McIntyre PB, Pemberton JH, Wolff BG, Dozois RR, Beart RW Jr. Indeterminate colitis. Long-term outcome in patients after ileal pouch-anal anastomosis. *Dis Colon Rectum* 1995;38:51–54.
 46. Foley EF, Schoetz DJ Jr, Roberts PL, Marcello PW, Murray JJ, Collier JA, Veidenheimer MC. Rediversion after ileal pouch-anal anastomosis. Causes of failures and predictors of subsequent pouch salvage. *Dis Colon Rectum* 1995;38:793–798.
 47. Belliveau P, Trudel J, Vasilevsky CA, Stein B, Gordon PH. Ileoanal anastomosis with reservoirs: complications and long-term results. *Can J Surg* 1999;42:345–352.
 48. Koltun WA, Schoetz DJ Jr, Roberts PL, Murray JJ, Collier JA, Veidenheimer MC. Indeterminate colitis predisposes to perineal complications after ileal pouch-anal anastomosis. *Dis Colon Rectum* 1991;34:857–860.
 49. Yu CS, Pemberton JH, Larson D. Ileal pouch-anal anastomosis in patients with indeterminate colitis: long-term results. *Dis Colon Rectum* 2000;43:1487–1496.
 50. Fazio VW, Ziv Y, Church JM, Oakley JR, Lavery IC, Milsom JW, Schroeder TK. Ileal pouch-anal anastomoses complications and function in 1005 patients. *Ann Surg* 1995;222:120–127.
 51. Ording OK, Juul S, Berndtsson I, Oresland T, Laurberg S. Ulcerative colitis: female fecundity before diagnosis, during disease, and after surgery compared with a population sample. *Gastroenterology* 2002;122:15–19.
 52. Fonager K, Sorensen HT, Olsen J, Dahlerup JF, Rasmussen SN. Pregnancy outcome for women with Crohn's disease: a follow-up study based on linkage between national registries. *Am J Gastroenterol* 1998;93:2426–2430.
 53. Mayberry JF, Weterman IT. European survey of fertility and pregnancy in women with Crohn's disease: a case control study by European collaborative group. *Gut* 1986;27:821–825.
 54. Woolfson K, Cohen Z, McLeod RS. Crohn's disease and pregnancy. *Dis Colon Rectum* 1990;33:869–873.
 55. Hahnloser D, Pemberton JH, Wolff BG, Larson D, Harrington J, Farouk R, Dozois RR. Pregnancy and delivery before and after ileal pouch-anal anastomosis (IPAA) for inflammatory bowel disease: immediate and long-term consequences and outcomes. *Dis Colon Rectum* 2004 (in press).
 56. Sagar PM, Dozois RR, Wolff BG, Kelly KA. Disconnection, pouch revision and reconnection of the ileal pouch-anal anastomosis. *Br J Surg* 1996;83:1401–1405.
 57. Galandiuk S, Scott NA, Dozois RR, Kelly KA, Ilstrup DM, Beart RW Jr, Wolff BG, Pemberton JH, Nivatvongs S, Devine RM. Ileal pouch-anal anastomosis. Reoperation for pouch-related complications. *Ann Surg* 1990;212:446–452.
 58. Zmora O, Efron JE, Noguera JJ, Weiss EG, Wexner SD. Reoperative abdominal and perineal surgery in ileoanal pouch patients. *Dis Colon Rectum* 2001;44:1310–1314.
 59. Saltzberg SS, DiEdwardo C, Scott TE, LaMorte WW, Stucchi AF, Becker JM. Ileal pouch salvage following failed ileal pouch-anal anastomosis. *J Gastrointest Surg* 1999;3:633–641.
 60. Poggioli G, Marchetti F, Selleri S, Laureti S, Stocchi L, Gozzetti G. Redo pouches: salvaging of failed ileal pouch-anal anastomoses. *Dis Colon Rectum* 1993;36:492–496.
 61. Herbst F, Sielezneck I, Nicholls RJ. Salvage surgery for ileal pouch outlet obstruction. *Br J Surg* 1996;83:368–371.
 62. Fonkalsrud EW, Bustorff-Silva J. Reconstruction for chronic dysfunction of ileoanal pouches. *Ann Surg* 1999;229:197–204.
 63. Fazio VW, Wu JS, Lavery IC. Repeat ileal pouch-anal anastomosis to salvage septic complications of pelvic pouches: clinical outcome and quality of life assessment. *Ann Surg* 1998;228:588–597.
