A systematic review and meta-analysis of thoracoscopic versus thoracotomy sleeve lobectomy

Yifan Zhong1*, Yang Wang1*, Xuefei Hu1*, Gege Wang2, Yunlang She1, Jiajun Deng1, Lei Zhang1, Qiao Peng1, Yuming Zhu1, Gening Jiang1, Minglei Yang1,3, Dong Xie1, Chang Chen1

1Department of Thoracic Surgery, Tongji University Affiliated Shanghai Pulmonary Hospital, Tongji University School of Medicine, Shanghai, China; 2Institute of Clinical Epidemiology and Evidence-based Medicine, Tongji University School of Medicine, Shanghai, China; 3Department of Thoracic Surgery, Ningbo No. 2 Hospital, Chinese Academy of Sciences, Ningbo, China

Contributions: (I) Conception and design: Y Zhong, Y Wang, X Hu; (II) Administrative support: M Yang, D Xie, C Chen; (III) Provision of study materials or patients: Y Zhu, G Jiang; (IV) Collection and assembly of data: J Deng, L Zhang, Q Peng; (V) Data analysis and interpretation: G Wang, Y She; (VI) Manuscript writing: All authors; (VII) Final approval of manuscript: All authors.

*These authors contributed equally to this work.

Correspondence to: Chang Chen, MD, PhD. Department of Thoracic Surgery, Tongji University Affiliated Shanghai Pulmonary Hospital, Tongji University School of Medicine, Shanghai 200443, China. Email: chenthoracic@163.com; Dong Xie, MD, PhD. Department of Thoracic Surgery, Tongji University Affiliated Shanghai Pulmonary Hospital, Tongji University School of Medicine, Shanghai 200443, China. Email: kongduxd@163.com; Minglei Yang, MD. Department of Thoracic Surgery, Ningbo No.2 Hospital, Chinese Academy of Sciences, Ningbo 315010, China. Email: doctoryml@126.com.

Background: Operative safety and oncologic adequacy of thoracoscopic sleeve lobectomy remain controversial. As such, the purpose of this meta-analysis was to evaluate evidence comparing thoracoscopic and thoracotomy in sleeve lobectomy for centrally located non-small cell lung cancer (NSCLC).

Methods: Electronic searches of PubMed and Web of Science databases were undertaken from inception to March 2020. Comparative studies about thoracoscopic and thoracotomy sleeve lobectomy, with evaluation for perioperative outcomes and oncological results were identified. The following outcomes were measured in this meta-analysis: operating time, blood loss, numbers of resected lymph nodes, chest drainage time, postoperative complication rate, mortality, overall survival (OS). The standardized difference (SMD), relative risk (RR) and hazard ratio (HR) with 95% confidence intervals (CI) were pooled using Stata software.

Results: Six studies generating 281 thoracoscopic and 369 thoracotomy cases were finally included. There was no significant difference in intraoperative blood loss, number of resected lymph nodes, chest drainage time, postoperative complication rate and mortality between two groups. However, thoracoscopic sleeve lobectomy was associated with longer operation time (SMD 0.59, 95% CI: 0.14 to 1.03, P=0.010). And shorter postoperative hospital stays (SMD −0.24, 95% CI: −0.51 to 0.03, P=0.078) were observed in the thoracotomy group with marginal significance. Furthermore, sleeve lobectomy via thoracotomy could achieve comparable OS compared to that via thoracoscopy (HR 0.69, 95% CI: 0.38 to 1.00; P<0.001). In addition, there were no evident publication bias in all observational outcomes.

Conclusions: Current evidence suggests that thoracoscopic sleeve lobectomy is a safe and efficient surgical procedure for centrally located NSCLC, with comparable perioperative outcomes and equivalent oncological results compared to thoracotomy sleeve lobectomy.

