



LA CHIRURGIA MINI-INVASIVA DELLE NEOPLASIE DEL CARDIAS E DELL'ESOFAGO

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Minimally Invasive Surgery for cardia and esophageal cancer

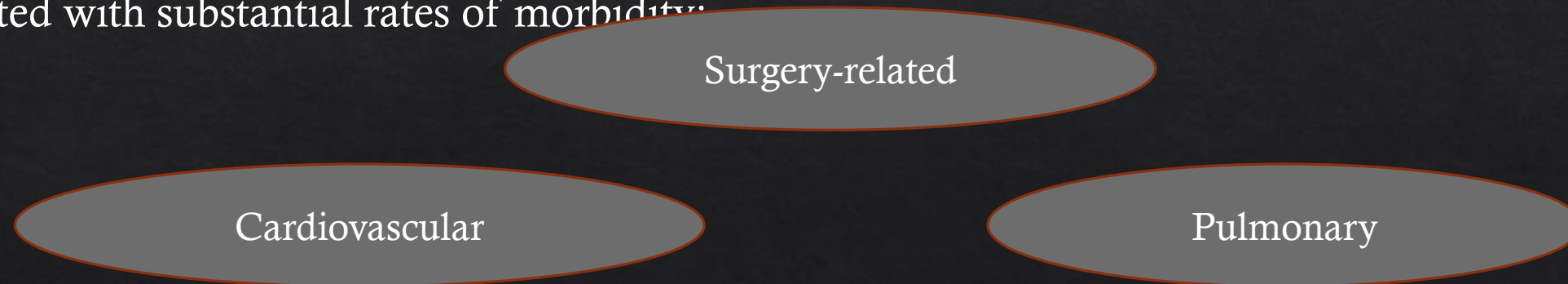
Background:

Surgical resection with radical lymphadenectomy, usually after the administration of neoadjuvant chemotherapy or chemoradiotherapy, remains the key component in the multimodality treatment of cardia/esophageal cancer.

The 5-year survival is 10 to 25% among all patients but increases to 40% among patients who undergo curative surgery.

Esophagectomy is a complex surgical procedure for which the mortality rates have historically been significant.

In modern practice, in high-volume centers with appropriate multidisciplinary teams, the mortality rate after esophageal resection has been reduced significantly. Despite this, it remains an operation associated with substantial rates of morbidity:



In studies with large cohort of patients receiving esophagectomy, mortality ranged between 2.7 % and 9.8 % with a morbidity of 17.9–57 %.

© Springer International Publishing Switzerland 2017 171 S. Giacobuzzi et al. (eds.), *Adenocarcinoma of the Esophagogastric Junction: From Barrett's Esophagus to Cancer*, DOI 10.1007/978-3-319-2878_18



MEDICAL

respiratory 21-27%
arrhythmias

SURGICAL

leaks 3,7-14%
chylotorax
necrosis of gastric conduit

HIGH VOLUME
CENTERS

MULTIDISCIPLINARY
TEAM

[Ann Surg](#), 2009 Nov;250(5):798-807. doi: 10.1097/SLA.0b013e3181bdd5a8.

Postoperative complications after transthoracic esophagectomy for cancer of the esophagus and gastroesophageal junction are correlated with early cancer recurrence: role of systematic grading of complications using the modified Clavien classification.

[Lerut T¹](#), [Moons J](#), [Coosemans W](#), [Van Raemdonck D](#), [De Leyn P](#), [Decaluwé H](#), [Decker G](#), [Nafteux P](#).

[World J Surg](#) (2015) 39:216–222
DOI 10.1007/s00268-014-2754-1

World Journal
of Surgery

ORIGINAL SCIENTIFIC REPORT

Major Post-Operative Complications Predict Long-Term Survival After Esophagectomy in Patients with Adenocarcinoma of the Esophagus

[Guillaume Luc](#) · [Marlène Durand](#) · [Laurence Chiche](#) · [Denis Collet](#)

By the early 1990s, some surgeons had developed and used protocols for thoracoscopic esophagectomy, initially restricting its use to T1 and T2 esophageal cancer without neoadjuvant chemoradiation.

With time indications for minimally invasive esophageal resection have been expanded to include more advanced disease, irrespective of whether patients have received neoadjuvant treatments.

THORACOSCOPIC EN BLOC TOTAL ESOPHAGECTOMY WITH RADICAL MEDIASTINAL LYMPHADENECTOMY

Takashi Akaishi, MD
Iwao Kaneda, MD
Norio Higuchi, MD
Yoshiki Kuriya, MD
Jun-ichi Kuramoto, MD
Tsuneo Toyoda, MD
Akio Wakabayashi, MD

Objective: Total esophagectomy with en bloc mediastinal lymphadenectomy for cancer carries a substantial morbidity and mortality rate. To investigate the feasibility of thoracoscopic technique, we carried out an extensive laboratory study. Encouraged by our excellent results, we conducted a clinical trial. **Methods:** From September 1994 to September 1995, 39 patients thoracic esophageal cancer lesions not invading surrounding organs underwent total esophagectomy with mediastinal lymphadenectomy by means of thoracoscopy. Ages ranged from 47 to 86 years. The procedures were conventional except for the thoracic portion, which was performed as a thoracoscopic procedure with six trocar holes instead of thoracotomy. All harvested lymph nodes were counted for each station. Spirometric data and plethysmographically determined vital capacity were measured before and after operation for all patients. **Results:** All procedures were accomplished as scheduled, and none was converted to open thoracotomy. The operating time was 200 ± 41 minutes (mean \pm standard deviation). Estimated blood loss was 270 ± 157 ml. The harvested lymph nodes numbered 19.7 ± 11.1 per patient. Seventeen patients (45%) had positive lymph nodes. There were no in-hospital deaths within 30 days. Twenty-two patients did not require postoperative ventilatory support. Vital capacity decreased to $85\% \pm 11\%$ of the preoperative values, and forced expiratory volume in 1 second decreased to $82\% \pm 16\%$. **Conclusions:** Thoracoscopic mediastinal lymphadenectomy is technically feasible, and its completeness is comparable to that of the open technique. The decline in pulmonary function is significantly less than that seen in our previous experience with the open technique. (J Thorac Cardiovasc Surg 1996;112:1533-41)

Endosc Surg Allied Technol. 1994 Feb;2(1):21-5.

Thoracoscopic subtotal oesophagectomy.

Cuschieri A¹.

Author information

Abstract

A technique for endoscopic-assisted oesophagectomy is described using either a lateral or prone position for the oesophageal dissection. In a consecutive series of 34 patients, two were found to have hepatic deposits at preliminary laparoscopy, and four were inoperable at **thoracoscopic** staging. A further two had fibrous obliteration of the pleural cavity. Of the 26 procedures, 20 were performed in the lateral position and six in the prone position. There was one conversion to open thoracotomy due to massive aortic bleeding. There were no deaths. Postoperative complications included pneumonia (n = 3), recurrent laryngeal nerve palsy (n = 2) and one anastomotic leak. The median postoperative stay was 12 days (range 9-30 days). It is suggested that the prone position has technical advantages and reduces the postoperative respiratory complications.

Session IV: Esophageal Disease

Thoracoscopic Esophagectomy: Technique and Initial Results

Dominique Gossot, MD, Pierre Fourquier, MD, and Michel Celerier, MD

Department of Surgery, Saint-Louis Hospital, Paris, France

To reduce the high morbidity rate associated with esophageal surgery, we have developed a technique of thoracoscopic esophagectomy. A feasibility study was first carried out in an animal model and a specific instrument was developed for this purpose. Esophagectomy using a right thoracoscopic approach was attempted in 15 patients, 13 males and 2 females whose average age was 48 years. Indications consisted of squamous cell carcinoma in 10 patients, adenocarcinoma in 1, and caustic stenosis in 4. We used a technique that consisted of double-lumen tracheal intubation and the creation of five ports. The whole esophagus was mobilized thoracoscopically and the esophagectomy was completed through the abdomen. The reconstruction was achieved using a gastric pull-through, and the anastomosis was made in the neck.

There were three failures: in 1 patient there was a large tumor, making the exposure unsafe, and, in 2 patients, incomplete lung collapse made exposure of the posterior mediastinum difficult. These 3 cases were converted into a thoracotomy. The thoracoscopic dissection was successful in the remaining 12 patients. The average time of the thoracoscopic stage was 125 minutes. The postoperative course was uneventful in 10 patients. Two patients had a left atelectasis. Although our series is limited, these initial results indicate that thoracoscopic esophagectomy is feasible. However, further evaluation of the technique is needed to assess its benefit in terms of respiratory morbidity.

(Ann Thorac Surg 1993;56:667-70)

Unlike other minimally invasive procedures, minimally invasive esophagectomy (MIE) has not been broadly adopted. No matter what approach is used, MIE remains a very complex operation

Advanced disease

Morbidity and mortality

Oncological radicality

Reproducibility

Cost



TOPICS UNDER DEBATE

The primary goal of MIE is to decrease surgical morbidity associated with the open approach. Present, the majority of data derives from retrospective nonrandomized series and suggests that mortality rates appear equivalent with some suggestion of benefit in terms of overall morbidity favoring a minimally invasive approach

Clinical outcomes of video-assisted thoracoscopic surgery esophagectomy for esophageal cancer: a propensity score-matched analysis

Duk Hwan Moon¹, Jong Mog Lee², Jae Hyun Jeon², Hee Chul Yang², Moon Soo Kim²

Indian J Surg (August 2017) 79(4):319–325
DOI 10.1007/s12262-016-1519-5



Review Article

ORIGINAL ARTICLE

Minimally Invasive Ivor-Lewis Esophagectomy (MIILE): A Single-Center Experience

Jun Wang¹ · Mei-qing Xu¹ · Ming-ran Xie¹ · Xiu-yu Mei¹

Surgical Endoscopy (2018) 32:4957–4965
<https://doi.org/10.1007/s00464-018-6257-2>



Total minimally invasive esophagectomy for esophageal adenocarcinoma reduces postoperative pain and pneumonia compared to hybrid esophagectomy

Felix Berth¹ · Patrick S. Plum¹ · Seung-Hun Chon¹ · Christian A. Gutschow² · Elfriede Bollschweiler¹ · Arnulf H. Hölscher³

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Surg Endosc (2015) 29:2614–2619
DOI 10.1007/s00464-014-3978-8



