

# Robotic versus laparoscopic gastrectomy with D2 lymph node dissection for advanced gastric cancer: a propensity score-matched analysis

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**Background:** Robotic gastrectomy (RG) is a new surgical method alternative for gastric cancer. However, few studies have evaluated the outcomes of RG for advanced gastric cancer (AGC). Thus, the aim of this study was to compare the short-and long-term outcomes of RG and laparoscopic gastrectomy (LG) with D2 lymph node dissection for AGC.

**Patients and methods:** We retrospectively evaluated 454 patients with AGC who underwent RG or LG with D2 lymph node dissection for AGC between August 2013 and March 2017. The short-and long-term outcomes were compared between the propensity score-matched groups.

**Results:** The RG group was associated with longer operation time, less intraoperative blood loss, and higher hospital cost. Additionally, there was a tendency favoring RG in terms of number of harvested lymph nodes, time to first flatus, time to first start diet, and postoperative hospital stay, although the differences were not statistically significant. The overall postoperative complication rate was 13.4% and 11.6% in the RG and LG groups, respectively, with no significant difference ( $P=0.686$ ). The 3-year overall survival and recurrence rates of the RG and LG groups were also comparable (78.6% vs 74.1%,  $P=0.483$ ; 18.8% vs 21.4%,  $P=0.617$ ; respectively).

**Conclusion:** RG with D2 lymph node dissection is safe and feasible for AGC in terms of both short- and long-term outcomes. High-volume randomized controlled trials with sufficient follow-up are needed to confirm this rationale.

**Keywords:** robotic gastrectomy, laparoscopic gastrectomy, advanced gastric cancer

## Introduction

Gastric cancer is a worldwide health concern and is the second leading cause of cancer-related deaths in China.<sup>1</sup> Gastrectomy with proper perigastric lymph node dissection remains the cornerstone of radical resection of potentially curable gastric cancer. With the advantages of minimally invasive and better short-term outcomes, laparoscopic gastrectomy (LG) for gastric cancer has garnered tremendous popularity over open gastrectomy.<sup>2,3</sup> Currently, extensive researches have reported that laparoscopy-assisted gastrectomy (LAG) is a safe and feasible procedure with better short-term outcomes and equivalent long-term survival outcomes compared with open gastrectomy (OG).<sup>4-6</sup> Robotic surgery has been introduced as a better operation method with several advantages, which could overcome the technical limitations of conventional laparoscopy. There is evidence that the robotic gastrectomy (RG) could help to overcome some technical difficulties encountered when performing laparoscopic surgery, such as lymph node dissection, handling deep-seated vessels, and intracorporeal anastomoses.<sup>7-10</sup> However, these studies mainly focused on early

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gastric cancer, and majority of them only evaluated short-term outcomes. There is still a lack of well-matched studies that report long-term outcomes of RG with D2 lymph node dissection for patients with advanced gastric cancer (AGC). We, therefore, designed this study to compare the short- and long-term outcomes of RG and LG for AGC using a propensity score-matching analysis.

## Patients and methods

### Patients

Patients with postoperative pathological diagnoses of AGC who underwent RG or LG were screened from the prospectively maintained gastric cancer database at the Department of Digestive Surgery, Xijing Hospital of Digestive Diseases, Fourth Military Medical University from August 2013 to March 2017. The inclusion criteria were as follows: 1) histologically confirmed adenocarcinoma by gastroscopy and pathological biopsy; 2) depth of invasion confined to pT2, pT3, or pT4a; 3) no distant metastasis or invasion to adjacent organs; 4) not combined with other malignancy; 5) no emergency operation; and 6) no preoperative chemotherapy or radiation therapy was performed. Ultimately, 454 patients were included in the analysis. The entire cohort included 125 and 329 patients who underwent RG and LG, respectively. To reduce the effect of potential confounding due to the limits of respective studies, we performed propensity score matching using a logistic regression model with the following variables: age, gender, comorbidities, tumor size, extent of resection, histologic type, pT stage, pN stage, and pTNM stage. We performed one-to-one matching using a 0.02 caliper width. Finally, the propensity score-matched cohort comprised 112 cases in each group. Pathologic staging was evaluated according to the 8th Union for International Cancer Control/American Joint Committee on Cancer staging system of gastric cancer.<sup>11</sup> Postoperative complications were recorded and classified according to the Clavien–Dindo classification system.<sup>12,13</sup>

### Surgical procedure

The patients chose the surgical procedure (RG vs LG) by their individual decision after they were informed of the advantages and risks of each procedure. Patients in the 2 groups underwent distal gastrectomy (DG) or total gastrectomy (TG) based on the location of tumor. D2 lymph node dissection was performed according to the Japanese Gastric Cancer Treatment Guidelines (version 3).<sup>14</sup> During TG, spleen-preserving No. 10 lymph node dissection and omentectomy were performed. The reconstruction type

(gastroduodenostomy or gastrojejunostomy for DG and Roux-en-Y esophagojejunostomy for TG) was selected according to the surgeon's preference. The extracorporeal reconstructions were mostly conducted in our center. We routinely administered postoperative adjuvant chemotherapy with 5-fluorouracil and cisplatin. This study was approved by the ethics committee of Xijing Hospital, and written informed consent was granted by the patients.

