# Uniportal versus three-port video-assisted thoracoscopic surgery for spontaneous pneumothorax: a meta-analysis

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**Background:** Whether or not uniportal video-assisted thoracoscopic surgery (VATS) is beneficial for spontaneous pneumothorax remains inconclusive. This meta-analysis aimed to summarize the available evidence to assess the feasibility and advantages of uniportal VATS for the treatment of spontaneous pneumothorax compared with three-port VATS.

**Methods:** Eligible publications were identified by searching the Cochrane Library, PubMed, EMBASE, Web of Science, China National Knowledge Infrastructure (CNKI), Wanfang Data databases and CQVIP. Odds ratios (OR) and standardized mean differences (SMD) with 95% confidence intervals (CI) were calculated to compare dichotomous and continuous variables, respectively.

**Results:** This meta-analysis was based on 17 studies and included a total of 988 patients with spontaneous pneumothorax. No death was reported during the perioperative period. Compared with three-port VATS groups, there was a statistically significant difference in uniportal VATS groups regarding postoperative hospital stay (SMD= -0.58; 95% CI: -1.04 to -0.12; P=0.01), paresthesia (OR=0.13; 95% CI: 0.07 to 0.24; P<0.00001), visual analogue pain score (VAS) at 24 hours (h) (SMD= -0.87; 95% CI: -1.07 to -0.68; P<0.00001), VAS at 72 h (SMD= -0.49; 95% CI: -0.68 to -0.30; P<0.00001), and patients satisfaction scale (PSS) at 24 h (SMD= -0.81; 95% CI: -1.21 to -0.41; P<0.0001), PSS at 48 h (SMD= -0.69; 95% CI: -1.08 to -0.29; P=0.0007). However there was no statistically significant difference on the recurrence (OR=0.79; 95% CI: 0.42 to 1.46; P=0.45), operative time (SMD= -0.23; 95% CI: -0.21 to 0.67; P=0.31), length of postoperative drainage (SMD= -0.17; 95% CI: -0.40 to -0.07; P=0.16), VAS at 48 h (SMD= -0.40; 95% CI: -1.47 to 0.67; P=0.46), and PSS at 72 h (SMD= -0.13; 95% CI: -0.52 to -0.25; P=0.50).

**Conclusions:** The results for mortality, recurrence, operative time, and length of postoperative drainage were similar between uniportal and three-port VATS. Uniportal VATS resulted in reduction in postoperative pain and paresthesia as well as an improvement in patients' satisfaction. This meta-analysis indicated that using uniportal VATS to treat spontaneous pneumothorax was safe and feasible, and it may be a better alternative procedure because of its advantage in reducing postoperative pain and paresthesia.

**Keywords:** Uniportal; three-port surgery; video-assisted thoracoscopic surgery (VATS); spontaneous pneumothorax; meta-analysis

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#### Introduction

Pneumothorax has been described as a situation in which air is inserted into the pleural space. Open thoracotomy and video-assisted thoracoscopic surgery (VATS) are the main surgical treatments for pneumothorax. Open thoracotomy with pleurectomy was once used as the gold standard technique as recent as 2003 (1). The treatment of spontaneous pneumothorax has been developed from open thoracotomy to VATS over recent decades. Because of the lesser invasion, compared with open thoracotomy, conventional VATS has been clearly shown to offer greater advantages in regard to postoperative stay, operative time, length of postoperative drainage and postoperative pain (2-4). However the recurrence and residual neurological symptoms (including paresthesia, postoperative pain) still frequently occurred in this minimally invasive surgery (5-8). The main reason for postoperative pain may be injury to the intercostal nerves, as open thoracotomy usually requires one large incision or costal resection, and conventional VATS usually requires one small incision for the scope and multiple small incisions, to complete the procedure of dissection, resection, and grasping.