Open versus minimally invasive esophagectomy: clinical outcomes for locally advanced esophageal adenocarcinoma

Juha Kauppi · Jari Räsänen · Eero Sihvo ·
Riikka Huuhtanen · Kaisa Nelskylä ·
Jarmo Salo

Minimally invasive esophagectomy for Barrett's adenocarcinoma

Emanuele Asti, Daniele Bernardi, Marco Sozzi, Luigi Bonavina

Surg Endosc (2010) 24:1621–1629
DOI 10.1007/s00464-009-0822-7

Is minimally invasive surgery beneficial in the management of esophageal cancer? A meta-analysis

Kamal Nagpal · Kamran Ahmed · Amit Vats ·
Danny Yakoub · David James · Hutan Ashrafian ·
Ara Darzi · Krishna Moorthy · Thanos Athanasiou

Table 2 Results of meta-analysis comparing MIE versus open and HMIE versus open esophagectomy

Outcome	No. of studies	No. of patients	OR/WMD	95% CI	P-value	HG χ^2	HG P-value
MIE versus open							
Operative time	5	312	5.91	-39.53, 51.34	0.80	29.13	<0.001
Blood loss	5	312	-268.53	-369.91, -167.16	<0.001	12.14	0.02
ICU stay	3	216	-0.97	-1.31, -0.63	<0.001	1.16	0.56
Length of stay	4	267	-2.75	-4.65, -0.86	0.004	2.01	0.57
GI bleeding	2	171	0.32	0.04, 2.81	0.30	0.01	0.91
Anastomotic leak	5	447	0.58	0.28, 1.20	0.14	3.43	0.49
Anastomotic stricture	4	379	1.58	0.30, 8.40	0.59	15.07	0.002
Gastric conduit ischemia	3	236	1.80	0.34, 9.60	0.49	1.88	0.39
Respiratory complications	5	447	0.58	0.35, 0.98	0.04	2.91	0.57
Cardiac complications	3	368	1.11	0.51, 2.45	0.79	0.42	0.81
Chyle leak	5	447	1.37	0.35, 5.47	0.65	2.73	0.44
Vocal cord palsy	5	447	0.76	0.19, 3.10	0.71	0.12	0.94
Total morbidity	5	447	0.52	0.32, 0.84	0.007	4.36	0.36
30-day mortality	5	447	0.55	0.19, 1.57	0.26	0.04	0.98
Number of LN retrieved	4	240	1.02	-0.84, 2.88	0.28	2.76	0.43
HMIE versus open							
Operative time	3	653	5.64	-31.77, 43.05	0.77	18.60	<0.001
Blood loss	3	653	-147.87	-297.40, 1.67	0.05	8.58	0.01
Anastomotic leak	4	806	0.51	0.28, 0.91	0.02	0.50	0.92
Respiratory complications	4	806	0.68	0.48, 0.96	0.03	1.12	0.77
Cardiac complications	4	806	0.67	0.42, 1.07	0.09	2.44	0.49
Chyle leak	2	562	1.48	0.23, 9.53	0.68	1.81	0.18
Vocal cord palsy	4	806	1.54	0.89, 2.67	0.13	1.45	0.69
Total morbidity	3	725	0.62	0.31, 1.25	0.18	5.17	0.08
30-day mortality	4	806	0.56	0.19, 1.60	0.28	0.92	0.34
Number of LN retrieved	3	632	-1.44	-6.61, 3.73	0.58	19.37	<0.001

MIE minimally invasive esophagectomy, HMIE hybrid minimally invasive esophagectomy, LN lymph node, ICU intensive care unit, OR odds ratio, WMD weighted mean difference, CI confidence interval, HG heterogeneity

Authors (year)	n	Approaches	Mortality n (%)	Overall morbidity n (%)
Law et al. (1997) [10]	22	MIE (TSO)	0	18 (81.8)
	63	Open	0	63 (100)
Nguyen et al. (2000) [11]	18	MIE (TLSO)	0	7 (38.9)
	36	Open	0	19 (52.8)
Osugi et al. (2003) [12]	77	MIE (VATS)	0	31 (40.3)
	72	Open	0	32 (44.4)
Kunisaki et al. (2004) [13]	15	MIE (VATS+HALS)	0	NS
	30	Open	0	NS
Van den Broek et al. (2004) [14]	25	MIE (THO)	0	14 (70)
	20	Open	0	18 (72)
Bresadola et al. (2006) [15]	14	MIE (THO and TLSO)	0	8 (57.1)
	14	Open	0	6 (42.9)
Bernabe et al. (2005) [16]	17	MIE (THO)	0	NS
	14	Open	0	NS
Shiraishi et al. (2006) [17]	116	MIE (TLSO)	3 (2.6)	NS
	37	Open	3 (8.1)	NS
Braghetto et al. (2006) [18]	47	MIE (VATS/LSO)	3 (6.3)	18 (38.2)
	119	Open	13 (10.9)	72 (60.5)
Smithers et al. (2007) [19]	332	MIE (TLSO)	7 (2.1)	207 (62.3)
	114	Open	3 (2.6)	76 (66.7)
Fabian et al. (2008) [9]	22	MIE (TLSE)	1 (4.5)	15 (68.2)
	43	Open	4 (9.8)	31 (72.1)
Zingg et al. (2009) [20]	56	MIE (TLSO)	2 (3.6)	19 (34.5)
	98	Open	6 (6.1)	20 (23.5)
Perry et al. (2009) [21]	21	MIE (LIO)	0	13 (62)
	21	Open	1 (5)	17 (81)
Parameswaran et al. (2009) [22]	50	MIE (TLSO)	1 (2)	24 (48)
	30	Open	1 (3)	15 (50)
Pham et al. (2010) [23]	44	MIE (TLSO)	3 (6.8)	NS
	46	Open	2 (4.3)	NS
Schoppeman et al. (2010) [24]	31	MIE (TLSO)	0	11 (35.5)
	31	Open	0	23 (74.2)
Singh et al. (2010) [25]	33	MIE (TLSO)	Values NS	Values NS
	31	Open	p=0.34	P=0.06
Mamidanna et al. (2012) [26]	1155	MIE (TLSO, HMIO)	46 (4.0)	NS
	6347	Open	274 (4.3)	NS
Ben-David et al. (2012) [27]	100	MIE (TLSO)	1 (1)	NS
	32	Open	2 (5)	NS
Briez et al. (2012) [28]	140	MIE (HMIO)	2.1	35.7
	140	Open	12.9	59.3
Xie et al. (2014) [29]	106	MIE (TLSO)	2 (1.9)	28 (26.4)
	163	Open	4 (2.5)	56 (34.4)
Hsu et al. (2014) [30]	66	MIE (TLSO)	5 (7.6)	NS
	63	Open	5 (7.9)	NS

The TRIALS

Biere et al. *BMC Surgery* 2011, 11:2
<http://www.biomedcentral.com/1471-2482/11/2>



STUDY PROTOCOL

Open Access

Traditional invasive vs. minimally invasive esophagectomy: a multi-center, randomized trial (TIME-trial)

Surya S A Y Biere¹, Kirsten W Maas¹, Luigi Bonavina³, Josep Roig Garcia⁴, Mark I van Berge Henegouwen⁵, Camiel Rosman⁶, Meindert N Sosef², Elly S M de Lange², H Jaap Bonjer¹, Miguel A Cuesta^{1*}, Donald L van der Peet^{1*}

Minimally invasive versus open oesophagectomy for patients with oesophageal cancer: a multicentre, open-label, randomised controlled trial

Surya S A Y Biere, Mark I van Berge Henegouwen, Kirsten W Maas, Luigi Bonavina, Camiel Rosman, Josep Roig Garcia, Suzanne S Gisbertz, Jean H G Klinkenbijl, Markus W Hollmann, Elly S M de Lange, H Jaap Bonjer, Donald L van der Peet, Miguel A Cuesta

RANDOMIZED CONTROLLED TRIAL

Minimally Invasive Versus Open Esophageal Resection Three-year Follow-up of the Previously Reported Randomized Controlled Trial: the TIME Trial

Jennifer Straatman, MD, PhD,* Nicole van der Wielen, MD,* Miguel A. Cuesta, MD, PhD,*
Freek Daams, MD, PhD,* Josep Roig Garcia, MD, PhD,† Luigi Bonavina, MD, PhD,‡
Camiel Rosman, MD, PhD,§ Mark I. van Berge Henegouwen, MD, PhD,¶
Suzanne S. Gisbertz, MD, PhD,¶ and Donald L. van der Peet, MD, PhD*

The TIME trial is prospective, multi-center, randomized study comparing traditional transthoracic esophageal resection with minimally invasive resection for esophageal cancer.



TIME TRIAL

June 1, 2009 - March 31, 2011

Amsterdam, Milan, Girona

18-75 years; cT1-3, N0-1, M0

At least 10 MIE

More than 30 oesophagectomies per year

56 open (laparotomy/thoracotomy); 59 MIE (thoracoscopy/laparoscopy)

Primary outcome: postoperative pulmonary infection within first 2 weeks of surgery and during the whole stay in hospital

Secondary outcomes: other complications (leaks, chylothorax, reoperation,...); intraoperative data (blood loss,...); postoperative data (ICU, VAS,...); histological data (lymph nodes, pStage, resection margins, ...); mortality