 64. Cohen Z, Smith D, McLeod R. Reconstructive surgery for pelvic pouches. *World J Surg* 1998;22:342–346.
 65. MacLean AR, O'Connor B, Parkes R, Cohen Z, McLeod RS. Reconstructive surgery for failed ileal pouch-anal anastomosis: a viable surgical option with acceptable results. *Dis Colon Rectum* 2002;45:880–886.
 66. Ogunbiyi OA, Korsgen S, Keighley MR. Pouch salvage. Long-term outcome. *Dis Colon Rectum* 1997;40:548–552.
 67. Peters WR. Laparoscopic total proctocolectomy with creation of ileostomy for ulcerative colitis: report of two cases. *J Laparoendosc Surg* 1992;2:175–178.
 68. Marcello PW, Milsom JW, Wong SK, Hammerhofer KA, Goormastic M, Church JM, Fazio VW. Laparoscopic restorative proctocolectomy: case-matched comparative study with open restorative proctocolectomy. *Dis Colon Rectum* 2000;43:604–608.
 69. Hasegawa H, Watanabe M, Baba H, Nishibori H, Kitajima M. Laparoscopic restorative proctocolectomy for patients with ulcerative colitis. *J Laparoendosc Adv Surg Tech A* 2002;12:403–406.

70. Dunker MS, Bemelman WA, Slors JF, van Duijvendijk P, Gouma DJ. Functional outcome, quality of life, body image, and cosmesis in patients after laparoscopic-assisted and conventional restorative proctocolectomy: a comparative study. *Dis Colon Rectum* 2001;44:1800–1807.
71. Reissman P, Salky BA, Pfeifer J, Edye M, Jagelman DG, Wexner SD. Laparoscopic surgery in the management of inflammatory bowel disease. *Am J Surg* 1996;171:47–50.
72. Schmitt SL, Cohen SM, Wexner SD, Nogueras JJ, Jagelman DG. Does laparoscopic-assisted ileal pouch anal anastomosis reduce the length of hospitalization? *Int J Colorectal Dis* 1994;9:134–137.
73. Wexner SD, Johansen OB, Nogueras JJ, Jagelman DG. Laparoscopic total abdominal colectomy. A prospective trial. *Dis Colon Rectum* 1992;35:651–655.
74. Hildebrandt U, Lindemann W, Kreissler-Haag D, Feifel G, Ecker KW. [Laparoscopically-assisted proctocolectomy with ileoanal pouch in ulcerative colitis]. *Zentralbl Chir* 1998;123:403–405.
75. Young-Fadok TM, Dozois E, Sandborn WJ, Tremaine WJ. A case matched study of laparoscopic proctocolectomy and ileal pouch-anal anastomosis (PC-IPAA) versus open PC-IPAA for ulcerative colitis. *Gastroenterology* 2001;2303:A-452.
76. Senagore AJ, Kilbride MJ, Luchtefeld MA, MacKeigan JM, Davis AT, Moore JD. Superior nitrogen balance after laparoscopic-assisted colectomy. *Ann Surg* 1995;221:171–175.
77. Hildebrandt U, Kessler K, Pistorius G, Lindemann W, Ecker KW, Feifel G, Menger MD. Granulocyte elastase and systemic cytokine response after laparoscopic-assisted and open resections in Crohn's disease. *Dis Colon Rectum* 1999;42:1480–1486.
78. Trokel MJ, Bessler M, Treat MR, Whelan RL, Nowygrod R. Preservation of immune response after laparoscopy. *Surg Endosc* 1994;8:1385–1387.
79. Liu CD, Rolandelli R, Ashley SW, Evans B, Shin M, McFadden DW. Laparoscopic surgery for inflammatory bowel disease. *Am Surg* 1995;61:1054–1056.
80. Santoro E, Carlini M, Carboni F, Feroce A. Laparoscopic total proctocolectomy with ileal J pouch-anal anastomosis. *Hepatogastroenterology* 1999;46:894–899.
81. Thibault C, Poulin EC. Total laparoscopic proctocolectomy and laparoscopy-assisted proctocolectomy for inflammatory bowel disease: operative technique and preliminary report. *Surg Laparosc Endosc* 1995;5:472–476.
82. Tucker JG, Ambroze WL, Orangio GR, Duncan TD, Mason EM, Lucas GW. Laparoscopically assisted bowel surgery. Analysis of 114 cases. *Surg Endosc* 1995;9:297–300.
83. Garrard CL, Clements RH, Nanney L, Davidson JM, Richards WO. Adhesion formation is reduced after laparoscopic surgery. *Surg Endosc* 1999;13:10–13.
