



Laparoscopic surgery contributes to a decrease in short-term complications in surgical ulcerative colitis patients during 2008–2017: a multicenter retrospective study in China

Zerong Cai^{1*}, Xiaosheng He^{1*}, Jianfeng Gong², Peng Du³, Wenjian Meng⁴, Wei Zhou⁵, Jinbo Jiang⁶, Bin Wu⁷, Weitang Yuan⁸, Qi Xue^{9,10}, Lianwen Yuan¹¹, Jinhai Wang¹², Jiandong Tai¹³, Jie Liang¹⁴, Weiming Zhu², Ping Lan¹, Xiaojian Wu¹

¹Department of Colorectal Surgery, Guangdong Institute of Gastroenterology, Guangdong Provincial Key Laboratory of Colorectal and Pelvic Floor Diseases, The Sixth Affiliated Hospital, Sun Yat-sen University, Guangzhou; ²Department of General Surgery, Jinling Hospital, Medical School of Nanjing University, Nanjing; ³Department of Colorectal Surgery, Xinhua Hospital, Shanghai Jiaotong University School of Medicine, Shanghai; ⁴Department of Gastrointestinal Surgery, West China Hospital, Sichuan University, Chengdu; ⁵Department of General Surgery, Sir Run Run Shaw Hospital, Zhejiang University School of Medicine, Hangzhou; ⁶Department of General Surgery, Qilu Hospital, Shandong University, Jinan; ⁷Department of General Surgery, Peking Union Medical College Hospital, Peking Union Medical College, Chinese Academy of Medical Sciences, Beijing; ⁸Department of Colorectal Surgery, The First Affiliated Hospital of Zhengzhou University, Zhengzhou; ⁹Department of General Surgery, Nanfang Hospital, Southern Medical University, Guangzhou; ¹⁰Department of General Surgery, Integrated Hospital of Traditional Chinese Medicine, Southern Medical University, Guangzhou; ¹¹Department of Geriatric Surgery, The Second Xiangya Hospital, Central South University, Changsha; ¹²Department of Colorectal Surgery, The First Affiliated Hospital, Zhejiang University College of Medicine, Hangzhou; ¹³Department of Colorectal and Anal Surgery, The First Hospital of Jilin University, Changchun, Jilin; ¹⁴State Key Laboratory of Cancer Biology and Institute of Digestive Diseases, Xijing Hospital of Digestive Diseases, Xi'an, China

Background/Aims: The aim of this study was to analyze the chronological changes in postoperative complications in surgical ulcerative colitis patients over the past decade in China and to investigate the potential parameters that contributed to the changes. **Methods:** Ulcerative colitis patients who underwent surgery during 2008–2017 were retrospectively enrolled from 13 hospitals in China. Postoperative complications were compared among different operation years. Risk factors for complications were identified by logistic regression analysis. **Results:** A total of 446 surgical ulcerative colitis patients were analyzed. Fewer short-term complications (24.8% vs. 41.0%, $P=0.001$) and more laparoscopic surgeries (66.4% vs. 25.0%, $P<0.001$) were found among patients who received surgery during 2014–2017 than 2008–2013. Logistic regression suggested that independent protective factors against short-term complications were a higher preoperative body mass index (odds ratio [OR], 0.870; 95% confidence interval [CI], 0.785–0.964; $P=0.008$), laparoscopic surgery (OR, 0.391; 95% CI, 0.217–0.705; $P=0.002$) and elective surgery (OR, 0.213; 95% CI, 0.067–0.675; $P=0.009$). The chronological decrease in short-term complications was associated with an increase in laparoscopic surgery. **Conclusions:** Our data revealed a downward trend of short-term postoperative complications among surgical ulcerative colitis patients in China during the past decade, which may be due to the promotion of minimally invasive techniques among Chinese surgeons. (Intest Res 2023;21:235-243)

Key Words: Colitis, ulcerative; Surgery; Complication; Laparoscopy

Received January 21, 2022. Revised March 28, 2022. Accepted April 1, 2022.

Correspondence to Xiaojian Wu, Department of Colorectal Surgery, The Sixth Affiliated Hospital, Sun Yat-sen University, 26# Yuancun Erheng Road, Guangzhou 510655, China. Tel: +86-20-38455325, Fax: +86-20-38254221, E-mail: wuxjian@mail.sysu.edu.cn

*These authors contributed equally to this study as first authors.