Keywords: Video-assisted thoracoscopic surgery (VATS); robotic-assisted thoracoscopic surgery (RATS); thoracotomy; sleeve lobectomy; non-small cell lung cancer (NSCLC)
Introduction

Centrally located non-small-cell lung cancer (NSCLC) is the second common type of lung neoplasms, and curative intent surgery is the preferred therapeutic strategy (1). Initially, pneumonectomy was introduced as the standard surgical procedure for centrally located NSCLC (2,3). However, over the past decades, sleeve lobectomy with more lung parenchyma sparing and without sacrificing the oncologic radicality has been accepted as an alternative to pneumonectomy for centrally located NSCLC (4,5). Its superiority in perioperative outcomes and long-term survival has been confirmed in many researches (4,5).

With the evolution of the minimally invasive technique, thoracoscopic surgery is extensively adopted in various pulmonary procedures (6-8). However, despite the advantages of thoracoscopic surgery in perioperative outcomes including shorter postoperative hospital stays, decreased chest drainage time and reduced postoperative complication rate (9-11), thoracotomy is still the preferred surgical approach in sleeve lobectomy considering the technical difficulty of thoracoscopic sleeve resection.

Since sleeve lobectomy via thoracoscopy was firstly attempted in 2002 (12), several researches reported the successful implementation of this procedure in hand of experienced surgeons (13-16). Recently, a series of comparative studies further identified the feasibility of thoracoscopic sleeve lobectomy for centrally located NSCLC (17-22). However, its operative safety and oncologic efficacy compared to thoracotomy sleeve lobectomy remain controversial. Therefore, the purpose of this meta-analysis was to evaluate evidence comparing thoracotomy and thoracotomy sleeve lobectomy in the treatment of centrally located NSCLC. We present the following article in accordance with the PRISMA reporting checklist (available at http://dx.doi.org/10.21037/jtd-20-1855).

Methods

Search strategy

This study was guided by the PRISMA protocol (23) and the recommendations from the Cochrane Collaboration (24). Computerized searches of PubMed and Web of Science databases were conducted from inception to March 2020. The Mesh terms included: sleeve lobectomy, thoracoscopic, thoracotomy, VATS, robotics, and centrally located lung cancer.

Study selection

Inclusion criteria were as follows: (I) patients who were diagnosed with centrally located NSCLC and underwent sleeve lobectomy; (II) randomized controlled trials and observational (cohort and case-control) studies comparing sleeve lobectomy via thoracoscopy versus thoracotomy; (III) if two or more researches included the same cohort, only the largest or the most recently published one was included; (IV) studies whose full texts were available and language was English. Literatures meeting the following criteria were excluded: publications with incomplete data, review articles, editorials, comments, letters, case reports, and animal experiments.

The selection of original studies was according to the process of reviewing titles, abstracts, and full texts. Two independent researchers (J Deng, Y Zhong) preliminarily identified the literatures based on the titles and abstracts according to the predefined selection criteria. A consensus was achieved after discussion in case of disagreement. Thereafter, full texts of potentially relevant literatures were reassessed to determine their conformity with the criteria.

Data extraction and quality assessment

Relevant data of included articles were initially extracted by two independent investigators (J Deng, Y Zhong) and verified by a senior thoracic surgeon with professional-clinical knowledge (C Chen). Retrieved data were input into a Microsoft Excel (Microsoft Corporation, Redmond, WA, USA, 2019) database.

Video-assisted thoracoscopic surgery (VATS) and robotic-assisted thoracoscopic surgery (RATS) cases were integrated into the thoracoscopic group for further comparison with the thoracotomy group. The following outcomes were compared between thoracoscopic and thoracotomy groups: operating time, blood loss, numbers of lymph node; postoperative hospital stay, chest drainage time, postoperative complication rate, mortality, overall survival (OS).

