

Application value of continuous seromuscular layer sutures in the reinforcement of esophagojejunostomy in total gastrectomy for gastric cancer: a retrospective comparative cohort study

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Background: The development process of gastrointestinal anastomosis is from complex to simple, from two layers to one layer, from extramucosal anastomosis to seromuscular anastomosis. With the rapid development of anastomosis instruments, the anastomosis process becomes more and more convenient. However, relevant studies have shown that related complications such as anastomotic leakage still occur. This study sought to investigate the feasibility and safety of seromuscular layer sutures in the reinforcement of esophagojejunostomy after open radical total gastrectomy.

Methods: This study retrospectively analyzed patients who underwent Roux-en-Y esophagojejunostomy after open radical total gastrectomy at The Third Department of Surgery, The Fourth Hospital of Hebei Medical University from April 2019 to May 2020. The inclusion criteria of patients were between 18 and 80 years old; pathology confirmed gastric adenocarcinoma; preoperative imaging showed no distant metastasis and did not receive neoadjuvant therapy; no complex infectious diseases; no blood transfusion was performed before operation. A total of 192 patients were included according to the inclusion criteria. They were divided into the following 2 groups based on whether seromuscular layer suturing of the anastomosis was performed: (I) group A (the simple anastomosis group, n=76); (II) and group B (the seromuscular layer suture group, n=116). The baseline data, surgical data and postoperative complications were compared between the two groups.

Results: All patients underwent esophagojejunostomy Roux-en-Y anastomosis after open radical total gastrectomy, and no perioperative deaths occurred. There was no significant difference in baseline data between the two groups. Group B had an earlier time to liquid diet than group A (4.23±0.76 vs. 4.57±0.58 days, P<0.001). The incidence of postoperative anastomotic leakage in group B (1.72%) was lower than that in group A (9.21%), and the difference was statistically significant (P=0.03). The incidence of pleural effusion was lower in group B (15.52%) than group A (32.89%), and the difference was statistically significant (P=0.005).

Conclusions: Compared to the simple anastomosis, seromuscular layer sutures after esophagojejunostomy may decrease the rates of postoperative anastomotic leakage and pleural effusion. This suture method is feasible and may provide a new option to increase surgical safety.

Keywords: Gastric neoplasms; total gastrectomy; seromuscular layer suture; Roux-en-Y esophagojejunostomy

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Introduction

Gastric cancer is one of the most common malignant tumors of the gastrointestinal tract, and its morbidity and mortality rates are high (1-4). Radical surgical resection is the primary treatment for early and advanced gastric cancer (5-8). Digestive tract reconstruction, especially esophagojejunostomy, plays an essential role in the surgery (9). With the development of surgical techniques, the safety of the surgery has gradually improved; however, surgery-related complications occur occasionally. The emergence of these complications significantly increases the suffering of patients, prolongs hospitalization time, increases hospitalization costs, and delays subsequent antitumor therapy (10,11). A common complication after total gastrectomy is esophagojejunostomy leakage, which is closely related to a variety of factors (6,12). The main influencing factor is the clinician's intraoperative operation, especially the suturing and reinforcement of the anastomosis (12-14). Thus, the incidence of postoperative complications needs to be reduced and the safety of the surgery enhanced.

In open total gastrectomy, the circular stapler is used for esophagojejunostomy. As the circular stapler is safe and easy to operate, it can achieve surgical margins and satisfactory anastomosis (15,16). However, the stapler also has a number of disadvantages, including that it is expensive, and prone to causing anastomotic stenosis. Even with a 3-row stapler, there may be bleeding from the anastomotic stoma. Further, the incidence of anastomotic stenosis has been reported to

Highlight box

Key findings

 Seromuscular layer suturing of the esophagojejunostomy may reduce the incidence of postoperative anastomotic leakage and pleural effusion.

What is known and what is new?

- With the rapid development of instrument anastomosis, the anastomosis process becomes more and more convenient.
 However, related complications such as anastomotic leakage still occur.
- We combined the characteristics of instrument anastomosis and hand-sewn sutures. After the instrument anastomosis was completed, we sutured the seromuscular layer of anastomosis continuously using knot-free barbed sutures.

