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Does minimally invasive oesophagectomy provide a benefit in hospital length of stay when compared with open oesophagectomy?

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Summary

A best evidence topic in thoracic surgery was written according to a structured protocol. The question addressed was: 'in patients undergoing oesophagectomy, does a minimally invasive approach convey a benefit in hospital length of stay (LOS), when compared to an open approach?' A total of 647 papers were identified, using an *a priori* defined search strategy; 24 papers represented the best evidence to answer the clinical question. The authors, journal, date, country of publication, patient group, study type, relevant outcomes and key results are tabulated. Of the studies identified, data from two randomized controlled trials were available. The first randomized study compared the use of open thoracotomy and laparotomy versus thoracoscopy and laparoscopy. Those undergoing minimally invasive oesophagectomy (MIO) left hospital on average 3 days earlier than those treated with the open oesophagectomy (OO) technique ($P = 0.044$). The other randomized trial, which compared thoracotomy with thoracoscopy and laparoscopy, demonstrated a reduction of 1.8 days in the LOS when employing the MIO technique ($P < 0.001$). With the addition of the remaining 22 non-randomized studies, comprising 3 prospective and 19 retrospective cohort studies, which are heterogeneous with regard to their design, study populations and outcomes; data are available representing 3173 MIO and 25 691 OO procedures. In total, 13 studies (including the randomized trials) demonstrate a significant reduction in hospital LOS associated with MIO; 10 suggest no significant difference between techniques; and only 1 suggests a significantly greater length of stay associated with MIO. The only two randomized trials comparing MIO and OO demonstrated a reduction in length of stay in the MIO group, without compromising survival or increasing complication rates. All but one of the non-randomized studies demonstrated either a significant reduction in length of stay with MIO or no difference. The benefit in reduced length of stay was not at the cost of worsened survival or increased complications, and conversion rates in all studies were low.

Keywords: Oesophagectomy • Minimally invasive • Open • Length of stay • Outcome

INTRODUCTION

A best evidence topic was constructed according to a structured protocol. This is fully described in the *ICVTS* [1].

THREE-PART QUESTION

In [patients undergoing oesophagectomy] does a [minimally invasive approach, compared to an open approach] convey benefit in [hospital length of stay]?

CLINICAL SCENARIO

A 65-year old male patient attends the outpatient department to discuss surgery for oesophageal cancer. You counsel him for open surgery, but he enquires about 'keyhole surgery', and thinks he

would recover more quickly. You resolve to search the literature to provide an evidence-based answer.

SEARCH STRATEGY

A Medline® search from 1950 to November 2014 was performed, using the Ovid interface, with the following terms: 'Oesophag*.mp OR Esophag*.mp' AND 'minimally invasive.mp OR Thoracic Surgery, Video Assisted/ OR vats.mp OR video assisted.mp OR mio.mp OR mie.mp OR thoroscop*.mp OR laparoscop*.mp' AND 'length of stay/ OR time to discharge.mp OR inpatient stay. mp OR patient stay.mp OR Patient Discharge/'.

SEARCH OUTCOMES

Six hundred and forty-seven papers were found using the search strategy. Where relevant, the full text versions of papers were