Since single-incision thoracoscopic surgery (SITS) for pulmonary wedge resection to treat pneumothorax was first reported by Yamamoto et al. in 1998 (9), sympathectomy, pulmonary resection, bronchial sleeve lobectomy, and bronchial reconstructive surgery have been performed via uniportal VATS (10-14). This offers another treatment option for spontaneous pneumothorax. With the application of uniportal VATS for spontaneous pneumothorax in a retrospective study, Huang et al. reported that the uniportal VATS group had a better outcomes in terms of operative time, intraoperative blood loss, and postoperative pain, but worse outcomes in regard to chest tube drainage time, postoperative length of hospital stay, postoperative leakage and recurrence than three-port VATS group (15). Uniportal VATS appears to have several disadvantages, such as limitation of the visual field, difficulty of use of VATS instruments, and ergonomic discomfort. The safety and feasibility of uniportal VATS seems to be controversial. Jutley et al. reported that uniportal VATS had an advantage in the reduction of postoperative pain and paresthesia compared to three-port VATS (16). However this was not reported by Yang et al. (17). These studies yielded different results. We performed the first meta-analysis to combine these comparative outcomes during the perioperative period, and investigate the safety, feasibility and advantages of uniportal VATS.

#### **Materials and methods**

#### Literature search strategy

A research for relevant publications was conducted in the Cochrane Library, PubMed, EMBASE, Web of Science, China National Knowledge Infrastructure (CNKI), Wanfang Data databases, and CQVIP, from their date of publication to April 2015. The search strategy used "uniportal video-assisted thoracoscopic surgery", "singleport video-assisted thoracoscopic surgery", "three-port video-assisted thoracoscopic surgery", "spontaneous pneumothorax", and "pneumothorax" as key words or MeSH terms. We also collected the relevant references of all retrieved articles and conducted further searches.

#### Criteria for inclusion and exclusion

Eligible studies included the following criteria: (I) the studies evaluated the effect of uniportal VATS versus three-port VATS; (II) human-based studies with spontaneous pneumothorax; (III) studies were published in English and Chinese; (IV) studies were limited to RCT, case-control or cohort studies; (V) date on one of the outcomes [mortality, complications, recurrence, postoperative stay, operative time, length of postoperative drainage, paresthesia, visual analogue pain score (VAS) and patients satisfaction scale (PSS)] was provided. The exclusion criteria were as follows: (I) studies published in a language other than English or Chinese; (II) nonhuman-bases studies, studies without controllable patients, overlapping studies, case reports, letters, reviews or meta-analyses; (III) studies in which the necessary data were not provided.

#### Data extraction

Two investigators (Shi-Lei Qin, Yan-Long Yang) independently performed the assessment and data extraction using the author, year of publication, number of patients, age, gender, affected side, mortality, complications, recurrence, operative time, postoperative stay, length of postoperative drainage, paresthesia, VAS, PSS, and duration of follow-up. We contacted the authors if these data were missing or incomplete in their publications. Disagreements were resolved by discussion and adjudication by a senior investigator (Lei Xian). The VAS was assessed according to Duncan (18).

#### Quality assessment

The quality of the included studies was independently

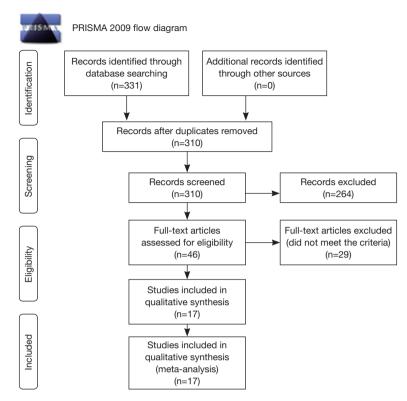


Figure 1 Flow diagram showing the procedure for selecting of studies.

assessed using the Newcastle-Ottawa scale (NOS) by two investigators (Shi-Lei Qin, Yan-Long Yang). Studies with five or more stars were defined as high quality studies. Disagreements were resolved by discussion.