A ‘star system’ based on Newcastle-Ottawa Scale (NOS) was adopted to estimate the quality of included studies, which is widely applied in the evaluation of nonrandomized studies and comprised of 3 dimensions: selection of subjects, comparability of study groups, and the ascertainment of either the exposure or outcome of interest for case-control or cohort studies (25). Studies with 7–9 stars were graded as high quality whereas those with <7 stars were defined as low quality. Any disagreement was resolved by a consensus after discussion.
Statistical analysis

Standardized difference (SMD), relative risk (RR) and hazard ratio (HR) with 95% confidence intervals (CI) were calculated for continuous and dichotomous variables, respectively. Heterogeneity among different studies was evaluated with utilization of Cochran’s Q test and Higgins $I^2$. Statistical heterogeneity was considered as $I^2$ statistic greater than 50% (26). A fixed-effect model was applied if the heterogeneity was acceptable ($P>0.10$, or $P\leq0.10$ but $I^2\leq50\%$). Otherwise, a random effect model was employed. Publication bias were checked by virtue of the funnel plots model and confirmed with Begg’s and Egger’s tests (27,28). The statistically significant P value was set at <0.05. All mentioned analyses were performed using Stata 14.0 (StataCorp, College Station, TX, USA).

Results

Study description and quality assessment

A total of 152 publications were initially discovered from the two aforementioned online databases up to March 2020. After step by step exclusion, 6 cohort studies (17-22) comparing thoracoscopy and thoracotomy sleeve lobectomy were finally included (Figure 1).

The study descriptions and outcomes of the quality assessment were summarized in Table 1. The study population was comprised of 281 thoracoscopic cases and 369 thoracotomy cases. The mean age of the cohort in every study was more than 60 years old and stage I was the most common pathological stage in the entire population. In addition, all researches were retrospective cohort studies conducted in China and published between 2015 and 2019. Of them, three studies were evaluated as high quality with 7 stars based on NOS.

Baseline characteristics

As displayed in Table 2, baseline characteristics of patients including age, sex, body mass index (BMI), smoking history, FEV1%, tumor location, histological type, pathological T stage and pathological N stage were all similar between the thoracoscopic and thoracotomy groups.
Operative time

Six studies provided information on operative time of sleeve lobectomy, and heterogeneity was identified in these included researches (P<0.001, I^2=83.3%). The meta-analysis indicated that patients undergoing thoracoscopic sleeve lobectomy experienced significantly longer intraoperative operation time (SMD 0.59, 95% CI: 0.14, 1.03, P=0.010) (Figure 2).

Blood loss

Blood loss was evaluated in 6 articles. Compared to thoracotomy, the intraoperative blood loss was similar between thoracoscopic and thoracotomy sleeve lobectomy (SMD −0.13, 95% CI: −0.43 to 0.18, P=0.416), with evidence of heterogeneity (P=0.014, I^2=65.1%) (Figure 3).

Number of resected lymph nodes

All included researches reported data of number of resected lymph nodes, sleeve lobectomy by thoracoscopic surgery could achieve equivalent performance in intraoperative lymph node resection compared with that by thoracotomy (SMD 0.02, 95% CI: −0.19 to 0.22, P=0.878), without evidence of heterogeneity (P=0.700, I^2=0%) (Figure 4).

Postoperative hospital stay

In regard to the postoperative hospital stay, the meta-analysis (6 studies estimated this data) revealed shorter postoperative hospital stays in the thoracoscopic sleeve lobectomy with marginal significance (SMD −0.24, 95% CI: −0.51 to 0.03, P=0.078). Heterogeneity was observed among included studies (P=0.053, I^2=54.3%) (Figure 5).

Chest drainage time

Five articles reported data of chest drainage time and our meta-analysis demonstrated that patients receiving sleeve lobectomy via thoracoscopic have similar chest drainage time to those via thoracotomy (SMD −0.26, 95% CI: −0.69 to 0.17, P=0.235), with specific heterogeneity (P<0.001, I^2=80.6%) (Figure 6).