INTRODUCTION

Ulcerative colitis (UC) is an immunologically mediated bowel disorder that progressively and chronically damages the colon and rectum.¹ Although medical treatment, including glucocor-

ticoids and biological agents, has led to higher rate of disease remission² and lower rate of colectomy in recent years, 8% to 24% of UC patients still need surgical management.^{3,4}

Restorative total proctocolectomy with ileal pouch anal anastomosis (IPAA) is a theoretically curative and sphincter-preserving procedure that was first reported in 1978 by Parks and Nicholls,⁵ and this procedure has been internationally accepted as the gold standard for most surgical UC patients at present. The first reported total proctocolectomy plus IPAA for UC patients in China was in 1999,⁶ and several studies have explored the clinical parameters impacting postoperative complications.⁷⁻⁹

As biologic agents have emerged for treating UC patients, some studies revealed that the emergent colectomy rate and postoperative complications in acute severe UC have decreased,^{10,11} although some other studies held opposite opinion.^{12,13} However, the reasons for the improved outcomes of UC surgery in the new era remain unclear. In addition, minimally invasive surgeries and individual IPAA stage procedures have been applied to improve surgical outcomes of UC patients.^{14,15} However, surgeons still have concerns about whether IPAA surgery would increase postoperative complications, and whether the introduction of surgical techniques such as laparoscopic surgery and 3-stage IPAA procedures would contribute to favorable outcomes for UC patients.

Therefore, we retrospectively analyzed the trend of postoperative complications for surgical UC patients during 2008–2017 at 13 inflammatory bowel disease (IBD) centers throughout China, and the clinical parameters that contributed to the changes were investigated.

METHODS

1. Study Subjects

Details of the 13 IBD centers are listed in the acknowledgment section. UC patients who underwent colectomy from January 2008 to December 2017 were retrospectively enrolled. The inclusion criteria were a clinical and pathological diagnosis of UC and undergoing colectomy because of an emergency or an elective indication. The exclusion criteria were lost to follow-up and lacking important complication data. This study was approved by the Institutional Review Board of the Sixth Affiliated Hospital of Sun Yat-sen University (IRB No. 2019ZS-LYE-197). All the methods were performed in accordance with the Declaration of Helsinki. Informed consent was not obtained from all the participants, and the need for consent of

this study was deemed unnecessary by the Institutional Review Board of the Sixth Affiliated Hospital of Sun Yat-sen University according to national regulations.

2. Definitions of the Parameters

Patient characteristics, including age, sex, disease extent and severity, the preoperative body mass index (BMI), albumin, C-reactive protein, and erythrocyte sedimentation rate, were collected. Details on surgical methods, operation year, operative approaches (open or laparoscopic), surgical procedures (2-stage or 3-stage IPAA), and surgical complications were retrieved.

The disease extent of the UC patients was defined by the maximal macroscopic extent on colonoscopy and was classified as proctitis (E1), left-side colitis (E2), or extensive colitis (E3) according to the Montreal classification.¹⁶ Disease severity was defined based on the Truelove and Witts classification, including an assessment of bloody stool, pulse, temperature, hemoglobin, C-reactive protein, and erythrocyte sedimentation rate level.¹⁷ Surgical methods were defined as an IPAA procedure (hand sewn or stapled) and sub/total proctocolectomy with ileorectal or ileoanal anastomosis, sub/total proctocolectomy with ileostomy, or segmental colectomy and diverting loop ileostomy without bowel resection, which were grouped as non-IPAA procedures. The operation year was defined as the time when the patients received the first-stage operation. Detailed definitions of the surgery timing (emergency or elective), operative approaches, and surgical stage of the IPAA are shown in Supplementary Table 1.

3. Outcome Measurement

The primary outcome of this study was postoperative complications, including short-term and long-term complications. Short-term complications were defined as complications that occurred within 30 days after any stage of the operation, including intra-abdominal hemorrhage (requiring blood transfusion or an invasive intervention), abdominal infection, ileus, pulmonary infection, urinary retention, urinary infection, anastomotic leakage, anastomotic hemorrhage, and enteritis. Long-term complications were defined as complications that occurred more than 30 days after any stage of the operation, which included anastomotic stricture (requiring instrumental dilatation or surgical intervention), pouchitis, abdominal infection (including abscess), enteral fistula, ileus, and urinary infection. We categorized complications into infectious and noninfectious complications basing on whether they were caused by

definite pathogenic microorganism. More detailed definitions of the complications are shown in Supplementary Table 1.

4. Statistical Methods

Continuous variables are presented as the mean ± standard deviation (SD) or median (range), and categorical data are presented as numbers (percentages). Student *t*-tests and Mann-Whitney *U* tests were used to compare continuous variables, and chi-square tests were used to compare categorical variables. Linear-by-linear associations test was used to assess the chronological changes in postoperative complications. Univariate and multivariable logistic regression models were used to identify the risk factors for postoperative complications. Statistical analyses were performed with SPSS software version 19.0 (IBM Corp., Armonk, NY, USA). A *P*-value < 0.05 was considered statistically significant.