Table 1: Best evidence papers

Author (year), journal and country, Study type (level of evidence)	n _{OO}	n _{MIO}	Outcomes	OO result	MIO result	Sig. (P-value)	Comments
Biere <i>et al.</i> (2012), Lancet, Netherlands, Spain, Italy [2] Multicentre, open-label randomized controlled trial (level I)	56	59	Inpatient LOS (days)	14	11	0.044	Groups are comparable for all clinical and demographic factors at baseline. Eight conversions from MIO to OO were required OO: thoracotomy and laparotomy MIO: thoracoscopy and laparoscopy
			ITU LOS (days)	1	1	0.706	
			Operative duration (min)	299	329	0.02	
			Estimated blood loss (ml)	475	200	<0.001	
			Lymph node yield	21	20	0.852	
			Mortality	0	1	NS	
			Reoperation	6	8	0.641	
			Anastomotic leak	4	7	0.39	
Conversion	N/A	8	<0.001				
Guo <i>et al.</i> (2013), Chin Ger J Clin Oncol, China [3] Open-label randomized controlled trial (level I)	111	111	Inpatient LOS (days)	11.4 ± 2.3	9.6 ± 1.7	<0.001	Insufficient data presented regarding randomization. Prospective study. In-hospital mortality follow-up only was available. Little data presented regarding baseline characteristics of groups OO: transthoracic MIO: thoracoscopy and laparoscopy
			ITU LOS (days)	3.2 ± 0.6	3.3 ± 0.7	0.256	
			Operative duration (min)	218.7 ± 91	272.3 ± 57.9	<0.001	
			Estimated blood loss (ml)	590 ± 324.4	219.7 ± 194.4	<0.001	
			Lymph node yield	19.2 ± 12.5	24.3 ± 21.0	<0.001	
			Mortality	0	0	NS	
			Reoperation	NR	NR	NR	
			Anastomotic leak	2	1	0.556	
Parameswaran <i>et al.</i> (2009), World J Surg, UK [4] Retrospective cohort study (level IIb)	30	50	Inpatient LOS (days)	10	12	0.01	Patients operated on sequentially; OO 2002–2003; MIO 2004–2006. More females underwent OO, otherwise groups comparable for baseline characteristics OO: Ivor-Lewis MIO: thoracoscopy and laparoscopy
			ITU LOS (days)	NR	NR	NR	
			Operative duration (min)	266	442	<0.01	
			Estimated blood loss (ml)	NR	NR	NR	
			Lymph node yield	10	23	<0.001	
			Mortality	NR	NR	NR	
			Reoperation	5	6	0.75	
			Anastomotic leak	1	4	NS	
Conversion	0	1	NR				
Pham <i>et al.</i> (2010), Am J Surg, USA [5] Retrospective cohort study (level IIb)	46	44	Inpatient LOS (days)	14	15	0.51	Groups comparable for age, ASA, BMI and stage of malignancy. A higher proportion of women underwent OO. OO: Ivor-Lewis MIO: thoracoscopy and laparoscopy
			ITU LOS (days)	4	5	0.35	
			Operative duration (min)	437	543	<0.01	
			Estimated blood loss (ml)	780	407	<0.01	
			Lymph node yield	8	13	<0.01	
			Mortality	2	3	0.34	
			Reoperation	NR	NR	NR	
			Anastomotic leak	4	5	0.78	
Gao <i>et al.</i> (2011), Interact CardioVasc Thorac Surg, China [6] Retrospective cohort study (level IIb)	78	96	Inpatient LOS (days)	17.5 ± 6.4	12.6 ± 8.8	<0.01	Mortality follow-up was available at 30 days. No conversions from MIO to OO. Groups comparable in all demographic, clinical and pathological characteristics OO: McKeown MIO: three-field MIO with thoracotomy and laparoscopy
			ITU LOS (h)	20.0 ± 4.2	19.2 ± 3.5	NS	
			Operative duration (min)	284.3 ± 31.1	330.2 ± 36.7	<0.01	
			Estimated blood loss (ml)	519.3 ± 47.7	346.7 ± 41.1	<0.01	
			Lymph node yield	28 ± 6.2	27.8 ± 5.6	NS	
			Mortality	3	2	NS	
			Reoperation	NR	NR	NR	
			Anastomotic leak	6	7	NS	
Schoppmann <i>et al.</i> (2010), Surg Endosc Austria [7]	31	31	Inpatient LOS (days)	29	15	0.001	Groups comparable in all demographic, clinical and pathological characteristics
			ITU LOS (days)	10	3	0.001	
			Operative duration (min)	400	411	NS	

Continued

Table 1: (Continued)