Postoperative complication rate

The type and proportion of complications with the thoracoscopic sleeve and thoracotomy sleeve lobectomy were summarized in Table S1. Of the 6 studies included, the meta-analysis demonstrated that postoperative complication rate estimated by the Forrest plot was similar between two groups (RR 0.74, 95% CI: 0.51–1.06, P=0.103), with no evidence of heterogeneity (P=0.743, I^2=0%) (Figure 7).

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**Table 1** Study description and quality assessment

<table>
<thead>
<tr>
<th>First author, year</th>
<th>Country</th>
<th>Study type</th>
<th>Treatment</th>
<th>Conversion rate</th>
<th>Number of patients</th>
<th>Mean age, year</th>
<th>Male vs. female</th>
<th>Pathological stage</th>
<th>Quality score</th>
</tr>
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<tbody>
<tr>
<td>Zhou, 2015</td>
<td>China</td>
<td>Cohort</td>
<td>Thoracoscopy</td>
<td>0%</td>
<td>10</td>
<td>60.5</td>
<td>9/1</td>
<td>6 2 2</td>
<td>6</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Thoracotomy</td>
<td>41</td>
<td>62.5</td>
<td>35/6</td>
<td>18 10 13</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Gu, 2018</td>
<td>China</td>
<td>Cohort</td>
<td>Thoracoscopy</td>
<td>1.7%</td>
<td>17</td>
<td>62.0</td>
<td>17/0</td>
<td>6 5 6</td>
<td>6</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Thoracotomy</td>
<td>86</td>
<td>61.1</td>
<td>80/6</td>
<td>36 29 21</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Gao, 2019</td>
<td>China</td>
<td>Cohort</td>
<td>Thoracoscopy</td>
<td>5.6%</td>
<td>54</td>
<td>60.7</td>
<td>44/10</td>
<td>23 19 12</td>
<td>7</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Thoracotomy</td>
<td>94</td>
<td>60.4</td>
<td>87/7</td>
<td>20 35 39</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Qiu, 2019</td>
<td>China</td>
<td>Cohort</td>
<td>Thoracoscopy</td>
<td>2.9%</td>
<td>122</td>
<td>60.4</td>
<td>108/14</td>
<td>NG NG NG</td>
<td>7</td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td>Thoracotomy</td>
<td>66</td>
<td>61.1</td>
<td>62/4</td>
<td>NG NG NG</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Wu, 2019</td>
<td>China</td>
<td>Cohort</td>
<td>Thoracoscopy</td>
<td>NG</td>
<td>21</td>
<td>62.0</td>
<td>19/2</td>
<td>8 4 9</td>
<td>7</td>
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<tr>
<td></td>
<td></td>
<td></td>
<td>Thoracotomy</td>
<td>21</td>
<td>61.0</td>
<td>19/2</td>
<td>7 6 8</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Xie, 2020</td>
<td>China</td>
<td>Cohort</td>
<td>Thoracoscopy</td>
<td>4.5%</td>
<td>112</td>
<td>62.7</td>
<td>91/21</td>
<td>48 35 29</td>
<td>7</td>
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<tr>
<td></td>
<td></td>
<td></td>
<td>Thoracotomy</td>
<td>251</td>
<td>62.1</td>
<td>234/17</td>
<td>104 72 75</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Quality assessment of included studies was performed using the modified Newcastle-Ottawa Scale (NOS); NG, not given.
### Table 2 Baseline characteristics between thoracoscopic and thoracotomy sleeve lobectomy