### Postoperative evaluation and follow-up

Postoperative complications were recorded as complications that occurred within 30 days after surgery. Patients were followed up every 3 months during the first 2 years and then every 6 months from 2 to 5 years. The length of follow-up was defined as the time from surgery to the final follow-up date of December 2017 or time of death. Cancer recurrence was diagnosed based on radiologic or histological signs of disease.

### Statistical analysis

The data were analyzed using SPSS, version 22.0 (SPSS Inc., Chicago, IL, USA). All continuous variables are presented as the mean  $\pm$  SD. The chi-square test was used to compare categorical variables between the 2 groups, and the independent sample *t*-test or Mann–Whitney *U*-test was used to compare continuous variables. Overall survival (OS) curves were calculated using the Kaplan–Meier method and analyzed by the log-rank test. *P*-values  $<0.05$  were considered statistically significant.

## Results

### Clinicopathologic characteristics

The clinicopathologic characteristics of the patients are summarized in Table 1. The background characteristics were well matched, with no significant differences in gender, age, body mass index (BMI), tumor size, extent of resection, histological type, pT stage, pN stage, pTNM stage, and comorbidities between the RG and LG groups.

### Surgical outcomes and postoperative complications

The surgical outcomes and postoperative complications are summarized in Table 2. The intraoperative blood loss was significantly reduced in the RG group compared with the LG group (179.2 vs 234.9 mL, *P*=0.000). However, the operative time was significantly longer in the RG group than that in the LG group (261.7 vs 227.8 min, *P*=0.000). Additionally, the total cost of hospitalization was significantly higher in the RG group than that in the LG group (92365.0 vs 69476.5

**Table 1** Characteristics of patients

Variables	Entire cohort		P-value	Propensity score-matched cohort		P-value
	RG (n=125)	LG (n=329)		RG (n=112)	LG (n=112)	
Age (years)	55.4±11.5	56.9±10.5	0.197	55.6±11.3	56.1±11.1	0.725
Gender (male/female)	84/41	229/100	0.621	78/34	79/33	0.884
Body mass index (kg/m <sup>2</sup> )	23.7±2.8	23.1±3.0	0.041	23.6±2.9	23.6±3.0	0.856
Tumor size (≤5.0/>5.0 cm)	91/34	239/90	0.973	83/29	84/28	0.878
Comorbidities (present/absent)	25/100	85/244	0.195	23/89	24/88	0.870
Histological type			0.000			0.494
Well/moderately	28	133		23	19	
Poorly/undifferentiated	97	196		89	93	
pT stage			0.006			0.960
T2	47	82		38	36	
T3	45	113		41	42	
T4a	33	134		33	34	
pN stage			0.048			0.821
N0	41	97		36	35	
N1	42	86		36	35	
N2	25	66		25	27	
N3a	13	41		12	9	
N3b	4	39		3	6	
pTNM stage			0.031			0.993
IB	24	37		20	18	
IIA	27	55		22	23	
IIB	24	72		22	21	
IIIA	26	64		25	25	
IIIB	17	53		16	19	
IIIC	7	48		7	6	

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

RMB,  $P=0.000$ ). No significant difference was found between the 2 groups in terms of number of harvested lymph nodes ( $29.5\pm 9.6$  vs  $27.8\pm 8.7$ ,  $P=0.149$ ), time to first flatus ( $2.6\pm 0.6$  vs  $2.8\pm 1.1$ ,  $P=0.124$ ), time to start liquid diet ( $1.6\pm 0.7$  vs  $1.7\pm 0.5$ ,  $P=0.320$ ), time to start soft diet ( $3.6\pm 1.6$  vs  $3.9\pm 2.0$ ,  $P=0.179$ ), and days of postoperative hospital stay ( $6.9\pm 2.3$  vs  $7.0\pm 3.8$ ,  $P=0.718$ ).

There was no intraoperative or 30-day postoperative mortality. The incidences of overall complications did not differ significantly between the RG and LG groups (13.4% vs 11.6%,  $P=0.686$ ). Moreover, no significant differences were noted in the minor (Clavien–Dindo grade II) and major complication (Clavien–Dindo grade  $\geq$  IIIa) rates between the 2 groups (6.3% vs 5.4%,  $P=0.775$ ; 7.1% vs 6.3%,  $P=0.789$ ; respectively). Regarding individual complications, the incidence of wound problem, anastomotic leakage, duodenal stump fistula, and abdominal infection were also similar between the 2 groups.