### Statistical analysis

The effective values of dichotomous variables or continuous variables were estimated by odds ratios (OR) or standardized mean differences (SMD) with their 95% CIs respectively. The Chi-square and I<sup>2</sup> tests were used to evaluate statistical heterogeneity. The significance level of heterogeneity was fixed at P<0.10. When P>0.1 and I<sup>2</sup><50%, heterogeneity was classified as acceptable (no heterogeneity: I<sup>2</sup>=0%-25%; low heterogeneity: I<sup>2</sup>=25%-50%; moderate heterogeneity: I<sup>2</sup>=50%-75%; high heterogeneity: I<sup>2</sup>=75%-100%). It was then tested using a fixed effects model. If I<sup>2</sup>>50%, a random effects model was conducted. For the source of heterogeneity, a sensitivity analysis of each study was used for secondary analysis. Publication bias was assessed by Egger's and Begg's test using STATA version 12.0 (Stata Corporation, College Station, TX). All P values were

two-sided. A P value <0.05 was considered statistically significant. All statistical analysis was performed by Review Manager V.5.3 (The Cochrane Collaboration, Software Update, Oxford, UK) and STATA version 12.0.

### **Results**

### Study characteristics

The present work followed the guidelines for systematic reviews and meta-analyses (PRISMA Flow Diagram). In total, 331 studies were identified by searching seven electronic databases. After removing duplicates and screening abstracts, the remaining potentially eligible for inclusion were further assessed by a screen of the full texts. After the exclusion of 29 studies that did not meet our inclusion criteria, 17 studies (15-17,19-32) were included in the final analysis. The process of article selection is summarized in *Figure 1*. Four studies were published in Chinese (15,22,27,28), and 13 studies were published in English (16,17,20,21,23-26,29-32). *Table 1* shows the NOS quality scale and all eligible studies scored highly (with five stars or more). All 17 studies were retrospective case-

control studies and included a total of 988 patients, of which 502 (50.8%) patients underwent uniportal VATS and 486 (49.2%) patients underwent three-port VATS. The main characteristics of all of the studies included are shown in *Table 2*.

### Perioperative mortality and recurrence

Seven studies (15,17,21,26,29,30) with 493 patients reported that no death occurred in either the uniportal or threeport VATS groups. Fifteen studies (15-17,19-25,27,29-32) assessed the postoperative recurrence rate, with a combined total of 852 patients included in this meta-analysis. No significant heterogeneity was detected (P=0.57,  $I^2$ =0%). A Fixed effects model was selected. There was no significant difference between the uniportal and three-port VATS groups (OR=0.79; 95% CI: 0.42 to 1.46; P=0.45). However the recurrence rates of the uniportal VATS groups seemed to be lower than that of the three-port VATS groups (uniportal *vs.* three-port: 4.34% *vs.* 4.79%). The results are summarized in *Figure 2*.

#### **Operative time**

Twelve studies (15,17,21-23,26-32) included comparable data on operative time with a combined total of 800 patients included in this meta-analysis. The mean of operative times of the uniportal and three-port VATS groups were 61.29 vs. 59.04 min, respectively. High heterogeneity was detected between the uniportal and three-port VATS groups (P<0.00001; I<sup>2</sup>=88%). A random effects model was used, and there was no significant difference between the uniportal and three-port VATS groups (SMD =-0.23; 95% CI: -0.21 to 0.67; P=0.31). The results are summarized in *Figure 3*.

# Length of postoperative drainage

A total of 11 studies (15,16,20,22,23,26-29,31,32) included comparable data on length of postoperative drainage, with a combined total of 744 patients included in this metaanalysis. The mean lengths of postoperative drainage for the uniportal and three-port VATS groups were 3.29 and 3.44 days, respectively. Moderate heterogeneity was observed among these studies (P=0.01;  $I^2=55\%$ ). A random effects model was used, and no significant difference between the uniportal and three-port VATS groups was found (SMD =-0.17; 95% CI: -0.40 to -0.07; P=0.16). The results are summarized in *Figure 4*.