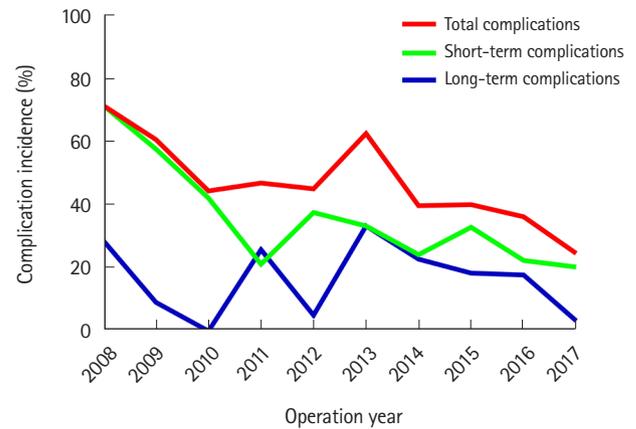


Fig. 1. Chronological change of postoperative complications including total complications, short-term complications, and long-term complications for surgical ulcerative colitis patients during 2008–2017 in China.

Table 1. Comparison of Postoperative Complications in Patients with Ulcerative Colitis between Subgroups of 2008–2013 and 2014–2017

Complications	Total (n = 446)	2008–2013 group (n = 144)	2014–2017 group (n = 302)	P-value
Short-term complications	134 (30.0)	59 (41.0)	75 (24.8)	0.001
Infectious complications	66 (14.8)	32 (22.2)	34 (11.3)	0.002
Abdominal infection	38 (8.5)	21 (14.6)	17 (5.6)	0.002
Pulmonary infection	25 (5.6)	10 (6.9)	15 (5.0)	0.400
Urinary infection	14 (3.1)	8 (5.6)	6 (2.0)	0.040
Noninfectious complications	102 (22.9)	43 (29.9)	59 (19.5)	0.015
Anastomotic leakage	13 (2.9)	9 (6.3)	4 (1.3)	0.004
Ileus	42 (9.4)	22 (15.3)	20 (6.6)	0.003
Intra-abdominal hemorrhage	23 (5.2)	8 (5.6)	15 (5.0)	0.790
Anastomotic hemorrhage	15 (3.4)	7 (4.9)	8 (2.6)	0.230
Urinary retention	13 (2.9)	4 (2.8)	9 (3.0)	0.910
Enteritis	7 (1.6)	1 (0.7)	6 (2.0)	0.310
Long-term complications ^a	66 (15.8)	21 (15.8)	45 (15.8)	0.990
Infectious complications	13 (3.1)	6 (4.5)	7 (2.5)	0.260
Abdominal infection	5 (1.2)	3 (2.3)	2 (0.7)	0.330
Urinary infection	4 (1.0)	0	4 (1.4)	0.310
Noninfectious complications	49 (11.8)	19 (14.3)	30 (10.6)	0.270
Enteral fistula	11 (2.6)	5 (3.8)	6 (2.1)	0.340
Pouchitis	28 (6.7)	12 (9.0)	16 (5.6)	0.200
Anastomotic stricture	14 (3.4)	3 (2.3)	11 (3.9)	0.560
Ileus	13 (3.1)	5 (3.8)	8 (2.8)	0.560

Values are presented as number (%).

^aLong-term complication information of 29 cases of patients was lost.

RESULTS

1. Short-term Complications for Surgical UC Patients Decreased during 2008–2017

A total of 446 UC patients who received surgical treatment during 2008–2017 were enrolled, with a mean age of 44.7 ± 14.2 years, and 58.7% of them were men. The five most common indications for surgical treatment were: ineffective medical therapy (63.5%), uncontrolled intestinal bleeding (11.6%), patients' personal demand (11.0%), dysplasia or cancer (5.7%), and toxic megacolon (4.1%). The number of surgical UC patients in China increased after 2013 (Supplementary Fig. 1). A total of 134 cases (30.0%) of short-term complications and 66 cases (15.8%) of long-term complications were observed after a follow-up of 30.1 months (range, 1.0–125.9 months). As shown in Fig. 1, the incidence of short-term complications in surgical UC patients showed a significant downtrend ($P < 0.001$), while the similar trend was not observed for the incidence of long-term complications.

Considering that laparoscopic surgery has become popular in China since 2014, we divided the patients into 2007–2013 and 2014–2017 subgroups according to their operation year. As shown in Table 1, patients who underwent surgery during 2014–2017 had lower rates of short-term complications, infectious short-term complications, abdominal infections, anastomotic leakage, urinary infections, and ileus (all $P < 0.05$). No significantly difference of long-term complications including pouchitis was found between the 2007–2013 and 2014–2017 subgroups ($P > 0.05$).

2. Higher BMI, Elective Surgery and Laparoscopic Surgery Protect against Short-term Complications

Clinical parameters were compared among patients with and without short-term postoperative complications. As shown in Table 2, patients who had short-term postoperative complications were characterized by more E1 disease (12.3% vs. 4.4%, $P = 0.025$), a lower preoperative BMI ($18.8 \pm 2.7 \text{ kg/m}^2$ vs. $20.1 \pm 3.1 \text{ kg/m}^2$, $P = 0.003$), and underwent more open surgery (57.5% vs. 42.4%, $P = 0.004$) and emergency surgery (17.1% vs. 7.2%, $P = 0.001$).

