World Journal of Gastrointestinal Surgery

World J Gastrointest Surg 2024 August 27; 16(8): 2365-2747





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World Journal of Gastrointestinal Surgery

Monthly Volume 16 Number 8 August 27, 2024 Contents **EDITORIAL** 2365 Immunotherapy for gastric cancer and liver metastasis: Is it time to bid farewell Dehal A 2369 Role of endoscopic ultrasound-guided biliary drainage for palliation of malignant biliary obstruction Singh S, Chandan S, Facciorusso A 2374 Consideration on immunotherapy of liver metastases of malignant tumors Jiang C, Zhang ZH, Li JX 2382 Beyond total mesorectal excision: The emerging role of minimally invasive surgery for locally advanced rectal cancer Perini D, Cammelli F, Scheiterle M, Martellucci J, Di Bella A, Bergamini C, Prosperi P, Giordano A 2386 Clinical application value of long non-coding RNAs signatures of genomic instability in predicting prognosis of hepatocellular carcinoma Xing XW, Huang X, Li WP, Wang MK, Yang JS 2393 Treatment strategy and therapy based on immune response in patients with gastric cancers Jacenik D, Fichna J **FRONTIER** 2396 Problems with repairing gut sphincters malfunctions Bortolotti M **REVIEW** Advancements in nutritional diagnosis and support strategies during the perioperative period for patients 2409 with liver cancer Li XQ, Liang Y, Huang CF, Li SN, Cheng L, You C, Liu YX, Wang T **ORIGINAL ARTICLE Case Control Study** 2426 Surgical resection and neoadjuvant therapy in patients with gastric cancer and ovarian metastasis: A realworld study Yan HP, Lu HR, Zhang YX, Yang L, Chen ZL 2436 Alteration of ascending colon mucosal microbiota in patients after cholecystectomy Fan MY, Jiang QL, Cui MY, Zhao MQ, Wang JJ, Lu YY



Contents

World Journal of Gastrointestinal Surgery

Monthly Volume 16 Number 8 August 27, 2024

Retrospective Cohort Study

Survival prognostic analysis of laparoscopic D2 radical resection for locally advanced gastric cancer: A 2451 multicenter cohort study

Sun XM, Liu K, Wu W, Meng C

2461 Benefits of jejunostomy feeding in patients who underwent gastrectomy for cancer treatment

Jaquet R, Rivkine E, De Souza N, Roudié J

2474 Application of ¹⁸F-fluorodeoxyglucose positron emission tomography/computed tomography imaging in recurrent anastomotic tumors after surgery in digestive tract tumors

Ge DF, Ren H, Yang ZC, Zhao SX, Cheng ZT, Wu DD, Zhang B

- 2484 Impact of minimally invasive surgery on immune function and stress response in gastric cancer patients Zhu RH, Li PC, Zhang J, Song HH
- 2494 Assessment of perianal fistulizing Crohn's disease activity with endoanal ultrasound: A retrospective cohort study

Hong N, Liu WY, Zhang JL, Qian K, Liu J, Ye XJ, Zeng FY, Yu Y, Zhang KG

2503 Lymph node dissection does not affect the survival of patients with tumor node metastasis stages I and II colorectal cancer

He F, Ou SP, Yuan Y, Oian K

Retrospective Study

2511 Energy spectrum computed tomography multi-parameter imaging in preoperative assessment of vascular and neuroinvasive status in gastric cancer

Wang J, Liang JC, Lin FT, Ma J

- Clinical significance of peripheral blood immune cells in patients with gastric cancer after surgery 2521 Wang QW, Zhu JW, Gong LZ
- 2528 Lone-Star retractor perineal exposure method for laparoscopic abdominal perineal resection of rectal cancer

Ma J, Tang DB, Tang YQ, Wang DT, Jiang P, Zhang YM

- 2538 Indication of conservative treatment by antibiotics for uncomplicated and complicated acute appendicitis Hosokawa Y, Moritani M, Makuuchi Y, Nagakawa Y
- 2546 Preoperative prediction of hepatocellular carcinoma microvascular invasion based on magnetic resonance imaging feature extraction artificial neural network

Xu JY, Yang YF, Huang ZY, Qian XY, Meng FH

2555 Transmembrane serine protease 4 expression in the prognosis of radical resection for biliary tract cancer Shibata Y, Sudo T, Tazuma S, Tanimine N, Onoe T, Shimizu Y, Yamaguchi A, Kuraoka K, Takahashi S, Tashiro H