<table>
<thead>
<tr>
<th>Variables</th>
<th>Number of publications</th>
<th>Test for overall effect</th>
<th>Test for heterogeneity</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td>Thoracoscopy vs.</td>
<td></td>
</tr>
<tr>
<td></td>
<td></td>
<td>thoracotomy</td>
<td></td>
</tr>
<tr>
<td></td>
<td></td>
<td>P value</td>
<td>I²</td>
</tr>
<tr>
<td>Age, SMD (95% CI)</td>
<td>5</td>
<td>−0.01 (−0.19, 0.16)</td>
<td>0.868</td>
</tr>
<tr>
<td>Sex (male), RR (95% CI)</td>
<td>6</td>
<td>1.00 (0.88, 1.13)</td>
<td>0.952</td>
</tr>
<tr>
<td>BMI, SMD (95% CI)</td>
<td>5</td>
<td>0.04 (−0.13, 0.21)</td>
<td>0.633</td>
</tr>
<tr>
<td>FEV1%, SMD (95% CI)</td>
<td>4</td>
<td>−0.16 (−0.37, 0.05)</td>
<td>0.145</td>
</tr>
<tr>
<td>Smoking (ever), RR (95% CI)</td>
<td>6</td>
<td>0.97 (0.83, 1.13)</td>
<td>0.666</td>
</tr>
<tr>
<td>Histological type (SCC), RR (95%CI)</td>
<td>6</td>
<td>1.00 (0.87, 1.15)</td>
<td>0.981</td>
</tr>
<tr>
<td>Location (left), RR (95% CI)</td>
<td>6</td>
<td>1.08 (0.89, 1.30)</td>
<td>0.450</td>
</tr>
<tr>
<td>pT1 Stage, RR (95% CI)</td>
<td>3</td>
<td>1.09 (0.68, 1.76)</td>
<td>0.713</td>
</tr>
<tr>
<td>pT2 Stage, RR (95% CI)</td>
<td>3</td>
<td>1.03 (0.86, 1.23)</td>
<td>0.772</td>
</tr>
<tr>
<td>pT3 Stage, RR (95% CI)</td>
<td>3</td>
<td>1.11 (0.40, 3.08)</td>
<td>0.839</td>
</tr>
<tr>
<td>pT4 Stage, RR (95% CI)</td>
<td>2</td>
<td>0.61 (0.21, 1.76)</td>
<td>0.361</td>
</tr>
<tr>
<td>pN0 Stage, RR (95% CI)</td>
<td>4</td>
<td>1.04 (0.85, 1.27)</td>
<td>0.724</td>
</tr>
<tr>
<td>pN1 Stage, RR (95% CI)</td>
<td>4</td>
<td>0.93 (0.63, 1.37)</td>
<td>0.712</td>
</tr>
<tr>
<td>pN2 Stage, RR (95% CI)</td>
<td>4</td>
<td>0.95 (0.69, 1.32)</td>
<td>0.775</td>
</tr>
</tbody>
</table>

Note: for propensity matched studies, only cases after matching were included. BMI, body mass index; SCC, squamous cell carcinoma.

**Figure 2** Meta-analysis: the operation time for thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.
Mortality

All 6 included publications provided data of mortality, the meta-analysis revealed that thoracoscopic surgery did not increase the mortality of sleeve lobectomy compared to conventional thoracotomy (30-day mortality: RR 0.42, 95% CI: 0.06–3.12, P=0.394; 90-day mortality: RR 0.55, 95% CI: 0.15–2.04, P=0.367) and no significant heterogeneity was identified among publications (30-day mortality: P=0.495, $I^2$=0%; 90-day mortality: P=0.428, $I^2$=0%) (Figure 8).

OS

OS was defined as the duration from surgery to the date...
of death or last follow-up. 6 studies with 650 patients were included in the analysis of OS. Sleeve lobectomy by thoracoscopy was associated with favorable OS (HR 0.69, 95% CI: 0.38–1.00; P<0.001), without evidence of heterogeneity (P=0.915, I²=0%) (Figure 9).

Publication bias
As displayed in Figure 10, no potential publication bias existed in the HRs of OS among studies according to the asymmetrical distribution of funnel plot, which was further identified by the publication bias test (Begg’s test, P=0.806;
Figure 7 Meta-analysis: the postoperative complication rate for thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.