84. Luciano AA, Maier DB, Koch EI, Nulsen JC, Whitman GF. A comparative study of postoperative adhesions following laser surgery by laparoscopy versus laparotomy in the rabbit model. *Obstet Gynecol* 1989;74:220–224.
85. Schippers E, Tittel A, Ottinger A, Schumpelick V. Laparoscopy versus laparotomy: comparison of adhesion-formation after bowel resection in a canine model. *Dig Surg* 1998;15:145–147.
86. Bemelman WA, Slors JF, Dunker MS, van Hogezaand RA, van Deventer SJ, Ringers J, Griffioen G, Gouma DJ. Laparoscopic-assisted vs. open ileocolic resection for Crohn's disease. A comparative study. *Surg Endosc* 2000;14:721–725.
87. Francois Y, Dozois RR, Kelly KA, Beart RW Jr, Wolff BG, Pemberton JH, Ilstrup DM. Small intestinal obstruction complicating ileal pouch-anal anastomosis. *Ann Surg* 1989;209:46–50.
88. Dunker MS, Stiggelbout AM, van Hogezaand RA, Ringers J, Griffioen G, Bemelman WA. Cosmesis and body image after laparoscopic-assisted and open ileocolic resection for Crohn's disease. *Surg Endosc* 1998;12:1334–1340.
89. Dunker MS, Bemelman WA, Slors JF, van Duijvendijk P, Gouma DJ. Functional outcome, quality of life, body image, and cosmesis in patients after laparoscopic-assisted and conventional restorative proctocolectomy: a comparative study. *Dis Colon Rectum* 2001;44:1800–1807.
90. Milsom JW, Hammerhofer KA, Bohm B, Marcello P, Elson P, Fazio VW. Prospective, randomized trial comparing laparoscopic vs. conventional surgery for refractory ileocolic Crohn's disease. *Dis Colon Rectum* 2001;44:1–8.
91. Ogunbiyi OA, Fleshman JW. Place of laparoscopic surgery in Crohn's disease. *Baillieres Clin Gastroenterol* 1998;12:157–165.
92. Bauer JJ, Harris MT, Grumbach NM, Gorfine SR. Laparoscopic-assisted intestinal resection for Crohn's disease. Which patients are good candidates? *J Clin Gastroenterol* 1996;23:44–46.
93. Bergamaschi R, Pessaux P, Arnaud JP. Comparison of conventional and laparoscopic ileocolic resection for Crohn's disease. *Dis Colon Rectum* 2003;46:1129–1133.
94. Canin-Endres J, Salky B, Gattorno F, Edye M. Laparoscopically assisted intestinal resection in 88 patients with Crohn's disease. *Surg Endosc* 1999;13:595–599.
95. Duepre HJ, Senagore AJ, Delaney CP, Brady KM, Fazio VW. Advantages of laparoscopic resection for ileocecal Crohn's disease. *Dis Colon Rectum* 2002;45:605–610.
96. Duepre HJ, Senagore AJ, Delaney CP, Fazio VW. Does means of access affect the incidence of small bowel obstruction and ventral hernia after bowel resection? *Laparoscopy versus laparotomy*. *J Am Coll Surg* 2003;197:177–181.
97. Hamel CT, Hildebrandt U, Weiss EG, Feifel G, Wexner SD. Laparoscopic surgery for inflammatory bowel disease. *Surg Endosc* 2001;15:642–645.
98. Hildebrandt U, Ecker KW, Feifel G. [Minimally invasive surgery and Crohn disease]. *Chirurg* 1998;69:915–921.
99. Liu CD, Rolandelli R, Ashley SW, Evans B, Shin M, McFadden DW. Laparoscopic surgery for inflammatory bowel disease. *Am Surg* 1995;61:1054–1056.
100. Ludwig KA, Milsom JW, Church JM, Fazio VW. Preliminary experience with laparoscopic intestinal surgery for Crohn's disease. *Am J Surg* 1996;171:52–55.
101. Milsom JW, Lavery IC, Bohm B, Fazio VW. Laparoscopically assisted ileocolic resection in Crohn's disease. *Surg Laparosc Endosc* 1993;3:77–80.
102. Reissman P, Salky BA, Pfeifer J, Edye M, Jagelman DG, Wexner SD. Laparoscopic surgery in the management of inflammatory bowel disease. *Am J Surg* 1996;171:47–50.
103. Reissman P, Salky BA, Edye M, Wexner SD. Laparoscopic surgery in Crohn's disease. Indications and results. *Surg Endosc* 1996;10:1201–1203.
