

Pulmonary function after segmentectomy versus lobectomy in patients with early-stage non-small-cell lung cancer: a meta-analysis Journal of International Medical Research 49(9) 1–14 © The Author(s) 2021 Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/03000605211044204 journals.sagepub.com/home/imr



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#### Abstract

**Objective:** Segmentectomy is widely performed for early-stage lung cancer. However, the effects of segmentectomy versus lobectomy on pulmonary function remain unclear. We performed a meta-analysis with the aim of comparing segmentectomy and lobectomy in terms of preservation of pulmonary function in patients with early-stage non-small-cell lung cancer (NSCLC).

**Methods:** We conducted a literature search of PubMed using the terms 'pulmonary function' AND 'segmentectomy' AND 'lobectomy'. The primary outcomes of interest were the forced expiratory volume in I second (FEVI), FEVI as percent of predicted (%FEVI), change in FEVI ( $\Delta$ %FEVI), and the ratio of postoperative to preoperative FEVI.

**Results:** Thirteen studies comprising 2027 patients met the inclusion and exclusion criteria and were included for analysis, including 787 patients in the segmentectomy group and 1240 patients in the lobectomy group. Patients in the segmentectomy group showed significantly better preservation of FEV1 and %FEV1 compared with the lobectomy group. The reduction in FEV1 after surgery was significantly less in the segmentectomy group compared with the lobectomy group, and  $\Delta$ %FEV1 was significantly higher in the segmentectomy group than in the lobectomy group.

**Conclusion:** Segmentectomy results in better preservation of pulmonary function compared with lobectomy in patients with early-stage NSCLC.

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## Keywords

Segmentectomy, lobectomy, lung cancer, forced expiratory volume in 1 second, pulmonary function, meta-analysis

Date received: 7 May 2021; accepted: 17 August 2021

# Introduction

Pulmonary lobectomy with systemic mediastinal lymph node dissection is a standard surgery for patients with early-stage nonsmall-cell lung cancer (NSCLC). A randomized trial reported by the Lung Cancer Study Group (LCSG) in 1995 comparing sublobar resection, including segmentectomy and resection, wedge suggested increased mortality and locoregional recurrence in the limited-resection group.<sup>1</sup> However, there were potential biases affecting the results of segmentectomy in the LCSG study, and several recent studies have suggested similar local recurrence rates following sublobar resection and lobectomy in patients with IA tumors  $<2 \text{ cm}.^{2-6}$  Computed tomography (CT) screening allows the detection of more and earlier-stage lung cancers.<sup>7</sup> Anatomic segmentectomy is widely used in clinical practice because it theoretically preserves more pulmonary parenchyma, leading to better preservation of pulmonary function compared with lobectomy. However, there have been conflicting results regarding the relative advantages of segmentectomy over lobectomy in terms of pulmonary function protection.<sup>1,2,8–13</sup> We therefore conducted a meta-analysis to compare postoperative pulmonary function between patients treated with segmentectomy and lobectomy, and to verify if segmentectomy had an advantage over lobectomy in terms of preserved pulmonary function.

# Survey methodology

# Study protocol

We carried out a systematic review and meta-analysis of published studies to compare pulmonary function in patients with early-stage NSCLC after segmentectomy or lobectomy. This study was reported according to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) 2020 Checklist.<sup>14</sup> Ethics approval and informed consent were not required due to the nature of this study (meta-analysis of published articles).

The protocol for this systematic review was registered on INPLASY (INPLASY202180050) and is available in full at inplasy.com (https://doi.org/10. 37766/inplasy2021.8.0050).

# Search strategy

We conducted a literature search of PubMed for all relevant studies published from the date of database inception to December 2020 using the following search terms in the title/abstract field: 'pulmonary function' AND 'segmentectomy' AND 'lobectomy'. Only articles published in English were included. The reference lists of relevant review articles were checked to identify additional relevant articles.