#### Postoperative hospital stay

Eight studies (15,21,23,28-32) yielded comparable data on postoperative hospital stay with a combined total of 635 patients included in this meta-analysis. The heterogeneity was significant (P<0.00001; I<sup>2</sup>=86%), and a random effects model was used. This meta-analysis suggested that the uniportal VATS groups were significantly associated with shorter postoperative hospital stay compared to the threeport VATS groups (SMD =-0.58; 95% CI: -1.04 to -0.12; P=0.01). However, the mean of lengths of postoperative hospital stay in the uniportal VATS groups were shorter by no more than one day than in the three-port VATS groups (5.71 vs. 5.84 days). Thus we still concluded that there was not great clinical difference between the two groups in terms of length of postoperative hospital stay. The results are summarized in *Figure 5*.

#### Paraesthesia

Six studies (16,17,25,26,28,30) included comparable data on paresthesia with a combined total of 261 patients included in this meta-analysis. The rates of paresthesia in the uniportal and three-port VATS groups were 19.04% and 58.77%, respectively. No significant heterogeneity (P=0.87; I<sup>2</sup>=0%) was seen in the fixed effects model. This meta-analysis suggested that the uniportal VATS groups were significantly associated with a lower rate of paresthesia compared to the three-port VATS groups (OR=0.13; 95% CI: 0.07 to 0.24; P<0.00001). The results are summarized in *Figure 6*.

#### Postoperative pain

The VAS at 24 h, 48 h and 72 h were assessed for postoperative pain in 6, 3 and 6 studies, respectively. The mean VAS at 24 h, 48 h, 72 h in the uniportal *vs.* three-port VATS groups were 3.57 *vs.* 4.49, 2.97 *vs.* 3.63, 2.41 *vs.* 2.91, respectively.

Heterogeneity was acceptable in terms of VATS at 24 h and 72 h (VAS at 24 h: P=0.57, I<sup>2</sup>=0%; VAS at 72 h: P=0.15, I<sup>2</sup>=39%). A fixed effects model was selected for a metaanalysis, which indicated that the uniportal VATS groups were significantly associated with lower VAS at 24 h and 72 h when compared to the three-port VATS groups (VAS at 24 h: SMD =-0.87; 95% CI: -1.07 to -0.68; P<0.00001; VAS at 72 h: SMD =-0.49; 95% CI: -0.68 to -0.30; P<0.00001). The results are showed in *Figures 7,8*.

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		Selection			Comparability	rability		Exposure		
					Comparability of	Comparability of cases		Somo mothood of		lotoT
	IS ITTLE CASE	Representativeness	Selection of Definition	Definition	cases and controls on and controls on the	and controls on the	Ascertainment		Non-response	lotal nointo
(autitor, year)		of the cases	controls	of controls	of controls the basis of the design basis of the design or	basis of the design or	of exposure		rate	builts
	adequate				or analysis (age)	analysis (others)		cases and controls		
Byun, 2013	*			*	*		*	*		5
Chen, 2012	*	*		*	*	*	*	*		7
Chen, 2011	*	*		*	*	*	*	*		7
Huang, 2014	*			*	*	*	*	*		9
Igai, 2014	*			*	*	*	*	*		9
Jutley, 2005	*			*	*	*	*	*		9
Kang, 2012	*			*	*	*	*			5
Kang, 2013	*			*	*	*	*	*		9
Kang, 2014	*			*	*	*	*	*		9
Li, 2014	*			*	*	*	*	*		9
Liang, 2013	*			*	*	*	*	*		9
Mao, 2014	*			*	*	*	*	*		9
Ocakcioglu, 2015	*			*	*	*	*	*		9
Salati, 2008	*	*		*	*		*	*		9
Song, 2015	*			*	*	*	*	*		9
Tsuboshima, 2015	*	*		*	*	*	*	*		7
Yang, 2013	*	*		*	*	*	*	*		7
RCT, randomized controlled trial	ntrolled trial.									