World Journal of Gastrointestin			
Conter	Monthly Volume 16 Number 8 August 27, 2024		
2565	Systemic immune-inflammation index combined with pediatric appendicitis score in assessing the severity and prognosis for paediatric appendicitis		
	Guo LM, Jiang ZH, Liu HZ		
2574	Establishment of predictive models and determinants of preoperative gastric retention in endoscopic retrograde cholangiopancreatography		
	Jia Y, Wu HJ, Li T, Liu JB, Fang L, Liu ZM		
2583	Prediction model establishment and validation for enteral nutrition aspiration during hospitalization in patients with acute pancreatitis		
	Hou P, Wu HJ, Li T, Liu JB, Zhao QQ, Zhao HJ, Liu ZM		
2592	New anti-mesenteric delta-shaped stapled anastomosis: Technical report with short-term postoperative outcomes in patients with Crohn's disease		
	Lee JL, Yoon YS, Lee HG, Kim YI, Kim MH, Kim CW, Park IJ, Lim SB, Yu CS		
2602	Construction of a predictive model for gastric cancer neuroaggression and clinical validation analysis: A single-center retrospective study		
	Lan YY, Han J, Liu YY, Lan L		
2612	Efficiency and safety of laparoscopic left hemihepatectomy: A study of intrathecal <i>vs</i> extrathecal Glissonean pedicle techniques		
	Kang LM, Xu L, Zhang FW, Yu FK, Lang L		
2620	Predictive utility of the Rockall scoring system in patients suffering from acute nonvariceal upper gastrointestinal hemorrhage		
	Han DP, Gou CQ, Ren XM		
	Observational Study		
2630	Nomogram predicting the prognosis of primary liver cancer after radiofrequency ablation combined with transcatheter arterial chemoembolization		
	Shen HH, Hong YR, Xu W, Chen L, Chen JM, Yang ZG, Chen CH		
2640	Relationship between postoperative rehabilitation style, gastrointestinal function, and inflammatory factor levels in children with intussusception		
	Wei XY, Huo HC, Li X, Sun SL, Zhang J		
	Prospective Study		
2649	Innovative integration of lung ultrasound and wearable monitoring for predicting pulmonary complic- ations in colorectal surgery: A prospective study		
	Lin C, Wang PP, Wang ZY, Lan GR, Xu KW, Yu CH, Wu B		
	Randomized Controlled Trial		
2662	Effects of fluid therapy combined with a preoperative glucose load regimen on postoperative recovery in patients with rectal cancer		
	Xia LC, Zhang K, Wang CW		

Contents

World Journal of Gastrointestinal Surgery

Monthly Volume 16 Number 8 August 27, 2024

Randomized Clinical Trial

Application value of dexmedetomidine in anesthesia for elderly patients undergoing radical colon cancer 2671 surgery

Bu HM, Zhao M, Ma HM, Tian XP

Basic Study

2679 Effect of growth hormone on colonic anastomosis after intraperitoneal administration of 5-fluorouracil, bleomycin and cisplatin: An experimental study

Lambrou I, Mantzoros I, Ioannidis O, Tatsis D, Anestiadou E, Bisbinas V, Pramateftakis MG, Kotidis E, Driagka B, Kerasidou O, Symeonidis S, Bitsianis S, Sifaki F, Angelopoulos K, Demetriades H, Angelopoulos S

SYSTEMATIC REVIEWS

2689 Management of distal cholangiocarcinoma with arterial involvement: Systematic review and case series on the role of neoadjuvant therapy

Hall LA, Loader D, Gouveia S, Burak M, Halle-Smith J, Labib P, Alarabiyat M, Marudanayagam R, Dasari BV, Roberts KJ, Raza SS, Papamichail M, Bartlett DC, Sutcliffe RP, Chatzizacharias NA

SCIENTOMETRICS

2702 Global research landscape of Peutz-Jeghers syndrome and successful endoscopic management of intestinal intussusception in patients with recurrent laparotomies

Sun Q, Wang XY, Guo GJ, Wang L, Meng LM, Guo YF, Sun T, Ning SB

CASE REPORT

2719 Ultrasound-guided peripheral nerve blocks for anterior cutaneous nerve entrapment syndrome after robot-assisted gastrectomy: A case report

Saito Y, Takeuchi H, Tokumine J, Sawada R, Watanabe K, Yorozu T

2724 Primary coexisting adenocarcinoma of the colon and neuroendocrine tumor of the duodenum: A case report and review of the literature

Fei S, Wu WD, Zhang HS, Liu SJ, Li D, Jin B

2735 Anorectal hemangioma, a rare cause of lower gastrointestinal bleeding, treated with selective embolization: A case report

Pospisilova B, Frydrych J, Krajina A, Örhalmi J, Kajzrlikova IM, Vitek P

LETTER TO THE EDITOR

Hepatic recompensation according to the Baveno VII criteria via a transjugular intrahepatic portosystemic 2742 shunt: Is this true?

Zhang JS

2745 Machine learning in predicting postoperative complications in Crohn's disease Zhang LF, Chen LX, Yang WJ, Hu B



Contents

World Journal of Gastrointestinal Surgery

Monthly Volume 16 Number 8 August 27, 2024

ABOUT COVER

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WJGS mainly publishes articles reporting research results and findings obtained in the field of gastrointestinal surgery and covering a wide range of topics including biliary tract surgical procedures, biliopancreatic diversion, colectomy, esophagectomy, esophagostomy, pancreas transplantation, and pancreatectomy, etc.

INDEXING/ABSTRACTING

The WJGS is now abstracted and indexed in Science Citation Index Expanded (SCIE, also known as SciSearch®), Current Contents/Clinical Medicine, Journal Citation Reports/Science Edition, PubMed, PubMed Central, Reference Citation Analysis, China Science and Technology Journal Database, and Superstar Journals Database. The 2024 Edition of Journal Citation Reports[®] cites the 2023 journal impact factor (JIF) for WJGS as 1.8; JIF without journal self cites: 1.7; 5-year JIF: 1.9; JIF Rank: 123/290 in surgery; JIF Quartile: Q2; and 5-year JIF Quartile: Q3.

RESPONSIBLE EDITORS FOR THIS ISSUE

Production Editor: Zi-Hang Xu; Production Department Director: Xiang Li; Cover Editor: Jia-Ru Fan.