Studies were included if they met the following criteria: direct comparison between segmentectomy and lobectomy for earlystage NSCLC; pulmonary function tests performed before and after surgery in both groups; and article in English. The exclusion criteria were: review articles; limited sublobar resection, including wedge resection; pulmonary function tests not performed during follow-up; pulmonary function determined by CT or singlephoton-emission CT (SPECT), other than spirometry tests; and no detailed pulmonary function data.

## Data extraction

Data were extracted independently by two authors (XW, HG). In the event of disagreement, consensus was achieved by discussion. The following data were extracted from each included article: first author, publication year, study design, patient age and sex, smoking history, operation techniques, pulmonary function parameters.  $\Delta\%$ FEV1 was defined as change in forced expiratory volume in 1 second (FEV1) between before and after surgery.

## Quality assessment

The quality of the included studies was assessed using the Newcastle–Ottawa scale (NOS), with 9 being the highest score and a score  $\geq 6$  indicating a high-quality study. The assessments were performed independently by two authors (QH and YY), with disagreements settled by a third author (BC) if necessary.

## Statistical methods

All statistical analyses were carried out using Review Manager version 5.3 (The Cochrane Collaboration, Copenhagen, Denmark). Heterogeneity was calculated by Q-test and I<sup>2</sup> statistics. Studies with an  $I^2 > 50\%$  were considered to show a high degree of heterogeneity. If heterogeneity existed, a random-effects model was adopted; otherwise, a fixed-effects model was used. Pooled analysis was performed using the inverse variance model and results were reported as odds ratios (ORs) with 95% confidence intervals (CIs). A p value <0.05 was considered statistically significant.

# Results

The literature search identified 345 studies for review. Based on the title and abstract, 28 studies were selected for full-text review. Of these, 15 studies were excluded because they measured pulmonary function using CT or SPECT (n=6), 12,13,15-18 lacked follow-up pulmonary function data (n=4), 9,11,19,20 lacked a control group of lobectomy (n = 2),<sup>21,22</sup> did not directly comsegmentectomy and lobectomy pare (n = 2),<sup>23,24</sup> or because they included surgery for benign pulmonary nodules (n=1).<sup>25</sup> Thirteen studies were finally included in the meta-analysis (Figure 1).2,8,10,26-35 All studies were retrospective observational studies. The 13 studies comprised 2027 patients, including 787 patients in the segmentectomy group and 1240 patients in the lobectomy group. The patients' baseline characteristics are summarized in Tables 1 and 2. The included studies were assessed using the NOS for quality and were all considered to be high quality (score  $\geq 6$ ), except for the study by Echavarria et al. (Table 3). Patients in 10 of the studies<sup>2,8,10,27,28,30,32-35</sup> were classified as clinical or pathological stage I, most patients (87%–93.7%) in two other studies<sup>29,31</sup> were classified as pathological stage I, and one study<sup>26</sup> did not report the precise cancer stage, but noted that all patients were clinically early-stage. There was no significant difference in the results of pulmonary function tests before surgery between the segmentectomy and lobectomy groups, except for two studies: Keenan et al.<sup>2</sup> and Echavarria et al.<sup>26</sup> showed significantly better FEV1 as percent of predicted (%FEV1) in the segmentectomy group. The average age of the patients ranged from 58.2 to 68.7 years, with no

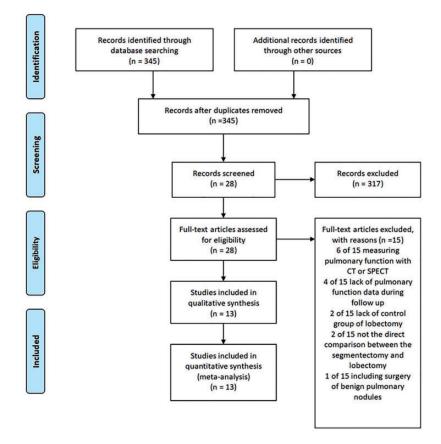


Figure 1. Flowchart of study selection. CT, computed tomography, SPECT, single-photon-emission computed tomography.

significant difference between the two groups. The percentage of females ranged from 33% to 56% in the segmentectomy group and 32.7% to 65% in the lobectomy group, with three studies having a significant difference in the percentage of females between the two groups. Five of the 13 studies provided data on smoking history, two of which showed a significant difference in smoking history between the two groups.

