

# Percutaneous endoscopic versus minimally invasive transforaminal lumbar interbody fusion for lumbar degenerative diseases: a meta-analysis

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## Abstract

**Introduction:** Transforaminal lumbar interbody fusion (TLIF) is commonly used in patients with lumbar degenerative disease (LDD). The most commonly used techniques include minimally invasive TLIF (MIS-TLIF) and percutaneous endoscopic TLIF (PE-TLIF).

**Aim:** To compare the safety and clinical effectiveness of PE-TLIF and MIS-TLIF in treating LDD.

**Material and methods:** We screened for related articles in multiple scientific databases, namely, PubMed, Embase, Cochrane Library, Wanfang, VIP, and CINK, and analyzed the relative outcomes.

**Results:** Based on our inclusion criteria, we selected 8 studies for meta-analysis. There are a total of 229 patients who underwent PE-TLIF and 258 patients who underwent MIS-TLIF. MIS-TLIF and PE-TLIF have similar effectiveness in relieving leg pain and improving the Oswestry Disability Index. However, PE-TLIF is superior in relieving back pain. The pooled data of fusion rates, postoperative analgesic, and complication rates are comparable between the 2 groups. The pooled operation and intra-operative fluoroscopic time are both significantly higher in the PE-TLIF group than the MIS-TLIF group. The pooled intra-operative blood loss, incision length, duration from surgery to ambulation, and hospital stay are significantly lower in the PE-TLIF group than the MIS-TLIF group. Most of the endpoints reveal significant heterogeneity. The endpoints of operation time and intra-operative blood loss reveal significant publication bias.

**Conclusions:** Both PE-TLIF and MIS-TLIF are safe and effective interventions for patients with LDD. When compared, although MIS-TLIF results in reduced operative time, less intra-operative blood loss and enhanced post-operative recovery can be achieved by PE-TLIF.

**Key words:** transforaminal lumbar interbody fusion, lumbar degenerative disease, Oswestry Disability Index, meta-analysis.

## Introduction

Lumbar degenerative diseases (LDD), which include lumbar spinal stenosis, lumbar disc herniation, spondylolisthesis, and lumbar instability, are common causes of back and leg pain [1–3]. Transforam-

inal lumbar interbody fusion (TLIF) is commonly used for patients with LDD and the purposes of TLIF are to relieve the symptoms and improve the quality of life [4–6]. Nevertheless, the classical open TLIF is correlated with iatrogenic injury of the paraspinal muscle, re-

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sulting in intractable postoperative low back pain [7]. At present, minimally invasive TLIF (MIS-TLIF) is widely used to minimize the soft tissue injury and intraoperative blood loss [8]. MIS-TLIF has shown significantly less blood loss compared with open TLIF [9]. However, MIS-TLIF is usually restricted by a limited working space and it is challenging to visualize deep surgical fields through the tubular retractor [10].

To overcome MIS-TLIF shortcomings, percutaneous endoscopic TLIF (PE-TLIF) has been commonly used for patients with LDD [11–13]. PE-TLIF makes possible a complete endoscopic discectomy, spinal canal and foramina decompression, and interbody fusion via the endoscopic and working portal [14–17]. Although numerous earlier investigations explored the relative efficacies of PE-TLIF and MIS-TLIF in treating LDD [11–18], most of these studies were retrospective in nature. Since multiple factors can sway the outcomes of a retrospective investigation, and introduce unintentional bias, a meta-analysis is recommended to extensively evaluate relevant literature, reduce bias risk, and enhance the overall statistical power of the collective study outcomes.

## Aim

Herein, our goal was to compare the safety and clinical effectiveness of PE-TLIF and MIS-TLIF in treating LDD.

## Material and methods

### Study selection

Our research abided by the guidelines of the Preferred Reporting Items for Systematic reviews and Meta-Analyses [18], and is registered at <https://inplasy.com/> (Number: INPLASY202220090).

We screened for relevant publications in scientific databases, namely, PubMed, Embase, Cochrane Library, Wanfang, VIP, and CINK from the date of database establishment to November 2021. Our search terminologies were as follows: ((Endoscopic Lumbar Interbody Fusion) OR (Endo-LIF)) AND ((Minimally Invasive Transforaminal Lumbar Interbody Fusion) OR (MIS-TLIF)).

The following criteria were used to select relevant articles for analysis:

- Studies: comparing between PE-TLIF and MIS-TLIF;
- Diseases: LDD;
- Language: Unlimited.

Studies with the following criteria were eliminated from the analysis: (a) single-arm investigations; (b) non-human investigations; and (c) studies without an English title and/or abstract.