Author (year), journal and country, Study type (level of evidence)	n_{OO}	n_{MIO}	Outcomes	OO result	MIO result	Sig. (P-value)	Comments
Retrospective cohort study (level IIb)			Estimated blood loss (transfusion units)	0.9 ± 1.9	0.52 ± 1.86	0.014	OO: thoracotomy and midline laparotomy MIO: thoracostomy and laparoscopy
			Lymph node yield	20.5 ± 12.6	17.9 ± 7.7	NS	
			Mortality	6	1	0.04	
			Reoperation	10	4	0.006	
			Anastomotic leak	8	1	0.024	
Sihag <i>et al.</i> (2012), Eur J Cardiothorac Surg, USA [8]	76	38	Inpatient LOS (days)	9 ± 4	7 ± 1	<0.001	Groups comparable for age, sex, tumour histology, clinical stage, preoperative comorbidities and neoadjuvant therapy
			ITU LOS (days)	1 ± 1	1 ± 0	0.001	
			Operative duration (min)	365.5 ± 124	360.5 ± 73	0.542	
			Estimated blood loss (ml)	250 ± 200	200 ± 100	<0.001	
			Lymph node yield	21 ± 9	19 ± 13	0.74	
Retrospective cohort study (level IIb)			Mortality	2	0	0.552	OO: Ivor-Lewis MIO: thoracoscopy/ laparoscopy
			Reoperation	NR	NR	NR	
			Anastomotic leak	0	2	0.552	
			Conversion	N/A	2		
Kinjo <i>et al.</i> (2012), Surg Endosc, Japan [9]	79	72	Inpatient LOS (days)	53	23	<0.001	Also evaluated hybrid MIO (thoracoscopic/ laparotomy-based) approach (data not included). Duration of thoracic procedure only is available
			ITU LOS (days)	1 ± 7	1 ± 2	0.114	
			Operative duration (min)	268 ± 80	308 ± 73	<0.001	
			Estimated blood loss (ml)	680	320	<0.001	
			Lymph node yield	18	28	0.002	
Prospective cohort study (level Ib)			Mortality	0	0	NS	OO: transthoracic MIO: thoracoscopy and laparoscopy
			Reoperation	3	5	0.344	
			Anastomotic leak	3	4	NS	
			Conversion	N/A	3	NR	
Sundaram <i>et al.</i> (2012), Surg Endosc, USA [10]	26	47	Inpatient LOS (days)	14 ± 10	16 ± 10	0.480	Evaluated both open transthoracic and transhiatal approaches (data presented for former approach only). Groups comparable for sex and other variables; patients undergoing MIO were older by 2 years
			ITU LOS (days)	5 ± 3	4 ± 3	0.101	
			Operative duration (min)	480 ± 180	420 ± 190	<0.001	
			Estimated blood loss (ml)	700 ± 550	0 ± 350	<0.001	
			Lymph node yield	19 ± 10	20 ± 13	NS	
Retrospective cohort study (level IIb)			Mortality	0	2	0.577	OO: thoracotomy and laparotomy MIO: modified Ivor-Lewis; thoracoscopy and laparoscopy with intrathoracic anastomosis
			Reoperation	NR	NR	NR	
			Anastomotic leak	4	4	0.183	
			Conversion	N/A	7	NR	
Ben-David <i>et al.</i> (2012), Surg Endosc, USA [11]	32	100	Inpatient LOS (days)	14	7.5	0.05	Different surgical modalities performed at different centres. No group-specific, baseline demographic or clinical data presented. No data on conversions presented
			ITU LOS (days)	NR	NR	NR	
			Operative duration (min)	NR	330 ± 210	NR	
			Estimated blood loss (ml)	NR	125 ± 200	NR	
			Lymph node yield	NR	NR	NR	
Retrospective cohort study (level IIb)			Mortality	2	1	0.04	OO: thoracotomy and laparotomy MIO: thoracoscopy and laparoscopy, or Ivor-Lewis
			Reoperation	NR	NR	NR	
			Anastomotic leak	4	4	0.04	
Dolan <i>et al.</i> (2013), Surg Endosc, USA [12]	64	71	Inpatient LOS (days)	13.5 ± 16.2	12.0 ± 8.0	0.024	Groups comparable for age, sex and stage of malignancy; more MIO patients received chemo-/radiotherapy. One conversion from MIO to OO was required
			ITU LOS (days)	3 ± 5	3 ± 4	0.688	
			Operative duration (min)	579 ± 98	554 ± 112	0.263	
			Estimated blood loss (ml)	500 ± 575	250 ± 150	<0.001	
			Lymph node yield	9.5 ± 10.2	18 ± 9.8	<0.001	
Retrospective cohort study (level IIb)			Mortality	3	2	0.459	OO: Ivor-Lewis with minor modifications
			Reoperation	NR	NR	NR	
			Anastomotic leak	10	8	0.285	
			Conversion	N/A	7	NR	