## Surgical outcomes and postoperative complications in different phases

To examine the learning curve effect, we performed subgroup analysis by dividing the patients into 2 phases. The initial

and late phase both comprised 56 patients in each group. Table 3 shows surgical outcomes and postoperative complications in different phase between the 2 groups. The operative time was significantly longer in the RG group than those in the LG group during the initial phase (286.0 vs 236.8 min,  $P=0.000$ ). In the late phase, no significant differences were noted between the two groups (237.4 vs 218.8 min,  $P=0.060$ ). The number of harvested lymph nodes, time to first flatus, time to start liquid diet, time to start soft diet, days of postoperative hospital stay, and incidence of overall postoperative complications were not significantly different between the RG and LG groups in the initial or late phase ( $P>0.05$ ).

## Subgroup analysis of different related factors

We evaluate the surgical outcomes of patients according to different related factors, including the extent of resection, age, BMI, depth of invasion, and tumor size. The surgical outcomes of subgroup analyses are summarized in Tables 4–8. The results indicated that the RG groups had less intraoperative blood loss in all subgroup analyses ( $P<0.05$ ). However, RG was associated with longer operation time and higher cost in all subgroup analysis. RG had more harvested lymph nodes

**Table 2** Surgical outcomes and postoperative complications

Characteristics	RG (n=112)	LG (n=112)	P-value
Operation time (min)	261.7±63.9	227.8±45.8	0.000
Estimated blood loss (mL)	179.2±66.8	234.8±139.5	0.000
No. of retrieved lymph nodes	29.5±9.6	27.7±8.7	0.149
Time to first flatus (d)	2.6±0.6	2.8±1.1	0.124
Time to start liquid diet (d)	1.6±0.7	1.7±0.5	0.320
Time to start soft diet (d)	3.6±1.6	3.9±2.0	0.179
Postoperative hospital stay (d)	6.9±2.3	7.0±3.8	0.718
Cost of hospitalization (RMB)	92,365.0±6015.0	69,476.5±5885.6	0.000
Overall complications (%)	15 (13.4)	13 (11.6)	0.686
Grade II (%)	7 (6.3)	6 (5.4)	0.775
Wound infection	2	1	1.000
Pneumonia	2	1	1.000
Pancreatitis	1	0	1.000
Severe anemia	2	4	0.683
Grade IIIa (%)	4 (3.6)	5 (4.5)	1.000
Wound problem	0	1	1.000
Abdominal infection	2	0	0.478
Anastomosis leakage	1	2	1.000
Duodenal stump fistula	1	2	1.000
Grade IIIb (%)	2 (1.8)	1 (0.9)	1.000
Intra-abdominal bleeding	0	1	1.000
Anastomosis bleeding	2	0	0.478
Grade IV (%)	2 (1.8)	1 (0.9)	1.000
Heart failure	1	0	1.000
Respiratory failure	1	1	1.000
Grade V (%)	0 (0.0)	0 (0.0)	–
Clavien-Dindo grade ≥ IIIa (%)	8 (7.1)	7 (6.3)	0.789

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** d, days; LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

**Table 3** Comparison of the 2 surgery methods in different phases

Variables	Initial phase			Late phase		
	RG (n=56)	LG (n=56)	P-value	RG (n=56)	LG (n=56)	P-value
Age (years)	54.4±11.1	56.4±11.7	0.351	56.8±11.4	55.9±10.6	0.643
Gender (male/female)	38/18	42/14	0.403	40/16	37/19	0.541
Body mass index	23.9±2.4	23.4±3.0	0.323	23.3±3.3	23.7±3.1	0.546
Comorbidity (present/absent)	10/46	13/43	0.483	13/43	11/45	0.645
Extent of resection (DG/TG)	31/25	33/23	0.703	35/21	33/23	0.699
Operation time (min)	286.0±57.7	236.9±48.7	0.000	237.4±60.9	218.8±41.0	0.060
Estimated blood loss (mL)	183.9±75.6	243.8±141.0	0.006	174.5±56.9	225.9±138.7	0.012
No. of retrieved lymph nodes	29.6±8.1	26.6±7.2	0.042	29.4±10.9	28.8±9.9	0.779
Time to first flatus (days)	2.6±0.6	2.8±1.1	0.124	2.6±0.5	2.7±0.4	0.261
Time to start liquid diet (days)	1.6±0.7	1.7±0.5	0.320	1.6±0.7	1.6±0.5	0.675
Time to start soft diet (days)	3.6±2.0	3.9±2.0	0.179	3.4±1.2	3.6±0.9	0.470
Postoperative hospital stay (days)	6.9±2.3	7.0±3.8	0.718	6.9±2.2	6.4±2.3	0.254
Cost of hospitalization (RMB)	92,906.1±5288.7	69,924.2±5980.0	0.000	91,824.0±6667.3	69,028.8±5808.8	0.000