### Data accumulation

Two separate authors screened eligible publications and collected baseline study data, namely, author, year of publication, country of publication, study format, and quality assessment; and patient data, namely, population size, age, gender, disk range, body mass index (BMI), and follow-up duration; and many treatment-related data. Disagreements were rectified through consultation with a third author.

### Quality evaluation

Randomized controlled trials (RCTs) were evaluated using the Cochrane risk of bias tool for the potential bias in performance, attrition, detection, screening, reporting, and so on. Non-RCTs were analyzed via the 9-point Newcastle-Ottawa scale (NOS) for potential selection bias, namely, selection (4 points), comparability (2 points), and exposure (3 points), carrying either high ( $\geq 7$ ), moderate (4–6), or low ( $< 4$ ) risk of bias [19].

### Endpoints

The endpoints of this meta-analysis included visual analog scale (VAS)-leg, VAS-back, Oswestry Disability Index (ODI), fusion rate, surgical duration, blood loss, complication rates, and postoperative hospitalization.

### Statistical analysis

RevMan v5.3 and Stata 12.0 were employed in this study. Dichotomous variables were accumulated depending upon odds ratios (ORs) with 95% confidence intervals (CIs), while continuous variables are accumulated depending upon mean difference (MD) with 95% CIs. Heterogeneity was examined via  $I^2$  tests, with  $I^2 > 50\%$  representing marked heterogeneity. Random-effects models were used to analyze marked heterogeneity, while fixed-effects models were used to analyze marked homogeneity. Heterogeneity sources are analyzed through sensitivity assessment. Funnel plot and Egger test are employed for evaluating publication bias risks.

## Results

### Included studies

We initially found 323 relative studies by using the research strategy. After step-by-step selection, a total of 8 studies were eligible for analysis (Figure 1, Table I). Seven studies were retrospective [12–18], and 1 study was prospective [11]. Seven studies were from China [11, 12, 14–18], and 1 study was from South Korea [13].

In these 8 included studies, 229 patients underwent PE-TLIF and 258 patients underwent MIS-TLIF (Table II). The mean age, gender ratio, and mean BMI were comparable between PE-TLIF and MIS-TLIF groups in the analyzed studies. The NOS of these studies ranged between 7 and 8.

### Improvement of VAS-leg

Six studies compared the VAS-leg before and after treatment [11–14, 16, 17]. The pooled  $\Delta$ VAS-leg values are comparable between the 2 groups (MD =  $-0.31$ ; 95% CI =  $-0.95-0.33$ ;  $p = 0.35$ , Figure 2 A). Significant heterogeneity is found for this endpoint ( $I^2 = 98\%$ ). Sensitivity analysis suggests that removing individual studies has no impact on the detected heterogeneity. No obvious publication bias is observed (Egger test,  $p = 0.597$ ).

### Improvement of VAS-back

Six studies compared the VAS-back before and after treatment [11–14, 16, 17]. The pooled  $\Delta$ VAS-back value is significantly higher in the PE-TLIF group