Continued

Table 1: (Continued)

Author (year), journal and country, Study type (level of evidence)	n _{OO}	n _{MIO}	Outcomes	OO result	MIO result	Sig. (P-value)	Comments
Noble <i>et al.</i> (2013), Dis Esophagus, UK [13] Prospective cohort study (level Ib)	53	53	Inpatient LOS (days)	12	12	0.358	MIO: three-field; with thoracoscopy and laparoscopy
			ITU LOS (days)	1	1	0.995	Groups comparable for age, gender, ASA, BMI, stage of malignancy and use of neoadjuvant therapy. Four conversions from MIO to OO were required. Mortality follow-up to a median of 17 months
			Operative duration (min)	240	300	0.0001	
			Estimated blood loss (ml)	400	300	0.021	
			Lymph node yield	19	18	0.584	
			Mortality	24	15	NS	
			Reoperation	5	6	0.75	
			Anastomotic leak	2	5	NS	
Conversion	N/A	4	NR				
Meng <i>et al.</i> (2014), J Thorac Dis, China [14] Retrospective cohort study (level IIb)	89	94	Inpatient LOS (days)	17.1 ± 10.2	13.9 ± 7.5	0.017	OO: Ivor-Lewis MIO: thoracoscopy and laparoscopy Groups comparable for demographic and pathological characteristics
			ITU LOS (days)	13 ± 14.6	9 ± 9.6	0.295	
			Operative duration (min)	247.8 ± 44.1	251.3 ± 45.4	0.617	
			Estimated blood loss (ml)	261.4 ± 87.2	182.6 ± 78.3	<0.001	
			Lymph node yield	17.4 ± 3.4	16.2 ± 3.1	0.132	
			Mortality	4	1	0.155	
			Reoperation	3	2	0.951	
			Anastomotic leak	7	6	0.696	
Fabian <i>et al.</i> (2008), Dis Esophagus, USA [15] Retrospective cohort study (level IIIb)	43	22	% discharged within 10 days	12	16	0.006	Groups comparable for age, sex, diagnosis, stage of malignancy and use of neoadjuvant therapy. One conversion of MIO to OO was required
			Inpatient LOS (days)	11	9.5	0.30	
			ITU LOS (days)	NR	NR	NR	
			Operative duration (min)	270 ± 87	333 ± 72	0.01	
			Estimated blood loss (ml)	356 ± 136	178 ± 96	<0.0001	
			Lymph node yield	8 ± 7	15 ± 6	0.0002	
			Mortality	4	1	0.45	
			Reoperation	NR	NR	NR	
Anastomotic leak	3	3	NS				
Lazzarino <i>et al.</i> (2010), Ann Surg, UK [16] Retrospective cohort study (level IIIb) (National-level data used)	17 974	699	Inpatient LOS (days)	16	15	0.96	Not comparable for socioeconomic status or comorbidity
			ITU LOS (days)	NR	NR	NR	
			Operative duration (min)	NR	NR	NR	
			Estimated blood loss (ml)	NR	NR	NR	
			Lymph node yield	NR	NR	NR	
			Mortality	566	9	0.55	
			Reoperation	NR	NR	NR	
			Anastomotic leak	NR	NR	NR	
Mamidanna <i>et al.</i> (2012), Ann Surg, UK [17] Retrospective cohort study (level IIIb) (National-level data used)	6347	1155	Inpatient LOS (days)	15	15	<0.001	Non-malignant underlying causes excluded. Groups comparable for demographic variables apart from socioeconomic deprivation
			ITU LOS (days)	NR	NR	NR	
			Operative duration (min)	NR	NR	NR	
			Estimated blood loss (ml)	NR	NR	NR	
			Lymph node yield	NR	NR	NR	
			Mortality	274	46	0.605	
			Reoperation	355	102	<0.001	
			Anastomotic leak	NR	NR	NR	
Bakhos <i>et al.</i> (2012), Ann Thorac Surg, USA [18]	121	99	Inpatient LOS (days)	11 ± 19.5	10 ± 8.8	0.06	Groups comparable for age, sex and receipt of neoadjuvant therapy. Six conversions from MIO to
			ITU LOS (days)	NR	NR	NR	
			Operative duration (min)	NR	NR	NR	
			Estimated blood loss (ml)	NR	NR	NR	

