



## Early Outcomes of Robot-Assisted Versus Thoracoscopic-Assisted Ivor Lewis Esophagectomy for Esophageal Cancer: A Propensity Score-Matched Study

Yajie Zhang, MD, PhD<sup>1</sup>, Yu Han, MD<sup>1</sup>, Qinyi Gan, MD<sup>1</sup>, Jie Xiang, MD<sup>1</sup>, Runsen Jin, MD, PhD<sup>1</sup>, Kai Chen, MD<sup>1</sup>, Jiaming Che, MD<sup>1</sup>, Junbiao Hang, MD<sup>1</sup>, and Hecheng Li, MD, PhD<sup>1</sup>

Department of Thoracic Surgery, Ruijin Hospital, Shanghai Jiaotong University School of Medicine, Shanghai, China

### ABSTRACT

**Background.** Both robot-assisted Ivor Lewis esophagectomy (RAILE) and conventional thoracoscopic-assisted Ivor Lewis esophagectomy (TAILE) are minimally invasive surgical techniques for the treatment of middle and distal esophageal cancer. However, no research studies comparing early outcomes between RAILE and TAILE have been reported.

**Methods.** A retrospective analysis was made of 184 patients, 76 in the RAILE group and 108 in the TAILE group, who underwent minimally invasive Ivor Lewis esophagectomy between December 2014 and June 2018. Propensity score-matched analysis was performed between the two groups based on demographics, comorbidities, American Society of Anesthesiologists score, tumor location, tumor size, and pathological stage. Perioperative outcomes were compared.

**Results.** Two conversions to thoracotomy occurred in the RAILE group. There was no 30-day in either group. Sixty-six matched pairs were identified for each group. Within the propensity score-matched cohorts, the operative time in the RAILE group was significantly longer than that in the TAILE group ( $302.0 \pm 62.9$  vs.  $274.7 \pm 38.0$  min,  $P = 0.004$ ). There was no significant difference in the blood loss [200.0 ml (interquartile range [IQR], 100.0–262.5 ml) vs. 200.0 ml (150.0–245.0 ml),  $P = 0.100$ ], rates of overall complications (28.8 vs. 24.2%,

$P = 0.554$ ), length of stay [9.0 days (IQR 8.0–12.3 days) vs. 9.0 days (IQR 8.0–11.3 days),  $P = 0.517$ ], the number of total dissected lymph nodes ( $19.2 \pm 9.2$  vs.  $19.3 \pm 9.5$ ,  $P = 0.955$ ), and detailed categories of lymph nodes.

**Conclusions.** RAILE demonstrated comparable early outcomes compared with TAILE and should be considered as an alternative minimally invasive option for treating esophageal cancer.

Esophageal cancer ranks the seventh in terms of incidence (572,000 new cases) and sixth in mortality overall (509,000 deaths) around the world in 2018.<sup>1</sup> In China, esophageal cancer is prominent, with an estimated 477,900 new cases and 375,000 deaths in 2015.<sup>2</sup> Despite improvements in neoadjuvant and adjuvant treatment regimens, surgical resection remains the main multimodality treatment for esophageal cancer.<sup>3</sup> Esophageal cancer is most likely to occur in the middle and distal esophagus.<sup>4</sup> Traditionally, an open Ivor Lewis esophagogastrectomy, which incorporates a laparotomy and a right thoracotomy, is one of the most commonly utilized approaches for allowing a subtotal esophagectomy as well as a thorough lymphadenectomy.<sup>5,6</sup> However, the mortality and morbidity associated with this procedure are high.<sup>7</sup> During the past two decades, the role of minimally invasive esophagectomy (MIE) has been well established. Thoracoscopic-assisted Ivor Lewis esophagectomy (TAILE) has become increasingly adopted.<sup>8</sup> Compared with open resections, TAILE results in shorter hospital stays and decreased postoperative morbidities, with comparable oncologic outcomes.<sup>9</sup> However, TAILE has some intrinsic limitations, including its two-dimensional view, reduced hand-eye coordination, and limited degree of freedom of

Yajie Zhang and Yu Han have contributed equally to this work.

© Society of Surgical Oncology 2019

First Received: 16 August 2018;  
Published Online: 6 March 2019

H. Li, MD, PhD  
e-mail: lihecheng2000@hotmail.com

movement, which make it difficult to perform key steps during the esophagectomy, including intrathoracic anastomosis and mediastinal lymph node dissection.

Recently, robot-assisted Ivor Lewis esophagectomy (RAILE) has been developed in some large thoracic centers. Robot systems were designed to provide some advantages over conventional minimally invasive surgery, including a magnified three-dimensional view and an improved dexterity, due to the surgical wrists and tremor filtration. Although we and other teams have demonstrated that RAILE is safe and feasible, the literature published in this area is still sparse.<sup>10–13</sup> More importantly, there are no studies comparing RAILE with TAILE to determine whether the RAILE technical advantages are reflected in surgical procedures. The purpose of this study was to compare the short-term outcomes and oncologic safety between RAILE and TAILE using a propensity score-matched (PSM) analysis.