NAME OF JOURNAL	INSTRUCTIONS TO AUTHORS			
World Journal of Gastrointestinal Surgery	https://www.wjgnet.com/bpg/gerinfo/204			
ISSN	GUIDELINES FOR ETHICS DOCUMENTS			
ISSN 1948-9366 (online)	https://www.wjgnet.com/bpg/GerInfo/287			
LAUNCH DATE	GUIDELINES FOR NON-NATIVE SPEAKERS OF ENGLISH			
November 30, 2009	https://www.wjgnet.com/bpg/gerinfo/240			
FREQUENCY	PUBLICATION ETHICS			
Monthly	https://www.wjgnet.com/bpg/GerInfo/288			
EDITORS-IN-CHIEF	PUBLICATION MISCONDUCT			
Peter Schemmer	https://www.wjgnet.com/bpg/gerinfo/208			
EDITORIAL BOARD MEMBERS	ARTICLE PROCESSING CHARGE			
https://www.wjgnet.com/1948-9366/editorialboard.htm	https://www.wjgnet.com/bpg/gerinfo/242			
PUBLICATION DATE	STEPS FOR SUBMITTING MANUSCRIPTS			
August 27, 2024	https://www.wjgnet.com/bpg/GerInfo/239			
COPYRIGHT	ONLINE SUBMISSION			
© 2024 Baishideng Publishing Group Inc	https://www.f6publishing.com			

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World J Gastrointest Surg 2024 August 27; 16(8): 2451-2460

DOI: 10.4240/wjgs.v16.i8.2451

Retrospective Cohort Study

ISSN 1948-9366 (online)

ORIGINAL ARTICLE

Survival prognostic analysis of laparoscopic D2 radical resection for locally advanced gastric cancer: A multicenter cohort study

Xiu-Ming Sun, Kui Liu, Wen Wu, Chao Meng

Specialty type: Gastroenterology and hepatology

Provenance and peer review: Unsolicited article; Externally peer reviewed.

Peer-review model: Single blind

Peer-review report's classification Scientific Quality: Grade A, Grade C, Grade C Novelty: Grade A, Grade B, Grade В Creativity or Innovation: Grade A, Grade B, Grade C Scientific Significance: Grade A,

Grade B, Grade C

P-Reviewer: Sato T; Shim CN

Received: January 27, 2024 Revised: June 7, 2024 Accepted: June 25, 2024 Published online: August 27, 2024 Processing time: 202 Days and 8.1 Hours



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Abstract

BACKGROUND

With the development of minimally invasive surgical techniques, the use of laparoscopic D2 radical surgery for the treatment of locally advanced gastric cancer (GC) has gradually increased. However, the effect of this procedure on survival and prognosis remains controversial. This study evaluated the survival and prognosis of patients receiving laparoscopic D2 radical resection for the treatment of locally advanced GC to provide more reliable clinical evidence, guide clinical decision-making, optimize treatment strategies, and improve the survival rate and quality of life of patients.

AIM

To investigate the survival prognosis and influencing factors of laparoscopic D2 radical resection for locally advanced GC patients.

METHODS

A retrospective cohort study was performed. Clinicopathological data from 652 patients with locally advanced GC in our hospitals from December 2013 to December 2023 were collected. There were 442 males and 210 females. The mean age was 57 ± 12 years. All patients underwent a laparoscopic D2 radical operation for distal GC. The patients were followed up in the outpatient department and by telephone to determine their tumor recurrence, metastasis, and survival. The follow-up period ended in December 2023. Normally distributed data are expressed as the mean ± SD, and normally distributed data are expressed as M (Q1, Q3) or M (range). Statistical data are expressed as absolute numbers or



percentages; the χ^2 test was used for comparisons between groups, and the Mann-Whitney *U* nonparametric test was used for comparisons of rank data. The life table method was used to calculate the survival rate, the Kaplan-Meier method was used to construct survival curves, the log rank test was used for survival analysis, and the Cox risk regression model was used for univariate and multifactor analysis.

RESULTS

The median overall survival (OS) time for the 652 patients was 81 months, with a 10-year OS rate of 46.1%. Patients with TNM stages II and III had 10-year OS rates of 59.6% and 37.5%, respectively, which were significantly different (P < 0.05). Univariate analysis indicated that factors such as age, maximum tumor diameter, tumor differentiation grade (low to undifferentiated), pathological TNM stage, pathological T stage, pathological N stage (N2, N3), and postoperative chemotherapy significantly influenced the 10-year OS rate for patients with locally advanced GC following laparoscopic D2 radical resection for distal stomach cancer [hazard ratio (HR): 1.45, 1.64, 1.45, 1.64, 1.37, 2.05, 1.30, 1.68, 3.08, and 0.56 with confidence intervals (CIs) of 1.15-1.84, 1.32-2.03, 1.05-1.77, 1.62-2.59, 1.05-1.61, 1.17-2.42, 2.15-4.41, and 0.44-0.70, respectively; P < 0.05]. Multifactor analysis revealed that a tumor diameter greater than 4 cm, low tumor differentiation, and pathological TNM stage III were independent risk factors for the 10-year OS rate in these patients (HR: 1.48, 1.44, 1.81 with a 95%CI: 1.19-1.84). Additionally, postoperative chemotherapy emerged as an independent protective factor for the 10-year OS rate (HR: 0.57, 95%CI: 0.45-0.73; P < 0.05].

CONCLUSION

A maximum tumor diameter exceeding 4 cm, low tumor differentiation, and pathological TNM stage III were identified as independent risk factors for the 10-year OS rate in patients with locally advanced GC following laparoscopic D2 radical resection for distal GC. Conversely, postoperative chemotherapy was found to be an independent protective factor for the 10-year OS rate in these patients.