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** DG, distal gastrectomy; LG, laparoscopic gastrectomy; RG, robotic gastrectomy; TG, total gastrectomy.

in patients who underwent DG (28.77±11.00 vs 24.8±7.0,  $P=0.014$ ) and patients with low BMI (29.0±8.6 vs 26.0±6.9,  $P=0.026$ ). However, no statistical significant differences were observed in the comparison of other subgroups. No differences were found between the 2 groups regarding the time to first flatus, time to start liquid diet, postoperative hospital stay, and incidence of overall postoperative complications.

## Follow-up result

Within a median follow-up period of 28 months (range 3–52 months), the recurrence rate was 18.8% (21/112) in the RG group and 21.4% (24/112) in the LG group, with no significant difference ( $P=0.617$ ). The 3-year OS rates of the RG and LG groups were 78.6% and 74.1% ( $P=0.483$ , Figure 1). Stratified analysis showed that the 3-year OS rates

**Table 4** Comparison of the 2 surgery methods in different resection extent

Variables	Distal gastrectomy			Total gastrectomy		
	RG (n=66)	LG (n=66)	P-value	RG (n=46)	LG (n=46)	P-value
Age (years)	55.2±11.6	54.1±54.1	0.593	56.2±10.8	59.0±10.5	0.360
Gender (male/female)	48/18	48/18	1.000	30/16	31/15	0.825
Body mass index	23.2±3.0	23.3±3.1	0.950	24.2±2.7	24.0±2.9	0.704
Comorbidity (present/absent)	14/52	10/56	0.367	9/37	14/32	0.229
Operation time (min)	220.5±29.9	198.1±25.0	0.000	320.8±52.1	270.5±33.1	0.000
Estimated blood loss (mL)	160.0±57.0	221.3±133.0	0.001	206.7±70.6	254.1±147.7	0.053
No. of retrieved lymph nodes	28.8±11.0	24.8±7.0	0.014	30.6±7.0	32.0±9.1	0.406
Time to first flatus (days)	2.5±0.7	2.6±1.1	0.489	2.8±0.5	3.1±1.1	0.086
Time to start liquid diet (days)	1.5±0.5	1.6±0.4	0.372	1.8±0.8	1.9±0.6	0.533
Time to start soft diet (days)	3.4±1.6	3.7±2.1	0.267	3.9±1.6	4.2±1.8	0.439
Postoperative hospital stay (days)	6.6±2.3	6.7±2.3	0.820	7.3±2.3	7.6±5.2	0.777
Overall complications (%)	7 (10.6)	7 (10.6)	1.000	8 (17.4)	6 (13.0)	0.562
Cost of hospitalization (RMB)	89,274.7±4848.6	66,178.0±4072.8	0.000	96,769.0±4602.7	74,209.1±4758.6	0.000

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

**Table 5** Comparison of the 2 surgery methods in different age groups

Variables	Age ≤60			Age >60		
	RG (n=69)	LG (n=69)	P-value	RG (n=43)	LG (n=43)	P-value
Gender (male/female)	46/23	49/20	0.581	32/11	30/13	0.631
Body mass index	23.7±3.0	24.0±3.0	0.546	23.5±2.7	22.8±3.0	0.264
Comorbidity (present/absent)	6/63	5/64	0.753	17/26	19/24	0.662
Operation time (min)	258.7±63.5	222.9±42.6	0.000	266.6±65.0	235.7±49.9	0.015
Estimated blood loss (mL)	174.9±61.3	208.8±96.9	0.015	186.1±75.0	276.5±182.6	0.004
No. of retrieved lymph nodes	27.9±10.2	26.4±7.7	0.334	32.1±7.9	29.8±9.8	0.252
Time to first flatus (days)	2.4±0.6	2.5±0.4	0.527	3.0±0.5	3.4±1.6	0.122
Time to start liquid diet (days)	1.5±0.7	1.5±0.4	0.908	1.8±0.5	2.0±0.6	0.112
Time to start soft diet (days)	3.4±1.4	3.4±0.5	0.929	3.9±1.8	4.7±3.0	0.129
Postoperative hospital stay (days)	6.7±2.1	6.4±2.1	0.437	7.1±2.7	8.0±5.4	0.367
Overall complications (%)	5 (7.2)	4 (5.8)	0.730	10 (23.3)	9 (20.9)	0.795
Cost of hospitalization (RMB)	91,567.8±5510.0	68,331.3±4918.3	0.000	93,644.2±6614.1	71,314.1±6839.0	0.000