# FEVI

Five studies<sup>8,26,27,31,35</sup> including 933 patients provided results for postoperative pulmonary function tests, including 309

patients in the segmentectomy group and 624 patients in the lobectomy group. There was no significant difference in preoperative FEV1 between the segmentectomy and lobectomy groups in five studies, including two studies<sup>27,35</sup> that carried out one test 6 months after surgery, one study<sup>26</sup> that did not mention the exact time of the test, and two studies<sup>8,31</sup> that recorded two to three pulmonary tests during the follow-up. The maximum follow-up was 10 years after surgery in one study. To minimize the effects of the heterogeneity in terms of the timing of pulmonary tests, tests carried out 1 year after surgery were included in the analysis of these two studies.

Age (y)		1	Female (%)	I.	Smoking history (%)	1	VATS (%)	I.	Number of resected segments	ed segments				
		λu			Хш		λu	S.	Segmentectomy		Lobectomy	۲. 		
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$68.4 \pm 2.0$ $66.7 \pm 0.7$ $51$ $52$	66.7±0.7 51		52		81 85	=	001	100 NR		1	NR	NR	NR	Stapler
$68.7 \pm 8.3$ $65.7 \pm 9.7$ $46$ $49$	<b>65.7 ± 9.7</b> 46		49							1	142	5	12 m	NR
$67.4 \pm 9.6$ $68.2 \pm 10.5$ 33 $65$	<b>68.2 ± 10.5 33</b>		55		NR NR		0	29	23	I.58	126	0	l, 6 m	Electrocautery
<b>6</b> 8± <b>9 6</b> 8± <b>8</b> 39 20	<b>68</b> ± <b>8</b> 39		20		39 20	0		NR 10		2.92	103	0	6 m	Electrocautery +stapler
70 69 35 35	35		35		NR NR	24	4 24	4	œ	1.82	17	0	4 T	Stapler
<b>67.7</b> ± 8.3 <b>66.9</b> ± 7.2 33 35	<b>66.9</b> ±7.2 33 3	.,	35			89	88	8 NR		/	33	0	7-12 m	NR
70 73 51 45	73 51	-	45		48 57		NR N	NR NR	۸ NR	1	47	0	6 m	NR
$63.0\pm9.9$ $62.9\pm10.0$ 47 49	<b>62.9</b> ±10.0 47		49		NR NR		001	100 NR		1	NR	NR	NR	Electrocautery
	48		32.7		NR NR		001	100 NR	۶ NR	/	NR	NR	NR	Stapler
70.3 69.7 39 39	39		39				001	100 38		I.89	74	0	6 m	Electrocautery +stapler
<b>66.4</b> ±8.1 <b>66.0</b> ±9.5 <b>56</b> 41	<b>66.0±9.5</b> 56		4		30 54	70	35	R	۸ NR	1	NR	NR	y, 5 y,  0 y	NR
$65.0 \pm 6.4$ $64.0 \pm 7.3$ $45$ $48$	<b>64.0</b> ± <b>7.3 45</b>	-	48		NR NR		NR	NR 24	16	I.625	40	0	2 w, I2 m	Stapler
$58.2 \pm 9.5 60.1 \pm 8.3 53 57$	60.1 ± 8.3 53		57		NR NR			100 15		1.91	75	0	6 m	Stapler

Table 1. Baseline characteristics of patients in included studies.