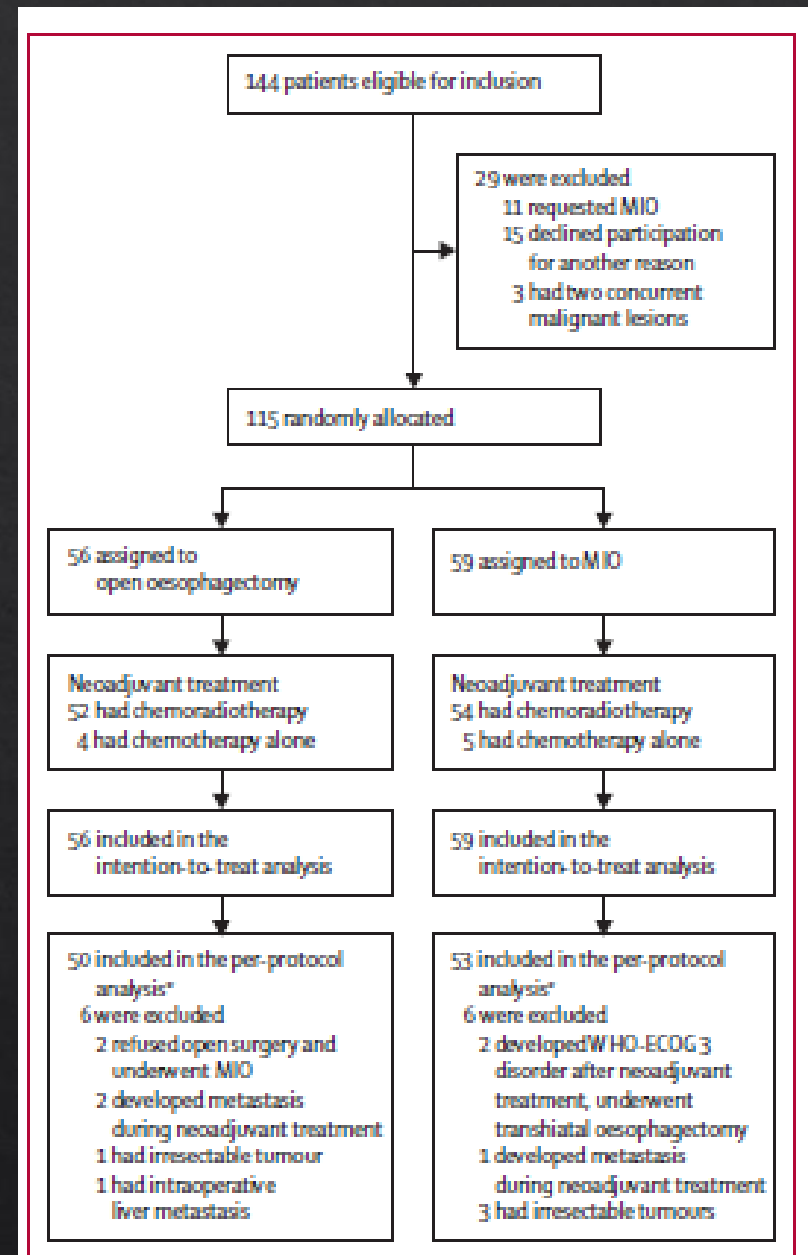


Figure: Trial profile

MIO= minimally invasive oesophagectomy. ECOG=Eastern Cooperative Oncology Group. *See appendix for per-protocol analysis of primary and secondary outcome parameters.

Short-term outcomes

	OO (N=56)	MIO (N=59)	p value
Primary outcomes			
Pulmonary infection within 2 weeks	16 (29%)	5 (9%)	0.005
Pulmonary infection in-hospital	19 (34%)	7 (12%)	0.005
Secondary outcomes			
Hospital stay (days)*	14 (1-120)	11 (7-80)	0.044
Short-term quality of life†			
SF 36†			
Physical component summary	36 (6; 34-39)	42 (8; 39-46)	0.007
Mental component summary	45 (11; 40-50)	46 (10; 41-50)	0.806
EORTC C30†			
Global health	51 (21; 44-58)	61 (18; 56-67)	0.020
OES 18‡			
Talking	37 (39; 25-49)	18 (26; 10-26)	0.008
Pain	19 (21; 13-26)	8 (11; 5-11)	0.002
Total lymph nodes retrieved*	21 (7-47)	20 (3-44)	0.852
Resection margin§			0.080
R0	47 (84%)	54 (92%)	-
R1	5 (9%)	1 (2%)	-
pStage¶			
0	0 (0%)	1 (2%)	-
I	4 (7%)	4 (7%)	-
Ila	16 (29%)	17 (29%)	-
Ilb	6 (11%)	9 (15%)	-
III	14 (25%)	11 (19%)	-
IV	5 (9%)	4 (7%)	-
No residual tumour or lymph-node metastasis	7 (13%)	9 (15%)	-
Mortality			
30-day mortality	0 (0%)	1 (2%)	-
In-hospital mortality	1 (2%)	2 (3%)	-

Data are n (%), median (range), or mean (SD, 95% CI), unless otherwise indicated. OO=open oesophagectomy. MIO=minimally invasive oesophagectomy. SF 36=Short Form 36 Health Survey (version 2). EORTC=European Organization for Research and Treatment of Cancer Quality of Life Questionnaires. *Skewed distribution, Mann-Whitney test applied. †Measures general aspects of health; scores range from 0 to 100, with higher scores representing better well-being. ‡Assesses several aspects of oesophageal function; scores range from 0 to 100, with lower scores indicating better function. Only statistically significant domains presented. §Defined as >1 mm from a resection margin. ¶Staging based on the American Joint Committee on Cancer, 6th ed; four patients in each group did not undergo resection due to metastasis or irresectability of the tumour. ||Death from any cause.

Table 2: Primary and secondary outcomes for the intention-to-treat population

	OO (N=56)	MIO (N=59)	p value
Intraoperative data			
Operative time (min)**†	299 (66-570)	329 (90-559)	0.002
Blood loss (mL)†	475 (50-3000)	200 (20-1200)	<0.001
Conversions‡	NA	8 (14%)	-
Level of anastomosis§			
Cervical	37 (66%)	38 (64%)	0.970
Thoracic	15 (27%)	17 (29%)	
Postoperative data			
ICU stay (days)†	1 (0-106)	1 (0-50)	0.706
VAS (10 days)¶	3 (2)	2 (2)	0.001
Epidural failure	11 (20%)	10 (17%)	0.734
Other complications			
Anastomotic leakage	4 (7%)	7 (12%)	0.390
Thoracic complications without anastomotic leakage**	2 (4%)	2 (3%)	0.958
Vocal-cord paralysis††	8 (14%)	1 (2%)	0.012
Pulmonary embolism	0 (0%)	1 (2%)	0.328
Reoperations	6 (11%)	8 (14%)	0.641

Data are median (range), n (%), or mean (SD), unless otherwise indicated. OO=open oesophagectomy. MIO=minimally invasive oesophagectomy. NA=not applicable. ICU=intensive-care unit. VAS= Visual Analogue Scale pain score. *Time from skin incision to skin closure. †Skewed distribution, Mann-Whitney test applied. ‡Six patients were converted to thoracotomy and two to laparotomy. §Four patients in the OO group and four in the MIO group did not undergo resection with subsequent anastomosis because of metastasis or irresectability of the tumour. ¶Linear mixed model. ||In the first 2 days after surgery. **Thoracic complications not related to leakage were mediastinitis, empyema, chyloous leakage needing reoperation, and hiatal herniation. ††Confirmed by laryngoscopy.

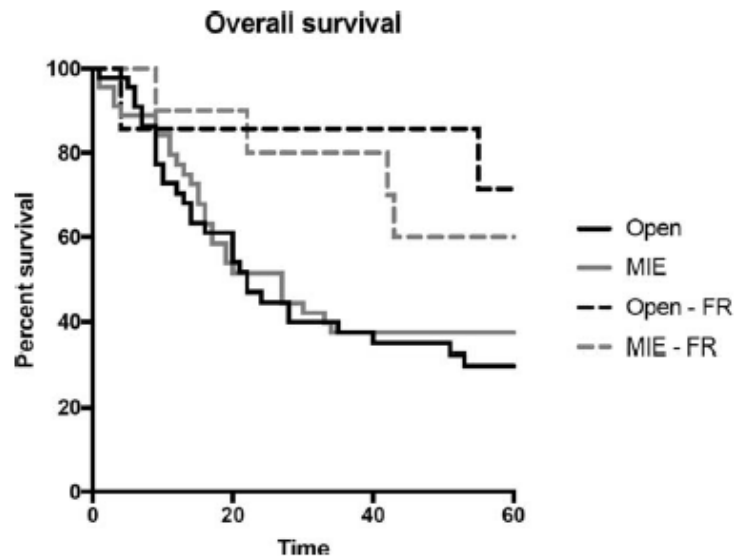
Table 3: Other outcomes of the intention-to-treat population

- Lower incidence of pulmonary infection
- Shorter hospital stay
- Better short-term QoL
- No compromise in the quality of resected specimen
- No significant differences in complications
- Less intraoperative bleeding

Final endpoints: long-term survival analysis: three-year follow-up

Minimally Invasive Versus Open Esophageal Resection
Three-year Follow-up of the Previously Reported Randomized Controlled Trial:
the TIME Trial

Jennifer Straatman, MD, PhD,* Nicole van der Wielen, MD,* Miguel A. Cuesta, MD, PhD,*
Freek Daams, MD, PhD,* Josep Roig Garcia, MD, PhD,† Luigi Bonavina, MD, PhD,‡
Camiel Rosman, MD, PhD,§ Mark I. van Berge Henegouwen, MD, PhD,¶
Suzanne S. Gisbertz, MD, PhD,¶ and Donald L. van der Peet, MD, PhD*



No differences
between OS
(41,2% open vs
42,9 MIS) and
DFS (37,3%
open vs 42,9%
MIS)

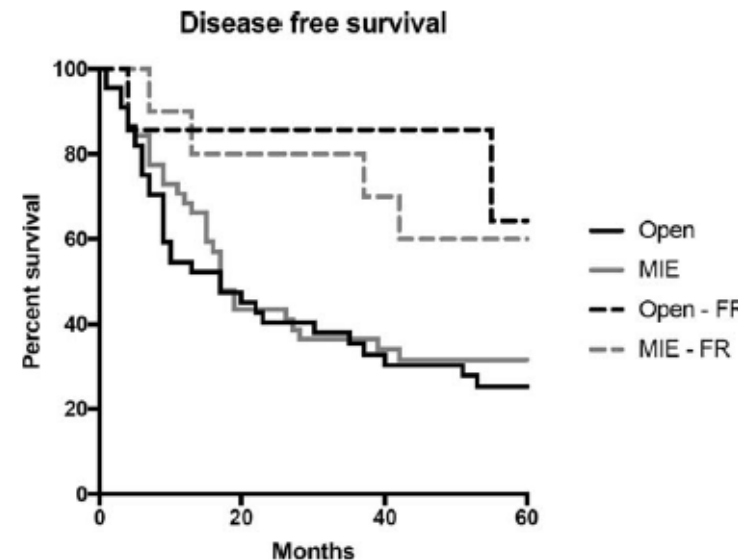


FIGURE 2. Kaplan-Meier curves for comparison of overall survival between open and minimally invasive esophagectomy (FR, full responders with no residual tumor).