Figure 8 Meta-analysis: the 30-day (A) and 90-day (B) mortality rate for thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.
Egger’s test, P=0.193). Funnel plots of the other results revealed no publication bias (Table S1, Figures S1-S7).

**Discussion**

Thoracoscopy has been widely adopted in the surgical management of lung cancer (6–8). Compared to conventional thoracotomy, favorable perioperative outcomes and uncompromising oncologic results could be achieved by this minimally invasive technique (9-11). Sleeve lobectomy, as the standard therapeutic strategy for centrally located NSCLC, was initially regarded as the contradiction for thoracoscopic surgery in consideration of its operative risk (29,30). Since the 21st century, sleeve lobectomy has been successfully performed via thoracoscopic platform (12-16), but only confined to experienced centers with a small sample size. The feasibility and safety of this procedure have not been fully investigated. Our meta-analysis proved thoracoscopic sleeve lobectomy is a feasible and safe procedure for centrally located NSCLC.

Thoracoscopic sleeve lobectomy has experienced evolution toward less operative trauma over the past decades. In 2002, Santambrogio et al. (12) firstly reported thoracoscopic sleeve bronchoplasty. Subsequently, several series of studies reported their experiences in sleeve lobectomy through video-assisted mini-thoracotomy (30-32). As experience of the thoracoscopic technique accumulated, sleeve lobectomy was successfully performed by complete VATS in 2008 (13). Thereafter, a series of case reports described complete VATS sleeve resection via three and four ports, the operative duration was reported from 176 to 287 minutes and the postoperative hospital stays were between 3 and 8 days (33-35). Over the years, VATS has evolved into a single incision access without rib spreading, which could offer better perioperative outcomes than multi-port surgery (36). In 2013, Gonzalez-Rivas et al. (14) reported the first case of uniportal VATS sleeve lobectomy. The operative time and postoperative hospital stays were 240 minutes and 5 days, respectively. With further advancements in operative technique, surgical equipment and anesthesia management, Shao et al. (16) firstly described complete VATS bronchial sleeve resection under non-intubated epidural anesthesia. In this case, the operative time was 165 minutes, the patient did not require assisted ventilation after surgery, he could drink and eat without any restriction at 4 hours postoperatively and became mobile at the first postoperative day. This patient was discharged after 6 days of operation and no complication was identified.

Nevertheless, thoracotomy was still the preferred surgical approach for sleeve lobectomy considering the high technical demand of sleeve lobectomy and the limitation of VATS in deep perception and maneuverability. The emergence of the RATS has provided the solution for the
deficiency of VATS. Schmid et al. (15) initially reported robotic sleeve resection, the operation duration was 364 minutes. And in the following researches, RATS was demonstrated as a safe approach for sleeve lobectomy with satisfactory perioperative outcomes (37,38).

Although the feasibility of thoracoscopic sleeve lobectomy has been proved in many reports, its efficacy compared to thoracotomy sleeve lobectomy remained controversial. Up to now, six studies investigated the operative safety and oncological adequacy of sleeve lobectomy via thoracoscopic surgery by comparing with that by thoracotomy. In these researches, intraoperative performance of thoracoscopic sleeve lobectomy varied in different centers. Four studies demonstrated that the VATS group experienced significantly longer operative duration than the thoracotomy group (17,19,20,22) and Qiu et al. (19) further identified that RATS can offer a significant advantage on operative duration over thoracotomy in bronchial sleeve lobectomy. While, similar operation time between the thoracoscopic and thoracotomy groups was identified in other two studies (18,21). Concerning on intraoperative blood loss, three articles suggested less estimated blood loss in thoracoscopic sleeve lobectomy (17,19,21). But no significant difference between two surgical approaches was observed in the remaining three publications (18,20,22). Discrepancy of operative expertise and impact of learning curve might contribute to the heterogeneity in intraoperative performance among medical centers. In larger and more experienced centers, sleeve lobectomy via thoracoscopic surgery could confer superiority in perioperative outcomes over that by thoracotomy (21,22). Radical lymphadenectomy is of great importance in reducing the late recurrence risk after surgical resection of tumors. For all six included researches, there was no significant difference in performance of intraoperative lymph node dissection between thoracoscopy and thoracotomy groups (17-22).