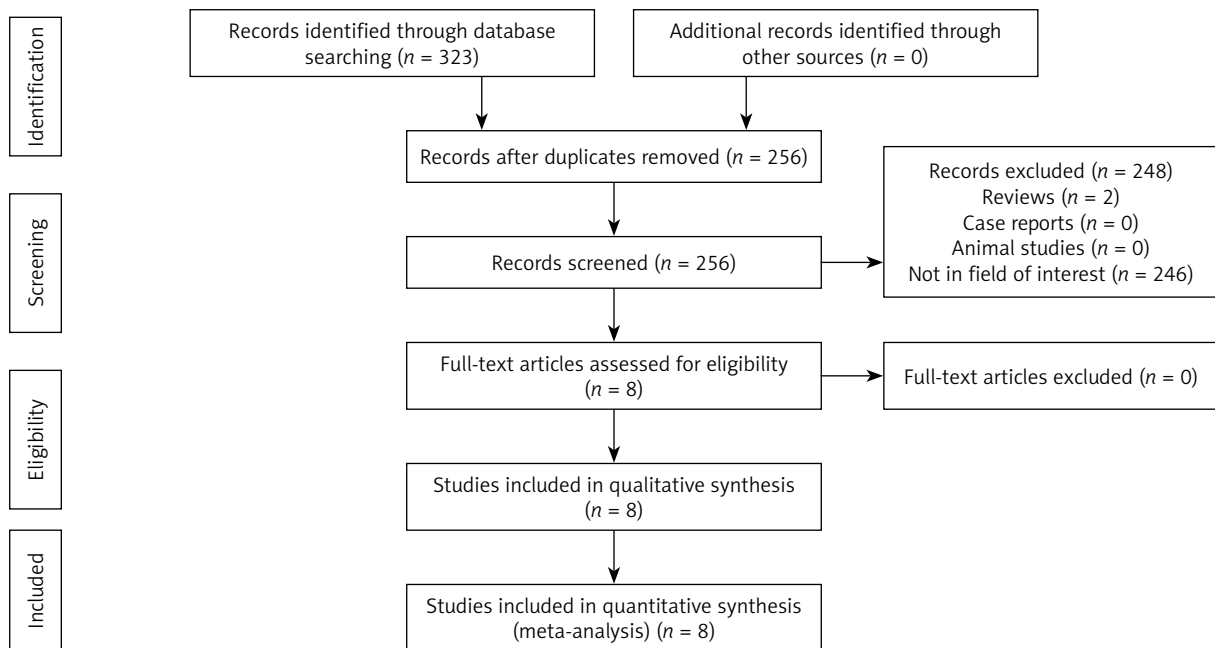


Figure 1. Flowchart of this study

Table I. Baseline data of the included studies

First author	Year	Countries	Design	Newcastle-Ottawa scale
Ao [11]	2020	China	Prospective	8
Dong [12]	2019	China	Retrospective	8
Kim [13]	2021	South Korea	Retrospective	7
Li [14]	2020	China	Retrospective	8
Wen [15]	2020	China	Retrospective	7
Xue [16]	2021	China	Retrospective	7
Zhang [17]	2021	China	Retrospective	8
Zhao [18]	2021	China	Retrospective	7

**Table II.** Patients' details in each study

Authors [ref.]	Disease types	Disk range	Groups	Number of patients	Age [years]	M/F	BMI	Follow-up [months]
Ao [11]	Degenerative spondylolisthesis	L3-S1	PE-TLIF	35	52.80	16/19	24.73	12
			MIS-TLIF	40	53.68	22/18	25.08	12
Dong [12]	Disc herniation, Spondylolisthesis, Spinal stenosis	L3-S1	PE-TLIF	28	58.3	17/11	23.3	24
			MIS-TLIF	25	54.8	9/16	23.0	24
Kim [13]	Degenerative/lytic spondylolisthesis	L2-S1	PE-TLIF	32	70.5	17/15	Unknown	27.2
			MIS-TLIF	55	67.3	25/30	Unknown	31.5
Li [14]	Disc herniation, Spondylolisthesis, Spinal stenosis	L4-S1	PE-TLIF	22	52.0	12/10	Unknown	14.9
			MIS-TLIF	30	50.7	18/12	Unknown	14.7
Wen [15]	Not given in detail	L4-S1	PE-TLIF	20	48.33	13/7	Unknown	12–20 for all
			MIS-TLIF	20	50.77	11/9	Unknown	
Xue [16]	Not given in detail	L3-5	PE-TLIF	20	46.3	11/9	Unknown	16.1
			MIS-TLIF	20	48.4	12/8	Unknown	15.8
Zhang [17]	Degenerative/isthmic spondylolisthesis	L3-S1	PE-TLIF	32	53.1	12/20	25.5	12
			MIS-TLIF	30	55.7	14/16	24.9	12
Zhao [18]	Not given in detail	L3-S1	PE-TLIF	40	56.93	37/17	Unknown	24
			MIS-TLIF	38	57.01	20/18	Unknown	24

BMI – body mass index, M – male, F – female, PE-TLIF – percutaneous endoscopic transforaminal lumbar interbody fusion, MIS-TLIF – minimally invasive transforaminal lumbar interbody fusion.

(MD = 1.62; 95% CI: 0.92–2.32;  $p < 0.00001$ , Figure 2 B). Significant heterogeneity is found for this endpoint ( $I^2 = 98%$ ). Sensitivity analysis suggests that removing individual studies has no impact on the detected heterogeneity. No obvious publication bias is observed (Egger test,  $p = 0.188$ ).

### Improvement of ODI

Six studies compared the ODI before and after treatment [11–14, 16, 17]. The pooled  $\Delta$ ODI values are comparable between the 2 groups (MD = 3.82; 95% CI: –1.53–9.18;  $p = 0.16$ , Figure 2 C). Significant heterogeneity is found for this endpoint ( $I^2 = 98%$ ). Sensitivity analysis suggests that removing individual studies has no impact on the detected heterogeneity. No obvious publication bias is observed (Egger test,  $p = 0.074$ ).