Viitbor 2000	No. of	Age (year)		Gender	er	Affected side	side	Diagnosed and		Follow-up (months)	(months)
Aurrior, year	patients (U/T)	T) U/T	P value	U:T (M/F)	P value	U:T (R/L)	P value	method		U/T	P value
Byun, 2013	12/12	N/A	N/A	N/A	N/A	N/A	N/A	Chest X-ray and CT	N/A	N/A	N/A
Chen, 2012	36/26	29.1/24.5	0.144	(27/9):(22/4)	0.359	(22/14):(13/13)	0.384	N/A	Mechanical	16.3/30.5	<0.001
Chen, 2011	10/20	20.50±5.54/29.32±12.5	0.246	(9/1):(19/1)	0.605	(3/17):(6/14)	-	Chest X-ray or CT	Mechanical	N/A	N/A
Huang, 2014	99/101	27.11±10.23/29.32±12.5	>0.05	(91/8):(92/9)	-	(55/44):(53/48)	0.673	Chest X-ray or CT	Chemical	6-8	N/A
lgai, 2014	31/72	23.8±6/23.4±6.2	0.72	(25/6):(69/3)	0.012	(11/20):(53/48)	0.22	Chest X-ray and CT	N/A	N/A	N/A
Jutley, 2005	16/19	28.9±15.4/32.1±10	0.46	(16/0):(14/5)	0.049	N/A	N/A	N/A	N/A	9.4/32.1	N/A
Kang, 2012	11/16	N/A	N/A	N/A	N/A	N/A	N/A	N/A	N/A	8.7±7.42	N/A
Kang, 2013	22/18	21.6/19.8	0.421	(19/3):(17/1)	0.397	(11/11):(6/12)	0.289	Chest X-ray	Sponge sticks or gauze peanuts 13.8/17.8	13.8/17.8	0.279
Kang, 2014	33/19	18/19	0.888	(27/6):(18/1)	0.189	(15/17):(6/13)	0.326	Chest CT	Sponge sticks or gauze peanuts 14.0/15.0	14.0/15.0	0.124
Li, 2014	9/16	19.6/21	N/A	(7/2):(14/2)	0.602	N/A	N/A	N/A	Mechanical	N/A	N/A
Liang, 2013	23/27	21.6/23.5	>0.05	(18/5):(20/7)	-	(13/10):(12/15)	0.571	Chest HRCT	Mechanical	6-24	N/A
Mao, 2014	21/22	23.1±3.7/22.7±2.9	>0.05	(19/2):(16/6)	0.24	(8/13):(11/11)	0.543	Chest X-ray and CT	Mechanical	N/A	N/A
Ocakcioglu, 2015	5 37/40	26.2/27.9	0.167	(32/5):(36/4)	0.257	(22/15):(23/17)	0.243	Chest X-ray and CT	Mechanical	21	N/A
Salati, 2008	28/23	24±6.3/26.4±6.4	N/A	N/A	N/A	N/A	N/A	Chest X-ray and CT	N/A	13/39	<0.0001
Song, 2015	37/23	22.0±6.3/22.0±6.3	0.36	(35/2):(23/0)	0.519	(16/21):(9/14)	0.148	Chest CT	Chemical	2.9/3.1	0.67
Tsuboshima, 2015	15 34/35	22.9±8.7/25.9±8.8	>0.05	(30/4):(30/5)	-	(15/19):(12/23)	0.465	Chest CT	N/A	6.7/19.9	<0.05
Yang, 2013	27/13	22.0±9.1/27.1±12.3	0.15	(20/7):(12/1)	0.24	(18/9):(10/3)	0.72	Chest CT	Chemical	3.5/4.6	0.13

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# Journal of Thoracic Disease, Vol 7, No 12 December 2015