Continued

Table 1: (Continued)

Author (year), journal and country, Study type (level of evidence)	<i>n</i> _{OO}	<i>n</i> _{MIO}	Outcomes	OO result	MIO result	Sig. (P-value)	Comments
Retrospective cohort study (level IIb)			Lymph node yield Mortality Reoperation Anastomotic leak Conversion	NR 5 NR 15 N/A	NR 3 NR 9 6	NR 0.73 NR 0.52 NR	OO were required. Focus of study was on pulmonary morbidity after oesophagectomy OO: various: Ivor-Lewis, McKeown, transhiatal and left thoracic/ thoracoabdominal MIO: thoracoscopy and laparoscopy
Dhamija <i>et al.</i> (2014), Eur J Cardiothorac Surg, USA [19]	64	61	Inpatient LOS (days) ITU LOS (days) Operative duration (min) Estimated blood loss (ml)	9 2 NR NR	8 3 NR NR	NS NS NR NR	Evaluated both open transthoracic and transhiatal approaches (data presented for latter approach only). Study focused on economic costs of OO versus MIO
Retrospective cohort study (level IIb)			Lymph node yield Mortality Reoperation Anastomotic leak Conversion	NR 2 NR NR N/A	NR 2 NR NR 1	NR NS NR NR NR	
Cash <i>et al.</i> (2014), Surg Endosc, USA [20]	33	60	Inpatient LOS (days) ITU LOS (days) Operative duration (min) Estimated blood loss (ml)	13 NR 274 NR	10 NR 275.5 NR	<0.0001 NR NS NR	Groups poorly comparable for age, sex, comorbidities and cancer stage. Two conversions from MIO to OO were required
Retrospective cohort study (level IIb)			Lymph node yield Mortality Reoperation Anastomotic leak Conversion	NR 1 6 8 NR	NR 0 0 3 2	NR 0.39 0.56 0.72 NR	
Hamouda <i>et al.</i> (2010), Surg Endosc, UK [21]	24	26	Inpatient LOS (days) ITU LOS (days) Operative duration (min) Estimated blood loss (transfusion units)	14 3 260 4.5	16 4 223 4	NS NS NS NS	Evaluated open Ivor-Lewis oesophagectomy versus laparoscopic Ivor-Lewis procedure in both early and late cohorts. Data from the late cohort are presented
Prospective cohort study (level Ib)			Lymph node yield Mortality Reoperation Anastomotic leak	24 0 1 2	23 0 4 4	NS NS NS NS	
Berger <i>et al.</i> (2011), J Am Coll Surg, USA [22]	53	65	Inpatient LOS (days) ITU LOS (days) Operative duration (min) Estimated blood loss (ml)	16 NR NR 619	9 NR NR 182	0.003 NR NR <0.0001	Groups comparable for demographic data. A few patients in the MIO group (15%) underwent thoracoscopy with laparotomy
Retrospective cohort study (level IIb)			Lymph node yield Mortality Reoperation Anastomotic leak	9 4 NR 6	20 5 NR 9	<0.0001 1.0 NR 1.0	
Smithers <i>et al.</i> (2007), Ann Surg, USA [23]	114	23	Inpatient LOS (days) ITU LOS (h) Operative duration (min) Estimated blood loss (ml)	14 23 300 600	11 19 330 300	0.03 0.03 0.01 0.017	Also evaluated a hybrid MIO (thoracoscopic/laparotomy-based) approach (data not