NR, not recorded; w, weeks; m, months; y, years.

preoperative FEVI (%) 87.3 ± 14.0 NR  $\mathbf{87}\pm\mathbf{11}$ Post/ NR 80.9 88.8 NR 81.7 ЯZ ЯX ЯZ ЯĽ NR 70.6 ± 16.8 NR 96.5±21.1 NR  $\textbf{86.8} \pm \textbf{19.5}$  $67.4 \pm 1.2$ **66.7 ± 1.8** %FEVI A X X X ЯЯ  $1.95 \pm 0.49$  $1.96 \pm 0.51$  $\textbf{I.83}\pm\textbf{0.04}$  $\textbf{I.83}\pm\textbf{0.06}$  $2.11 \pm 0.57$ FEVI (L) X X X X R R N N N  $-11.0 \pm 13.1$  $-8.1\pm10.6$ AFEVI (%)  $\textbf{-20.6}\pm\textbf{0.4}$  $\textbf{-18.9}\pm\textbf{9.9}$ Lobectomy NR --21.0 R R r- 7 NR -12 12 ЛR preoperative 93.3 ± 10.3 NR FEVI (%)  $\mathbf{95}\pm\mathbf{10}$ 89.8 Post/ 87.8 91.9 RR ЯS ЛR ЛR R N ЛR 100.2 ± 25.0 NR **73.2±19.9** 52.2 ± 2.8 NR NR NR  $\textbf{93.5}\pm\textbf{18.5}$  $\mathbf{69.8} \pm \mathbf{3.3}$ %FEVI Pulmonary function postoperative ЯĽ Жź Ж  $2.04 \pm 0.06$  $2.15 \pm 0.61$  $1.99 \pm 0.63$  $2.07 \pm 0.94$  $\textbf{I.90}\pm\textbf{0.10}$ FEVI (L) ж х х х ЯЯ ЯÄ Segmentectomy  $-12.9 \pm 8.7\%$ **ΔFEVI (%)**  $-8.9 \pm 10.8$ +1.0 (9.8)  $-8.9\pm0.7$ -15.5% NR - 18.4 NR 12% ЛR ЯЯ R R 109.4 ± 18.3 NR  $113.2 \pm 22.7$  $107.0 \pm 24.1$  $100.7 \pm 17.8$ 78.7 ± 17.4  $86.8 \pm 19.5$  $\mathbf{85.2} \pm \mathbf{1.5}^{*}$  $\textbf{74.8} \pm \textbf{9.0}$  $75.1 \pm 2^{*}$  $107\pm19$ %FEVI RR 4 NR 2.36 ± 0.66  $2.25 \pm 0.63$  $2.43 \pm 0.61$  $\textbf{2.34} \pm \textbf{0.60}$  $\mathbf{2.11} \pm \mathbf{0.62}$  $\textbf{2.25}\pm\textbf{0.07}$ Lobectomy  $\mathbf{2.32} \pm \mathbf{0.05}$ 2.11 ± 0.57 FEVI (L)  $\textbf{2.3}\pm\textbf{0.6}$ Ξ R R Pulmonary function preoperative  $\begin{array}{c} 101.6\pm24.0\\ 72.2\ \pm18.3\end{array}$ 1 05.0 ± 24.6 NR  $\mathsf{I}\,\mathsf{I}\,\mathsf{I}\,\mathsf{I}\,\mathsf{B}\,\pm\,\mathsf{20.5}$  $109.4 \pm 20.8$  $\mathbf{93.5} \pm \mathbf{18.5}$ 107 ± 23 45  $\textbf{73.3}\pm\textbf{9.0}$  $76.4 \pm 3.5$  $55.3 \pm 3$ %FEVI ЛR Segmentectomy  $\textbf{2.10} \pm \textbf{0.58}$  $2.22 \pm 0.55$  $\textbf{2.15}\pm\textbf{0.53}$  $\textbf{2.24} \pm \textbf{0.07}$  $2.51 \pm 1.03$  $2.15 \pm 0.61$  $\textbf{2.13}\pm\textbf{0.61}$  $2.08 \pm 0.1$ FEVI (L)  $\textbf{2.3}\pm\textbf{0.6}$ ЯZ Жĸ Helminen et al.<sup>28</sup> Tane et al.<sup>35</sup> Takizawa et al.<sup>8</sup> Gu et al.<sup>27</sup> Nomori et al.<sup>33</sup> Hwang et al.<sup>29</sup> Suzuki et al.<sup>34</sup> Keenan et al.<sup>2</sup> Kashiwabara et al.<sup>30</sup> Saito et al.<sup>10</sup> Martin-Ucar et al.<sup>32</sup> et al.<sup>26</sup> et al.<sup>31</sup> Echavarria **Sobayashi** Author Publication 2016 2016 2009 2020 2019 2018 2014 2017 2005 2014 2016 666 I year 2004

Table 2. Preoperative and postoperative pulmonary function in patients in included studies.