What is the implication, and what should change now?

 This suture method is feasible and may provide a new option to increase surgical safety. be correlated with the use of circular staplers (17,18).

Hand-sewn suturing is a basic operation technique of surgery, and it is mostly used for laparoscopic radical gastrectomy. It enables the surgeon to suture according to the specific situation of the anastomosis. The surgeon can also reduce the tension in the sutures based on their own experience. Hand-sewn suturing changes from double layer anastomosis to single layer anastomosis and then to seromuscular layer anastomosis. Double layer anastomosis may lead to blood supply disorder, and prone to anastomotic stenosis. Compared with double-layer anastomosis, singlelayer anastomosis has fewer sutures and less foreign body reaction at the anastomosis, but the sutures on the mucosa are prone to intestinal inflammation and affect healing. The seromuscular layer anastomosis requires only the serous layer, the muscular layer and the submucosa to be connected, without compressing the submucosal vessels and with slight damage to the blood supply to the intestinal wall. However, either type of hand-sewn suturing prolongs the operative time, and does not reduce the anastomotic-related complications. Thus, both mechanical stapled anastomosis and hand-sewn sutures have certain disadvantages.

Our center combined the characteristics of mechanical stapled anastomosis and hand-sewn sutures. After the esophagojejunostomy was completed using a circular stapler, the seromuscular layer was continuously sutured with knot-free barbed sutures at the anastomosis. This study aimed to further explore and compare the safety and short-term efficacy of simple anastomosis and seromuscular layer continuous sutures. We present the following article in accordance with the STROBE reporting checklist (available at https://jgo.amegroups.com/article/view/10.21037/jgo-22-1068/rc).

Methods

General information

This study adopted a retrospective cohort method. To be eligible for inclusion in the study, the patients had to meet the following inclusion criteria: (I) have been hospitalized for the first time at The Third Department of Surgery, The Fourth Hospital of Hebei Medical University, from April 2019 to May 2020 and be aged 18–80 years; (II) have adenocarcinoma, which was pathologically confirmed by preoperative gastroscopic biopsy, and a tumor located in the upper part of the stomach or at the esophagogastric junction; (III) have no distant metastasis as observed in the

Table 1 Comparison of the baseline data between the simple anastomosis group and the seromuscular layer suture group

Baseline data	Group A (n=76)	Group B (n=116)	Statistics	P value
Age, years	63.57±2.79	63.47±3.61	t=0.216	0.829
Gender			$\chi^2 = 0.883$	0.347
Male	58 (76.32)	95 (81.90)		
Female	18 (23.68)	21 (18.10)		
ECOG score			$\chi^2 = 0.278$	0.598
0	62 (81.58)	91 (78.45)		
1	14 (18.42)	25 (21.55)		
Tumor size			$\chi^2 = 0.104$	0.747
<5 cm	56 (73.68)	83 (71.55)		
≥5 cm	20 (26.32)	33 (28.45)		
pTNM stage			$\chi^2 = 0.03$	0.984
1	6	9		
II	22	35		
III	48	72		
BMI, kg/m ²	23.88±2.95	24.00±2.65	t=0.303	0.762
Follow-up time, months	16	12		

Data are presented as n, n (%) or mean ± standard deviation. Group A: the simple anastomosis group; Group B: the seromuscular layer suture group. ECOG score: Quality of life scores [Zubrod /ECOG/WHO (ZPS)]. ECOG, Eastern Cooperative Oncology Group; BMI, body mass index; pTNM, pathological tumor-node-metastasis.

preoperative imaging examination. (IV) have not received any neoadjuvant therapy before surgery; and (V) have no complicated infectious diseases and have not received any blood transfusions before surgery. Patients were excluded from the study if they met any of the following exclusion criteria: (I) were aged >80 years; (II) had liver cirrhosis or diabetes; (III) had advanced adenocarcinoma that was obviously invading the perigastric organs or tissue; and/or (IV) had positive margins that were confirmed pathologically during the operation.