Key Words: Gastric neoplasms; Chinese Laparoscopic Gastrointestinal Surgery Study Group; Laparoscopic surgery; Locally advanced gastric cancer; D2 lymph node dissection

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Core Tip: Data from multicenter patients with locally advanced gastric cancer (GC) were collected to compare the differences in survival and prognosis between laparoscopic D2 radical surgery and traditional open surgery. The contents of the study included postoperative survival rate, postoperative complication rate, postoperative recovery and quality of life. By comparing and analyzing the efficacy and safety of the two surgical methods, this study aimed to evaluate the actual effect of laparoscopic D2 radical surgery in the treatment of locally advanced GC, provide evidence for the selection of clinical surgical methods, and promote the application of minimally invasive surgical techniques in the treatment of GC.

Citation: Sun XM, Liu K, Wu W, Meng C. Survival prognostic analysis of laparoscopic D2 radical resection for locally advanced gastric cancer: A multicenter cohort study. *World J Gastrointest Surg* 2024; 16(8): 2451-2460 URL: https://www.wjgnet.com/1948-9366/full/v16/i8/2451.htm DOI: https://dx.doi.org/10.4240/wjgs.v16.i8.2451

INTRODUCTION

Gastric cancer (GC) is a malignant tumor with the fifth highest incidence and the fourth highest fatality rate in the world. In 2020, the number of GC deaths worldwide exceeded 760000[1-3]. Surgery is an important means of comprehensive treatment for GC[4]. The safety and long-term oncological efficacy of laparoscopy in the treatment of early (stage I) GC have been confirmed by high-level, evidence-based medical evidence[5]. Up to 80% of GC patients in China are locally advanced at first diagnosis and require more difficult D2 radical surgery. In 2009, the Chinese Laparoscopic Gastrointestinal Surgery Study Group was established in China, and in 2010, the first multicenter clinical database of laparoscopic GC surgery with the largest sample size was established in China. The CLASS-01 study began in 2012 and revealed that laparoscopic D2 radical gastrectomy had the same long-term survival benefit as open surgery for locally advanced GC[6-8]. In fact, it was better than open surgery in many ways. Laparoscopic minimally invasive surgery was recommended for the first time in the National Comprehensive Cancer Network Clinical Practice Guidelines for GC (2nd Edition, 2021) as an option for the treatment of locally advanced GC[9].

At 11 years of follow-up, a Dutch study revealed that D2 lymph node dissection did not provide any significant benefits. However, at 15 years of follow-up, D2 lymph node dissection significantly decreased the rate of local recurrence and death from GC, and the overall survival (OS) rate tended to increase.

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MATERIALS AND METHODS

General information

A retrospective cohort study was used. The clinicopathological data of 652 patients with locally advanced GC in our hospitals in the laparoscopic GC surgery multicenter database of the CLASS research group were collected. There were 442 males and 210 females. The mean age was 57 ± 12 years. The body mass index (BMI) of the 652 patients was 21 ± 3 kg/m². There were 396, 232, and 24 patients with Eastern Cooperative Oncology Group (ECOG) scores of 0, 1, and \geq 2, respectively. Of the 652 patients, 130 had preoperative comorbidities. Tumors were located in the lower part of the stomach in 517 patients and in the middle part of the stomach in 135 patients, and the patients and their families signed informed consent forms.

Inclusion criteria

(1) Preoperative gastroscopy and postoperative histopathological examination both confirmed primary gastric adenocarcinoma; (2) Patients who had not previously undergone preoperative chemoradiotherapy or other antitumor therapies; (3) Laparoscopic-assisted radical resection of distal GC D2 or D2+ was performed; and (4) Preoperative gastroscopy and whole-abdominal enhanced computed tomography (CT) or whole-body positron emission tomography/CT were performed, and postoperative histopathological examination confirmed the diagnosis of stage II or III (stage T2N1-3M0, stage T3-4N0-3M0) GC.

Exclusion criteria

The exclusion criteria for patients were as follows: (1) Underwent laparotomy or laparoscopic conversion to laparotomy; (2) Had a history of gastrectomy; (3) Had a history of other malignant tumors or concurrent malignant tumors; (4) Had distant metastasis before and during the operation; (5) Had a postoperative histopathological examination confirming stage T4b disease; (6) Had no R0 excision; (7) Had a postoperative survival time less than 1 month; and (8) Were lost to follow-up at the first postoperative visit.

Treatment method

All patients underwent a laparoscopic D2 radical operation for distal GC. The surgical procedures and methods used are described in the references. For basic principles, refer to the relevant guidelines. The overall surgical principles, lymph node dissection standards, and surgical resection scope were in accordance with relevant GC guidelines and operating norms of the period, including the 2nd and 3rd editions of the GC Treatment Guidelines of the Japan GC Society and the 2007 edition of the Laparoscopic and Endoscopic Surgery Group of the Chinese Medical Association Surgery Society. The specific reconstruction method depends on the operative experience and personal preference of the surgeon. Adjuvant chemotherapy is recommended after surgery.

Observation indices

(1) Surgical information included the following parameters: maximum tumor diameter, number of lymph node dissections, number of lymph node metastases, tumor proximal incisional margin, tumor distal incisal margin, tumor Borrmann classification, tumor pathological T stage, pathological N stage, and pathological TNM stage; (2) Follow-up information: The number of patients followed up, follow-up time, postoperative recurrence and metastasis, and overall postoperative survival were obtained; and (3) Analysis of prognostic factors included sex, age, BMI, ECOG score, intraoperative blood transfusion, complications, maximum tumor diameter, tumor location, Borrmann classification, tumor differentiation degree, pathological TNM stage, pathological T stage, pathological N stage, method of digestive tract reconstruction, lymph node dissection, proximal resection margin of tumor, distal resection margin of tumor, combined organ resection, postoperative chemotherapy, number of lymph node dissections, complications, and Clavien-Dindo grade \geq III complications.