 $^{*}$ p < 0.05 %FEV l between segmentectomy and lobectomy.

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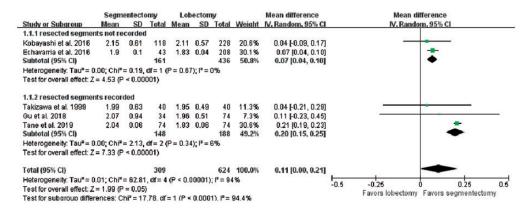
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		Selection					Outcome			
Publication year	Study	Representativeness of exposed cohort	Selection of nonexposed cohort	Ascertainment of exposure	Outcome of interest absent at start of study	Comparability (based on design and analysis)	Follow-up lo Assessment enough for of outcome outcomes to	ng occur	Adequacy of follow-up	Total score
2016	Echavarria et al. <sup>26</sup>	_	_	_	_	0	_	0	0	2
2004	Keenan et al. <sup>2</sup>	_	_	_	_	0	_	_	_	7
2014	Saito et al. <sup>10</sup>	_	_	_	_	_	_	_	_	8
2017	Nomori et al. <sup>33</sup>	_	_	_	_	_	_	_	_	8
2005	Martin-Ucar et al. <sup>32</sup>	_	_	_	_	_	_	_	0	7
2016	Suzuki et al. <sup>34</sup>	_	_	_	_	_	_	_	_	8
2009	Kashiwabara et al. <sup>30</sup>	_	_	_	_	_	_	_	_	8
2014	Hwang et al. <sup>29</sup>	_	_	_	_	_	_	0	_	7
2020	Helminen et al. <sup>28</sup>	_	_	_	_	_	_	0	_	7
2019	Tane et al. <sup>35</sup>	_	_	_	_	_	_	_	_	8
2016	Kobayashi et al. <sup>31</sup>	_	_	_	_	_	_	_	0	7
666 I	Takizawa et al <sup>8</sup>	_	_	_	_	_	_	_	_	8
2018	Gu et al. <sup>27</sup>	_	_	_	_	_	_	_	0	7

The segmentectomy group showed significantly better preservation of FEV1 comwith the lobectomy pared group (OR = 0.11, 95% CI, 0.00, 0.21, p = 0.05)(Figure 2). However, there was high heterogeneity  $(I^2 = 94\%)$  among these studies, indicating that the results need to be interpreted with caution. Three of the five studies<sup>8,27,35</sup> provided detailed information on the number of resected segments in the segmentectomy group, and the average number of resected segments ranged from 1.63 to 1.91. Subgroup analysis of these three studies still demonstrated a significantly better FEV1 in the segmentectomy compared with the lobectomy group (OR = 0.20, 95% CI, 0.15, 0.25, p < 0.001)(Figure 2). An  $I^2$  value of 6% indicated low heterogeneity across the studies. Three of the five studies<sup>26,27,35</sup> performed segmentectomy and lobectomy with complete video-assisted thoracoscopic surgery (VATS). Pooled analysis of these three studies indicated that FEV1 was better preserved in the segmentectomy compared with the lobectomy group (OR = 0.14, 95CI%, 0.01, 0.27, p = 0.04) (Figure 3). However, there was high heterogeneity  $(I^2 = 97\%)$  among these studies, indicating the need for caution. Four of five studies<sup>8,26,27,35</sup> divided the intersegmental plane with a stapler during segmentectomy, and pooled analysis of these four studies showed better preservation of FEV1 in the segmentectomy compared with the lobectomy group (OR = 0.12, 95CI%, 0.01, 0.24, p = 0.04) (Figure 4). However, the high heterogeneity ( $I^2 = 95\%$ ) among these studies again highlights the need for caution when interpreting the results.