Logistic regression analysis was performed to explore risk factors for short-term complications of surgical UC patients. As shown in Table 3, a higher preoperative BMI (odds ratio [OR], 0.864; 95% confidence interval [CI], 0.783–0.954; $P = 0.004$), laparoscopic surgery (OR, 0.546; 95% CI, 0.362–0.822; $P = 0.004$), and elective surgery (OR, 0.366; 95% CI, 0.196–0.684; $P = 0.002$)

Table 2. Compared of Clinical and Surgical Parameters between Surgical Ulcerative Colitis Patients with and without Short-term Postoperative Complications

Characteristic	Without complications	With complications	P-value
Sex			0.953
Female	129 (41.3)	55 (41.0)	
Male	183 (58.7)	79 (59.0)	
Age (yr)	44.0 ± 13.9	46.2 ± 14.7	0.192
Disease extent			0.025
E1	10 (4.4)	14 (12.3)	
E2	44 (19.5)	18 (15.8)	
E3	172 (76.1)	82 (71.9)	
Disease severity			0.227
Mild	23 (10.0)	6 (5.4)	
Moderate	51 (21.3)	22 (19.6)	
Severe	152 (68.8)	84 (75.0)	
Preoperative BMI (kg/m^2)	20.1 ± 3.1	18.8 ± 2.7	0.003
Preoperative ALB level (g/L)	32.7 ± 6.4	31.8 ± 6.3	0.170
Preoperative CRP level (mg/L)	12.2 (0.3–172.1)	14.8 (0.2–171.0)	0.120
Preoperative ESR (mm/hr)	23 (1–129)	29 (2–140)	0.396
Preoperative corticosteroid			0.415
No	114 (38.3)	44 (34.1)	
Yes	184 (61.7)	85 (65.9)	
Preoperative infliximab			0.992
No	266 (89.3)	116 (89.2)	
Yes	32 (10.7)	14 (10.8)	
Operation approach (vs.)			0.004
Open surgery	132 (42.4)	77 (57.5)	
Laparoscopic surgery	179 (57.6)	57 (42.5)	
Elective vs. emergency			0.001
Elective	290 (92.9)	111 (82.8)	
Emergency	22 (7.1)	23 (17.2)	
IPAA surgery			0.953
No	122 (39.1)	52 (38.8)	
Yes	190 (60.9)	82 (61.2)	
IPAA stage			0.088
2	142 (74.7)	69 (84.1)	
3	48 (25.3)	13 (15.9)	

Values are presented as number (%), mean \pm standard deviation, or median (range).

BMI, body mass index; ALB, albumin; CRP, C-reaction protein; ESR, erythrocyte sedimentation rate; IPAA, ileal pouch anal anastomosis.

Table 3. Univariate and Multivariable Logistic Regression Analyses of Risk Factor for Short-term Complications in Surgical Ulcerative Colitis Patients

Characteristic	Univariate logistic regression		Multivariable logistic regression	
	OR (95% CI)	P-value	OR (95% CI)	P-value
Sex (female vs. male)	1.013 (0.671–1.531)	0.950	NA	
Age (per year)	1.011 (0.995–1.027)	0.190	NA	
Disease extent (E1, E2, E3)	0.725 (0.504–1.043)	0.080	NA	
Disease severity (mild, moderate, severe)	1.386 (0.949–2.026)	0.090	NA	
Preoperative BMI (per kg/m ²)	0.864 (0.783–0.954)	0.004	0.870 (0.785–0.964)	0.008
Preoperative ALB level (per g/L)	0.977 (0.946–1.010)	0.170	NA	
Preoperative CRP level (per mg/L)	1.005 (0.998–1.012)	0.190	NA	
Preoperative ESR (per mm/hr)	1.006 (0.996–1.016)	0.280	NA	
Preoperative corticosteroid (no vs. yes)	1.197 (0.777–1.845)	0.420	NA	
Preoperative infliximab (no vs. yes)	1.003 (0.516–1.950)	0.990	NA	
Laparoscopy vs. open surgery	0.546 (0.362–0.822)	0.004	0.391 (0.217–0.705)	0.002
Elective vs. emergency	0.366 (0.196–0.684)	0.002	0.213 (0.067–0.675)	0.009
Non-IPAA vs. IPAA	0.953 (0.668–1.534)	0.950	2.015 (0.995–4.080)	0.052
IPAA stages (2 vs. 3)	0.557 (0.283–1.097)	0.090	NA	

OR, odds ratio; CI, confidence interval; BMI, body mass index; ALB, albumin; CRP, C-reaction protein; ESR, erythrocyte sedimentation rate; IPAA, ileal pouch anal anastomosis; NA, not available.

were protective factors against short-term complications in UC patients who underwent surgery, according to the univariate logistic regression analysis. After adjusting for confounding factors, including preoperative BMI, operative approach (laparoscopy vs. open surgery), surgery timing (elective vs. emergency) and operation method (non-IPAA vs. IPAA), multivariable logistic regression analysis showed that independent protective factors for short-term complications were a higher preoperative BMI (OR, 0.870; 95% CI, 0.785–0.964; $P=0.008$), laparoscopic surgery (OR, 0.391; 95% CI, 0.217–0.705; $P=0.002$) and elective surgery (OR, 0.213; 95% CI, 0.067–0.675; $P=0.009$).