The tumor-node-metastasis (TNM) staging system of the American Joint Committee on Cancer (AJCC) (8th edition) was used for the pathological staging of the patients. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by the Ethics Committee of The Fourth Hospital of Hebei Medical University (No. 2021KS011). Individual consent for this retrospective analysis was waived.

Based on the above inclusion and exclusion criteria, the data of 192 patients with gastric cancer who underwent open radical total gastrectomy and Roux-en-Y esophagojejunostomy were collected. Based on whether the seromuscular layer suture was used at the anastomosis, the patients were divided into group A (the simple anastomosis group, n=76) and group B (the seromuscular layer suture group, n=116). There were 76 patients in group A, of whom 58 were male and 18 were female. The patients in group A had a mean age of 63.57±2.79 years, and a mean BMI of 23.88±2.95 kg/m². There were 116 patients in group B, of whom 95 were male and 21 were female. The patients had a mean age of 63.47±3.61 years, and a mean BMI of 24.00±2.65 kg/m². There were no significant differences between the 2 groups in terms of age, gender, BMI, Eastern Cooperative Oncology Group (ECOG) score, tumor size, or pathological tumor-node-metastasis (pTNM) (P>0.05). Thus, the 2 groups of patients were comparable (see Table 1 for further details).

Patients

All the patients underwent preoperative abdominal contrast-enhanced competed tomography examinations.

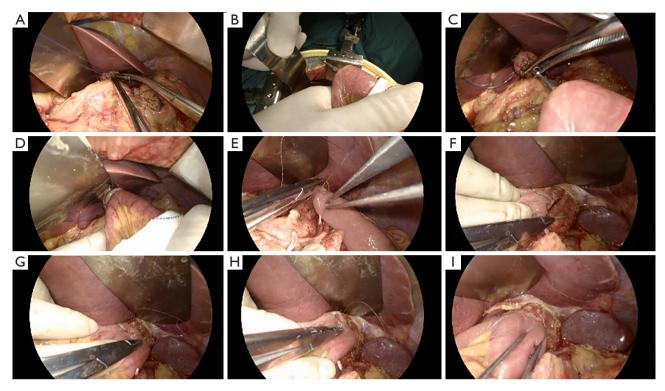


Figure 1 Anastomosis and suture operation process. (A-I) Please refer to the text for detailed steps.

No cases of distant metastasis were found, and gastroscopic biopsies were conducted to confirm the diagnosis of adenocarcinoma. Before the operation, the chief surgeon explained the advantages and disadvantages of the 2 methods to the patients and their families. Seromuscular layer suture may increase the safety of anastomosis and reduce the incidence of anastomotic complications, but it will prolong the operation time and increase the operation cost. On the contrary, simple anastomosis has shorter operation time and lower operation cost. The patients, their families, and the surgeon jointly decided if the seromuscular layer suture would be applied in esophagusjejunostomy anastomosis during the operation. The preoperative preparation for the 2 groups was the same.

Operation procedures for patients

All the operation procedures were performed by specialized gastrointestinal surgeons with more than 10 years of experience. The 2 groups of patients underwent tracheal intubation under general anesthesia. An abdominal exploration and the exfoliated cytology were performed first. After completing the D2 lymph node dissection and

total gastric dissection, the duodenum was severed with a linear stapler. The esophagus was dissociated 2-3 cm above the pre-cut line, and the bilateral vagus nerves were severed. After clamping the esophagus with purse-string forceps, a temporary purse string was placed. After the esophagus was cut with an electrosurgical knife, an anvil seat was placed, and the purse string was tightened and knotted (Figure 1A). The mesentery was incised at a distance of 15 cm from the Trevor ligament, and the jejunum was severed with a linear stapler. After the mesenteric vascular arch was ligated, the distal jejunum was inserted into the main body of the circular stapler, until it protruded on the opposite side of the mesangial border (Figure 1B). Next, the anvil seat and the stapler were connected and tightened for 15 seconds to complete the end-to-side anastomosis of the esophagus and jejunum (Figure 1C,1D), and the linear stapler was used to close the jejunum stump. At this point, the esophagojejunostomy was completed.