### Postoperative analgesic

Two studies reported the dose of postoperative analgesic [11, 12]. The pooled dose of postoperative analgesic is comparable between the 2 groups (MD = –32.45; 95% CI: –82.74–17.83;  $p = 0.21$ , Figure 2 D).

Significant heterogeneity is found for this endpoint ( $I^2 = 99%$ ). Sensitivity analysis cannot be performed because there are only 2 studies. The funnel plot does not reveal any publication bias.

### Fusion rates

Seven studies reported the fusion rates after treatment [11–14, 16–18]. The pooled fusion rates are comparable between the 2 groups (90.4% vs. 90.3%;  $p = 0.86$ , Figure 2 E). No significant heterogeneity is found for this endpoint ( $I^2 = 0%$ ). No obvious publication bias is observed (Egger test,  $p = 0.797$ ).

### Operation time

All studies reported the operation time. The pooled operation time is significantly lower in the MIS-TLIF group than in the PE-TLIF group (MD = 22.41; 95% CI: 6.48–38.34;  $p = 0.006$ , Figure 2 F). Significant heterogeneity is found for this endpoint ( $I^2 = 95%$ ). Sensitivity analysis suggests that removing individual studies has no impact on the detected heterogeneity. We observed marked publication bias (Egger test,  $p = 0.049$ ).

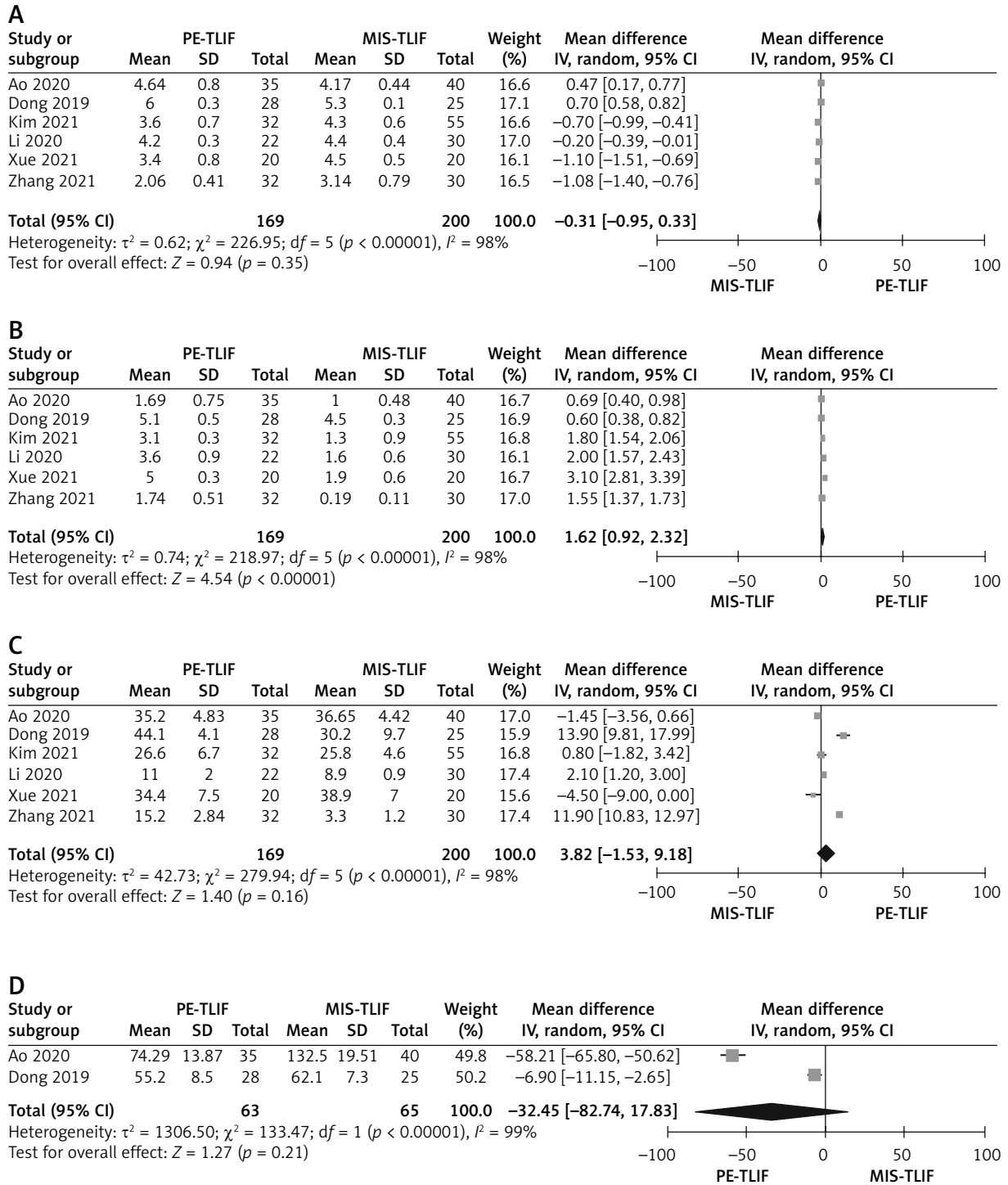
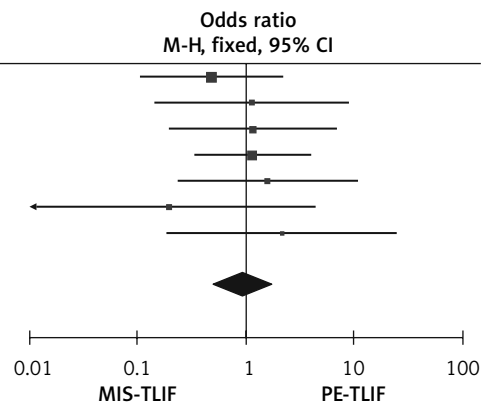