In terms of postoperative outcomes, five included studies revealed no statistical difference in postoperative complication rate between the thoracoscopic and thoracotomy groups (17-20,22). Only one research proved lower complication rate after thoracoscopic sleeve resection (21). And for all included studies, mortality rate was similar between the thoracoscopic and thoracotomy group (17-22). In addition, postoperative hospital stays and chest drainage time were significantly shorter in patients receiving thoracoscopic sleeve lobectomy, which was proved by most included articles (17,19,21,22). Regarding oncologic results, all six studies demonstrated that thoracoscopic sleeve lobectomy can achieve equivalent OS compared to thoracotomy sleeve lobectomy (17-22). The main reason for the hesitation to perform sleeve lobectomy is the concern about the local recurrence. Unfortunately, the results of local recurrence were only obtainable in two articles (19,21), which limited the implementation of meta-analysis based on this factor, these two studies showed no significant difference in local recurrence rate between thoracoscopic and thoracotomy sleeve lobectomy.

To the best of our knowledge, this is the first study evaluating the efficacy of thoracoscopic sleeve lobectomy by virtue of meta-analysis. In this study, comparable perioperative outcomes and oncologic results were identified in the patients undergoing sleeve lobectomy via thoracoscopic surgery compared to thoracotomy, which confirmed the efficacy of thoracoscopic sleeve lobectomy. The thoracoscopic group experienced longer operation time than the thoracotomy group, which might arise from the operative difficulty of sleeve lobectomy and nonproficiency of the thoracoscopic technique. Proficiency of thoracoscopic sleeve lobectomy varied among different centers, the intraoperative performance was usually better in more experienced centers. Also, there was heterogeneity of experience on thoracoscopic sleeve lobectomy among surgeons, most included studies did not adjust the bias of surgeons, which affected the intraoperative outcomes between thoracoscopic and thoracotomy groups. In general, with the increasing proficiency, thoracoscopic...
surgery could achieve equivalent operative performance to thoracotomy (39-41). Additionally, as displayed in Figure 9, our research revealed thoracoscopic sleeve lobectomy can offer a significant advantage on OS over thoracotomy sleeve lobectomy (P<0.001, HR 0.38–1.00). However, in the original results of all included studies, no significant difference was observed in OS between two groups, which might be the consequence of the limited sample size of thoracoscopic sleeve lobectomy.

Several limitations of this meta-analysis should be acknowledged. Firstly, all included researches were retrospective cohort studies which have certain inevitable bias. We evaluated the quality of publications based on a widely adopted instrument for observational studies and judged most included researches at high quality. Additionally, the funnel plot and bias test of most variables did not indicate evident publication bias among included studies, which was also reassuring. And we hope a randomized controlled trial in future can validate our conclusions.

Secondly, the quantity of included publications in this study was relatively small and every included study was single-center based on Chinese population. Currently, only experienced medical institutions can routinely perform thoracoscopic sleeve lobectomy in view of the technical difficulty of this surgical procedure. Further, China has the largest population of lung cancer in the world and many centers in China have accumulated adequate experiences in thoracoscopic sleeve lobectomy. Still, a large-scale international study in future is warranted to verify results in our research.

Thirdly, there were limitations in the scope of analysis process. Systematic analyses could not be conducted for variables only described in a small number of researches such as recurrence-free survival and local recurrence. Additionally, limited by the number of studies investigating thoracoscopic sleeve lobectomy, the uniportal VATS, multiportal VATS and RATS cases were analyzed as a whole when compared with the thoracotomy group and subgroup analyses based on surgical approaches could not be performed, so the potential bias resulting from the differences among these three surgical approaches are inevitable. Furthermore, the follow-up time of the survival data in all included studies was relatively short, which weakened the level of evidence for the oncologic efficacy of thoracoscopic sleeve lobectomy.