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

**Table 6** Comparison of the 2 surgery methods in different body mass index

Variables	BMI <25			BMI ≥25		
	RG (n=72)	LG (n=68)	P-value	RG (n=40)	LG (n=44)	P-value
Age (years)	56.8±11.2	57.8±11.0	0.621	53.4±11.2	53.6±10.8	0.360
Gender (male/female)	50/22	49/19	0.734	28/12	30/14	0.857
Comorbidity (present/absent)	19/53	14/54	0.419	4/36	10/34	0.118
Operation time (min)	261.9±68.6	224.5±46.5	0.000	261.3±55.3	233.0±44.7	0.011
Estimated blood loss (mL)	181.0±70.2	232.7±133.1	0.004	176.0±60.8	238.2±150.3	0.017
No. of retrieved lymph nodes	29.0±8.6	26.0±6.9	0.026	30.5±11.2	30.4±10.4	0.066
Time to first flatus (days)	2.6±0.6	2.8±1.1	0.187	2.7±0.7	2.9±1.1	0.440
Time to start liquid diet (days)	1.6±0.7	1.7±0.6	0.680	1.6±0.5	1.7±0.5	0.231
Time to start soft diet (days)	3.6±1.6	4.0±2.3	0.260	3.5±1.5	3.8±1.4	0.442
Postoperative hospital stay (days)	7.1±2.6	6.7±2.6	0.378	6.5±1.5	7.5±5.1	0.217
Overall complications (%)	10 (13.9)	8 (11.8)	0.707	5 (12.5)	5 (11.4)	0.860
Cost of hospitalization (RMB)	92,660.0±5997.1	69,447.0±6138.0	0.000	91,834.0±6086.9	69,522.1±5541.9	0.000

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** BMI, body mass index; LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

for the patients after RG and LG were as follows: a total of 90.0 % vs 94.4 % for the patients with stage IB ( $P=0.436$ , Figure 2A), 81.8% vs 77.3 % for the patients with stage II ( $P=0.667$ , Figure 2B), and 70.8 % vs 64.0 % for the patients with stage III ( $P=0.687$ , Figure 2C). Comparisons of the OS rates did not show significant difference for any of the stages.

**Table 7** Comparison of the 2 surgery methods in different depth of invasion

Variables	No serosa invasion			Serosa invasion		
	RG (n=38)	LG (n=36)	P-value	RG (n=74)	LG (n=76)	P-value
Age (years)	55.8±12.5	56.2±12.6	0.890	55.5±10.7	56.1±10.4	0.732
Gender (male/female)	26/12	24/12	0.872	52/22	55/21	0.776
Body mass index	24.1±2.8	24.0±2.7	0.989	23.4±2.9	23.3±3.2	0.866
Comorbidity (present/absent)	7/31	9/27	0.492	58/16	61/15	0.776
Operation time (min)	280.7±71.1	242.4±51.3	0.010	252.0±58.0	221.0±41.5	0.000
Estimated blood loss (mL)	178.2±65.8	258.3±159.0	0.006	179.7±67.7	223.7±128.9	0.010
No. of retrieved lymph nodes	30.5±10.7	30.8±9.0	0.896	29.0±9.0	26.3±8.3	0.055
Time to first flatus (days)	2.7±0.8	2.8±0.7	0.465	2.6±0.6	2.8±1.3	0.176
Time to start liquid diet (days)	1.8±1.0	1.7±0.5	0.881	1.6±0.4	1.7±0.6	0.096
Time to start soft diet (days)	3.6±2.2	3.7±1.2	0.771	3.6±1.2	4.0±2.2	0.154
Postoperative hospital stay (days)	7.2±3.2	7.0±4.0	0.777	6.7±1.7	7.1±3.7	0.462
Overall complications (%)	7 (18.4)	4 (11.1)	0.376	8 (10.8)	9 (11.8)	0.842
Cost of hospitalization (RMB)	92,932.9±6380.2	70,763.1±5555.6	0.000	92,073.4±5841.9	68,867.0±5974.4	0.000