Figure 2. Pooled results of: A – improvement of VAS-leg, B – improvement of VAS-back, C – improvement of ODI, D – postoperative analgesic

**E**

Study or subgroup	PE-TLIF		MIS-TLIF		Weight (%)	Odds ratio M-H, fixed, 95% CI
	Events	Total	Events	Total		
Ao 2020	29	34	36	39	36.1	0.48 [0.11, 2.19]
Dong 2019	26	28	23	25	9.2	1.13 [0.15, 8.68]
Kim 2021	30	32	51	55	12.4	1.18 [0.20, 6.81]
Li 2020	16	22	21	30	25.7	1.14 [0.34, 3.87]
Xue 2021	18	20	17	20	9.0	1.59 [0.24, 10.70]
Zhang 2021	30	32	30	30	12.6	0.20 [0.01, 4.34]
Zhao 2021	39	40	36	38	4.9	2.17 [0.19, 24.93]

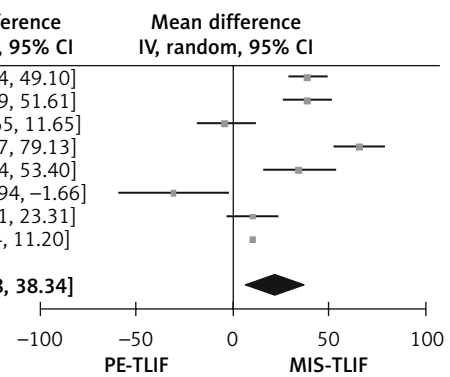
**Total (95% CI)** 208 237 100.0 **0.94 [0.49, 1.80]**  
 Total events 188 214  
 Heterogeneity:  $\chi^2 = 2.64$ ;  $df = 6$  ( $p = 0.85$ ),  $I^2 = 0\%$   
 Test for overall effect:  $Z = 0.17$  ( $p = 0.86$ )



**F**

Study or subgroup	PE-TLIF			MIS-TLIF			Weight (%)	Mean difference IV, random, 95% CI
	Mean	SD	Total	Mean	SD	Total		
Ao 2020	143	24.2	35	103.63	17.79	40	13.3	39.37 [29.64, 49.10]
Dong 2019	184.6	24.4	28	145.4	21.7	25	12.9	39.20 [26.79, 51.61]
Kim 2021	169.5	24.9	32	173	47.1	55	12.5	-3.50 [-18.65, 11.65]
Li 2020	187.7	27	22	121.6	18.2	30	12.8	66.10 [53.07, 79.13]
Wen 2020	150.45	35.87	20	115.48	21.97	20	11.8	34.97 [16.54, 53.40]
Xue 2021	140.3	35.6	20	170.6	54.8	20	9.7	-30.30 [-58.94, -1.66]
Zhang 2021	202.6	31.4	32	192.1	18.9	30	12.9	10.50 [-2.31, 23.31]
Zhao 2021	100.92	1.34	40	90.45	1.87	38	14.0	10.47 [9.74, 11.20]

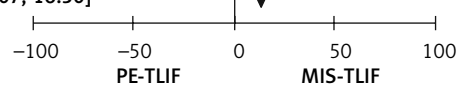
**Total (95% CI)** 229 258 100.0 **22.41 [6.48, 38.34]**  
 Heterogeneity:  $\tau^2 = 470.24$ ;  $\chi^2 = 141.15$ ;  $df = 7$  ( $p < 0.00001$ ),  $I^2 = 95\%$   
 Test for overall effect:  $Z = 2.76$  ( $p = 0.006$ )