## %FEV1

Five studies<sup>2,8,26,28,31</sup> recorded the results of pulmonary tests in terms of %FEV1 after surgery. These studies included 976 patients, with 305 patients in the



**Figure 2.** Funnel plot analysis of segmentectomy versus lobectomy on postoperative FEV1. SD, standard deviation; IV, inverse variance; CI, confidence interval.

	Segm	entecto	omy	Lob	ecton	IV		Mean difference	Mean difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl	
Echavarria et al. 2016	1.9	0.1	43	1.83	0.04	208	44.2%	0.07 [0.04, 0.10]		
Gu et al. 2018	2.07	0.94	34	1.96	0.51	74	10.9%	0.11 [-0.23, 0.45]		
Tane et al. 2019	2.04	0.06	74	1.83	0.06	74	44.9%	0.21 [0.19, 0.23]		
Fotal (95% CI)			151			356	100.0%	0.14 [0.01, 0.27]	+	
Heterogeneity: Tau <sup>2</sup> = 0	.01; Chi#	= 58.19	, df = 2	(P < 0.0	0001)	; I* = 97	796		-1 -0.5 0 0.5	-
Test for overall effect: Z	= 2.10 (P	= 0.04	)					12	-1 -0.5 0 0.5 Favors lobectomy Favors segmentectomy	ï

Figure 3. Funnel plot analysis of segmentectomy versus lobectomy with complete video-assisted thoracoscopic surgery on FEV1.

SD, standard deviation; IV, inverse variance; CI, confidence interval.

	Segm	entecto	omy	Lob	ecton	IY		Mean difference	Mean difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	IV, Random, 95% Cl
Echavarria et al. 2016	1.9	0.1	43	1.83	0.04	208	37.9%	0.07 [0.04, 0.10]	
Gu et al. 2018	2.07	0.94	34	1.96	0.51	74	9.3%	0.11 [-0.23, 0.45]	
Takizawa et al. 1999	1.99	0.63	40	1.95	0.49	40	14.3%	0.04 [-0.21, 0.29]	
Tane et al. 2019	2.04	0.06	74	1.83	0.06	74	38.5%	0.21 [0.19, 0.23]	
Total (95% CI)			191			396	100.0%	0.12 [0.01, 0.24]	-
Heterogeneity: Tau <sup>=</sup> = 0				(P < 0.0	0001)	;  * = 95	5%		-1 -0.5 0 0.5 1
Test for overall effect: Z	= 2.06 (P	= 0.04	)						Favors lobectomy Favors segmentectomy

Figure 4. Funnel plot analysis of segmentectomy dividing the intersegmental plane with stapler versus lobectomy on postoperative FEV1.

SD, standard deviation; IV, inverse variance; CI, confidence interval.

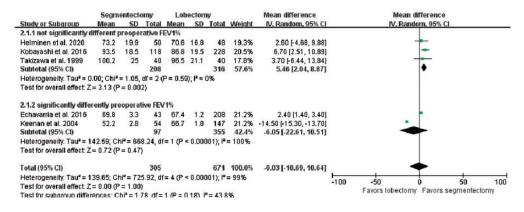
segmentectomy group and 671 patients in the lobectomy group. %FEV1 was significantly better in patients in the lobectomy group compared with the segmentectomy group according to Keenan et al.<sup>2</sup> and Echavarria et al.,<sup>26</sup> but there was no significant difference between the two groups in the other three studies. Two studies<sup>26,28</sup> did not report the exact time of the test, one study<sup>2</sup> did the test 12 months after surgery, and two studies<sup>8,31</sup> included two to three tests during the follow-up period of 2 weeks to 10 years after surgery. The test carried out at 12 months after surgery was included in the analysis. Pooled analysis of these five studies concluded that there was no significant difference between the lobectomy and segmentectomy groups in terms of postoperative %FEV1 reserve (OR = -0.03, 95%CI, -10.69, 10.64) (Figure 5). After excluding the studies by Keenan et al.<sup>2</sup> and Echavarria et al.,<sup>26</sup> pooled analysis of the remaining three studies demonstrated a better %FEV1 in the segmentectomy group compared with the lobectomy group (OR = 5.46, 95%CI, 2.04, 8.87, p = 0.002) (Figure 5). An I<sup>2</sup> value of 0 indicated no potential heterogeneity across the studies.