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

**Table 8** Comparison of the 2 surgery methods in different tumor size

Variables	Tumor size ≤5 cm			Tumor size >5 cm		
	RG (n=83)	LG (n=84)	P-value	RG (n=29)	LG (n=28)	P-value
Age (years)	54.7±11.6	54.9±11.0	0.917	58.1±10.1	59.8±10.8	0.545
Gender (male/female)	57/26	62/22	0.463	21/8	17/11	0.349
Body mass index	23.7±3.0	23.7±3.0	0.953	23.4±2.5	23.1±3.2	0.621
Comorbidity (present/absent)	15/68	16/68	0.871	8/21	8/20	0.934
Operation time (min)	248.8±60.8	219.5±41.3	0.000	298.5±58.8	252.8±50.1	0.003
Estimated blood loss (mL)	171.5±61.9	232.6±141.6	0.000	201.4±76.0	241.4±135.4	0.172
No. of retrieved lymph nodes	28.2±9.9	27.0±8.5	0.399	33.2±7.4	30.0±9.1	0.140
Time to first flatus (days)	2.6±0.7	2.7±1.1	0.347	2.8±0.5	3.1±1.2	0.144
Time to start liquid diet (days)	1.6±0.7	1.7±0.5	0.953	1.6±0.4	1.9±0.7	0.052
Time to start soft diet (days)	3.6±1.8	3.8±1.3	0.389	3.4±0.7	4.1±2.1	0.119
Postoperative hospital stay (days)	6.9±2.6	6.8±3.1	0.810	6.8±1.5	7.7±5.4	0.378
Overall complications (%)	11 (13.3)	8 (9.5)	0.448	4 (13.8)	5 (17.9)	0.954
Cost of hospitalization (RMB)	91,300.9±6265.8	68,616.0±5675.8	0.000	95,410.6±3935.9	72,058.0±5845.2	0.000

**Note:** Data are presented as mean ± standard deviation.

**Abbreviations:** LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

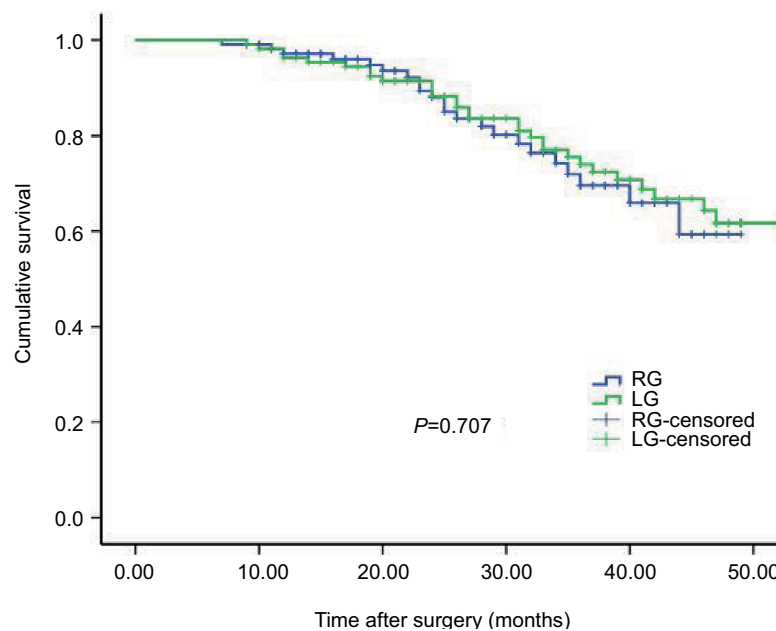
## Discussion

To date, a considerable number of studies have investigated the short-term outcomes of RG for gastric cancer.<sup>15–18</sup> However, the majority of these studies included patients at a relatively early stage, and few of them evaluated the outcomes of RG for AGC. Thus, we designed this study to compare the short- and long-term outcomes of RG and LG for AGC. Additionally, we used the propensity score-matching method to reduce bias.

In the present study, we observed that the RG group was associated with less blood loss compared with the LG group, which is consistent with the previous studies. Moreover, this advantage still exists when subgroup analysis was conducted. Junfeng et al<sup>19</sup> reported that the RG group had less intraoperative blood loss (101.4 vs 131.4 mL,  $P=0.017$ ) and more harvested lymph nodes ( $34.6±10.9$  vs  $32.7±11.2$ ,  $P=0.013$ )

compared with the LG group. This could be attributed to the fact that RG has tremor filtration, the ability to scale motion, and is stereoscopic, which could improve a surgeon's dexterity, allow precise dissection, and avoid injury blood vessels. Our result showed a tendency favoring RG in terms of the number of harvested lymph nodes, although no significant difference was observed between the RG and LG groups.

Lymph node dissection is one of the key elements of radical gastrectomy. It has been reported that the number of lymph node dissection was related to patients' prognosis.<sup>20–23</sup> From the standpoint of AGC curability, gastrectomy with D2 lymph node dissection is required according to the Japanese Gastric Cancer Association guidelines.<sup>24–26</sup> However, the superiority of more harvested lymph nodes was not consistent among all studies. Pugliese et al<sup>27</sup> and Song et al<sup>28</sup> reported that the number of harvested lymph