**G**

Study or subgroup	PE-TLIF			MIS-TLIF			Weight (%)	Mean difference IV, fixed, 95% CI
	Mean	SD	Total	Mean	SD	Total		
Dong 2019	68.5	15.1	28	53.7	6.4	25	11.9	14.80 [8.67, 20.93]
Zhang 2021	46.3	5.1	32	32.2	3.9	30	88.1	14.10 [11.85, 16.35]

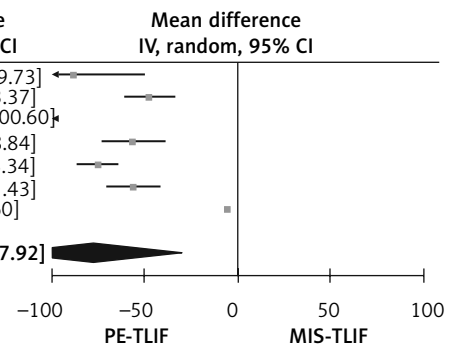
**Total (95% CI)** 60 55 100.0 **14.18 [12.07, 16.30]**  
 Heterogeneity:  $\chi^2 = 0.04$ ;  $df = 1$  ( $p = 0.83$ ),  $I^2 = 0\%$   
 Test for overall effect:  $Z = 13.15$  ( $p < 0.00001$ )



**H**

Study or subgroup	PE-TLIF			MIS-TLIF			Weight (%)	Mean difference IV, random, 95% CI
	Mean	SD	Total	Mean	SD	Total		
Ao 2020	84.29	44.34	35	171.79	112.27	40	13.4	-87.50 [-125.27, -49.73]
Dong 2019	191.1	30.8	28	238.2	19.5	25	14.4	-47.10 [-60.83, -33.37]
Li 2020	49.5	12.1	22	267.7	47.1	30	14.3	-218.20 [-235.80, -200.60]
Wen 2020	90.68	22.3	20	146.78	32.45	20	14.3	-56.10 [-73.36, -38.84]
Xue 2021	65.6	15.3	20	140.5	21.5	20	14.5	-74.90 [-86.46, -63.34]
Zhang 2021	73	26.4	32	129	31.7	30	14.4	-56.00 [-70.57, -41.43]
Zhao 2021	60.56	0.36	40	65.47	0.91	38	14.6	-4.91 [-5.22, -4.60]

**Total (95% CI)** 197 203 100.0 **-77.48 [-127.05, -27.92]**  
 Heterogeneity:  $\tau^2 = 4384.26$ ;  $\chi^2 = 839.42$ ;  $df = 6$  ( $p < 0.00001$ ),  $I^2 = 99\%$   
 Test for overall effect:  $Z = 3.06$  ( $p = 0.002$ )



**Figure 2.** Cont. **E** – fusion rates, **F** – operation time, **G** – intra-operative fluoroscopic time, **H** – intra-operative blood loss