TABLE 4. Multivariable Cox Regression Analysis for Overall Survival and Disease-free Survival

	Overall Survival			Disease-free Survival		
	HR	95% CI		HR	95% CI	
Procedure						
Open vs MI	0.961	0.585	1.579	0.946	0.585	1.531
Stage						
I-II vs 0	2.620	1.100	6.243	2.699	1.136	6.415
III-IV vs 0	3.202	1.294	7.922	3.586	1.461	8.802
Gender						
Man vs women	1.557	0.815	2.974	1.592	0.841	3.013
Age						
Per year increase	1.016	0.927	1.045	1.018	0.991	1.047

Recurrence or death, were observed (p=0,602)
In the minimally invasive group, 7 patients had
metastasis. In the open group, 8 patients had
metastasis. No differences were observed

MIRO TRIAL

The MIRO trial is a prospective multicentre controlled randomised phase III trial

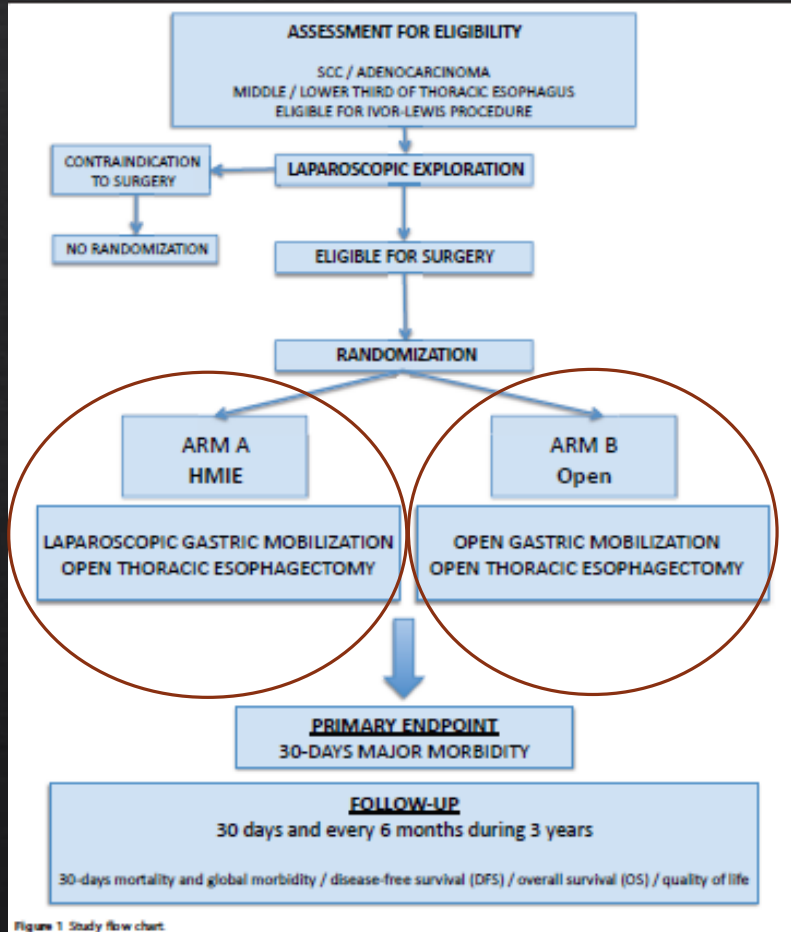


Figure 1 Study flow chart.

STUDY PROTOCOL

Open Access

Open versus laparoscopically-assisted oesophagectomy for cancer: a multicentre randomised controlled phase III trial - the MIRO trial

Nicolas Briez^{1,2}, Guillaume Piessen^{1,2}, Franck Bonnetain³, Cécile Brigand⁴, Nicolas Carrere⁵, Denis Collet⁶, Christophe Doddoli⁷, Renaud Flamein⁸, Jean-Yves Mabrut⁹, Bernard Meunier¹⁰, Simon Msika¹¹, Thierry Perniceni¹², Frédérique Peschaud¹³, Michel Prudhomme¹⁴, Jean-Pierre Triboulet^{1,2} and Christophe Mariette^{1,2*}

13 centers in France
at least 25 procedures

“...We hypothesise that HMIO based on the laparoscopic procedure and an open thoracic approach may provide a significant decrease in major postoperative complications without leading to any negative impact on oncological outcomes...”

Secondary end points: 30 days mortality; 30 days overall complications; major pulmonary complications; DFS; OS.

RESULTS

ORIGINAL ARTICLE

Hybrid Minimally Invasive Esophagectomy for Esophageal Cancer

C. Mariette,* S.R. Markar, T.S. Dabakuyo-Yonli, B. Meunier, D. Pezet, D. Collet, X.B. D'Journo, C. Brigand, T. Perniceni, N. Carrère, J.-Y. Mabrut, S. Msika, F. Peschard, M. Prudhomme, F. Bonnetain,* and G. Piessen, for the Fédération de Recherche en Chirurgie (FRENCH) and French Eso-Gastric Tumors (FREGAT) Working Group†

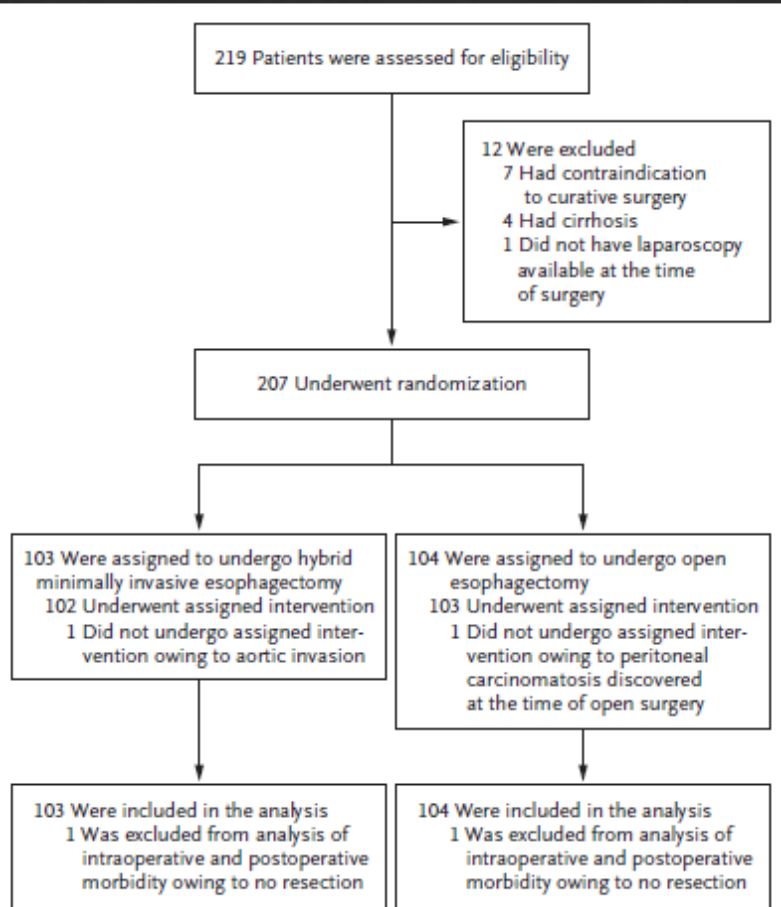


Figure 1. Screening, Randomization, and Follow-up of the Patients.

- **Primary end-point (major intraoperative and postoperative morbidity at 30 days):** 36% HMIE vs 64% open ($p < 0,001$). 77% lower risk of major intraoperative and postoperative complications after esophagectomy
- **Secondary end-point:** no differences in:
 - ✓ mortality at 30 days
 - ✓ intraoperative and postoperative overall morbidity BUT...

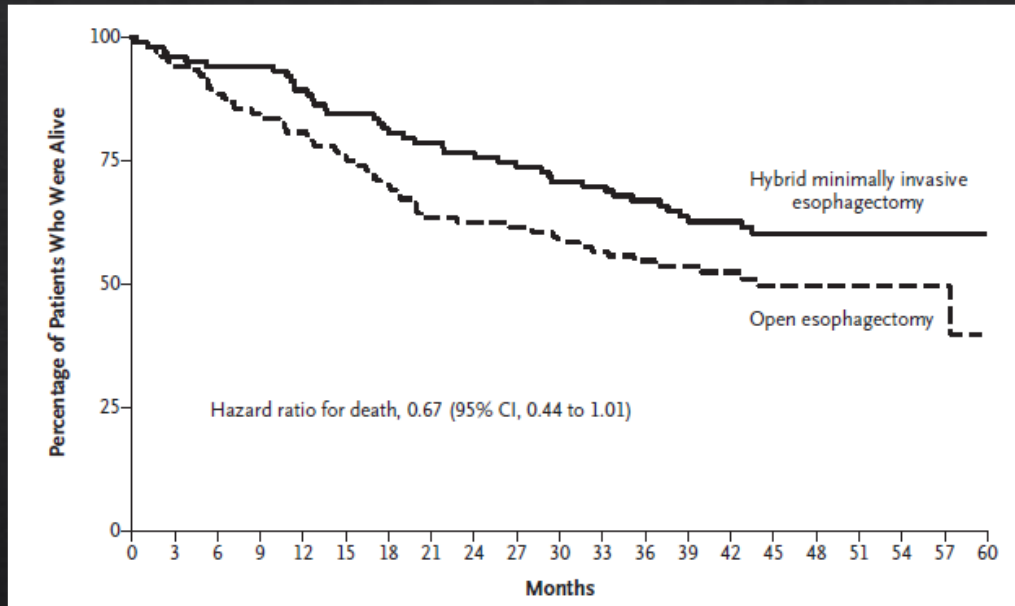


HYBRID MINIMALLY INVASIVE SURGERY WAS ASSOCIATED WITH A LOWER INCIDENCE OF MAJOR PULMONARY COMPLICATIONS WITHIN 30 DAYS (18% vs 30%). 50% lower risk of major pulmonary complications.