## PATIENTS AND METHODS

### *Patient Selection and Data Collection*

We performed a retrospective study of patients who underwent curative MIE including both RAILE and TAILE by a single surgeon (Dr. H.C.L.) between December 2014 and June 2018 at Ruijin Hospital affiliated to Shanghai Jiaotong University. TAILE was performed in our institution from December 2014 to May 2015. After that, our institution began the RAILE program. The inclusion criteria were: (a) endoscopically and pathologically diagnosed carcinoma of the mid-lower thoracic esophagus; (b) a tumor judged to be resectable based on a preoperative contrast-enhanced computed tomography of the neck, chest, and abdomen; and (c) patients without distant metastasis. The exclusion criteria were: (a) preoperative neoadjuvant therapy for the reason that the neoadjuvant chemoradiation was not fully realized in the course of this study in our institution; (b) potential surgical contraindications for MIE; and (c) previous history of gastrointestinal surgery or major right thoracic surgery. All patients chose the surgical approach according to their preference and signed informed consent forms. This study was approved by our institutional review board.

There was no difference in the preoperative preparation and postoperative treatment protocol between RAILE and TAILE. The basic demographics, pathological results, and short-term outcomes were retrospectively collected. The pathological stages were evaluated using the American Joint Committee on Cancer (AJCC) 7th edition TNM staging system.<sup>14</sup> To examine further details, the dissected lymph nodes were classified into the abdominal and

thoracic categories as well as those along the bilateral recurrent laryngeal nerves (RLNs).

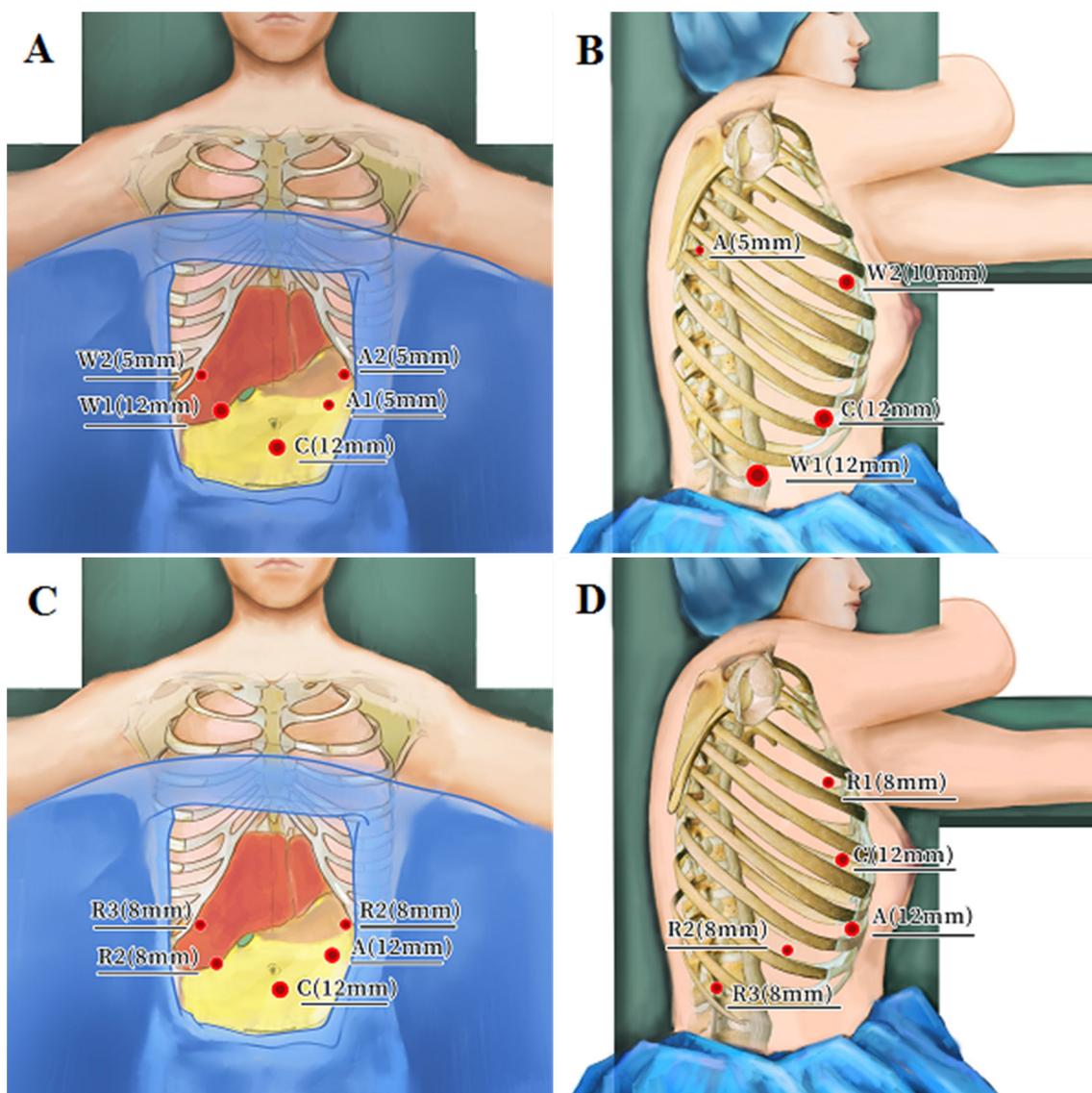
### *Surgical Technique*

RAILE was performed using a da Vinci Surgical System (Model S; Intuitive Surgical, Inc., Sunnyvale, CA), and TAILE was completed using a laparoscopic-thoracoscopic system (Karl Storz, Tuttlingen, Germany). The patient was initially supine with a reverse Trendelenburg position for the abdominal phase. The port positions for the abdominal part of RAILE and TAILE were similar (Fig. 1a and c, respectively). Both the abdominal phases of the two surgical approaches included the liver suspension, gastric mobilization with abdominal lymphadenectomy, intracorporeal gastric conduit formation, and laparoscopic feeding jejunostomy.