Evaluation criteria

Postoperative complications were assessed according to the Clavien-Dindo surgical complication grading system. The pathological TNM staging criteria refer to the 7th edition of the American Cancer Society Tumor Staging.

Follow-up visit

The patients were followed up in the outpatient department and by telephone to determine their tumor recurrence, metastasis, and survival. The follow-up period ends in December 2023.

Statistical analysis

SPSS 26.0 statistical software and R language software (ver. 3.4.3) were used for analysis. Normally distributed data are expressed as the mean ± SD, and normally distributed data are expressed as M (Q1, Q3) or M. Statistical data are expressed as absolute numbers or percentages; the χ^2 test was used for comparisons between groups, and the Mann-Whitney U nonparametric test was used for comparisons of rank data. The Cox risk regression model was used for univariate and multivariate analyses, and P < 0.05 was considered to indicate statistical significance.

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RESULTS

Operation situation

Of the 652 patients, 617 underwent D2 lymph node dissection, and 35 underwent D2 + lymph node dissection. Billroth II anastomosis was used in 348 patients, Billroth I anastomosis in 218 patients, Roux-en-Y anastomosis in 25 patients, and other anastomoses in 61 patients. Twelve patients underwent combined organ resection (1 patient who underwent combined partial transverse colon resection, 1 patient who underwent combined partial small intestine resection, 3 patients who underwent combined gallbladder resection, and 7 patients who underwent combined gallbladder resection). There were 569 patients who received intraoperative transfusions and 83 patients who did not receive intraoperative transfusions.

Pathological examination after the operation

The maximum tumor diameter of the 652 patients was 4.5 ± 2.0 cm. There were 26 (19, 35) lymph node dissections, of which 570 were > 15 and 82 were \leq 15. The number of lymph node metastases was 4 (1, 9). The proximal incisal margin of the tumor was 4.8 ± 1.6 cm, and the distal incisal margin of the tumor was 4.5 ± 1.5 cm. Among the 652 patients, 255 had Borrmann type I to II, 334 had Borrmann type III to IV, and 63 lacked Borrmann type data. The degree of tumor differentiation was high (medium-differentiated) in 171 patients, low (undifferentiated) in 430 patients, and absent in 51 patients. There were 123, 253, and 276 patients with T2, T3, and T4a disease in the T stage of tumor pathology; 116, 131, 214, and 191 patients with N0, N1, N2, and N3 disease in the N stage of pathology; and 260 and 392 patients with II and III diseases in the TNM stage, respectively.

Postoperative recovery and complications

For 652 patients, the first time of getting out of bed was 3 (2, 4) days, the first time of anal exhaust was 4 (3, 5) days, the first time of eating liquid food was 5 (4, 6) days, and the postoperative hospital stay was 10 (9, 13) days. Among the 652 patients, 69 had complications after surgery. Of these, 54 had complications related to the operation, 20 had complications related to the body as a whole, and 5 had both operation-related and systemic complications.

Additionally, 60 patients had Clavien-Dindo grade I to II complications, and 3, 5, and 1 patients had Clavien-Dindo grade I to II complications. Among the patients with grade I to II complications, there were 19 cases of duodenal residual fistula, 18 cases of respiratory complications, 11 cases of gastroparesis, 6 cases of poor wound healing and bleeding, 5 cases of abdominal infection, 4 cases of chylorrhea, 3 cases of digestive tract bleeding, and 1 case each of anastomotic fistula, intestinal obstruction, and urinary system complications III.

Among the patients with Grade A complications, there were 2 cases of abdominal infection and 1 case of duodenal residual fistula, chylorrhagia, or intestinal obstruction. There were 2 cases of grade IIIb complications, 2 of intestinal obstruction, 1 of anastomotic fistula, one of anastomotic stenosis, one of abdominal hemorrhage, and one of digestive tract hemorrhage; one patient with a grade A complication had a respiratory system complication. Multiple complications may be associated with the same patient. Among surgical complications and systemic complications, the most common were duodenal residual fistulas (3.07%, 20/652) and respiratory complications (2.91%, 19/652), respectively. All 69 patients were successfully cured and discharged from the hospital after treatment.

Follow-up situation

All 652 patients were followed up for 110-193 months, and the median follow-up time was 124 months. Among the 298 patients with postoperative recurrence and metastasis, 255 patients had recurrence and metastasis ≤ 5 years after surgery, and 43 patients had recurrence and metastasis > 5 years after surgery. There was no significant difference in the recurrence or metastasis type between the two patients (P > 0.05). There was no significant difference between patients with distant metastasis \leq 5 years and those with distant metastasis > 5 years after surgery, peritoneal metastasis, local recurrence, multiple recurrence and metastasis, or recurrence and metastasis at other sites (P > 0.05). There was a significant difference in TNM stage between patients with recurrence and metastasis \leq 5 years and those with recurrence and metastasis > 5 years after surgery (P < 0.05).

There was no statistically significant difference in the pathological T stage (P > 0.05). Further analysis revealed no statistically significant difference in the T2 stage or T3 stage of pathology between the two groups (P > 0.05), and there was a statistically significant difference in the T4a stage between the two groups (P < 0.05). The comparison of pathological N stages revealed statistically significant differences (P < 0.05), as shown in Table 1.