3. The Decrease in Short-term Complications Was Associated with Laparoscopic Surgery

We then compared clinical and surgical characteristics between patients who received surgery during 2008–2013 and 2014–2017. As shown in Table 4, more men (62.6% vs. 50.7%, $P=0.020$), more laparoscopic surgery (66.4% vs. 25.0%, $P<0.001$) and more 3-stage IPAA procedures (27.8% vs. 10.6%, $P=0.002$) were found in patients who underwent surgery during 2014–2017. Then, patients were divided into IPAA and non-IPAA cohorts, and the complications were compared (Table 5). For patients who received IPAA surgery, the difference in overall short-term postoperative and short-term infectious complica-

tions between open surgery and laparoscopic surgery was statistically significant (short-term complications, $P<0.001$; infectious complications, $P=0.009$), while among patients who received non-IPAA surgery, the benefit of laparoscopic surgery was not statistically significant ($P>0.05$).

DISCUSSION

In this article, we analyzed the chronological changes in postoperative complications for surgical UC patients during 2008–2017, and it was revealed that short-term complications, including infectious complications, decreased in the past decade in China. The decrease in complications was consistent with the increased application of laparoscopic surgery. Logistic regression analysis revealed that laparoscopic surgery was an independent protective factor for short-term postoperative complications.

The incidence of UC in China has increased, with a current incidence of 0.42 per 1,000,000 person-years.¹⁸ To our knowledge, this retrospective study is the first published nationwide multicenter investigation on surgical treatment of UC in China. As the IBD centers developed in China, more and more UC patients were diagnosed and managed by medical treatments and operations. Our results showed that the number of

Table 4. Compared of Clinical and Surgical Parameters between Surgical Ulcerative Colitis Patients Received Operation during 2008–2013 and 2014–2017

Characteristic	2008–2013	2014–2017	P-value
Sex			0.020
Female	71 (49.3)	113 (37.4)	
Male	73 (50.7)	189 (62.6)	
Age (yr)	44.7 ± 14.5	44.7 ± 14.0	0.960
Disease extent			0.820
E1	10 (7.6)	14 (6.7)	
E2	22 (16.6)	40 (19.2)	
E3	100 (75.8)	154 (74.1)	
Disease severity			0.380
Mild	8 (6.3)	21 (10.0)	
Moderate	31 (24.2)	42 (20.0)	
Severe	89 (69.5)	147 (70.0)	
Preoperative BMI (kg/m ²)	20.1 ± 3.4	19.6 ± 2.9	0.260
Preoperative ALB level (g/L)	32.1 ± 7.2	32.6 ± 5.9	0.500
Preoperative CRP level (mg/L)	11.2 (0.21–161.0)	12.8 (0.2–227.0)	0.762
Preoperative ESR (mm/hr)	27 (2–129)	28 (1–140)	0.557
Preoperative corticosteroid			0.380
No	54 (40.0)	104 (35.6)	
Yes	81 (60.0)	188 (64.4)	
Preoperative infliximab			0.420
No	122 (91.0)	260 (88.4)	
Yes	12 (9.0)	34 (11.6)	
Operation approach (vs.)			<0.001
Open surgery	108 (75.0)	101 (33.6)	
Laparoscopic surgery	36 (25.0)	200 (66.4)	
Elective vs. emergency			0.400
Elective	132 (91.7)	269 (89.1)	
Emergency	12 (8.3)	33 (10.9)	
IPAA surgery			0.560
No	59 (41.0)	115 (38.1)	
Yes	85 (59.0)	187 (61.9)	
IPAA stage			0.002
2	76 (89.4)	135 (72.2)	
3	9 (10.6)	52 (27.8)	

Values are presented as number (%), mean ± standard deviation, or median (range).

BMI, body mass index; ALB, albumin; CRP, C-reaction protein; ESR, erythrocyte sedimentation rate; IPAA, ileal pouch anal anastomosis.

patients who underwent surgery increased after 2014, but the short-term postoperative complications among surgical UC

Table 5. Compared of Short-term Postoperative Complications in Surgical Ulcerative Colitis Patients between Subgroups of Open Surgery and Laparoscopic Surgery

	Open surgery	Laparoscopic surgery	P-value
IPAA patient			
Short-term complications	49 (42.2)	33 (21.3)	<0.001
Infectious complications	20 (17.2)	11 (7.1)	0.009
Non-infectious complications	39 (33.6)	31 (20.0)	0.011
Non-IPAA patient			
Short-term complications	28 (30.1)	24 (29.6)	0.950
Infectious	19 (20.4)	16 (19.8)	0.910
Noninfectious	15 (16.1)	17 (21.0)	0.410

Values are presented as number (%).