The patient was then placed in the left-lateral decubitus position for the thoracic phase with single-lung ventilation. For TAILE, four ports were positioned as shown in Fig. 1b. A 12-mm camera port was inserted in the 8th intercostal space (ICS) at the anterior axillary line. A 12-mm and a 5-mm port were inserted in the 10th ICS at the posterior axillary line and in the 5th ICS at the anterior axillary line, respectively, for the first and second working ports. Finally, a 5-mm assistant port was inserted at the triangle of auscultation. With regard to RAILE (Fig. 1d), a 12-mm camera port was inserted in the 5th ICS at the anterior axillary line. Three 8-mm robotic ports were inserted in the 3rd ICS at the anterior axillary line, the 8th ICS at the posterior axillary line, and the 10th ICS posterior to the posterior axillary line, respectively. Finally, a 12-mm assistant port was created in the 7th ICS near the costal margin. The robot was positioned on the dorsocranial side, with one assistant on the anterior side. CO<sub>2</sub> insufflation at a pressure of 8–10 mmHg was used for both RAILE and TAILE. The esophagus was mobilized en bloc from the thoracic inlet to the gastroesophageal junction, along with all periesophageal lymph nodes. The lymph nodes along the bilateral RLNs were dissected carefully. For TAILE, the conduit was pulled up through the hiatus, and an intrathoracic end to end anastomosis was performed with a circular stapler. For RAILE, both a circular stapled anastomosis and a double-layered, completely hand-sewn intrathoracic anastomosis were used as we previously described.<sup>10</sup>

### *Statistical Analysis*

To minimize the bias caused by the nonrandomized selection of patients, we performed a PSM analysis to control the confounding factors using R Project Software (v.2.14.1; <http://www.r-project.org>). Each patient's



**FIG. 1** Port positions for minimally invasive esophagectomy. (A) The abdominal phase in TAILE. (B) The thoracic phase in TAILE. C, camera port; W1 and W2, the first and second working ports; A1 and A2, the first and second assistant ports. (C) The

abdominal phase in RAILE. (D) The thoracic phase in RAILE. C, camera port; R1, R2, and R3, the first, second, and third robotic ports; A, the assistant port

propensity score was derived from a multivariable logistic model with covariates, including age, gender, body mass index (BMI), American Society of Anesthesiologists (ASA) score, comorbidity, tumor location, tumor size, and pTNM stage. Patients treated with RAILE were matched 1:1 with no replacement to patients treated with TAILE using the nearest-neighbor method with a caliper width of 0.01.

The statistical analysis was performed using SPSS version 20.0 (SPSS Inc., Chicago, IL). Student's *t* test or Wilcoxon rank-sum test was applied to compare the continuous variables between the groups, and the Chi squared test or Fisher's exact test was applied for comparing the

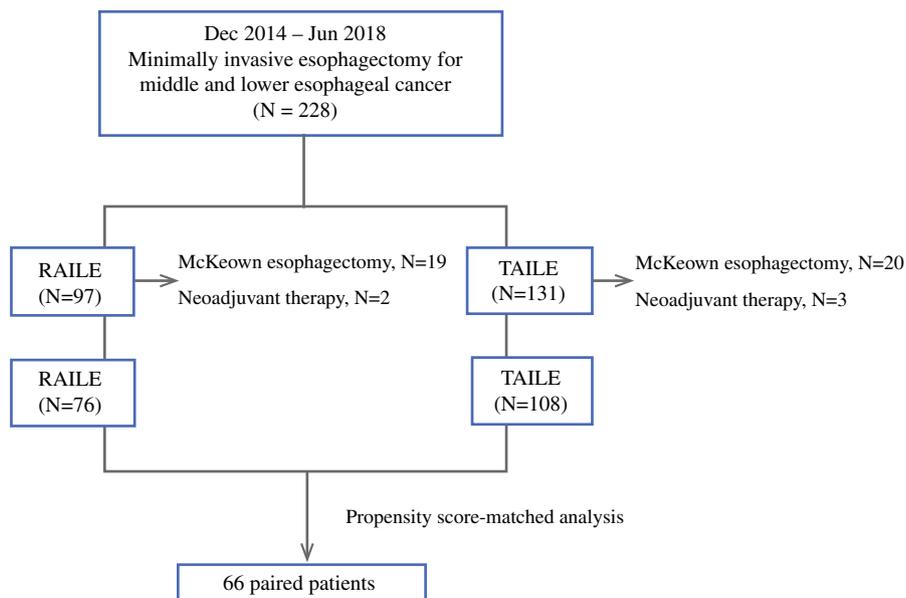
categorical data. A two-sided *P* value < 0.05 was considered statistically significant.