The median OS time for the 652 patients was 81 months, with a 10-year OS rate of 46.1%. The 10-year OS rates for patients with TNM stages II and III were 59.6% and 37.5%, respectively, showing a statistically significant difference (χ^2 = 35.29, P < 0.001) (Figure 1A). Further analysis revealed that the 10-year OS rates for patients with TNM stages IIA, IIB, IIIA, IIIB, and IIIC were 65.6%, 55.8%, 46.9%, 37.1%, and 24.0%, respectively ($\chi^2 = 55.06$, P < 0.001) (Figure 1B). For patients with stage T2, T3, or T4a disease, the 10-year OS rates were 55.2%, 46.5%, and 41.5%, respectively, which were significantly different (χ^2 = 8.39, P = 0.014) (Figure 1C). Additionally, the 10-year OS rates for patients with stages N0, N1, N2, and N3 disease were 63.7%, 56.2%, 48.5%, and 26.4%, respectively, which were statistically significant ($\gamma^2 = 54.89$, $P < 10^{-10}$ 0.001) (Figure 1D).

Analysis of prognostic factors

Sex, BMI, ECOG score, intraoperative blood transfusion, comorbidities, tumor location, Borrmann classification, digestive tract reconstruction mode, lymph node dissection, proximal resection margin of the tumor, distal resection margin of the tumor, combined organ resection, number of lymph nodes dissected, complications, and Clavien-Dindo grade \geq III



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Table 1 Comparison between 298 locally advanced gastric cancer patients with the time to postoperative recurrence and metastasis ≤ 5 years and > 5 years after laparoscopic assisted D2 radical distal gastrectomy

Clinical pathological factors	Recurrence and metastasis within \leq 5 years after surgery, <i>n</i> = 255	Recurrence and metastasis after surgery > 5 years, <i>n</i> = 43	Statistical value	P value
Types of recurrence and metastasis	-	-	$\chi^2 = 5.52$	0.238
Distant metastasis	21	5	$\chi^2 = 0.53$	0.466
Peritoneal metastasis	69	9	$\chi^2 = 0.72$	0.398
Local recurrence	37	10	$\chi^2 = 2.12$	0.146
Multiple relapses and metastases	52	4	$\chi^2 = 2.9$	0.085
Recurrence and metastasis in other parts	76	15	$\chi^2 = 0.45$	0.503
Pathological TNM staging	-	-	-	-
Stage II	62	23	$\chi^2 = 15.36$	< 0.001
Stage III	193	20		
Pathological T-staging	-	-	Z = -1.80	0.07
T2	42	9	$\chi^2 = 0.52$	47
Т3	95	21	$\chi^2 = 2.08$	0.150
T4a	118	13	$\chi^2 = 3.84$	0.049
Pathological N-staging	-	-	Z = -3.34	0.001
NO	19	12	$\chi^2 = 16.52$	< 0.001
N1	44	5	$\chi^2 = 0.85$	0.357
N2	85	18	$\chi^2 = 1.18$	0.277
N3	107	8	$\chi^2 = 8.47$	0.004

complications did not affect the D2 root diameter of laparoscopic distal GC patients. Factors related to the 10-year OS rate of patients with locally advanced GC after treatment were identified (P > 0.05) (Table 2).

The results of the multifactor analysis showed that the maximum diameter of the tumor was > 4 cm, and the degree of tumor differentiation was low (P < 0.05). After laparoscopic D2 radical resection for distal GC, chemotherapy after surgery was a protective factor for an OS rate greater than 10 years for people with locally advanced GC (P < 0.05) (Table 3).

DISCUSSION

The findings of the GASTRIC study group show that disease-free survival can be used as a substitute for total survival in GC clinical studies, but total survival is always the gold standard for evaluating long-term oncological efficacy. The latest follow-up results from the CLASS-01 study evaluated long-term oncological outcomes using 5-year OS. Based on the above evaluation methods of domestic and foreign research groups, our team used the 10-year OS rate to evaluate the long-term oncological efficacy of this study [10-14].

The 10-year survival rates of 652 patients at corresponding stages collected in this study from December 2013 to December 2023 were 46.1%, 65.6%, 55.8%, 46.9%, 37.1%, and 24.0%, respectively, which were slightly greater than those in the above study on open GC[15]. The reason may be that this study included patients with stage T4b disease (611 patients), stage M1 disease (427 patients), and nonradical resection (631 patients), which affected the long-term survival outcome of the whole group and of patients in each stage[16]. The results of a study that included the United States SEER database and a large sample of GC patients in China showed that the 10-year OS rate of patients with GC in these two countries was 32.2% [17-20]. The above indicators were all lower than the 10-year survival rates of patients at each corresponding stage in this study. The author believes that the most likely reason is that the SEER database lacks detailed records of patients' adjuvant chemotherapy and D2 lymph node dissection, the incidence of GC is low in the United States, neoadjuvant chemotherapy is more common, and more GC patients undergo surgery with a low degree of radical treatment[21]. Therefore, some patients who did not receive adjuvant chemotherapy or standard D2 lymph node dissection were included in the study, resulting in poor long-term survival outcomes. Another Brazilian study showed that the 10-year OS rate was 30.6% in 526 patients who underwent laparotomy with D2 radical resection for GC; however, the study included 62 stage IV patients[22]. This study concluded that the inclusion of stage IV patients was the main