Besides no differences in :

- ✓ histologic findings; pathological tumor or node stage
- ✓ total number of nodes resected or positive
- ✓ Resection margin involvement

LONG- TERM RESULTS:

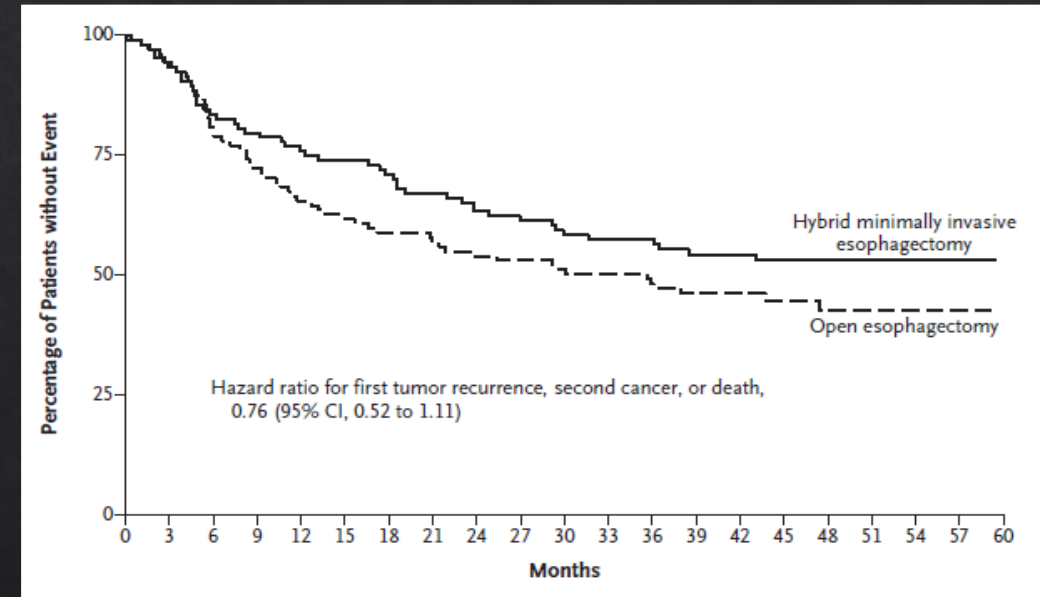


Median Overall Survival:

52,2 months among the 103 patients in hybrid-procedure vs 47,6 among the 104 patients in open-procedure (not significant difference).

OS 3 years: 67% hybrid vs 55% open

OS 5 years: 60% hybrid vs 40% open



Desease-free survival:

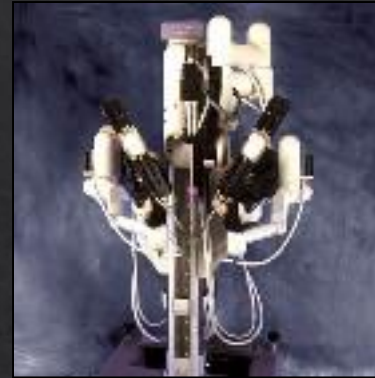
no differ significantly between two groups

DFS 3 years: 57% hybrid vs 48% open

DFS 5 years: 53% hybrid vs 43 % open



Surgeon console



Surgical Arm Cart



InSite™ Image Processing Equipment



InSite™ High-resolution 3-D Endoscope



EndoWrist™ Instruments



Robotic surgery benefits from a stable 3-dimensional, magnified view and articulated instruments enabling precise dissection with 7 degrees of freedom of movement.

First experience with robot-assisted thoracoscopic esophagolymphadenectomy for esophageal cancer

R. van Hillegersberg,¹ J. Boone,¹ W. A. Draaisma,¹ I. A. M. J. Broeders,¹ M. J. M. M. Giezeman,² I. H. M. Borel Rinkes¹

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Received: 4 October 2005/Accepted: 27 January 2006/Online publication: 15 May 2006

Original article

Robot-assisted thoracoscopic oesophagectomy for cancer

J. Boone¹, M. E. I. Schipper², W. A. Moojen¹, I. H. M. Borel Rinkes¹, G. J. E. Cromheecke³ and R. van Hillegersberg¹

Departments of ¹Surgery, ²Pathology and ³Anaesthesiology, University Medical Centre Utrecht, Utrecht, The Netherlands

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Robotic-assisted minimally invasive esophagectomy versus the conventional minimally invasive one:
a meta-analysis and systematic review

Dacheng Jin, MMed^{1,2,3,4†}, Liang Yao, MMed^{6†}, Jun Yu, MMed², Rong Liu, MD⁵, Tiankang Guo, MD³,
Kehu Yang, MD^{3,4*}, Yunjiu Gou, MD^{2*}

Clinical Study

Robot-Assisted Hybrid Esophagectomy Is Associated with a Shorter Length of Stay Compared to Conventional Transthoracic Esophagectomy: A Retrospective Study

Hans C. Rolff, Rikard B. Ambrus, Mohammed Belmouhand, Michael P. Achiam, Marianne Wegmann, Mette Siemsen, Steen C. Kofoed, and Lars B. Svendsen

THE INTERNATIONAL JOURNAL OF MEDICAL ROBOTICS AND COMPUTER ASSISTED SURGERY

Int J Med Robotics Comput Assist Surg 2016; 12: 421–426.

Published online 29 September 2015 in Wiley Online Library (wileyonlinelibrary.com) DOI: 10.1002/rcs.1705

ORIGINAL ARTICLE

Robot-sewn Ivor-Lewis anastomosis: preliminary experience and technical details

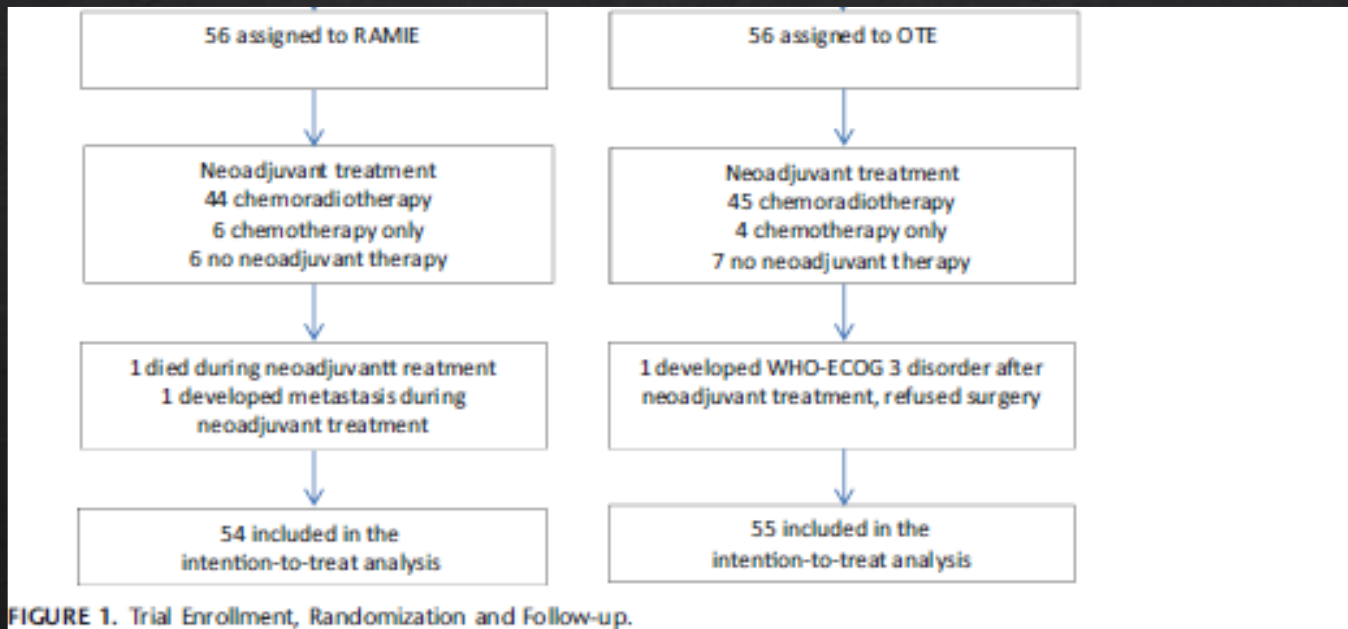
Robot-assisted minimally invasive thoraco-laparoscopic esophagectomy versus open transthoracic esophagectomy for resectable esophageal cancer, a randomized controlled trial (ROBOT trial)

Pieter C van der Sluis^{1*}, Jelle P Ruurda¹, Sylvia van der Horst¹, Roy JJ Verhage¹, Marc GH Besselink¹, Margriet JD Prins¹, Leonie Haverkamp¹, Carlo Schippers¹, Inne HM Borel Rinkes¹, Hans CA Joore², Fiebo JW ten Kate³, Hendrik Koffijberg⁴, Christiaan C Kroese², Maarten S van Leeuwen⁵, Martijn PJK Lolkema⁷, Onne Reerink⁸, Marguerite B Schipper⁸, Elles Steenhagen⁹, Frank P Weggaar⁹, Emile E Voest⁸, Peter D Siersema⁹ and Richard van Hillegersberg^{1*}

RANDOMIZED CONTROLLED TRIAL

Robot-assisted Minimally Invasive Thoracolaparoscopic Esophagectomy Versus Open Transthoracic Esophagectomy for Resectable Esophageal Cancer

A Randomized Controlled Trial



Primary end-point: % of overall surgery related postoperative complications

Secondary end-points: pulmonary complications, cardiac complications and post-operative bleeding

Were recorded: in-hospital mortality; mortality within 30 and 60 days, length of ICU, hospital stay; operation time, blood loss

OS; DFS; post-operative functional recovery; QoL questionnaire

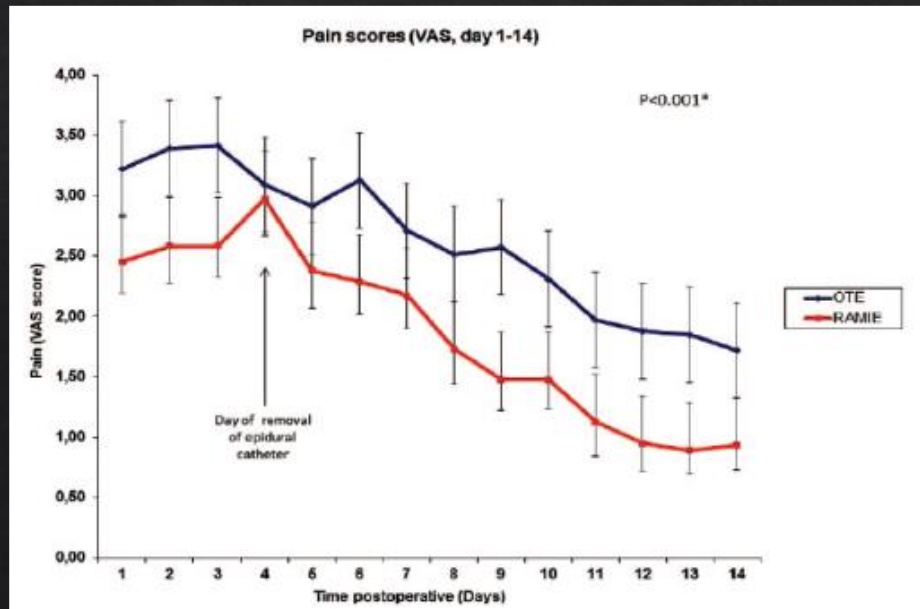
Robot-assisted thoraco-laparoscopic esophagectomy
VS
Open transthoracic esophagectomy