## RESULTS

### *Patient Characteristics*

A total of 184 patients who underwent minimally invasive Ivor-Lewis esophagectomy met the inclusion criteria between December 2014 and June 2018 (Fig. 2). Of them, 108 patients underwent TAILE with stapled anastomosis, and 76 patients were treated with RAILE, including 40 cases of stapled anastomosis and 36 cases of hand-sewn

**FIG. 2** Flow of the patients through the study. *RAILE* robot-assisted Ivor-Lewis esophagectomy; *TAILE* thoracoscopic-assisted Ivor-Lewis esophagectomy



anastomosis. After PSM, 66 paired patients were matched from the cohort. The baseline characteristics of the study population before and after PSM are summarized in Table 1. The two groups were comparable in patient characteristics, including age, sex, BMI, smoking history, comorbidity, ASA score, tumor location, tumor size, pTNM stage, or pathology.

### Perioperative Outcomes

Table 2 presents the perioperative outcomes between RAILE and TAILE. Before PSM, there was no conversion in TAILE, whereas two patients in RAILE were converted to thoracotomy due to severe intraoperative atrial fibrillation and severe chest adhesion, respectively. There was no 30-day mortality in either group. One patient in RAILE and two patients in TAILE died within 90 days after surgery ( $P = 1.000$ ). Compared with the TAILE group, the RAILE group experienced a significantly longer operative time ( $303.5 \pm 60.9$  vs.  $277.2 \pm 42.2$  min,  $P = 0.001$ ). The intraoperative blood loss was similar between the two groups [200.0 ml (IQR 100.0–300.0 ml) vs. 200.0 ml (IQR 150.0–250.0 ml),  $P = 0.059$ ]. The overall complication rate was comparable (31.6 vs. 33.3%,  $P = 0.803$ ), and the detailed complications, including the anastomotic leakage, pneumonia, pleural effusion, chylothorax, vocal cord paralysis, cardiac complications, and wound infection, were all similar between the two groups. There was no significant in the length of stay (LOS) between the two groups [9.0 days (IQR 8.0–12.0 days) vs. 9.0 days (IQR 8.0–11.8 days),  $P = 0.436$ ]. As for the pathological outcomes, the R0 resection was achieved in all but one patient in the TAILE group. There was no statistically significant

difference in the number of total dissected lymph nodes ( $19.7 \pm 9.8$  vs.  $20.3 \pm 9.7$ ,  $P = 0.689$ ), as well as the number of thoracic ( $10.6 \pm 7.0$  vs.  $12.5 \pm 8.7$ ,  $P = 0.107$ ), abdominal ( $9.1 \pm 6.8$  vs.  $7.8 \pm 6.2$ ,  $P = 0.186$ ), right RLN ( $1.4 \pm 1.6$  vs.  $1.3 \pm 2.3$ ,  $P = 0.899$ ), and left RLN lymph nodes ( $1.6 \pm 2.2$  vs.  $1.0 \pm 2.0$ ,  $P = 0.081$ ).

Next, we analyzed the perioperative outcomes in well matched patients. After PSM, the operative time in RAILE was still longer than that in TAILE ( $302.0 \pm 62.9$  vs.  $274.7 \pm 38.0$  min,  $P = 0.004$ ). There was no significant difference between the two groups with respect to the blood loss [200.0 ml (IQR 100.0–262.5 ml) vs. 200.0 ml (150.0–245.0 ml),  $P = 0.100$ ], 90-day mortality rates (1.5% vs. 1.5%,  $P = 1.000$ ), rates of overall complications (28.8 vs. 24.2%,  $P = 0.554$ ), LOS [9.0 days (IQR 8.0–12.3 days) vs. 9.0 days (IQR 8.0–11.3 days),  $P = 0.517$ ], as well as the number of total dissected lymph nodes ( $19.2 \pm 9.2$  vs.  $19.3 \pm 9.5$ ,  $P = 0.955$ ) and detailed categories of lymph nodes.

### DISCUSSION

To the best of our knowledge, this is the first study to focus on both RAILE and TAILE and compare the short-term outcomes between the two different surgical approaches. Notably, the present study includes a relatively large number of patients and applies a PSM study design to reduce the potential bias that could probably affect the perioperative outcomes. The results demonstrated that although RAILE experienced a significant longer operative time, the two surgical approaches were comparable in terms of the intraoperative blood loss, postoperative complication rate, LOS, and number of dissected lymph nodes.