In group B, surgeons sutured the seromuscular layer of anastomosis continuously using knot-free barbed sutures. In this process, the needle distance was 4–5 mm, and the margin was 3 mm to avoid full-thickness sutures (*Figure 1E-1H*). The operation ensured that the anastomotic

stoma had no tension, distortion, and good blood supply (Figure 1I).

The jejunojejunostomy was performed at about 40 cm below the esophagojejunostomy. The feeding tube was placed 15 cm distal to the jejuno-jejunostomy, and the gastric tube was placed 5 cm below the esophagojejunostomy. Finally, the anastomotic stoma was carefully observed to ensure that there was no leakage and no active bleeding in the operative field. Abdominal drainage tubes were placed under the esophagojejunostomy and in the spleen fossa.

Observation indicators

The intraoperative observation indicators were operation time and intraoperative blood loss. The postoperative observation indicators were time to liquid diet, time to first exhaust, and length of postoperative hospital stay. The postoperative complications included anastomotic leakage, anastomotic bleeding, anastomotic stenosis, and pleural effusion.

Evaluation criteria and definitions

The postoperative complications included any adverse conditions requiring conservative treatment or surgical intervention after surgery. Anastomotic leakage was diagnosed if the patient had intestinal contents flowing out of the drainage tube after surgery, or blue liquid was drained out after the oral administration of methylene blue, or contrast agent leakage was observed in the upper gastrointestinal angiography. Anastomotic stenosis included postoperative symptoms (e.g., difficulty eating, vomiting, or swallowing). An obstruction was diagnosed by angiography or gastroscopy. All the postoperative complications were comprehensively examined based on auxiliary examinations, blood indicators, symptoms, and signs. The operation time was calculated as the time from the opening of the abdomen to the closing of the abdomen (as noted on the anesthesia record sheet). The intraoperative blood loss was calculated based on the amount of blood in the suction bag minus the amount of irrigation. The time to liquid diet was determined based on the time recorded in the doctor's order sheet. The time to first exhaust was based on the medical records.

Postoperative management and follow-up

The 2 groups' postoperative nutritional support,

and symptomatic intervention were the same. In both groups, following the patients' smoot recovery, gastroenterography was checked 1 week after operation. In the gastroenterography process, the patients repeatedly adjusted their body positions to the upright, supine, and lateral positions. It is convenient for doctors to carefully observe anastomotic-related complications, especially anastomotic stenosis and anastomotic leakage. After discharge, the patients were followed up for anastomotic-related complications. The patients underwent electronic gastroscopy to check for anastomotic stenosis 6 months after the operation.

Statistical analysis

SPSS (version 26.0) software was used for the statistical analysis. The measurement data are expressed as the $(\bar{x}\pm s)$, and the T-test was used for comparisons between groups. The count data are expressed as the rate (%), and the χ^2 test or Fisher's exact test was used for comparisons. A P value <0.05 was considered statistically significant.

Results

Comparison of surgical data

All the 192 patients successfully underwent total radical gastrectomy, and there were no perioperative deaths. In group A, the mean operation time was 231.93±26.35 min, and the mean intraoperative blood loss was 61.09±8.04 mL. In group B, the mean operation time was 237.04±21.91 min, and the mean intraoperative blood loss was 62.51±6.50 mL. In relation to the intraoperative situation, there were no significant differences between the 2 groups (P>0.05). In relation to the postoperative conditions, in terms of the time to first exhaust and the length of postoperative hospital stay there were no significant differences between the 2 groups (P>0.05); however, group B had a significantly earlier time to liquid diet than group A (4.23±0.76 vs. 4.57±0.58 days, P<0.001) (see *Table 2* for details).