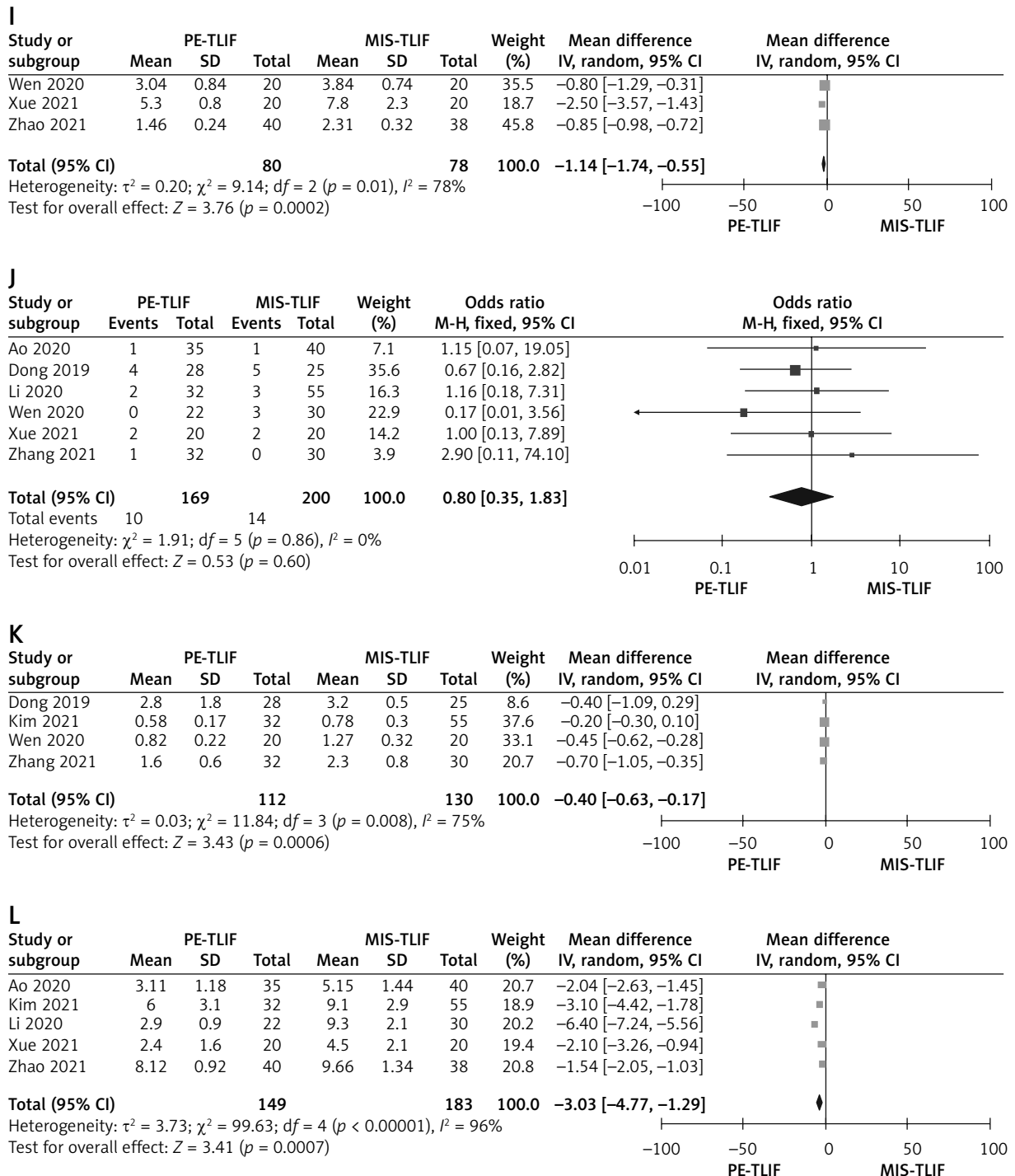


Figure 2. Cont. I – length of incision, J – complication rates, K – time to ambulation, L – hospital stay

### Intra-operative fluoroscopic time

Two studies reported the intra-operative fluoroscopic time [12, 17]. The pooled intra-operative fluoroscopic time is significantly lower in the MIS-TLIF

group than in the PE-TLIF group (MD = 14.18; 95% CI: 12.07–16.30;  $p < 0.00001$ , Figure 2 G). No significant heterogeneity is found for this endpoint ( $I^2 = 0\%$ ). The funnel plot does not reveal any publication bias.

### Intra-operative blood loss

Seven studies reported on the intra-operative blood loss [11, 12, 14–18]. The pooled intra-operative blood loss is drastically lower in the PE-TLIF group than in the MIS-TLIF group (MD = -77.48; 95% CI: -127.05–27.92;  $p = 0.002$ , Figure 2 H). Significant heterogeneity is found for this endpoint ( $I^2 = 99\%$ ). Sensitivity analysis suggests that removing individual studies has no impact on the detected heterogeneity. We observed marked publication bias (Egger test,  $p = 0.021$ ).

### Length of incision

Three studies reported the length of incision [15–17]. The pooled length of incision is significantly lower in the PE-TLIF group than in the MIS-TLIF group (MD = -1.14; 95% CI: -1.74–0.55;  $p = 0.0002$ , Figure 2 I). Significant heterogeneity is found for this endpoint ( $I^2 = 78\%$ ). The significant heterogeneity disappears ( $I^2 = 0\%$ ) when removing Xue's study [16]. No obvious publication bias is observed (Egger test,  $p = 0.683$ ).

### Complication rates

Six investigations reported the complication rates after treatment [11, 12, 14–17]. The collective complication rates are similar between the 2 groups (5.9% vs. 7.0%;  $p = 0.60$ , Figure 2 J). No significant heterogeneity is found for this endpoint ( $I^2 = 0\%$ ). No marked publication bias is observed (Egger test,  $p = 0.733$ ).