IPAA, ileal pouch anal anastomosis.

patients decreased. Although postoperative complications were variable over time and geographical area in previous studies, the risk of postoperative complications for UC patients worldwide has decreased in recent decades.^{19,20} A systematic review including UC patients from China suggested that postoperative complications improved after 2010, but the author did not investigate the possible factors causing this effect.²¹

Recently, there has been some concern that complications among UC patients who receive IPAA procedures may be affected by the introduction of biological reagents such as infliximab or vedolizumab, although this issue remains controversial.²² Preoperative corticosteroid treatment was regarded as a risk factor for postoperative complications for IBD patients,²³ and according to our investigation, preoperative treatment with glucocorticoids did not increase the risk of postoperative complications. This might be due to the individual surgical management such as avoiding emergent IPAA operation and effective rescue therapy for patients who received high-dose corticosteroid treatment. As the introduction of infliximab for moderate-severe active UC into China in 2007, more than 10% of surgical patients from this study received anti-tumor necrosis factor agents before surgery, and it was suggested that preoperative infliximab treatment did not affect the postoperative complications, which was been proved previously.²⁴ Lower BMI was found to be an independent risk factor for short-term complications in surgical UC patients, while it was reported that higher BMI was associated with frequent complications in patients receiving IPAA procedure.²⁵ According to our data, approximately 60% of UC patients had a BMI of <20 kg/m². Obesity did not become a routine problem for UC patients in

China. On the contrast, lower BMI appeared to increase complications in surgical UC patients. This clinical feature was quite different between Asia and Caucasian patients.

Laparoscopic surgery was a significant factor that contributed to the decline in postoperative complications. Our previous meta-analysis concluded that UC patients who underwent laparoscopic surgery had a lower total complication rate, although laparoscopic surgery prolonged the operation duration,²⁶ and it has already been widely accepted that the laparoscopic approach is associated with fewer complications and lower morbidity and mortality.²⁷ Moreover, this minimally invasive technique resulted in favorable long-term outcomes for UC patients, including fewer complications and a more balanced emotional condition.²⁸ The popularization of laparoscopic techniques for Chinese UC patients has helped improve their surgical outcomes. On the other hand, the benefit of laparoscopic surgery was only found to be statistically significant in the IPAA surgery subgroup, while among patients who received non-IPAA surgery, laparoscopic surgery did not reduce postoperative complications. Due to the technically demanding IPAA surgery and concerns about its risks of complications, colorectal surgeons do not perform IPAA surgery for all UC patients. The advantage of laparoscopic surgery for IPAA surgery includes a better view of the pelvis, neural preservation, and a clear anastomosis, and our results support the advantages of using laparoscopic surgery for IPAA.

The 3-stage IPAA, defined as subtotal colectomy and followed by excising remaining rectum and IPAA at the second time, and finally close ileostomy, was regarded as a safer procedure for high-risk surgical UC patients.²⁹ In our data, similar short-term complications were found between subgroups of 2-stage and 3-stage IPAA procedures. However, the patients who received the 3-stage IPAA procedure in this study were more likely to be at high risk for short-term complications, including a lower BMI, more preoperative infliximab treatment and more emergency operations (data not shown). This result is consistent with the conclusion by Mège et al.²⁹ and indicated that the 3-stage IPAA procedure is safer for patients with acute severe disease, although some other studies suggested that 2-stage IPAA also appeared to be safe and appropriate even for high-risk patients.³⁰ Therefore, the issue concerning IPAA stage still needs to be investigated in prospective research in the future.

No significant difference in long-term complications including pouchitis was found in the chronological analysis. According to Table 4, gender, operation approach and IPAA stage were

significant variables between 2008–2013 and 2014–2017 subgroups. It was reported that 3-stage IPAA and biologic agents were protective against long-term complications such as pouchitis and septic complications,³¹ while those complications did not show significant difference in our data although more 3-stage IPAA procedures were operated in the later years. Moreover, long-term complication information was lost in the 29 cases (6.5%) of overall patients. Thus, the change of the long-term complication needs to be investigated as more biological agents would be used in the future.

Our study has a few limitations. First, this is a multicenter retrospective study, and heterogeneity of the patient characteristics and surgical methods was inevitable. Second, long-term outcomes such as surgical failure, quality of life, and sexual function were not investigated in this study due to insufficient follow-up time, and the long-term cancer rate of the residual rectum or pouch is unknown. Further investigation is necessary. Third, we did not identify institutional or individual learning curves or surgeon/hospital volumes in this study, and the exact influence of surgical techniques from various surgeons on the outcome of surgical UC patients also needs to be determined.

Above all, our data revealed a downward trend of short-term postoperative complications among surgical UC patients in China during the past decade, which may be due to the promotion of minimally invasive techniques among Chinese surgeons.

ADDITIONAL INFORMATION

Funding Source

This work was supported by National Natural Science Foundation (grant number 81800484), Science & Technology Planning Project of Guangdong Province (grant numbers 20160916, 2015B020229001) and National Key Clinical Discipline in general surgery.