**TABLE 1** Patient characteristics

Characteristics <sup>a</sup>	All patients			Matched patients		
	RAILE ( <i>n</i> = 76)	TAILE ( <i>n</i> = 108)	<i>P</i> value	RAILE ( <i>n</i> = 66)	TAILE ( <i>n</i> = 66)	<i>P</i> value
Age, yr	61.8 ± 7.7	61.3 ± 7.7	0.665	62.3 ± 7.8	62.0 ± 7.8	0.776
Gender			0.862			1.000
Male	59 (77.6)	85 (78.7)		50 (75.8)	50 (75.8)	
Female	17 (22.4)	23 (21.3)		16 (24.2)	16 (24.2)	
BMI, kg/m <sup>2</sup>	22.8 ± 3.1	22.7 ± 3.9	0.850	22.9 ± 3.1	23.1 ± 4.5	0.782
ASA score			0.248			0.756
1	36 (47.4)	38 (35.2)		30 (45.5)	26 (39.4)	
2	37 (48.7)	64 (59.3)		33 (50.0)	36 (54.5)	
3	3 (3.9)	6 (5.6)		3 (4.5)	4 (6.1)	
History of smoking	39 (51.3)	70 (64.8)	0.067	33 (50.0)	42 (63.6)	0.175
Comorbidity	32 (42.1)	51 (47.2)	0.492	28 (42.4)	32 (48.5)	0.597
Tumor location			0.158			0.678
Middle thoracic	30 (39.5)	54 (50.0)		29 (43.9)	26 (39.4)	
Lower thoracic	46 (60.5)	54 (50.0)		37 (56.1)	40 (60.6)	
Tumor size, cm	3.1 ± 1.4	3.2 ± 1.4	0.835	3.1 ± 1.4	3.0 ± 1.4	0.438
pT stage			0.220			0.151
Tis	4 (5.3)	4 (3.7)		4 (6.1)	4 (6.1)	
T1	20 (26.3)	24 (22.2)		15 (22.7)	22 (33.3)	
T2	14 (18.4)	32 (29.6)		13 (19.7)	19 (16.0)	
T3	36 (47.4)	48 (44.4)		32 (48.5)	21 (31.8)	
T4a	2 (2.6)	0		2 (3.0)	0	
pN stage			0.294			0.066
N0	47 (64.8)	74 (68.5)		41 (62.1)	53 (80.3)	
N1	16 (21.1)	19 (17.6)		13 (19.7)	9 (13.6)	
N2	9 (11.8)	14 (13.0)		9 (13.6)	4 (6.1)	
N3	4 (5.3)	1 (0.9)		3 (4.5)	0	
TNM stage			0.215			0.164
0	4 (5.3)	4 (3.7)		4 (6.1)	4 (6.1)	
I	18 (23.7)	21 (19.4)		15 (22.7)	21 (31.8)	
II	29 (38.2)	58 (53.7)		24 (36.4)	33 (50.0)	
III	25 (32.9)	25 (23.1)		23 (34.8)	8 (12.1)	
Pathology			0.570			1.000
Squamous cell carcinoma	74 (97.4)	107 (99.1)		64 (97.0)	65 (98.5)	
Adenocarcinoma	0	0		0	0	
Adenosquamous carcinoma	2 (2.6)	1 (0.9)		2 (3.0)	1 (1.5)	

RAILE robot-assisted Ivor-Lewis esophagectomy, TAILE thoracoscopic-assisted Ivor-Lewis esophagectomy, BMI body mass index, ASA American Society of Anesthesiologists

<sup>a</sup>Categorical data are expressed as number (%) and continuous data as mean ± SD

Since the initial robot-assisted minimally invasive esophagectomy (RAMIE) was introduced in 2003 by Giulianotti et al., three main surgical approaches, namely, transhiatal esophagectomy, McKeown esophagectomy, and Ivor-Lewis esophagectomy, have been reported with robotic assistance.<sup>15</sup> van der Sluis et al. has recently published the results of the ROBOT trial, which is the first

randomized trial of RAMIE to date, and demonstrated that compared to open esophagectomy, RAMIE resulted in a lower percentage of overall surgery-related and cardiopulmonary complications, lower postoperative pain, better short-term quality of life, and a better short-term postoperative functional recovery with comparable long-term oncological results.<sup>16</sup> However, so far, most of the

**TABLE 2** Perioperative outcomes

Characteristics <sup>a</sup>	All patients			Matched patients		
	RAILE ( <i>n</i> = 76)	TAILE ( <i>n</i> = 108)	<i>P</i> value	RAILE ( <i>n</i> = 66)	TAILE ( <i>n</i> = 66)	<i>P</i> value
Operative time, min	303.5 ± 60.9	277.2 ± 42.2	0.001	302.0 ± 62.9	274.7 ± 38.0	0.004
Blood loss, ml	200.0 (100.0–300.0)	200.0 (150.0–250.0)	0.059	200.0 (100.0–262.5)	200.0 (150.0–245.0)	0.100
Conversion to open	2 (2.6)	0	0.169	1 (1.5)	0	1.000
Overall complications	24 (31.6)	36 (33.3)	0.803	19 (28.8)	16 (24.2)	0.554
Leakage	7 (9.2)	6 (5.6)	0.410	5 (7.6)	3 (4.5)	0.727
Pneumonia	5 (6.6)	10 (9.3)	0.513	4 (6.1)	5 (7.6)	1.000
Pleural effusion	0	5 (4.6)	0.078	0	2 (3.0)	0.476
Chylothorax	1 (1.3)	3 (2.8)	0.503	0	1 (1.5)	1.000
Vocal cord paralysis	5 (6.6)	7 (6.5)	0.979	4 (6.1)	3 (4.5)	1.000
Cardiac complications	5 (6.6)	4 (3.7)	0.587	5 (7.6)	2 (3.0)	0.375
Wound infection	1 (1.3)	1 (0.9)	1.000	1 (1.5)	0	1.000
In-hospital mortality	0	0		0	0	
30-day mortality	0	0		0	0	
90-day mortality	1 (1.3)	2 (1.9)	1.000	1 (1.5)	1 (1.5)	1.000
LOS, day	9.0 (8.0–12.0)	9.0 (8.0–11.8)	0.436	9.0 (8.0–12.3)	9.0 (8.0–11.3)	0.517
R0 Surgery	76 (100)	107 (99.1)	1.000	66 (100)	66 (100)	
Number of dissected LN	19.7 ± 9.8	20.3 ± 9.7	0.689	19.2 ± 9.2	19.3 ± 9.5	0.955
Abdominal phase	9.1 ± 6.8	7.8 ± 6.2	0.186	8.9 ± 6.7	7.3 ± 5.9	0.198
Thoracic phase	10.6 ± 7.0	12.5 ± 8.7	0.107	10.3 ± 5.8	11.9 ± 8.3	0.137
Right RLN	1.4 ± 1.6	1.3 ± 2.3	0.899	1.4 ± 1.6	1.6 ± 2.8	0.597
Left RLN	1.6 ± 2.2	1.0 ± 2.0	0.081	1.3 ± 1.9	0.9 ± 1.9	0.235