Comparison of the incidence of complications between the 2 groups

In group A, there were 7 (9.21%) cases of anastomotic leakage and 25 (32.89%) cases of pleural effusion. In group B, there were 2 (1.72%) cases of anastomotic leakage and 18 (15.52%) cases of pleural effusion. The incidence of

Table 2 Comparison of the surgical data between the simple anastomosis group and the seromuscular layer suture group

Surgical data	Group A (n=76)	Group B (n=116)	Statistics	P value
Operation time, min	231.93±26.35	237.04±21.91	t=-1.457	0.147
Intraoperative blood loss, mL	61.09±8.04	62.51±6.50	t=-1.285	0.163
Time to liquid diet, days	4.57±0.58	4.23±0.76	t=3.55	< 0.001
Time to first exhaust, days	4.11±0.80	3.99±1.02	t=0.883	0.378
Postoperative hospital stay, days	7.00±1.13	6.98±1.06	t=0.112	0.911

Data are presented as mean ± standard deviation. Group A: the simple anastomosis group; Group B: the seromuscular layer suture group.

Table 3 Comparison of complications between the simple anastomosis group and the seromuscular layer suture group

Complication	Group A (n=76)	Group B (n=116)	Statistics	P value
Anastomotic leakage	7 (9.21)	2 (1.72)	-	0.040
Anastomotic bleeding	0	0	-	-
Anastomotic stenosis	0	0	_	-
Pleural effusion	25 (32.89)	18 (15.52)	$\chi^2 = 7.98$	0.005

Data are presented as n (%). Group A: the simple anastomosis group; Group B: the seromuscular layer suture group.

postoperative anastomotic leakage in group B (1.72%) was lower than that in group A (9.21%), and the difference was statistically significant (P=0.03). The incidence of pleural effusion in group B (15.52%) was lower than that in group A (32.89%), and the difference was statistically significant (P=0.005). No anastomotic stenosis or anastomotic bleeding was observed in the 2 groups during the follow-up period (see *Table 3* for details). In this study, the median follow-up time was 16 months for patients in group A and 12 months for patients in group B (see *Table 1* for further details).

Discussion

As mature surgical procedures, radical total gastrectomy and Roux-en-Y esophagojejunostomy are widely used in clinical practice (19). With the continuous optimization of the surgical technique and the improvement of the anastomosis method, the overall complication rate continues to gradually decrease. The success or failure of total gastrectomy lies in the safety and effectiveness of the operation, and the focus of the operation is the reconstruction of the digestive tract. Anastomotic-related complications include anastomotic leakage, anastomotic bleeding, and anastomotic stenosis (20). Anastomotic leakage is one of the most severe complications, with an incidence rate of 1–11.5% (12). It seriously affects the postoperative recovery and short-term

and long-term curative effect of patients and is even related to the survival of tumor patients. Thus, it is imperative to find and actively implement ways to reduce anastomotic leakage.

Clinicians have engaged in extensive work to reduce complications after digestive tract reconstruction. Previous scholars have proposed a variety of esophagojejunostomy methods. The main techniques include reverse puncture, the Orvil and purse-string suture method using circular stapler or the esophagojejunostomy (overlap) method, and the functional end-to-end anastomosis method using linear stapler (16,21-23). With the continuous updating of stapler technology, the safety of mechanical stapled anastomosis has improved, and complications related to anastomosis have been reduced compared to those associated with hand-sewn anastomosis (24). Under the stapler method, 2 or 3 staggered staples are generally implanted into the tissue to achieve the double or triple cross suturing of the tissue. The suturing is tight, and leakage is reduced to a large extent. The staples of the stapler are also arranged neatly. The spacing and tightness of the stitching can be effectively controlled through its scale. Conversely, handsewn suturing is challenging to execute. The staples can also effectively promote tissue healing. When the patient's general condition is good, and the operation is not complex, the patient can recover relatively smoothly. However, if the

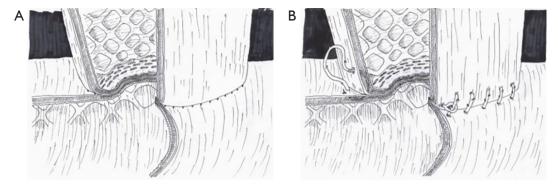


Figure 2 Seromuscular layer sutures. (A) Complete instrumental anastomosis of the esophagus and jejunum. (B) Continuous sutures of the seromuscular layer of the anastomosis.

patient has unfavorable factors, such as an advanced age, anemia, malnutrition, diabetes, high anastomotic tension, and unexpected tension (such as hiccups), the risk of surgery increases.