#### Qin et al. Unportal vs. three-port VATS for spontaneous pneumothorax

	Experimen	tal	Contro	ol		Odds ratio	Odds ratio
Study or subgroup	Events	Total	Events	Total	Weight	M-H, fixed, 95% CI	M-H, fixed, 95% Cl
Byun 2013	0	12	0	12		Not estimable	
Chen, C. H. 2012	1	35	2	24	10.2%	0.32 [0.03, 3.79]	
Chen, P. R. 2012	2	8	0	20	1.0%	15.77 [0.67, 372.41]	
Huang 2014	3	96	4	97	17.1%	0.75 [0.16, 3.44]	
lgai 2014	0	31	0	72		Not estimable	
Jutley 2005	0	16	1	18	6.1%	0.35 [0.01, 9.31]	
Kang 2012	0	11	0	16		Not estimable	
Kang 2014	2	20	1	17	4.3%	1.78 [0.15, 21.51]	
Li 2014	0	9	0	16		Not estimable	
Liang 2013	0	23	0	27		Not estimable	
Ocakcioglu 2015	0	37	4	36	20.0%	0.10 [0.00, 1.86]	
Salati 2008	3	25	3	18	13.6%	0.68 [0.12, 3.84]	
Song 2015	2	35	0	23	2.5%	3.51 [0.16, 76.46]	
Tsuboshima 2015	4	30	5	30	19.3%	0.77 [0.19, 3.20]	
Yang 2013	1	26	1	12	5.8%	0.44 [0.03, 7.69]	
Total (95% CI)		414		438	100.0%	0.79 [0.42, 1.46]	•
Total events	18		21				
Heterogeneity: Chi <sup>2</sup> =7	.62, df =9 (P=	0.57); I <sup>2</sup> =	:0%				0.002 0.1 1 10 500
Test for overall effect: 2	Z =0.76 (P=0.	45)					
							Favours [experimental] Favours [control]

Figure 2 Forest plot of OR and its 95% CI of recurrence between uniportal and three-port VATS groups. OR, odds ratios; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

		erimenta			Control			Std. mean difference	Std. mean difference
Study or subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV. random, 95% CI	IV. random, 95% Cl
Chen, P. R. 2012	80.5	20.74	10	85.5	21.87	20	7.6%	-0.23 [-0.99, 0.54]	
Huang 2014	54.6	19.4	99	64.9	24.3	101	9.4%	-0.47 [-0.75, -0.19]	
lgai 2014	55.2	15.5	31	35.9	14	72	8.9%	1.32 [0.86, 1.78]	
Kang 2014	41.21	9.1	33	41.58	9.14	19	8.5%	-0.04 [-1.57, 0.12]	
Li 2014	40.2	18.1	9	52.4	15.3	16	7.2%	-0.04 [-0.59, 0.52]	
Liang 2013	51.63	19.57	23	52.42	21.07	27	8.5%	3.08 [2.17, 3.99]	
Mao 2014	108	6	21	78	12	22	6.9%		
Ocakcioglu 2015	61.7	13.1	37	63.8	22.8	40	8.9%	-0.11 [-0.56, 0.34]	
Salati 2008	72.3	31.8	28	68.7	25.5	23	8.5%	0.12 [-0.43, 0.67]	<b>_</b>
Song 2015	57.1	18.6	37	60.7	18.4	23	8.6%	–0.19 [–0.71, 0.33]	
Tsuboshima 2015	67.9	19.4	34	62.5	15.4	35	8.8%	0.31 [-0.17, 0.78]	+
Yang 2013	74.6	22.8	27	72.4	20.3	13	8.0%	0.10 [-0.56, 0.76]	
Total (95% CI)			389			411	100.0%	0.23 [-0.21, 0.67]	◆
Heterogeneity: Tau <sup>2</sup> =	0.52, Ch	i <sup>2</sup> =91.6	9, df=1	1 (P<0.0	0001); I	<sup>2</sup> =88%		-	
Test for overall effect:	Z =1.02	(P=0.31	)		,,				-2 -1 0 1 2
		. 0.01	,						Favours [experimental] Favours [control]