**Figure 1** Kaplan–Meier curves of overall survival for all patients.

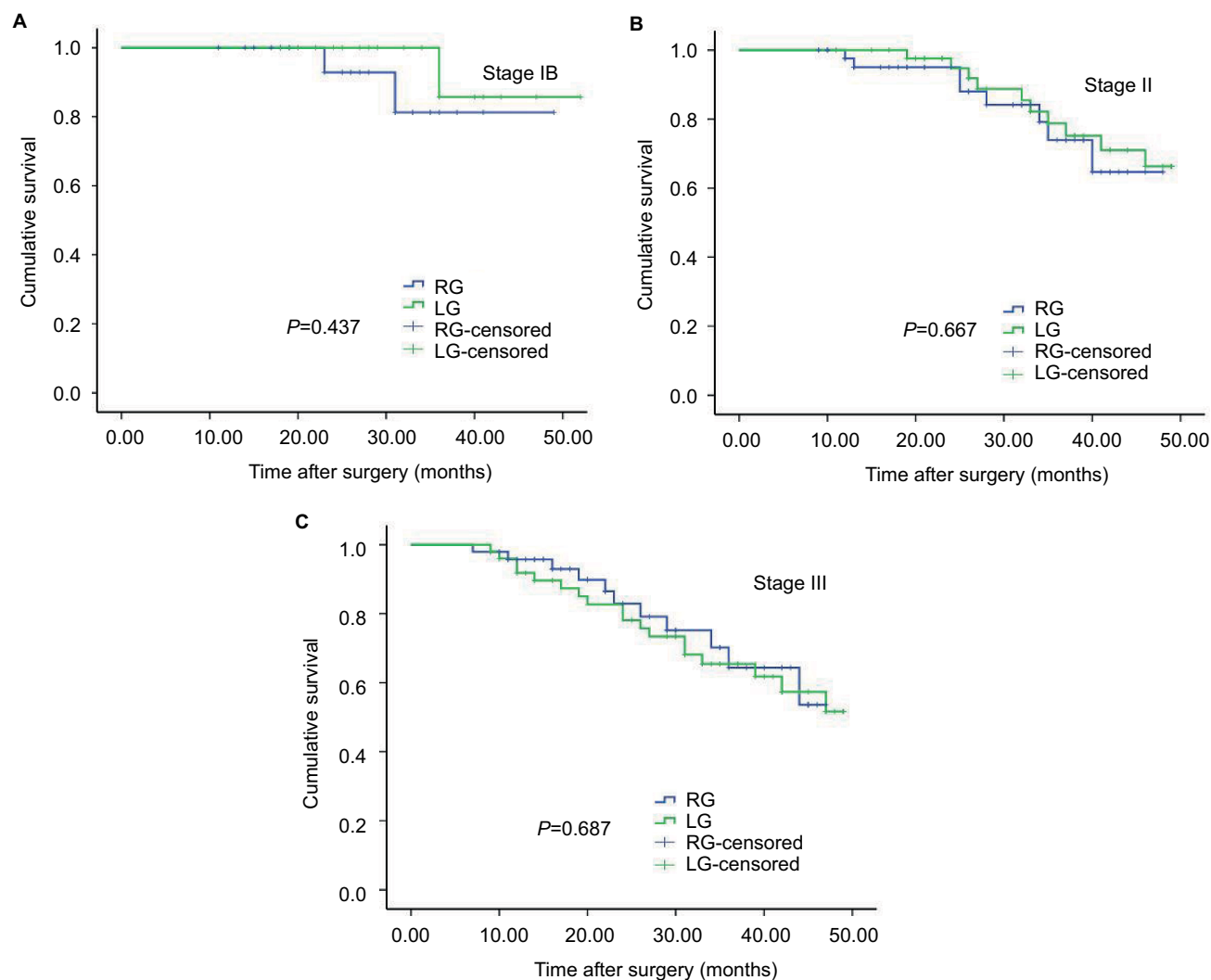
**Abbreviations:** LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

nodes in the RG group are less than those in the LG group, but these studies are in small sample size and the result was drawn from their initial experience. In addition, studies have demonstrated that surgeons become skilled in RG with the accumulation of experience of about 20–30 cases, which provides a short learning curve compared with LG.<sup>29–33</sup> Several meta-analyses revealed that harvested lymph nodes for RG and LG were similar, but they all showed a tendency favoring RG.<sup>17,34,35</sup> According to our experience, the advantage of RG is that it could enable the surgeons to perform more precise dissection and anastomoses. These superiorities are more obvious for difficult lymph node dissection (station 8a, 10, 11p, 12a), which remains the cornerstone of D2 lymphadenectomy. Moreover, the robotic system can provide a much more stable view and better exposure in narrow operating field. Regarding more complex surgery, such as the resection of gastric stump cancer, the robotic surgical system has the advantages of clear vision, flexible operation, and stable traction during the operation procedure, which is superior to the laparoscopic technique.