ROBOT TRIAL: RESULTS

Primary endpoint (overall surgery-related complications): 32/54 (59%) after RAMIE vs 44/55 (80%) after OTE ($p=0,02$)

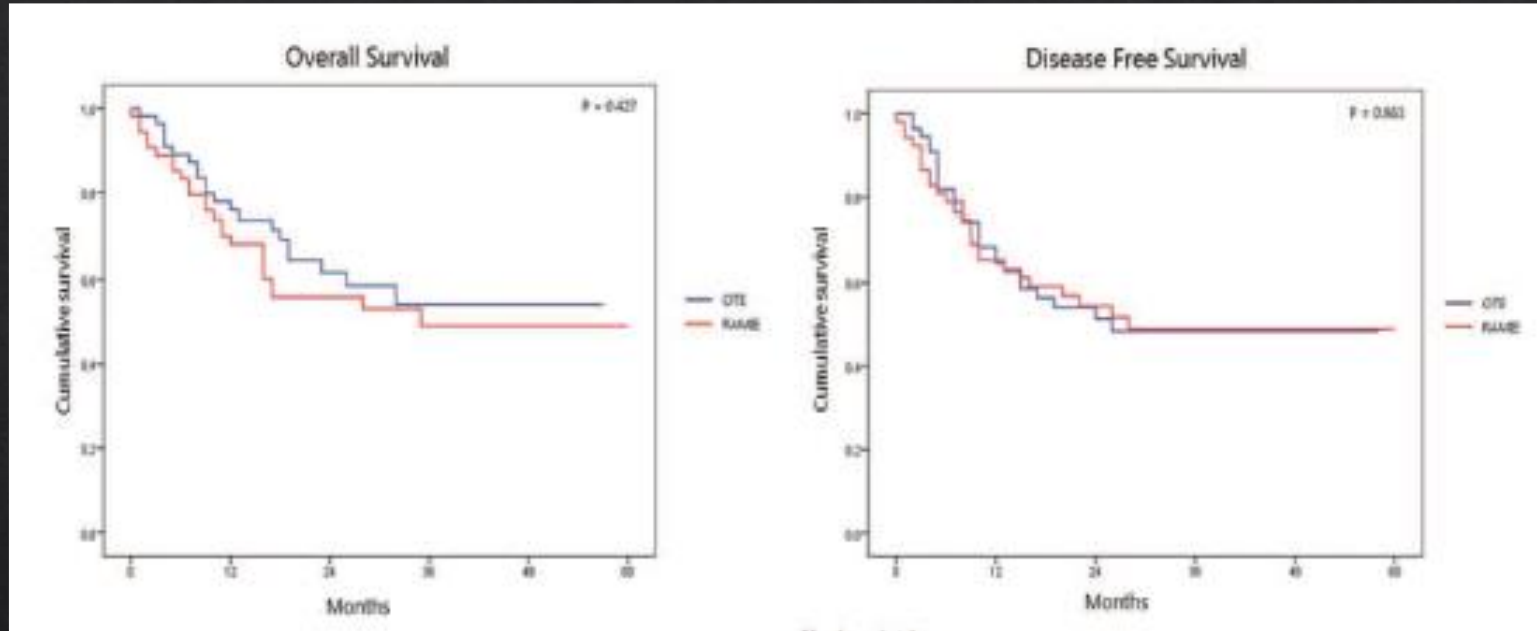
Others:

- Pulmonary complications: 32% RAMIE vs 58% OTE ($p=0,005$)
- Cardiac complications: 22% RAMIE vs 47% OTE ($p=0,006$)
- No significant differences in mortality and in all other complications
- Functional recovery at postoperative day 14 significantly better in RAMIE group (70%) vs OTE group (51%)
 $p=0.04$
- Short-term QoL higher in RAMIE both at discharge and 6 weeks post-discharge





Mean postoperative pain (VAS) during first 14 days significantly lower after RAMIE ($p < 0,001$)

- Intraoperative outcomes:
 - operative time : $p < 0,001$
 - blood loss: $p < 0,001$
- No differences in radical resection, lymph nodes retrieved and all pathologic outcomes



No differences in OS and in DFS

Robot-assisted minimally invasive esophagectomy (RAMIE) compared to conventional minimally invasive esophagectomy (MIE) for esophageal cancer: a propensity-matched analysis

E. Tagkalos,^{1,*} L. Goense ^{2,*} M. Hoppe-Lotichius,¹ J.P. Ruurda,² B. Babic,¹ E. Hadzijusufovic,¹ W. Kneist,¹ P.C. van der Sluis,¹ H. Lang,¹ R. van Hillegersberg,² P. P. Grimmering ¹

¹Department of General, Visceral and Transplant Surgery, University Medical Center of the Johannes Gutenberg University, Mainz, Germany and ²Department of Surgery, University Medical Center Utrecht, Utrecht, the Netherlands

SUMMARY. Robot-assisted minimally invasive esophagectomy (RAMIE) is increasingly being applied as treatment for esophageal cancer. In this study, the results of 50 RAMIE procedures were compared with 50 conventional minimally invasive esophagectomy (MIE) operations, which had been the standard treatment for esophageal cancer prior to the robotic era. Between April 2016 and March 2018, data of 100 consecutive patients with esophageal carcinoma undergoing modified Ivor Lewis esophagectomy were prospectively collected. All operations were performed by the same surgeon using an identical intrathoracic anastomotic reconstruction technique with the same perioperative management and pain control regimen. Intra-operative and postoperative complications were graded according to definitions stated by the Esophagectomy Complications Consensus Group. Data analysis was carried out with and without propensity score matching. Baseline characteristics did not show significant differences between the RAMIE and MIE group. Propensity score matching of the initial group of 100 patients resulted in two equal groups of 40 patients for each surgical approach. In the RAMIE group, the median total lymph node yield was 27 (range 13–84) compared to 23 in the MIE group (range 11–48), $P = 0.053$. Median intensive care unit (ICU) stay was 1 day (range 1–43) in the RAMIE group compared to 2 days (range 1–17) in the MIE group ($P = 0.029$). The incidence of postoperative complications was not significantly different between the two groups ($P = 0.581$). In this propensity-matched study comparing RAMIE to MIE, ICU stay was significantly shorter in the RAMIE group. There was a trend in improved lymphadenectomy in RAMIE.

KEY WORDS: esophageal cancer, Ivor Lewis, MIE, minimally invasive, RAMIE.

	MIE		RAMIE		P-value
	N		N		
Cumulative op time	321	(224–519)	383	(255–488)	0.001
Duration of the abdominal part	126	(66–335)	151	(80–250)	0.007
Duration of the thoracic part	202	(158–304)	223	(163–320)	0.068
Blood loss	350	±196	331	±207	0.670
Anastomotic leakage	9	18%	6	12%	0.575
Pneumonia	9	18%	6	12%	0.575
Wound infection	2	4%	0	0%	0.448
30-d mortality	1	2%	0	0%	0.315
90-d mortality	1	2%	2	4%	0.560
ICU stay	2.5	(1–25)	1	(1–43)	0.020
Hospital stay	19	(9–55)	12	(7–59)	0.413
T status					0.001
T0	14	28%	5	10%	
T1	14	28%	5	10%	
T2	10	20%	10	20%	
T3 + T4	12	24%	30	60%	
N status					0.169
N0	27	54%	21	42%	
N1	10	20%	9	18%	
N2	10	20%	10	20%	
N3	3	6%	10	20%	
M status					0.992
M0	49	98%	48	96%	
M1	1	2%	2	4%	
R0 status	49	98%	46%	92%	0.359
Number of resected lymph nodes	23	(11–48)	27	(13–84)	0.043
(b) After propensity matching					
Cumulative op time	321	(224–519)	388	(255–475)	0.001
Time of the abdominal part	125	(66–335)	151	(80–250)	0.001
Time of the thoracic part	201	(158–295)	223	(170–320)	0.052
Blood loss	343	±181	339	±198	0.930
Anastomotic leakage	5	12.5%	5	12.5%	0.966
Pneumonia	7	17.5%	6	15%	0.763
Wound infection	1	2.5%	0	0%	0.298
30-d mortality	1	2.5%	0	0%	0.311
90-d mortality	1	2.5%	2	5%	0.479
ICU stay	2	(1–17)	1	(1–43)	0.029
Hospital stay	12.5	(9–54)	12	(7–59)	0.685
T status					0.004
T0	13	32.5%	4	10%	
T1	11	27.5%	4	10%	
T2	6	15%	9	22.5%	
T3 + 4	10	25%	22	57.5%	
N status					0.210
N0	23	57.5%	16	40%	
N1	6	15%	8	20%	
N2	9	22.5%	8	20%	
N3	2	5%	8	20%	
M status					0.990
M0	39	97.5%	40	100%	
M1	1	2.5%	0	0%	
R0 status	39	97.5%	38	95%	1.000
Number of resected lymph nodes	23	(11–48)	27	(13–84)	0.053

Robot-assisted minimally invasive esophagectomy (RAMIE) compared to conventional minimally invasive esophagectomy (MIE) for esophageal cancer: a propensity-matched analysis

E. Tagkalos,^{1,*} L. Goense^{1,2,*} M. Hoppe-Lotichius,¹ J.P. Ruurda,² B. Babic,¹ E. Hadzijusufovic,¹ W. Kneist,¹ P.C. van der Sluis,¹ H. Lang,¹ R. van Hillegersberg,² P.P. Grimmering¹