RAILE robot-assisted Ivor Lewis esophagectomy, TAILE thoracoscopic-assisted Ivor-Lewis esophagectomy, LOS length of stay

<sup>a</sup>Categorical data are expressed as number (%) and continuous data as mean ± SD or median (interquartile range)

published literature focused on the robot-assisted transhiatal esophagectomy and McKeown esophagectomy.<sup>17–19</sup> Only a few centers have reported on RAILE, mostly with relatively small number of patients and different manner of intrathoracic anastomosis.<sup>11, 20–22</sup> In addition, the literature directly comparing RAILE with conventional TAILE is still scarce. One such study, which was conducted by Park et al. in 2016, compared the short-term outcomes of 62 robot-assisted and 43 thoracoscopic-assisted esophagectomy.<sup>23</sup> However, this study cohort involved two different surgical approaches, specifically the Ivor-Lewis and McKeown procedures.

In this comparative study, despite the mean operative time in RAILE was similar to the published studies, the only difference we observed between the two surgical approaches was that RAILE took longer operative time, which is consistent with the most other studies comparing the operative time of robot-assisted and thoracoscopic esophagectomy.<sup>11,12,24</sup> We analyzed the reasons for this longer operative time. First, although the apparent advantages of robotic system over conventional minimally invasive surgery, including high-definition, three-dimensional view, wristed instrumentation with seven degrees of

freedom and tremor-filtering facilitate meticulous dissection and thereby can theoretically decrease the operative time, the potential drawbacks of robotic system also inevitably affect the operative time. For example, during RAILE, the robot must be docked twice and moved from the head side to the dorsocranial side when turning from the abdominal phase to the thoracic phase. In addition, the first, second, and third robotic arms need to be exchanged frequently according to the approach angle, which all contribute to the prolonged operation time. However, these shortcomings have been greatly overcome with the application of the most advanced Da Vinci Xi Surgical system, which has revolutionary improvements in its docking, targeting, and operating procedures. Second, although we did not compare the operative time of the abdominal phase and thoracic phase separately between the two surgical approaches, we have an impression that robotic surgery has no clear advantages in the abdominal phase due to the need of excessive visceral manipulation and multiple applications of mechanical devices by assistants. Moreover, even in robotic surgery, the final stage of abdominal surgery, known as a feeding jejunostomy, must be performed laparoscopically.

Lymph node dissection plays a crucial role in ensuring patient survival after a radical esophagectomy.<sup>25</sup> Mediastinal lymph node dissection, especially along the bilateral RLNs, requires a meticulous technique to achieve the complete removal of lymphatic chains and intact preservation of nerves. Compared with conventional thoracoscopy, the robot system has a better exposure of the upper mediastinum due to its ability to offer a three-dimensional, self-controlled magnified view and readily adjustable tractions on esophagus and trachea by the full use of the third robotic arm, which would otherwise be the work of an experienced assistant in TAILE.<sup>26</sup> In the abovementioned study comparing the short-term outcomes between RAILE and TAILE conducted by Park et al., they found that the robot system enabled a more radical lymphadenectomy, especially in the upper mediastinum.<sup>23</sup> The early experience of RAILE from the University of Pittsburgh suggested increased lymph node counts compared with the large cohort of conventional TAILE.<sup>27</sup> Chao et al. also showed that compared with thoracoscopic-assisted esophagectomy, robotic surgery resulted in a higher lymph node yield along the left RLN without increasing morbidity.<sup>28</sup> However, some other comparative studies together with our study demonstrated that the number of total dissected lymph nodes as well as the lymph nodes along the bilateral RLNs was comparable between the two surgical approaches.<sup>29,30</sup> These controversial results may be related to the surgeon's personal experience and the number of patients enrolled, which deserve further comparative analysis.

Although no significant advantages were shown in our study for the use of robotics during Ivor-Lewis esophagectomy, questions remain whether robotic surgery has brought some contributions to MIE. In TAILE, the intrathoracic anastomosis is the most challenging and difficult step during the thoracic phase due to the limited flexibility of the instrument tip and the two-dimensional visualization. Stapled intrathoracic anastomosis was mostly employed including transthoracic or transoral placement of the anvil and introduction of the circular stapler through a small thoracotomy.<sup>31</sup> This process requires close coordination within the surgical team, including the surgeon, assistant, and camera holder within the crowded surgical area. Even in RAILE, this process is sometimes difficult due to the rigid stapler in limited thoracic space and the placement of multiple robotic arms. As for hand-sewn anastomosis, few cases have been published to date with this technique during TAILE due to its high technical requirements of suturing ability.<sup>32,33</sup> However, we found this technique simple and reliable with robotic assistance, which has become our dominant anastomotic technique in RAILE. The magnified three-dimensional view and the angulation given to the needle holder by the robotic system make it feasible to perform a two-layer continuous or

interrupted suture without technical difficulties. In our recent study, we have demonstrated that the robotic hand-sewn anastomosis was feasible and safe with no increase of the operative time compared with stapled anastomosis.<sup>10</sup> Some other reports also indicated the feasibility and safety of this technique during RAILE.<sup>34–36</sup> The application of the hand-sewn anastomosis may be an example to demonstrate the clinical benefit of robot system in the future.