Conflict of Interest

No potential conflict of interest relevant to this article was reported.

Data Availability Statement

Not applicable.

Author Contributions

Conceptualization: Cai Z, He Z, Lan P, Wu X. Data curation: Cai

Z, He X, Gong J, Du P, Meng W, Zhou W, Jiang J, Wu B, Yuan W, Xue Q, Yuan L, Wang J, Tai J, Liang J, Zhu W. Funding acquisition: Cai Z, Lan P, Wu X. Investigation: Cai Z, He X. Methodology: Cai Z, He X. Writing - original draft: Cai Z. Writing - review & editing: all authors. Approval of final manuscript: all authors.

Additional Contributions

Collaborative IBD centers are as follows: Jinling Hospital (Nanjing University), Xinhua Hospital (Shanghai Jiao Tong University), West China Hospital (Sichuan University), the Sixth Affiliated Hospital (Sun Yat-sen University), Sir Run Run Shaw Hospital (Zhejiang University), the First Affiliated Hospital (Zhejiang University), Qilu Hospital (Shandong University), Peking Union Medical College Hospital, the First Affiliated Hospital (Zhengzhou University), Nanfang Hospital (Southern Medical University), the Second Xiangya Hospital (Central South University), First Hospital (Jilin University), and Xijing Hospital (Fourth Military Medical University).

We thank Lei Lian, Xianrui Wu, Bin Zheng, Xuanhui Liu, Xuming Huang, Wenyu Li, Xiaoling Li, and Futian Luo for their assistance in the study.

ORCID

Cai Z	https://orcid.org/0000-0002-4514-7116
He X	https://orcid.org/0000-0002-9441-4153
Gong J	https://orcid.org/0000-0001-6536-9990
Du P	https://orcid.org/0000-0002-3466-8020
Meng W	https://orcid.org/0000-0002-3247-9007
Zhou W	https://orcid.org/0000-0002-4856-4919
Jiang J	https://orcid.org/0000-0002-2809-9511
Wu B	https://orcid.org/0000-0003-0565-9587
Yuan W	https://orcid.org/0000-0001-7162-0360
Xue Q	https://orcid.org/0000-0003-4603-6634
Yuan L	https://orcid.org/0000-0003-3652-6680
Wang J	https://orcid.org/0000-0001-7565-0532
Tai J	https://orcid.org/0000-0003-3494-8783
Liang J	https://orcid.org/0000-0002-3273-2316
Zhu W	https://orcid.org/0000-0001-5994-4573
Lan P	https://orcid.org/0000-0001-5206-5533
Wu X	https://orcid.org/0000-0001-5610-2530

Supplementary Material

Supplementary materials are available at the Intestinal Research website (<https://www.irjournal.org>).

REFERENCES

1. Rubin DT, Ananthakrishnan AN, Siegel CA, Sauer BG, Long MD. ACG clinical guideline: ulcerative colitis in adults. *Am J Gastroenterol* 2019;114:384-413.
2. Hindryckx P, Jairath V, D'Haens G. Acute severe ulcerative colitis: from pathophysiology to clinical management. *Nat Rev Gastroenterol Hepatol* 2016;13:654-664.
3. Jeuring SF, Bours PH, Zeegers MP, et al. Disease outcome of ulcerative colitis in an era of changing treatment strategies: results from the Dutch population-based IBDSL cohort. *J Crohns Colitis* 2015;9:837-845.
4. Parragi L, Fournier N, Zeitz J, et al. Colectomy rates in ulcerative colitis are low and decreasing: 10-year follow-up data from the Swiss IBD cohort study. *J Crohns Colitis* 2018;12:811-818.
5. Parks AG, Nicholls RJ. Proctocolectomy without ileostomy for ulcerative colitis. *Br Med J* 1978;2:85-88.
6. Fu W, Zhang T, Zhang Z. The application of total proctocolectomy and ileal pouch-anal anastomosis in the treatment of ulcerative colitis. *Chin J Gastrointest Surg* 1999;4:208-210.
7. Han HQ, Liu G, Liu T, Lü YC, Fu Q. Clinical evaluation of restorative proctocolectomy with ileal pouch-anal anastomosis for ulcerative colitis. *Zhonghua Yi Xue Za Zhi* 2011;91:3205-3209.
8. Cui L. Treatment of ulcerative colitis by total colectomy ileal pouch-anal anastomosis. *Zhonghua Wei Chang Wai Ke Za Zhi* 2012;15:1221-1223.
9. Liu G, Han H, Liu T, Fu Q, Lyu Y. Clinical outcome of ileal pouch-anal anastomosis for chronic ulcerative colitis in China. *Chin Med J (Engl)* 2014;127:1497-1503.
10. Candido FD, Fiorino G, Spadaccini M, Danese S, Spinelli A. Are surgical rates decreasing in the biological era in IBD? *Curr Drug Targets* 2019;20:1356-1362.
11. Wong DJ, Roth EM, Feuerstein JD, Poylin VY. Surgery in the age of biologics. *Gastroenterol Rep (Oxf)* 2019;7:77-90.
12. Abelson JS, Michelassi F, Mao J, Sedrakyan A, Yeo H. Higher surgical morbidity for ulcerative colitis patients in the era of biologics. *Ann Surg* 2018;268:311-317.
13. Justiniano CF, Aquina CT, Becerra AZ, et al. Postoperative mortality after nonelective surgery for inflammatory bowel disease patients in the era of biologics. *Ann Surg* 2019;269:686-691.
14. Bikhchandani J, Polites SF, Wagie AE, Habermann EB, Cima RR. National trends of 3- versus 2-stage restorative proctocolectomy for chronic ulcerative colitis. *Dis Colon Rectum* 2015;58:

- 199-204.
15. de Buck van Overstraeten A, Mark-Christensen A, Wasmann KA, et al. Transanal versus transabdominal minimally invasive (completion) proctectomy with ileal pouch-anal anastomosis in ulcerative colitis: a comparative study. *Ann Surg* 2017; 266:878-883.
 16. Satsangi J, Silverberg MS, Vermeire S, Colombel JF. The Montreal classification of inflammatory bowel disease: controversies, consensus, and implications. *Gut* 2006;55:749-753.
 17. Truelove SC, Witts LJ. Cortisone in ulcerative colitis; final report on a therapeutic trial. *Br Med J* 1955;2:1041-1048.
 18. Ng SC, Shi HY, Hamidi N, et al. Worldwide incidence and prevalence of inflammatory bowel disease in the 21st century: a systematic review of population-based studies. *Lancet* 2017; 390:2769-2778.
 19. Ordás I, Domènech E, Mañosa M, et al. Post-operative morbidity and mortality of a cohort of steroid refractory acute severe ulcerative colitis: nationwide multicenter study of the GETECCU ENEIDA Registry. *Am J Gastroenterol* 2018;113: 1009-1016.
 20. Peyrin-Biroulet L, Germain A, Patel AS, Lindsay JO. Systematic review: outcomes and post-operative complications following colectomy for ulcerative colitis. *Aliment Pharmacol Ther* 2016;44:807-816.
 21. Yu Q, Mao R, Lian L, et al. Surgical management of inflammatory bowel disease in China: a systematic review of two decades. *Intest Res* 2016;14:322-332.
 22. Masaki T, Kishiki T, Kojima K, Asou N, Beniya A, Matsuoka H. Recent trends (2016-2017) in the treatment of inflammatory bowel disease. *Ann Gastroenterol Surg* 2018;2:282-288.
 23. Subramanian V, Saxena S, Kang JY, Pollok RC. Preoperative steroid use and risk of postoperative complications in patients with inflammatory bowel disease undergoing abdominal surgery. *Am J Gastroenterol* 2008;103:2373-2381.
 24. Zittan E, Milgrom R, Ma GW, et al. Preoperative anti-tumor necrosis factor therapy in patients with ulcerative colitis is not associated with an increased risk of infectious and noninfectious complications after ileal pouch-anal anastomosis. *Inflamm Bowel Dis* 2016;22:2442-2447.
 25. Kurnool S, Nguyen NH, Proudfoot J, et al. High body mass index is associated with increased risk of treatment failure and surgery in biologic-treated patients with ulcerative colitis. *Aliment Pharmacol Ther* 2018;47:1472-1479.
 26. Wu XJ, He XS, Zhou XY, Zou YF, Lan P. Safety and feasibility of laparoscopic surgery and open surgery in ulcerative colitis: a meta-analysis. *Zhonghua Wei Chang Wai Ke Za Zhi* 2008;11: 408-413.
 27. Causey MW, Stoddard D, Johnson EK, et al. Laparoscopy impacts outcomes favorably following colectomy for ulcerative colitis: a critical analysis of the ACS-NSQIP database. *Surg Endosc* 2013;27:603-609.
 28. Tajti J Jr, Látos M, Farkas K, et al. Effect of laparoscopic surgery on quality of life in ulcerative colitis. *J Laparoendosc Adv Surg Tech A* 2018;28:833-838.
 29. Mège D, Figueiredo MN, Manceau G, Maggiori L, Bouhnik Y, Panis Y. Three-stage laparoscopic ileal pouch-anal anastomosis is the best approach for high-risk patients with inflammatory bowel disease: an analysis of 185 consecutive patients. *J Crohns Colitis* 2016;10:898-904.
 30. Lee GC, Deery SE, Kunitake H, et al. Comparable perioperative outcomes, long-term outcomes, and quality of life in a retrospective analysis of ulcerative colitis patients following 2-stage versus 3-stage proctocolectomy with ileal pouch-anal anastomosis. *Int J Colorectal Dis* 2019;34:491-499.
 31. Lavryk OA, Stocchi L, Hull TL, et al. Impact of preoperative duration of ulcerative colitis on long-term outcomes of restorative proctocolectomy. *Int J Colorectal Dis* 2020;35:41-49.