¹Department of General, Visceral and Transplant Surgery, University Medical Center of the Johannes Gutenberg University, Mainz, Germany and ²Department of Surgery, University Medical Center Utrecht, Utrecht, the Netherlands

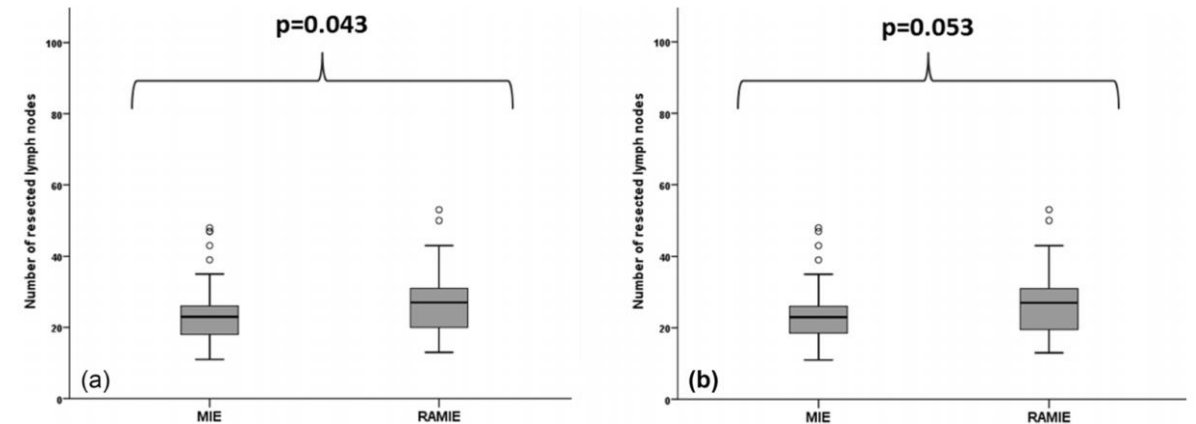


Fig. 1 Box-plot chart presenting the number of resected lymph nodes of MIE and RAMIE; (a) prior to pair matching; (b) after pair matching. The unmatched group (a) reveals a statistical significance in favor of the RAMIE group ($P = 0.043$). Nonetheless the match-paired group (b) shows a trend ($P = 0.053$) for the RAMIE group.

STUDY PROTOCOL

Open Access

Distribution of lymph node metastases in esophageal carcinoma [TIGER study]: study protocol of a multinational observational study



Abstract

Background: An important parameter for survival in patients with esophageal carcinoma is lymph node status. The distribution of lymph node metastases depends on tumor characteristics such as tumor location, histology, invasion depth, and on neoadjuvant treatment. The exact distribution is unknown. Neoadjuvant treatment and surgical strategy depends on the distribution pattern of nodal metastases but consensus on the extent of lymphadenectomy has not been reached. The aim of this study is to determine the distribution of lymph node metastases in patients with resectable esophageal or gastro-esophageal junction carcinoma in whom a transthoracic esophagectomy with a 2- or 3-field lymphadenectomy is performed. This can be the foundation for a uniform worldwide staging system and establishment of the optimal surgical strategy for esophageal cancer patients.

(Continued on next page)

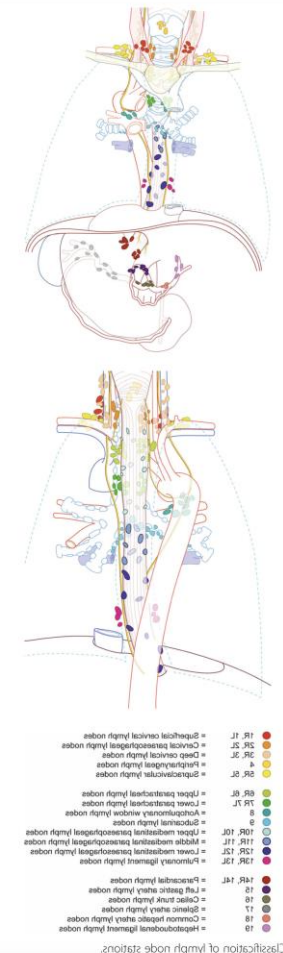
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Methods: The TIGER study is an international observational cohort study with 50 participating centers. Patients with a resectable esophageal or gastro-esophageal junction carcinoma in whom a transthoracic esophagectomy with a 2- or 3-field lymphadenectomy is performed in participating centers will be included. All lymph node stations will be excised and separately individually analyzed by pathological examination. The aim is to include 5000 patients. The primary endpoint is the distribution of lymph node metastases in esophageal and esophago-gastric junction carcinoma specimens following transthoracic esophagectomy with at least 2-field lymphadenectomy in relation to tumor histology, tumor location, invasion depth, number of lymph nodes and lymph node metastases, pre-operative diagnostics, neo-adjuvant therapy and (disease free) survival.

Discussion: The TIGER study will provide a roadmap of the location of lymph node metastases in relation to tumor histology, tumor location, invasion depth, number of lymph nodes and lymph node metastases, pre-operative diagnostics, neo-adjuvant therapy and survival. Patient-tailored treatment can be developed based on these results, such as the optimal radiation field and extent of lymphadenectomy based on the primary tumor characteristics.

Trial registration: [NCT03222895](https://www.clinicaltrials.gov/ct2/show/study/NCT03222895), date of registration: July 19th, 2017.

Keywords: Esophageal cancer, Lymph node metastases, Lymphadenectomy, Esophagectomy,



OUR EXPERIENCE
in Minimally Invasive
Esophagectomy

Chirurgia Robotica

Dipartimento di Scienze Chirurgiche – Università di Torino

Clinica Chirurgica I prof. M. Morino

Gennaio 2002 – Gennaio 2005

65 pazienti – 68 procedure

27 M - 38 F – Età media 46.2 (range 18 - 72)

<i>Procedure Robotiche</i>	<i>N° pazienti</i>	<i>N° conversioni</i>
Plastica anti-reflusso	26*	1 (4.3%)
Surrenalectomia	22 (6right-8left)	4 (28.6%)
Cholecistectomia	10	0
Miotomia sec. Heller + plastica antireflusco sec Dor	4	0
Emicolectomia dx	3	1 (33%) <i>open</i>

* +3 *colecistectomie associate*

DIPARTIMENTO DI SCIENZE CHIRURGICHE
Clinica Chirurgica I – Università di Torino (Prof. M. Morino)

BY-PASS GASTRICO LAPAROSCOPICO

ROBOT-ASSISTITO

novembre 2007 – luglio 2019



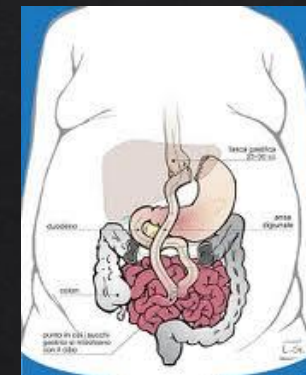
358 pazienti (306 F/ 52 M)

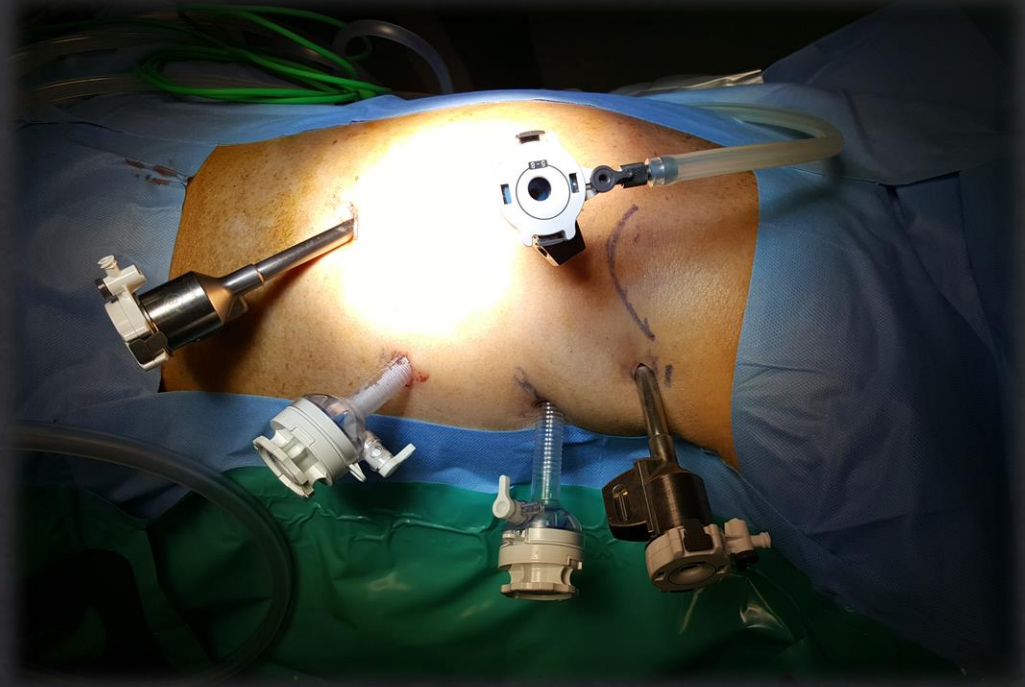
ETÀ 41.2 ± 8.2 (22 – 62) anni

PESO 115.3 ± 13.9 (83 – 146) Kg

BMI 43.4 ± 3.4 (33.7 – 49.8) kg/m²

EW% 98.2 ± 18.1 (55.1 – 129.1) %





From 23th April 2018 we performed 25 esophagectomies with minimally invasive approach.

**CLINICA CHIRURGICA I – DIPARTIMENTO DI SCIENZE CHIRURGICHE
UNIVERSITA' di TORINO**

Prof. Morino M.