This study has some limitations. First, it was conducted in a nonrandomized, retrospective manner. The use of robot system was based on patients' economic levels and their own preferences, which may cause selection bias. Even with the PSM analysis, this selection bias could not be ruled out. Second, this is still an early experience for comparisons between RAILE and TAILE, which included 76 cases of RAMIE and 108 cases of MIE, respectively. Previous studies demonstrated that the learning curve of RAMIE takes approximately 24–70 cases, and approximately 35–40 procedures are needed to reach proficiency for conventional MIE cases.<sup>37,38</sup> The current results of this early comparative study were definitely partly influenced by the learning curve. Moreover, the goal of the study was to gain short-term perioperative outcomes, and a longer follow-up is necessary to compare the oncology results between these two approaches.

In conclusion, we reported the comparative perioperative outcomes between RAILE and TAILE. As such, RAILE should be considered as an alternative minimally invasive method to TAILE for treating esophageal cancer.

**ACKNOWLEDGMENT** This work was supported by grants from the Shanghai Municipal Education Commission-Gaofeng Clinical Medicine Grant Support (20172005) and the Shanghai Jiao Tong University Cooperation Grant of Medicine, Science and Engineering (YG2015QN39). The authors thank Dr. Maosheng Huang, statistician from the Department of Epidemiology, the University of Texas MD Anderson Cancer Center for review of the manuscript.

**DISCLOSURE** The authors declare no conflicts of interest.

## REFERENCES

1. Bray F, Ferlay J, Soerjomataram I, Siegel RL, Torre LA, Jemal A. Global cancer statistics 2018: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries. *CA Cancer J Clin.* 2018;68(6):394–424.
2. Chen W, Zheng R, Baade PD, et al. Cancer statistics in China, 2015. *CA Cancer J Clin.* 2016;66(2):115–32.
3. Lagergren J, Smyth E, Cunningham D, Lagergren P. Oesophageal cancer. *Lancet.* 2017;390(10110):2383–96.
4. Napier KJ, Scheerer M, Misra S. Esophageal cancer: a review of epidemiology, pathogenesis, staging workup and treatment modalities. *World J Gastrointest Oncol.* 2014;6(5):112–20.
5. Lewis I. The surgical treatment of carcinoma of the oesophagus; with special reference to a new operation for growths of the middle third. *Br J Surg.* 1946;34:18–31.