Continued

Table 1: (Continued)

Author (year), journal and country, Study type (level of evidence)	n _{OO}	n _{MIO}	Outcomes	OO result	MIO result	Sig. (P-value)	Comments
study (level IIb)			Lymph node yield	16	17	NS	included). Significant intergroup differences in demographic and clinical variables. Two conversions of MIO to OO
			Mortality	3	0	NS	
			Reoperation	NR	NR	NR	
			Anastomotic leak	10	1	NS	
			Conversion	NR	12	NR	
							OO: open thoracotomy and laparotomy. MIO: total thoracoscopic and laparoscopic approach
Zingg <i>et al.</i> (2009), Ann Thorac Surg, Austria [24] Retrospective cohort study (level IIb)	98	56	Inpatient LOS (days)	21.9	19.7	0.463	To minimize selection bias, only patients who met the selection criteria for the MIO approach were included in the OO group
			ITU LOS (days)	6.8	3.0	0.022	
			Operative duration (min)	209.4	250.2	<0.001	
			Estimated blood loss (ml)	857	320	<0.001	
			Lymph node yield	6.7	5.7	0.144	
			Mortality	6	2	0.467	
			Reoperation	9	5	NS	
			Anastomotic leak	11	11	0.341	
			Conversion	NR	5	NR	
Blom <i>et al.</i> (2012), J Thorac Dis, Netherlands [25] Retrospective cohort study (level IIb)	49	41	Inpatient LOS (days)	13	11	0.072	Groups comparable for baseline characteristics. Two patients required conversion from MIO to OO. In-hospital mortality follow-up only was available
			ITU LOS (days)	1	1	0.188	
			Operative duration (h)	5.2	6	<0.001	
			Estimated blood loss (ml)	500	100	<0.001	
			Lymph node yield	25	19	<0.001	
			Mortality	0	1	0.456	
			Reoperation	NR	NR	NR	
			Anastomotic leak	1	4	0.173	
			Conversion	NR	2	NR	

All values presented are median ± interquartile range, unless otherwise specified.

ASA: American Society of Anesthesiologists classification; BMI: body mass index; ITU: intensive treatment unit; LOS: length of stay; MIO: minimally invasive oesophagectomy; ml: millilitres; NR: not reported; NS: not significant; OO: open oesophagectomy; U: units.

retrieved, and reference lists cross-checked. Studies with the level of evidence I and II (e.g. randomized controlled trials, meta-analyses or well-designed, controlled, observational studies) with ≥50 participants were included. This process yielded 24 papers, which were deemed to offer the best evidence. The key data from these papers are detailed in Table 1.

RESULTS

Of the 24 studies identified, data from two randomized controlled trials were available. Biere *et al.* [2] randomized 115 patients to undergo either open oesophagectomy (OO; thoracotomy and laparotomy, $n = 56$) or minimally invasive oesophagectomy (MIO; thoracoscopy and laparoscopy, $n = 59$). An open-label, multicentre design, with randomization via computer-generated sequence, and rigorous intention-to-treat analysis, was employed. The authors report that those who underwent MIO left hospital on average 3 days earlier than those treated with the open technique (11 vs 14 days, respectively; $P = 0.044$). A further randomized

controlled trial, by Guo *et al.* [3], enrolled 221 patients, who were similarly randomized in a single-blind, 1:1 fashion, to receive either thoracotomy with thoracoscopy ($n = 111$) or laparoscopy ($n = 111$). The authors observed an average reduction of 1.8 days in the LOS in patients managed by the MIO strategy, as opposed to an OO approach (9.6 vs 11.4 days, respectively; $P < 0.001$). Both studies demonstrated no differences in mortality or the overall complication rate; however, in the study by Guo *et al.*, there was a trend towards fewer pulmonary infections in the MIO group ($n = 3$) compared with the OO group ($n = 9$, $P = 0.072$) [3].