#### $\Delta$ %FEV1

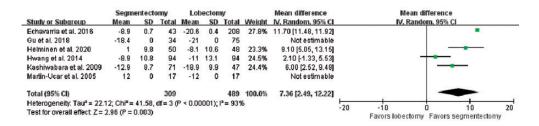
Six studies<sup>26–30,32</sup> including 700 patients calculated the change in FEV1 from before to after surgery. There were 259 patients in the segmentectomy group and 441 patients in the lobectomy group. There was no significant difference in FEV1 before surgery between the two groups in these six studies. Three of the six studies<sup>26,28,29</sup> did not record the exact time of the pulmonary tests, two studies<sup>27,30</sup> tested the FEV1 at 6 months after surgery, and one study<sup>32</sup> tested the FEV1 at 4 months after surgery. Pooled analysis showed a significantly smaller change in FEV1 between before and after surgery in the segmentectomy group than in the lobectomy group (OR = 7.36, 95%CI, 2.49. 12.22. p = 0.003) (Figure 6). However, there was high heterogeneity  $(I^2 = 93\%)$  among these studies, indicating the need for cautious interpretation.

# Ratio of postoperative to preoperative FEV1

Five studies<sup>8,10,33–35</sup> including 682 patients calculated the ratio of postoperative to



**Figure 5.** Funnel plot analysis of segmentectomy versus lobectomy on postoperative FEV1%. SD, standard deviation; IV, inverse variance; CI, confidence interval.



**Figure 6.** Funnel plot analysis of segmentectomy versus lobectomy on  $\Delta$ FEV1%. SD, standard deviation; IV, inverse variance; CI, confidence interval.

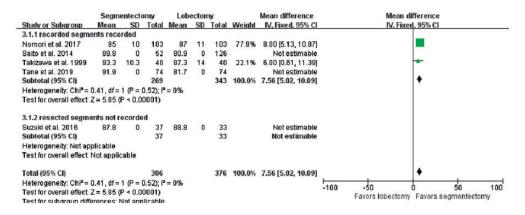


Figure 7. Funnel plot analysis of segmentectomy versus lobectomy on ratio of postoperative to preoperative FEV1.

SD, standard deviation; IV, inverse variance; CI, confidence interval.

preoperative FEV1, including 306 patients in the segmentectomy group and 376 patients in the lobectomy group. There was no significant difference in preoperative FEV1 between the segmentectomy and lobectomy groups in these five studies. The timing of pulmonary tests after surgery ranged from 2 weeks to 12 months, and two of the five studies<sup>8,10</sup> tested twice during follow-up. Tests carried out at 6 months postoperatively in one study and 12 months postoperatively in another study were included in the analysis. Pooled analvsis demonstrated a significantly better postoperative to preoperative FEV1 ratio in the segmentectomy group compared with the lobectomy group (OR = 7.56, 95%CI. 5.02, 10.09, p < 0.001) (Figure 7). Four of the five studies<sup>8,10,33,35</sup> recorded the number of resected segments in the segmentectomy group in detail, and the average number of resected segments ranged from 1.58 to 2.92. Subgroup analysis showed that the better postoperative to preoperative FEV1 ratio was preserved in the segmentectomy compared with the lobectomy group in these four studies (OR = 7.56, 95%CI, 5.02, 10.09, p < 0.001) (Figure 7). An I<sup>2</sup> value of 0 indicated no potential heterogeneity across the studies.