### Time to ambulation

Four studies reported the data of time to ambulation [12, 13, 15, 17]. The pooled time to ambulation is significantly lower in the PE-TLIF group than in the MIS-TLIF group (MD = -0.40; 95% CI: -0.63–0.17;  $p = 0.0006$ , Figure 2 K). Significant heterogeneity is found for this endpoint ( $I^2 = 75\%$ ). The significant heterogeneity disappears ( $I^2 = 0\%$ ) when removing Kim's study [13]. No marked publication bias is observed (Egger test,  $p = 0.665$ ).

### Hospital stay

Five studies reported the data of time to ambulation [11, 13, 14, 16, 17]. The pooled hospital stay time is significantly lower in the PE-TLIF group than in the MIS-TLIF group (MD = -3.03; 95% CI: -4.77–

1.29;  $p = 0.0007$ , Figure 2 L). Significant heterogeneity is found for this endpoint ( $I^2 = 96\%$ ). Sensitivity analysis suggests that removing individual studies has no impact on the detected heterogeneity. No marked publication bias is observed (Egger test,  $p = 0.400$ ).

## Discussion

This meta-analysis compared the clinical efficacy between PE-TLIF and MIS-TLIF for patients with LDD. The purpose of PE-TLIF and MIS-TLIF treatments for LDD is relieving the pain and enhancing quality of life. All the included studies found that VAS-leg, VAS-back, and ODI could be significantly improved by both PE-TLIF and MIS-TLIF. Our meta-analysis reveals that the degree of improvement of VAS-leg and ODI is comparable between PE-TLIF and MIS-TLIF. Furthermore, the need for a postoperative analgesic is also comparable between the 2 groups. These findings may indicate that both PE-TLIF and MIS-TLIF can achieve appropriate clinical effectiveness in patients with LDD. However, the PE-TLIF may achieve a faster recovery of back pain than MIS-TLIF does. This finding may be attributed to the fact that PE-TLIF intervention is associated with reduced surgical trauma to surrounding soft and bone tissues, compared to the MIS-TLIF intervention [16].

Fusion rate is also an important endpoint in this meta-analysis. We found that the collective fusion rate of PE-TLIF is comparable to MIS-TLIF, at approximately 90%. Hence, both PE-TLIF and MIS-TLIF achieve suitable interbody fusion. However, Ao *et al.* [11] found that MIS-TLIF can achieve a significantly higher fusion rate than PE-TLIF. This finding may suggest inadequate bone graft and expandable cage application [10]. Nonetheless, the results from the retrospective study should be further proven by RCTs.

The pooled operation and intra-operative fluoroscopic time are both significantly higher in the PE-TLIF group. The curette-mediated endplate preparation in PE-TLIF took a considerable amount of time. Since it relies on instrument palpations, without direct vision, and with sequential endoscopic guidance, the endplate preparation required more time than MIS-TLIF [20–22].

Even though PE-TLIF is usually time consuming, it can result in significantly less intra-operative blood loss and better post-operative recovery. These re-



sults may be attributed to the shorter length of incision during PE-TLIF.

This meta-analysis shows comparable pooled complication rates between the 2 groups. Furthermore, the pooled complication rates after PE-TLIF and MIS-TLIF are only 5.9% and 7.0%, respectively. These findings indicate that both PE-TLIF and MIS-TLIF are safe for patients with LDD. If some patients experience the severe complication that is an abscess, external drainage is an effective method to treat it [23].

A similar meta-analysis reported insights into unilateral bi-portal endoscopy (UBE) in the treatment of lumbar diseases [24]. The researchers found that UBE surgery could significantly relieve the VAS scores of leg and back [24]. However, that meta-analysis only included single-arm studies [24]. Therefore, we could not find results about the comparison of UBE with other techniques. Our meta-analysis contained 2 commonly used techniques, and this is an advantage over the previous meta-analysis.

Our meta-analysis faced certain limitations. First, our analysis did not include any RCT. Furthermore, the multiple types of diseases and different disk ranges also increase the risk of bias. Therefore, further RCTs should be conducted. Second, there was a lack of homogeneity in surgical interventions and number of channels, which may influence our results. Many endpoints in this meta-analysis demonstrate significant heterogeneity. Third, there is a lack of cost-effective analysis among the included studies. Fourth, all eligible investigations were based in Asia. Therefore, additional comprehensive, global studies are warranted once stent usage becomes more prevalent worldwide.

## Conclusions

Both PE-TLIF and MIS-TLIF are safe and effective strategies for patients with LDD. When compared, although MIS-TLIF results in reduced surgical duration and intra-operative blood loss, rapid post-operative recovery can be achieved by PE-TLIF.

## Acknowledgments

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## Conflict of interest

The authors declare no conflict of interest.

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