Finally, significant heterogeneity existed among some variables in our study. Every center adopted specific inclusion and evaluation criteria for postoperative complications, resulting in the difference of complication rate among studies. Moreover, the indication of thoracoscopic sleeve lobectomy and experience for this surgical procedure varied among surgeons, which might contribute to the heterogeneity in intraoperative performance and oncological results. We tried to identify the sources of heterogeneity from original researches, however most articles did not provide relevant information.

**Conclusions**

Current evidence suggests that thoracoscopic sleeve lobectomy is a safe and efficient surgical procedure for centrally located NSCLC, with uncompromising perioperative outcomes and equivalent oncological results compared to thoracotomy sleeve lobectomy. This conclusion requires confirmation by more large-scale and high-quality researches.

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**Footnote**

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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References

### Table S1 The type and proportion of complications with the thoracoscopic sleeve and thoracotomy sleeve lobectomy

<table>
<thead>
<tr>
<th>Types of complication</th>
<th>All included cases (n=650)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Thoracoscopy (n=281), %</td>
</tr>
<tr>
<td>Prolonged air leak</td>
<td>1.83</td>
</tr>
<tr>
<td>Cardiac arrhythmia</td>
<td>2.14</td>
</tr>
<tr>
<td>Atelectasis</td>
<td>1.21</td>
</tr>
<tr>
<td>Pneumonia</td>
<td>4.47</td>
</tr>
<tr>
<td>Chylothorax</td>
<td>1.07</td>
</tr>
<tr>
<td>Pulmonary embolus</td>
<td>0.71</td>
</tr>
<tr>
<td>Bronchopleural fistula</td>
<td>2.06</td>
</tr>
<tr>
<td>Subcutaneous emphysema</td>
<td>1.47</td>
</tr>
<tr>
<td>Pyothorax</td>
<td>0.71</td>
</tr>
<tr>
<td>Bronchial anastomosis bleeding</td>
<td>0.36</td>
</tr>
<tr>
<td>Anastomosis bursting</td>
<td>0</td>
</tr>
<tr>
<td>Multiple organ failure</td>
<td>0.36</td>
</tr>
<tr>
<td>Recurrent laryngeal nerve injury</td>
<td>0</td>
</tr>
<tr>
<td>Anastomotic stenosis/obstruction</td>
<td>0.44</td>
</tr>
<tr>
<td>Postoperative tracheotomy</td>
<td>0</td>
</tr>
<tr>
<td>Pneumothorax</td>
<td>1.20</td>
</tr>
<tr>
<td>Hemothorax</td>
<td>0.35</td>
</tr>
<tr>
<td>Pleural effusion</td>
<td>0.59</td>
</tr>
<tr>
<td>Acute myocardial infarction</td>
<td>0.29</td>
</tr>
<tr>
<td>Respiratory failure</td>
<td>0</td>
</tr>
</tbody>
</table>

For propensity matched studies, only cases after matching were included.
**Figure S1** The funnel plot and publication bias tests for operative time of thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.

**Figure S2** The funnel plot and publication bias tests for blood loss of thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.

**Figure S3** The funnel plot and publication bias tests for number of resected lymph nodes of thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.

**Figure S4** The funnel plot and publication bias tests for postoperative hospital stays of thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.
**Figure S5** The funnel plot and publication bias tests for chest drainage time of thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.

**Figure S6** The funnel plot and publication bias tests for postoperative complication rate of thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.

**Figure S7** The funnel plot and publication bias tests for 30-day (A) and 90-day (B) mortality of thoracoscopic and thoracotomy sleeve lobectomy. Note: for propensity matched studies, only cases after matching were included.