Compared to a stapler, it is difficult to uniformize the density and depth of hand-sewn digestive-tract suture; however, hand-sewn sutures have a broader range of applications. For example, vascular anastomosis, pancreasjejunum anastomosis, and sometimes thinner bile duct anastomosis require hand-sewn sutures, and a stapler cannot be used for these anastomoses (25). In addition, hand-sewn sutures are also flexible and reliable (26). The surgeon can appropriately change the amount of sutured tissue and the spacing and tension of the sutures according to their own experience and skills. Hand-sewn sutures can avoid large blood vessels, hemostatic sutures can be applied to the bleeding point, and the non-smooth tissue surface can be embedded to reduce postoperative adhesions (27). Additionally, the sutures used in hand-sewn suturing are constantly developing and progressing. The surgical sutures currently used have good compatibility and flexibility, and some varieties have absorbability and antibacterial properties. The application of barbed sutures in laparoscopic gastroduodenal ulcer perforation repair is safe and effective. Barbed sutures can shorten the suture and operation time, reduce the difficulty of laparoscopic suturing, and shorten the learning curve (28-31).

Successful anastomosis and subsequent healing depend on several factors, including the tension between the 2 connected parts of the gastrointestinal tract, the healthy blood supply to the surrounding tissue, and the strength of the final anastomosis. Thus, we combined the characteristics of instrument anastomosis and hand-sewn sutures. After the instrument anastomosis was completed, we sutured the seromuscular layer of anastomosis continuously using knot-free barbed sutures, as shown in *Figure 2*. In relation to the postoperative situation, we found that the time to liquid diet of group B patients was earlier than that of group A patients. Thus, group B was more stable than group A in the early postoperative period and patients in group A could eat earlier and more safely. Patients in group B also had a lower incidence of anastomotic leakage than those in group A, which may be related to the seromuscular layer suture closing the pinhole of the stapler. Thus, tissue inflammatory response was alleviated, tissue healing was accelerated, and the occurrence of anastomotic leakage was reduced.

There was no significant difference in anastomotic stenosis between group B and group A; thus, that continuous suture did not increase the incidence of anastomotic stenosis. Additionally, there was no anastomotic bleeding in either group, which may be related to the application of instrument anastomosis and the selection of the correct stapler, or to the small sample size of this study. The continuous suturing of the seromuscular layer reduces anastomotic exudation, reduces stimulation to the diaphragm and lower posterior mediastinum, and further reduces the occurrence of pleural effusion. Thus, the incidence of pleural effusion in group B (15.52%) was lower than in group A (32.89%), and the difference was statistically significant. Seromuscular layer sutures can compress the hemostasis, reduce the occurrence of anastomotic leakage, and do not increase the occurrence of anastomotic stenosis. The operation time was prolonged, but the safety of the operation was improved, and the patients were discharged smoothly with few complications.

The present study had a few limitations. First, this study

was retrospective, and it had a small sample size. Second, there was selection bias in this study. Third, the surgical methods were all open-surgery methods. Fourth, this study included few postoperative observation indicators. Thus, a multicenter, prospective, randomized controlled trial should be conducted in the future, and the surgical methods included in the laparoscopic surgery and the postoperative follow-up time should be appropriately extended.

Conclusions

Seromuscular layer suturing of the esophagojejunostomy can allow patients to drink water and eat safely in the early postoperative stage and may reduce the incidence of postoperative anastomotic leakage and pleural effusion. This suture method is feasible and may provide a new option to increase surgical safety.

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Footnote

Reporting Checklist: The authors have completed the STROBE reporting checklist. Available at https://jgo.amegroups.com/article/view/10.21037/jgo-22-1068/rc

Data Sharing Statement: Available at https://jgo.amegroups.com/article/view/10.21037/jgo-22-1068/dss

Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at https://jgo.amegroups.com/article/view/10.21037/jgo-22-1068/coif). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by the Ethics

Committee of The Fourth Hospital of Hebei Medical University (No. 2021KS011). Individual consent for this retrospective analysis was waived.

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