Within the remaining 22 non-randomized studies, data for a total of 3003 minimally invasive and 25 524 open oesophagectomies were available, in the form of 3 prospective and 19 retrospective cohort studies.

Eleven studies compared thoracotomy and laparotomy with thoracoscopy and laparoscopy [4–14], with 8 noting a significant reduction in hospital LOS in the MIO group, and 2 noting no difference [5–14]. The study by Parameswaran *et al.* [4] demonstrated a statistically longer MIO LOS (12 days) compared with the OO LOS (10 days, $P = 0.01$). The authors explain that the technique was

new to their unit (which covered a large geographic patient catchment area) and they wished to retain patients to ensure there were no late complications with the technique. In those non-randomized studies where demographic data were available, groups were generally comparable in age, American Society of Anesthesiologists classification and stage of malignancy.

A further study, by Fabian *et al.* [15], compared their surgeons' own preferred open method with a thorascopic procedure with or without laparoscopic assistance, observing a non-significant reduction of 1.5 days spent in hospital within the MIO group ($P = 0.3$).

Four groups compared the use of any open procedure with laparoscopy and/or thoracoscopy, analysing data for over 26 000 patients [16–19]. However, only one of these studies showed a statistically significant reduction in hospital length of stay (LOS) [17].

A further study compared the use of laparotomy with laparoscopy, noting a significant reduction in LOS of 3 days in the MIO group ($P < 0.0001$) [20]. This was a non-randomized study [20] with patients undergoing MIO being older and with significantly more comorbidities. However, these factors would be expected to lengthen, rather than shorten, LOS, and as such the significance of the MIO approach may have been underestimated.

Three studies compared the use of hybrid procedures with both thoracotomy and laparotomy, and thoracoscopy and laparoscopy. Hamouda *et al.* [21] compared OO and MIO procedures with a hybrid procedure consisting of a thoracotomy with laparoscopic assistance, finding a non-significant increase in the LOS of 2 days in the MIO group. Those treated with the hybrid procedure also required an intensive care unit stay 1 day longer than those in the open group, though this was non-significant, and the authors did not pass any comment on the potential reasons for the difference. Berger *et al.* [22] compared OO and MIO techniques while performing an additional small number of hybrid procedures consisting of thoracoscopically assisted laparotomy, which they grouped with the MIO cohort. They observed that the use of MIO led to a 7-day shorter hospital stay ($P = 0.003$). Smithers *et al.* [23] also compared the use of thoracoscopic-assisted laparotomy with OO and MIO techniques, noting reductions in stay of 3 and 1 day(s) with the MIO and hybrid techniques, respectively ($P = 0.03$).

The remaining two studies chose to compare the use of thoracotomy alone with a combined thoracoscopic and laparoscopic approach. There were no significant differences in LOS observed in either study [24, 25].

Of the 22 non-randomized studies, 6 studies noted a significant reduction in the complication rate using the MIO technique, while only 1 noted an increase, with the remaining 15 noting no difference. One study noted a significant reduction in the 30-day mortality using the MIO technique, while the remaining 21 noted no difference.

Importantly, for those 1011 patients whose data were available, conversion from an MIO to an open technique was required in 7.5% of cases.

CLINICAL BOTTOM LINE

The only two randomized trials comparing MIO and OO demonstrated a reduction in the LOS in the MIO group, without compromising survival or increasing complication rates. All except one of the non-randomized studies demonstrated either a significant reduction in LOS with MIO or no difference. The benefit in

reduced LOS was not at the cost of worsened survival or increased complications.

Conflict of interest: none declared.

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