6. Boone J, Livestro DP, Elias SG, Borel Rinkes IH, van Hillegersberg R. International survey on esophageal cancer: part I surgical techniques. *Dis Esophagus*. 2009;22(3):195–202.
7. Hulscher JB, Tijssen JG, Obertop H, van Lanschoot JJ. Transthoracic versus transhiatal resection for carcinoma of the esophagus: a meta-analysis. *Ann Thorac Surg*. 2001;72(1):306–13.
8. Lazzarino AI, Nagpal K, Bottle A, Faiz O, Moorthy K, Aylin P. Open versus minimally invasive esophagectomy: trends of utilization and associated outcomes in England. *Annals Surg*. 2010;252(2):292–8.
9. Moon DH, Lee JM, Jeon JH, Yang HC, Kim MS. Clinical outcomes of video-assisted thoracoscopic surgery esophagectomy for esophageal cancer: a propensity score-matched analysis. *J Thorac Dis*. 2017;9(9):3005–12.
10. Zhang Y, Xiang J, Han Y, et al. Initial experience of robot-assisted Ivor-Lewis esophagectomy: 61 consecutive cases from a single Chinese institution. *Dis Esophagus*. 2018;31(12):doy048.
11. Cerfolio RJ, Bryant AS, Hawn MT. Technical aspects and early results of robotic esophagectomy with chest anastomosis. *J Thorac Cardiovasc Surg*. 2013;145(1):90–6.
12. Hodari A, Park KU, Lace B, Tsiouris A, Hammoud Z. Robot-assisted minimally invasive ivor lewis esophagectomy with real-time perfusion assessment. *Ann Thorac Surg*. 2015;100(3):947–52.
13. Wee JO, Bravo-Iniguez CE, Jaklitsch MT. Early experience of robot-assisted esophagectomy with circular end-to-end stapled anastomosis. *Ann Thorac Surg*. 2016;102(1):253–9.
14. Talsma K, van Hagen P, Grotenhuis BA, et al. Comparison of the 6th and 7th editions of the UICC-AJCC TNM classification for esophageal cancer. *Ann Surg Oncol*. 2012;19(7):2142–8.
15. Giulianotti PC, Coratti A, Angelini M, et al. Robotics in general surgery: personal experience in a large community hospital. *Arch Surg*. 2003;138(7):777–84.
16. van der Sluis PC, van der Horst S, May AM, et al. Robot-assisted minimally invasive thoracoscopic esophagectomy versus open transthoracic esophagectomy for resectable esophageal cancer: a randomized controlled trial. *Ann Surg*. (2018). <https://doi.org/10.1097/SLA.00000000000030313>.
17. van der Sluis PC, Ruurda JP, Verhage RJ, et al. Oncologic long-term results of robot-assisted minimally invasive thoraco-laparoscopic esophagectomy with two-field lymphadenectomy for esophageal cancer. *Ann Surg Oncol*. 2015;22 Suppl 3:S1350–6.
18. Kernstine KH. The first series of completely robotic esophagectomies with three-field lymphadenectomy: initial experience. *Surg Endosc*. 2008;22(9):2102.
19. Dunn DH, Johnson EM, Morpew JA, Dilworth HP, Krueger JL, Banerji N. Robot-assisted transhiatal esophagectomy: a 3-year single-center experience. *Dis Esophagus* 2013;26(2):159–66.
20. de la Fuente SG, Weber J, Hoffe SE, Shridhar R, Karl R, Meredith KL. Initial experience from a large referral center with robotic-assisted Ivor Lewis esophagogastrectomy for oncologic purposes. *Surg Endosc*. 2013;27(9):3339–47.
21. Sarkaria IS, Rizk NP, Grosser R, et al. Attaining proficiency in robotic-assisted minimally invasive esophagectomy while maximizing safety during procedure development. *Innovations*. 2016;11(4):268–73.
22. Sarkaria IS, Rizk NP, Finley DJ, et al. Combined thoracoscopic and laparoscopic robotic-assisted minimally invasive esophagectomy using a four-arm platform: experience, technique and cautions during early procedure development. *Eur J Cardiothorac Surg*. 2013;43(5):e107–15.
23. Park S, Hwang Y, Lee HJ, Park IK, Kim YT, Kang CH. Comparison of robot-assisted esophagectomy and thoracoscopic esophagectomy in esophageal squamous cell carcinoma. *J Thorac Dis*. 2016;8(10):2853–61.
24. Deng HY, Huang WX, Li G, et al. Comparison of short-term outcomes between robot-assisted minimally invasive esophagectomy and video-assisted minimally invasive esophagectomy in treating middle thoracic esophageal cancer. *Dis Esophagus*. 2018; 31(8):doy012.
25. Tsurumaru M, Kajiyama Y, Udagawa H, Akiyama H. Outcomes of extended lymph node dissection for squamous cell carcinoma of the thoracic esophagus. *Ann Thorac Cardiovasc Surg*. 2001;7(6):325–9.
26. van der Horst S, Weijs TJ, Ruurda JP, et al. Robot-assisted minimally invasive thoraco-laparoscopic esophagectomy for esophageal cancer in the upper mediastinum. *J Thorac Dis*. 2017;9(Suppl 8):S834–42.
27. Okusanya OT, Sarkaria IS, Hess NR, et al. Robotic assisted minimally invasive esophagectomy (RAMIE): the University of Pittsburgh Medical Center initial experience. *Ann Cardiothorac Surg*. 2017;6(2):179–85.
28. Chao YK, Hsieh MJ, Liu YH, Liu HP. Lymph node evaluation in robot-assisted versus video-assisted thoracoscopic esophagectomy for esophageal squamous cell carcinoma: a propensity-matched analysis. *World J Surg*. 2018;42(2):590–8.
29. He H, Wu Q, Wang Z, et al. Short-term outcomes of robot-assisted minimally invasive esophagectomy for esophageal cancer: a propensity score matched analysis. *J Cardiothorac Surg*. 23 2018;13(1):52.
30. Weksler B, Sharma P, Moudgill N, Chojnacki KA, Rosato EL. Robot-assisted minimally invasive esophagectomy is equivalent to thoracoscopic minimally invasive esophagectomy. *Dis Esophagus*. 2012;25(5):403–9.
31. Maas KW, Biere SS, Scheepers JJ, et al. Minimally invasive intrathoracic anastomosis after Ivor Lewis esophagectomy for cancer: a review of transoral or transthoracic use of staplers. *Surg Endosc*. 2012;26(7):1795–802.
32. Elshaer M, Gravante G, Tang CB, Jayanthi NV. Totally minimally invasive two-stage esophagectomy with intrathoracic hand-sewn anastomosis: short-term clinical and oncological outcomes. *Dis Esophagus*. 2018;31(3):dox150.
33. Cadiere GB, Dapri G, Himpens J, Fodderie L, Rajan A. Ivor Lewis esophagectomy with manual esogastric anastomosis by thoracoscopy in prone position and laparoscopy. *Surg Endosc*. 2010;24(6):1482–5.
34. Diez Del Val I, Loureiro Gonzalez C, Larburu Etxaniz S, et al. Contribution of robotics to minimally invasive esophagectomy. *J Robotic Surg*. 2013;7(4):325–32.
35. Trugeda S, Fernandez-Diaz MJ, Rodriguez-Sanjuan JC, Palazuelos CM, Fernandez-Escalante C, Gomez-Fleitas M. Initial results of robot-assisted Ivor-Lewis oesophagectomy with intrathoracic hand-sewn anastomosis in the prone position. *Int J Med Robot*. 2014;10(4):397–403.
36. Bongiolatti S, Anneschiarico M, Di Marino M, et al. Robot-sewn Ivor-Lewis anastomosis: preliminary experience and technical details. *Int J Med Robot*. 2016;12(3):421–426.
37. van der Sluis PC, Ruurda JP, van der Horst S, Goense L, van Hillegersberg R. Learning curve for robot-assisted minimally invasive thoracoscopic esophagectomy: results from 312 cases. *Ann Thorac Surg*. 2018;106(1):264–71.
38. Tapias LF, Morse CR. Minimally invasive Ivor Lewis esophagectomy: description of a learning curve. *J Am Coll Surg*. 2014;218(6):1130–40.