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Table 2 Univariate analysis of overall survival rate of locally advanced gastric cancer patients undergoing laparoscopic D2 radical distal gastrectomy						
Clinical pathological factors	$\boldsymbol{\beta}$ value	SE	Wald value	Hazard ratio	95%CI	P value
Sex as female/male	0.03	0.11	0.07	1.03	0.82-1.30	0.788
Age	0.37	0.12	9.58	1.45	1.15-1.84	0.002
Body mass index	-0.25	0.19	1.76	0.78	0.54-1.13	0.19
Scoring for EUSSCG	-	-	1.97	-	-	0.37
1-0	-0.04	0.12	0.09	0.97	0.77-1.21	0.77
≥2	0.36	0.28	1.71	1.44	0.84-2.47	0.19
Intraoperative blood transfusion	0.14	0.16	0.82	1.15	0.85-1.56	0.37
Hejing syndrome	0.10	0.13	0.59	1.11	0.85-1.44	0.44
Maximum diameter of the tumor in > 4 cm to \leq 4 cm	0.49	0.11	20.35	1.64	1.32-2.03	< 0.001
Tumor location	0.00	0.14	0.06	0.97	0.74-1.27	0.81
Remmunm typing	-	-	2.16	-	-	0.34
Type III-IV is more common than Type I-II	0.07	0.12	0.32	1.07	0.85-1.34	0.57
Unknown ratio of type I to type II	-0.23	0.21	1.22	0.80	0.53-1.19	0.27
degree of tumor differentiation	-	-	5.99	-	-	0.05
Low undifferentiated to high medium differentiated	0.31	0.13	5.57	1.37	1.05-1.77	0.02
Unknown high to medium differentiation	0.10	0.23	0.20	0.11	0.71-0.73	0.65
Pathological TNM staging	0.72	0.12	35.29	2.05	1.62-2.59	< 0.001
Pathological T-staging	0.26	0.11	5.69	1.30	1.05-1.61	0.02
Pathological N-staging	-	-	55.20	-	-	< 0.001
N1	0.26	0.21	0.48	1.30	0.85-1.97	0.22
N2	0.52	0.19	7.88	1.68	.17-2.42	0.01
N3	1.13	0.18	37.78	3.08	2.15-4.41	< 0.00
Gastrointestinal reconstruction methods	-	-	4.49	-	-	0.21
Billroth II matches better than Billroth I matches	0.08	0.12	0.41	1.08	0.85-1.37	0.52
Roux-en-Y matches better than Billroth I matches	0.02	0.29	0.00	1.02	0.58-1.82	0.94
Other matches better than Billroth I	0.40	0.19	4.40	1.50	1.03-2.18	0.13
Lymph node dissection	-0.17	0.24	0.46	0.85	0.53-1.36	0.94
Tumor proximal margin	0.00	0.01	0.02	1.00	0.99-1.0	0.88
Distal margin of tumor resection	0.00	0.00	0.02	1.00	0.99-1.0	0.90
Combined organ resection	0.77	0.58	1.78	2.17	0.70-6.75	0.18
Postoperative chemotherapy	-0.59	0.12	24.40	0.56	0.44-0.70	< 0.001
Number of lymph node dissection	0.11	0.17	0.44	1.12	0.80-1.57	0.51
Complications	0.09	0.17	0.26	0.10	0.78-1.52	0.61
Clavien-Dindo ≥ Grade III complications	-0.18	0.50	0.13	0.84	0.31-2.24	0.72

reason for the lower 10-year OS rate of patients who underwent laparoscopic D2 radical gastrectomy for GC.

If laparoscopic surgery is used to treat locally advanced GC, especially for pathological T4a tumors with serous membrane involvement, there is a chance that the tumor cells will spread, and the peritoneum will become implanted under the pneumoperitoneum[23]. Therefore, long-term observation of the survival, recurrence, and metastasis characteristics of patients with this type of GC after laparoscopic surgery has important clinical significance and is also the main reason why this study focused on the 10-year survival of patients with locally advanced GC in pathological TNM stages II and III. Among the 652 patients in this study, 298 had postoperative recurrence and metastasis, among which peritoneal metastasis accounted for the greatest proportion (26.17%, 78/298). Peritoneal metastasis is the most common type of postoperative recurrence of GC, which is consistent with the recurrence and metastasis reported in many traditional open

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Table 3 Multivariate analysis of overall survival rate of locally advanced gastric cancer patients undergoing laparoscopic D2 radical distal gastrectomy

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Clinical pathological factors	$\boldsymbol{\beta}$ value	SE	Wald value	Hazard ratio	95%CI	P value
Ages of > 65 years to \leq 65 years	0.24	0.12	3.73	1.27	0.99-1.61	0.053
Maximum diameter of the tumor of > 4 cm to < 4 cm	0.39	0.11	12.19	1.48	1.19-1.84	< 0.001
Degree of tumor differentiation			10.08			0.006
Low undifferentiated to high medium differentiated	0.37	0.14	7.36	1.44	1.11-1.88	0.007
Unknown high to medium differentiation	-0.07	0.23	0.10	0.93	0.60-1.46	0.753
Pathological TNM staging of stage III compared to stage II	0.59	0.12	23.41	1.81	1.42-2.30	< 0.001
Postoperative chemotherapy	-0.56	0.12	20.81	0.57	0.45-0.73	< 0.001