# Discussion

Numerous retrospective studies have compared postoperative pulmonary function between patients undergoing segmentectomy and lobectomy; however, the benefits of segmentectomy in terms of preserving pulmonary function remain unclear. To the best of our knowledge, the current meta-analysis was the first to compare postoperative pulmonary function between patients treated with segmentectomy and lobectomy. A low FEV1 is an independent risk factor for complications after lung surgeries, including lobectomy and segmentectomy.<sup>36</sup> The current results showed that segmentectomy provided better preservation of pulmonary function in terms of FEV1, %FEV1,  $\Delta$ %FEV1, and the ratio of postoperative to preoperative FEV1. A pooled analysis of five studies found no significant difference in postoperative %FEV1 between the lobectomy and segmentectomy groups; however, after excluding the studies by Keenan et al. and Echavarria et al., segmentectomy proved to be better at preserving %FEV1, possibly because preoperative %FEV1 was significantly better in the lobectomy group in these two excluded studies. The better performance of segmentectomy in protecting pulmonary function could partly be attributed to the fewer segments resected compared with lobectomy. The average number of segments resected in the included studies ranged from 1.58 to 2.92. Postoperative pulmonary function recovery differed among the measured variables. Vital capacity recovered within 1 month after surgery and FEV1 recovered within 3 months, because of surgical pain and injury.<sup>37-39</sup> Pulmonary function then reached a stable level.<sup>40</sup> The pulmonary tests included in the current analysis were carried out at 6 or 12 months after surgery, and were representative of postoperative pulmonary function.

Five of the 13 included studies used a stapler to divide the intersegmental plane. Pooled analysis of these studies demonstrated that segmentectomy still preserved pulmonary function better than lobectomy in these five studies. VATS reduces the loss of pulmonary function after lobectomy as a result of decreased pain, improved chest wall mechanics, and reduced inflammation.<sup>41,42</sup> The surgical procedures were performed with complete VATS in five of the 13 studies in this meta-analysis, and segmentectomy preserved pulmonary function significantly better than lobectomy in patients undergoing complete VATS. according to the pooled analysis. The predicted preoperative %FEV1 was >70% in all the included studies, except for one which had a mean preoperative %FEV1 of 55.3% in the segmentectomy group, which was significantly worse than the the lobectomy 75.1% in group. Kashiwabara et al.<sup>30</sup> suggested that segmentectomy should be considered in patients with a normal (>80%) predicted postoperative FEV1, given that patients with an estimated postoperative FEV1/predicted normal FEV1 ratio <70% gained no functional benefit from segmentectomy. However, their conclusion was based on a small sample of 50 patients and should thus

be translated cautiously in clinical practice. A large-scale prospective study is needed to explore the benefits of segmentectomy in patients with poor pulmonary function. Nomori et al.<sup>43</sup> reported that postoperative pulmonary function after left upper division segmentectomy was similar to that after left upper lobectomy. Similarly, the function of the right middle lobe was significantly decreased after right upper lobectomy compared with that after segmentectomy of the right up lobe.<sup>18,44,45</sup> The current study could not draw any conclusions regarding whether or not postoperative pulmonary function was affected by the location of the resected segments, because relevant detailed information was not provided by the included studies.

This study had several limitations. First, the meta-analysis may have had a publication bias because we limited the search to studies published in English. However, we were unable to analyze publication bias because of the small number of studies. Second, pulmonary function parameters were restricted to FEV1 in our analysis, because information on other potentially relevant parameters was not available in all the studies. Third, we only included studies that compared pulmonary function using spirometry tests other than CT or SPECT.

# Conclusion

This was the first meta-analysis to evaluate and compare postoperative pulmonary function after segmentectomy and lobectomy in patients with early-stage NSCLC. The results suggested that segmentectomy preserved pulmonary function in these patients better than lobectomy. However, further well-designed, large-scale randomized prospective studies are needed to confirm our findings in clinical practice.

## **Declaration of conflicting interest**

The authors declare that there is no conflict of interest.

# Funding

The authors disclosed receipt of the following financial support for the research, authorship and/or publication of this article: This work was supported by Taizhou Municipal Science and Technology Bureau (grant number 1801ky09) and the Medical Health Science and Technology Project of Zhejiang Provincial Health Commission (grant number 2019KY773).

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