CASISTICA ESOFAGECTOMIA MINI-INVASIVA (MIE)

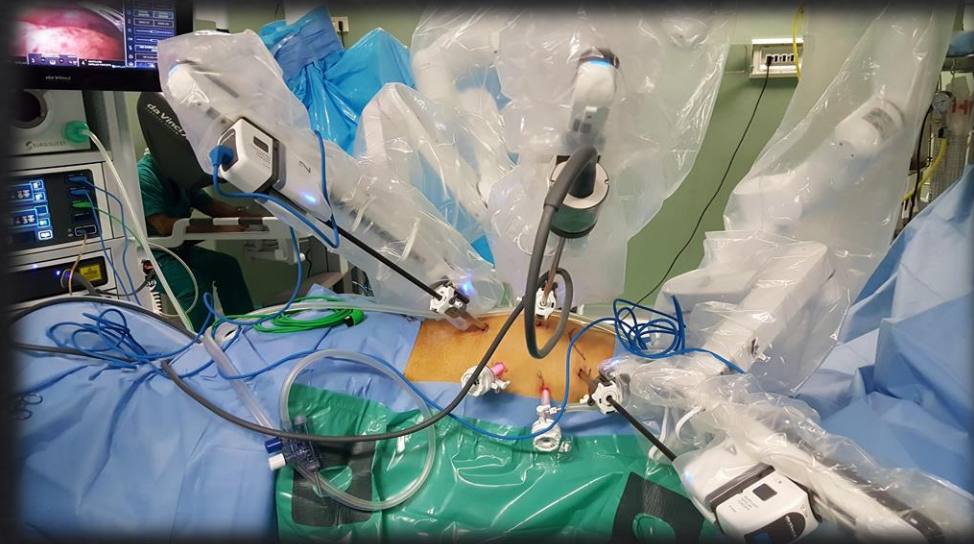
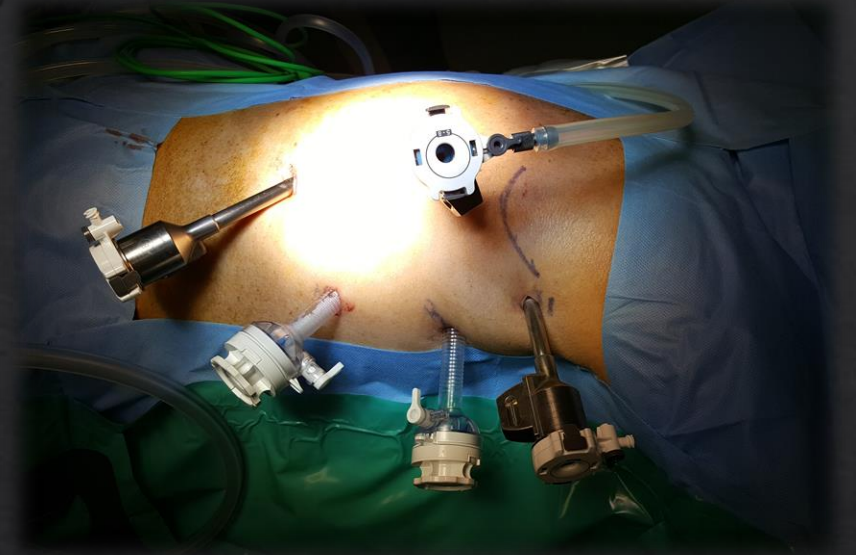
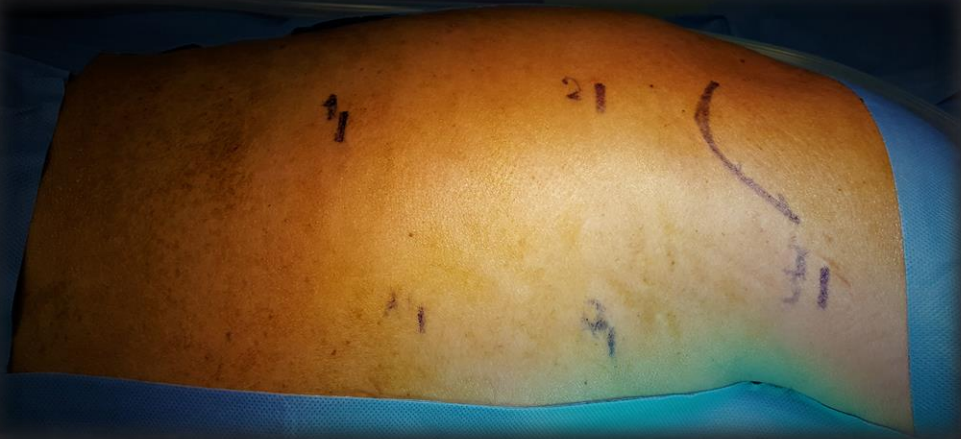
Aprile 2018 – Luglio 2019						
N° pazienti	25					
Maschi/Femmine	22/3					
Età	69.5 (range 45-82)					
Tipo istologico	AdenoCa 22		Ca squamoso 3			
Sede: Cardias	Siewert 1	8 casi	Siewert 2	6 casi	Siewert 3	1 caso
Esofago prossimale	2 casi					
Esofago medio	4 casi					
Esofago distale	4 casi					
Trattamenti preoperatori:	19					
Chemio:	8					
Rx	2					
Chemio + Rx	9					

**CLINICA CHIRURGICA I – DIPARTIMENTO DI SCIENZE CHIRURGICHE
UNIVERSITA' di TORINO
Prof. Morino M.**

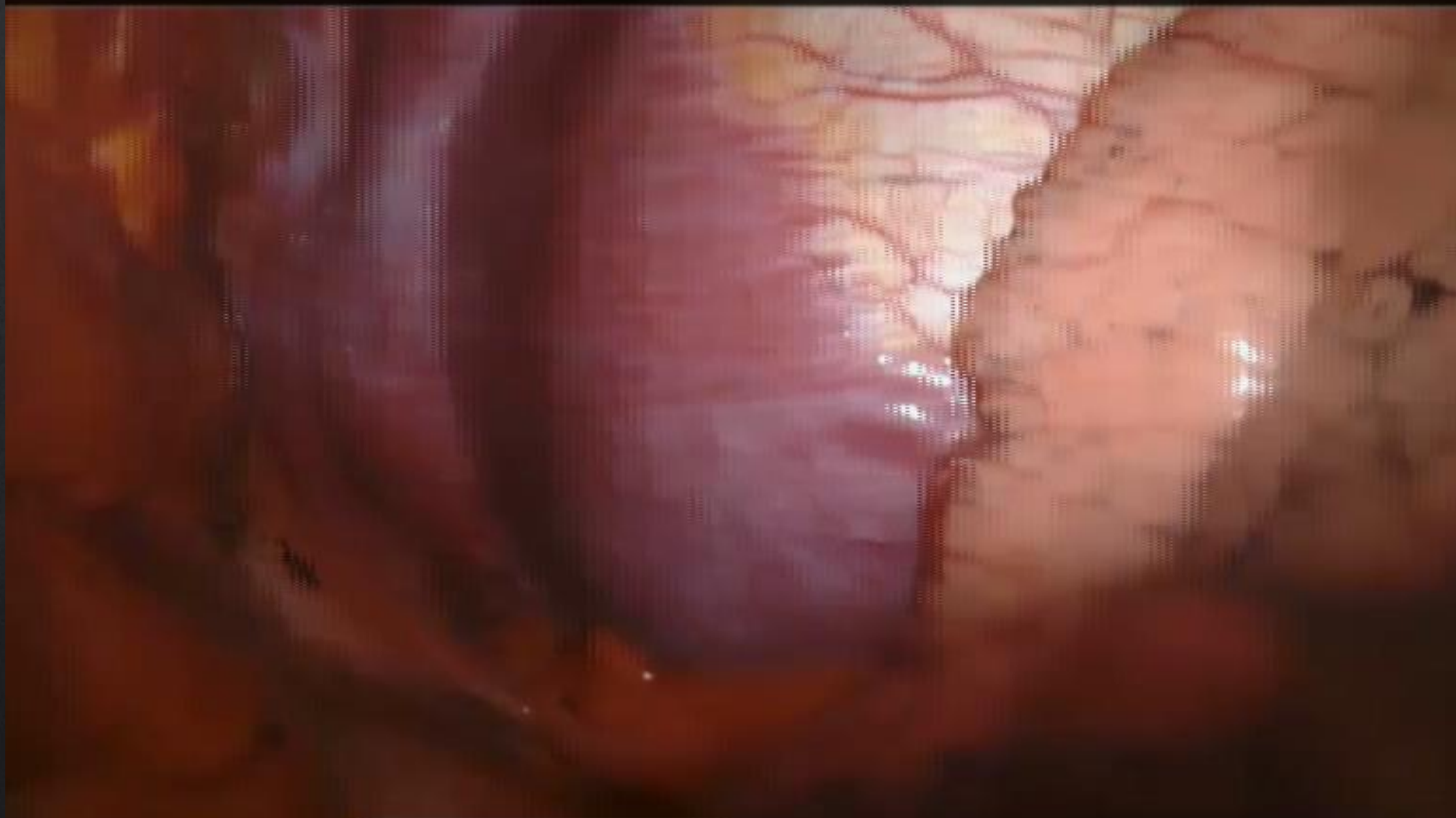
CASISTICA ESOFAGECTOMIA MINI-INVASIVA (MIE)

Aprile 2018 – Luglio 2019	
Tipo di intervento:	
Ivor Lewis (laparo addome + robotica toracica)	23
Mc Keown (robotica toracica, laparo addominale, cervicotomica)	3
Tempi operatori	
Conversioni:	tempo addominale: 1 caso di conversione open tempo toracico : 0

PORT PLACEMENT IN ROBOT-ASSISTED THORACOSCOPY IN IVOR-LEWIS PROCEDURE









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UNIVERSITA' di TORINO**

Prof. Morino M.

CASISTICA ESOFAGECTOMIA MINI-INVASIVA (MIE)

Aprile 2018 – Luglio 2019	
Tempi operatori:	320 (244-520) min
Complicanze maggiori	Fistola anastomotica: 1 Emotorace dx: 1
Complicanze minori	2 casi FA cardiovertita farmacologicamente 1 ritenzione urinaria
Mortalità a 30 gg	0
Reinterventi:	2
Degenza in ICU:	2,3 gg (range 1-7)
Degenza totale:	15.5 gg (range 10-38)
Dolore post-operatorio (mean score VAS):	1,8

**CLINICA CHIRURGICA I – DIPARTIMENTO DI SCIENZE CHIRURGICHE
UNIVERSITA' di TORINO**

Prof. Morino M.

CASISTICA ESOFAGECTOMIA MINI-INVASIVA (MIE)

Aprile 2018 – Luglio 2019	
Stadio TNM:	PT1N0 3 casi pT2N0 5 casi pT3N0 8 casi pT3N1 5 casi pT3N2 4 casi
N° linfonodi asportati	23.9 (range 10-32)
Margini di resezione:	Indenni: 24 Margine circonferenziale < 1 mm: 1