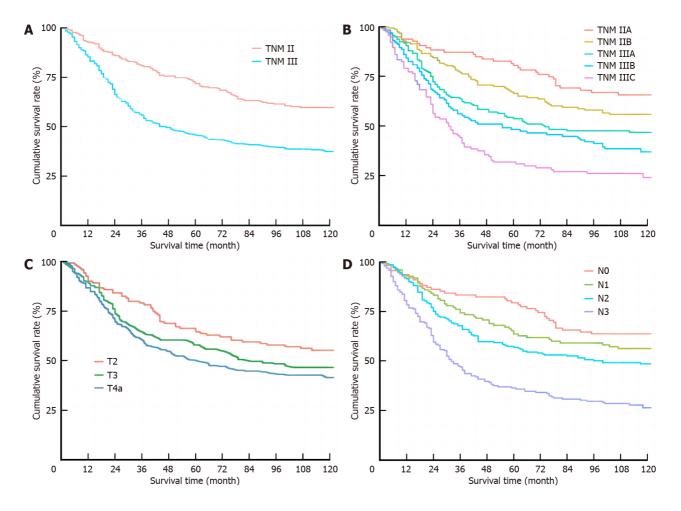


Figure 1 Survival outcome of laparoscopic D2 radical surgery for locally advanced gastric cancer. A: Overall survival curve of 652 patients with locally advanced gastric cancer in pathological TNM stages II and III after laparoscopic D2 radical treatment for distal gastric cancer; B: Overall survival curve of patients with locally advanced gastric cancer at pathological TNM stage IA, IIB, IIIA, IIIB, IIIC after laparoscopic D2 radical treatment for distal gastric cancer; C: Overall survival curve of patients with T2, T3 and T4a locally advanced gastric cancer after laparoscopic D2 radical resection for distal gastric cancer; D: Overall survival curve of patients with stage N0, stage N1, stage N2 and stage N3 locally advanced gastric cancer after laparoscopic D2 radical resection for distal gastric cancer; C: cancer.

GC studies. In this study, 43.96% (131/298) of patients with postoperative recurrence and metastasis had stage T4a disease according to pathology[24-26]. The high incidence of recurrence and metastasis in patients with late pathological T stage disease may be the main reason for the large proportion of peritoneal metastasis cases[26-28].

No recurrence or metastasis at 5 years after radical surgery is generally considered "clinically cured." However, due to the lack of > 5 years of survival data, few studies have reported the risk and characteristics of recurrence and metastasis > 5 years after laparoscopic surgery for advanced GC[29]. In this study, 43 patients experienced recurrence or metastasis > 5 years after surgery. This suggests that even if there is no tumor survival for 5 years, regular follow-up is still necessary to

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guard against tumor recurrence and metastasis. Further analysis revealed no significant difference in the incidence of GC recurrence or metastasis between the two groups[30]. However, it is worth noting that the patients who experienced recurrence and metastasis \leq 5 years after surgery were mainly in pathological TNM stage III, T4a, and N3. The patients with recurrence and metastasis > 5 years after surgery were mainly in TNM stages II and N0. The above results showed that patients with locally advanced GC with serous membrane involvement, more lymph node metastasis, and a later TNM stage were more likely to experience recurrence and metastasis in the early postoperative period[31]. However, patients with locally advanced GC without lymph node metastasis and an early pathological TNM stage have a greater risk of recurrence and metastasis after 5 years of survival without a tumor[32-34].

The results of the CLASS-01 study showed that the 5-year OS rates of GC patients with different pathological TNM stages in the laparoscopic group were 91% in stage A, 81% in stage B, 73% in stage A, 47% in stage B, and 28% in stage C [35]. This study investigated how long people with different pathological TNM stages had lived for 10 years. The OS rate was 65.6% for stage A, 55.8% for stage B, 46.9% for stage A, 37.1% for stage B, and 24.0% for stage C. The later the stage of the tumor was, the smaller the difference between the 5-year OS rate and the 10-year OS rate, and the more gradual the decline in the 5- to 10-year postoperative OS rate. In other words, for patients who have successfully survived for 5 years, the later the tumor stage, the greater the probability of continuing to survive for 5 years, and the less obvious the decline in survival rate after 5 to 10 years. Conditional survival can explain this phenomenon[36]. The traditional survival assessment is based on clinicopathological indicators determined at the time of diagnosis and operation and does not consider the influence of the existing survival time on the subsequent survival time[37]. Conditional survival refers to the possibility of surviving for additional years and/or months on the basis of having survived for a certain period of time, which fully considers the dynamic change in survival time and changes in survival time[38].

CONCLUSION

Laparoscope-assisted D2 radical resection for locally advanced GC has a satisfactory 10-year oncologic effect. Patients with TNM stage III, T4a, and N3 disease had a high recurrence and metastasis ratio \leq 5 years after surgery, while those with TNM stage II and N0 disease had a high recurrence and metastasis ratio > 5 years after surgery. The largest tumor was more than 4 cm in size, and it was not very differentiated. The long-term oncological efficacy of laparoscopic treatment for locally advanced GC needs to be further verified by prospective randomized controlled studies with large sample sizes.

FOOTNOTES

Author contributions: Sun XM wrote the manuscript; Liu K and Wu W collected the data; and Meng C guided the study. All authors reviewed, edited, and approved the final manuscript and revised it critically for important intellectual content, gave final approval of the version to be published, and agreed to be accountable for all aspects of the work.

Institutional review board statement: This study was reviewed and approved by the Ethics Committee of the First Affiliated Hospital of Naval Medical University.

Informed consent statement: Informed consent was obtained from the patients and their families for this study.

Conflict-of-interest statement: The authors declare no conflicts of interest.

Data sharing statement: The data of this study can be provided upon reasonable request and can only be shared after the application is submitted and approved by the corresponding author.

STROBE statement: The authors have read the STROBE Statement-checklist of items, and the manuscript was prepared and revised according to the STROBE Statement-checklist of items.

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S-Editor: Qu XL L-Editor: Filipodia P-Editor: Zhang L

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August 27, 2024 Volume 16 Issue 8

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Sun XM et al. Laparoscopic D2 radical resection in GC

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