Postoperative complication is an important factor for evaluating the safety and feasibility of a surgical procedure. We evaluate the postoperative complications according the Clavien–Dindo classification system, which has proven to be a standardized classification for assessment for complications in many types of surgery. In the current study, we found that the incidences of overall complications were comparable between the RG and LG groups (13.4% vs 11.6%,  $P=0.686$ ).

Moreover, no significant differences were noted in the minor (Clavien–Dindo grade II) and major complication (Clavien–Dindo grade  $\geq$  IIIa) rates between the 2 groups (6.8% vs 5.4%,  $P=0.775$ ; 4.8% vs 6.8%,  $P=0.453$ ; respectively). Furthermore, analyses of specified complications revealed that the rate of wound problem, anastomotic leakage, duodenal stump fistula, and abdominal infection were also similar between the 2 groups.

The extent of resection, age, BMI, depth of invasion, and tumor size could be factors associated with the short-term outcomes.<sup>36–38</sup> In this study, we performed analyses by grouping the patients according to different factors. We found that patients in the RG group were all associated with lower blood loss. Intraoperative blood loss has been shown to be associated with prognosis of patients in gastric cancer.<sup>39,40</sup> Kamei et al reported that curative resection with lower blood loss can contribute to improved survival.<sup>41</sup> Moreover, the less intraoperative blood loss may promote postoperative recovery, especially for elderly patients. Our result revealed that RG patients had a tendency to recover fast after surgery, although no significant differences were found in respect of these indicators. Lee et al<sup>7</sup> reported that the superiority of RG was more obvious in high BMI patients when performing DG. In the present study, among patients in the high BMI group, RG was also associated with less intraoperative blood loss. Meanwhile, the RG group was comparable with the LG group in terms of other surgical outcomes. According to the present findings, it seems that optimal surgical



**Figure 2** Kaplan–Meier curves of overall survival (A) patients in stage IB (B) patients in stage II (C) patients in stage III. **Abbreviations:** LG, laparoscopic gastrectomy; RG, robotic gastrectomy.

outcomes may have already been achieved with LG, leaving little room for improvement via RG. Enhanced recovery after surgery protocols has been routinely applied to accelerate the postoperative recovery of patients with gastric cancer in our center since 2012, which could explain that the time to first flatus, time to start diet, and postoperative hospital stay were superior to those reported in previous studies.<sup>17,42</sup>

Long-term survival outcomes are key indicator for assessing oncological safety. Therefore, the long-term outcomes are also important and necessary for the wide application of RG in AGC. Junfeng et al<sup>19</sup> reported that the 3-year OS rates of patients with gastric cancer in RG and LG groups were 68.1% and 63.7%, respectively, with a median follow-up of 17 months; the difference was not statistically significant. In a propensity-matched analysis of RG vs LG for gastric cancer from 2005 to 2009, no significant differences were found in 5-year OS rate (93.2% vs 94.2%,  $P=0.521$ ) and

disease-free survival rate (90.7% vs 92.6%,  $P=0.229$ ) with a median follow-up of 85 months. However, patients in early stage account for 83.4% in the study reported by Obama et al.<sup>43</sup> In the present study, patients in stages II and III account for 83.0% of all cases in propensity score-matched cohort, indicating a poor prognosis for these cases. The current study showed that the 3-year OS rate after RG was similar with those in LG with a median follow-up of 28 months (78.6% vs 74.1%,  $P=0.483$ ). Regarding stratified stages, the 3-year OS was also comparable between the 2 groups. These results indicated that RG and LG had comparable long-term survival outcomes for AGC.

Financial cost may also be an important factor in driving the selection of surgery type.<sup>44,45</sup> To examine cost-effectiveness of the RG and LG, we retrospectively reviewed the overall hospital expenditure and found that the cost of RG group was about 23,000 RMB more than that of the LG group. A recent



meta-analysis reported that the RG groups cost about \$ 4000 more than the LG groups. In our clinical practice, the higher cost, longer operation time, and limited benefit to patients are the major stumbling block to the wide application of RG. Meanwhile, the long-term benefits of less intraoperative blood loss have not been proven yet. Additionally, there is also no published study about the quality of life after RG, which is often used to assess the surgical outcomes.<sup>46,47</sup> We believe that these problems will be solved in the near future with the accumulation of surgeon's experience, reduced costs of surgery, and more high-quality research on this topic.

We admit that our study has several limitations. First, this is a retrospective study conducted at a single center with limited duration of follow-up. Additionally, the allocation of the operation method to either RG or LG may include a selection bias, which could not be offset by propensity score-matching method.

## Conclusion

RG with D2 lymph node dissection is safe and feasible for AGC in terms of both short- and long-term outcomes. The higher cost, longer operation time, and limited benefit are the major stumbling blocks to the wide application of RG. High-volume randomized controlled trials with sufficient follow-up are needed to confirm this rationale.

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## Disclosure

The authors report no conflicts of interest in this work.

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