104. Wu JS, Birnbaum EH, Kodner IJ, Fry RD, Read TE, Fleshman JW. Laparoscopic-assisted ileocolic resections in patients with Crohn's disease: are abscesses, phlegmons, or recurrent disease contraindications? *Surgery* 1997;122:682–688.
105. Williams JG, Wong WD, Rothenberger DA, Goldberg SM. Recurrence of Crohn's disease after resection. *Br J Surg* 1991;78:10–19.
106. Lock MR, Farmer RG, Fazio VW, Jagelman DG, Lavery IC, Weakley FL. Recurrence and reoperation for Crohn's disease: the role of disease location in prognosis. *N Engl J Med* 1981;304:1586–1588.
107. Hashemi M, Novell JR, Lewis AA. Side-to-side stapled anastomosis may delay recurrence in Crohn's disease. *Dis Colon Rectum* 1998;41:1293–1296.
108. 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.
109. Munoz-Juarez M, Yamamoto T, Wolff BG, Keighley MR. Wide-lumen stapled anastomosis vs. conventional end-to-end anas-

- tomosis in the treatment of Crohn's disease. *Dis Colon Rectum* 2001;44:20–25.
110. Spencer MP, Nelson H, Wolff BG, Dozois RR. Strictureplasty for obstructive Crohn's disease: the Mayo experience. *Mayo Clin Proc* 1994;69:33–36.
 111. Dietz DW, Laureti S, Strong SA, Hull TL, Church J, Remzi FH, Lavery IC, Fazio VW. Safety and longterm efficacy of strictureplasty in 314 patients with obstructing small bowel Crohn's disease. *J Am Coll Surg* 2001;192:330–337.
 112. Yamamoto T, Bain IM, Allan RN, Keighley MR. An audit of strictureplasty for small-bowel Crohn's disease. *Dis Colon Rectum* 1999;42:797–803.
 113. Schwartz DA, Pemberton JH, Sandborn WJ. Diagnosis and treatment of perianal fistulas in Crohn disease. *Ann Intern Med* 2001;135:906–918.
 114. White RA, Eisenstat TE, Rubin RJ, Salvati EP. Seton management of complex anorectal fistulas in patients with Crohn's disease. *Dis Colon Rectum* 1990;33:587–589.
 115. Takesue Y, Ohge H, Yokoyama T, Murakami Y, Imamura Y, Sueda T. Long-term results of seton drainage on complex anal fistulae in patients with Crohn's disease. *J Gastroenterol* 2002;37:912–915.
 116. Sugita A, Koganei K, Harada H, Yamazaki Y, Fukushima T, Shimada H. Surgery for Crohn's anal fistulas. *J Gastroenterol* 1995;30(Suppl 8):143–146.
 117. Scott HJ, Northover JM. Evaluation of surgery for perianal Crohn's fistulas. *Dis Colon Rectum* 1996;39:1039–1043.
 118. Koganei K, Sugita A, Harada H, Fukushima T, Shimada H. Seton treatment for perianal Crohn's fistulas. *Surg Today* 1995;25:32–36.
 119. Pearl RK, Andrews JR, Orsay CP, Weisman RI, Prasad ML, Nelson RL, Cintron JR, Abcarian H. Role of the seton in the management of anorectal fistulas. *Dis Colon Rectum* 1993;36:573–577.
 120. Schwartz DA, Pemberton JH, Sandborn WJ. Diagnosis and treatment of perianal fistulas in Crohn disease. *Ann Intern Med* 2001;135:906–918.
 121. Pearl RK, Andrews JR, Orsay CP, Weisman RI, Prasad ML, Nelson RL, Cintron JR, Abcarian H. Role of the seton in the management of anorectal fistulas. *Dis Colon Rectum* 1993;36:573–577.
 122. Ricart E, Panaccione R, Loftus EV, Tremaine WJ, Sandborn WJ. Infliximab for Crohn's disease in clinical practice at the Mayo Clinic: the first 100 patients. *Am J Gastroenterol* 2001;96:722–729.
 123. Topstad DR, Panaccione R, Heine JA, Johnson DR, MacLean AR, Buie WD. Combined seton placement, infliximab infusion, and maintenance immunosuppressives improve healing rate in fistulizing anorectal Crohn's disease: a single center experience. *Dis Colon Rectum* 2003;46:577–583.
 124. Regueiro M, Mardini H. Treatment of perianal fistulizing Crohn's disease with infliximab alone or as an adjunct to exam under anesthesia with seton placement. *Inflamm Bowel Dis* 2003;9:98–103.
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