Figure 3 Forest plot of SMD and its 95% CI of operative time between uniportal and three-port VATS groups. SMD, standardized mean differences; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

However, the heterogeneity was unacceptable in terms of VATS at 48 h (P=0.0002;  $I^2$ =88%), and a random effects model was selected. There was no significant difference in the VAS at 48 h between the two VATS groups (SMD =-0.40; 95% CI: -1.47 to 0.67; P=0.46). After performing of sensitivity analysis, one study was excluded (29). Heterogeneity was not found (P=0.49;  $I^2$ =0%), and a fixed effects model was selected. However, a statistical difference

was not-found (SMD =0.13; 95% CI: -0.37 to 0.63; P=0.62). The results were summarized in *Figure 9*.

#### Patients satisfaction scale (PSS)

Only two studies (21,29) provided comparable data on PSS with a combined total of 107 patients included in this meta-analysis. Summary statistics found that the mean

#### Journal of Thoracic Disease, Vol 7, No 12 December 2015

	Expe	erimenta	al		Control			Std. mean difference	Std. mean difference
Study or subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV. random, 95% CI	IV. random, 95% CI
Chen, P. R. 2012	3	0.94	10	3.55	1.5	20	6.1%	-0.40 [-1.16, 0.37]	
Huang 2014	4.7	1.8	99	5.5	2.9	101	14.1%	-0.33 [-0.61, -0.05]	
lgai 2014	1.2	0.5	31	1	0.3	72	11.1%	0.54 [0.11, 0.96]	
Jutley 2005	4.6	2	16	3.	2.1	19	7.2%	-0.33 [-0.34, 1.00]	
Kang 2014	2.82	0.92	33	3.79	1.44	19	8.4%	-0.84 [-1.43, -0.25]	
Li 2014	3.8	1.8	9	4.5	1.6	16	5.6%	-0.40 [-1.23, 0.42]	
Liang 2013	3.93	1.69	23	4.52	1.98	27	8.8%	-0.31 [-0.87, 0.25]	
Mao 2014	1.92	0.48	21	2	0.45	22	8.2%	-0.17 [-0.77, 0.43]	
Ocakcioglu 2015	3	0.88	37	3.5	1.55	40	10.7%	-0.39 [-0.84, 0.06]	
Song 2015	3.6	2.3	37	4	1.1	23	9.4%	0.20 [-0.73, 0.32]	
Tsuboshima 2015	1.3	0.9	34	1.2	0.5	35	10.3%	0.14 [-0.34, 0.61]	
Total (95% CI)						394	100.0%	-0.17 [-0.40, 0.07]	•
Heterogeneity: Tau <sup>2</sup> =	0.08, Chi	<sup>2</sup> =22.3	0, df=1	0 (P<0.0	01); I <sup>2</sup> =5	55%		-	
Test for overall effect:	Z =1.40	(P=0.16	6)						-1 -0.5 0 0.5 1 Favours [experimental] Favours [control]

Figure 4 Forest plot of SMD and its 95% CI of length of postoperative drainage between uniportal and three-port VATS groups. SMD, standardized mean differences; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

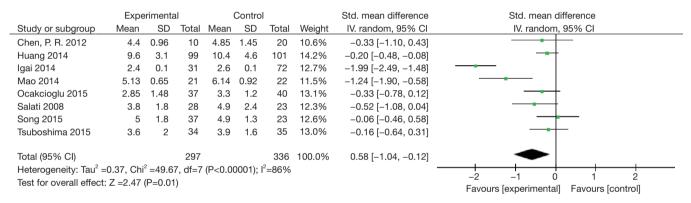


Figure 5 Forest plot of SMD and its 95% CI of postoperative hospital stay between uniportal and three-port VATS groups. SMD, standardized mean differences; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

	Experime	ental	Cont	rol		Odds ratio			Odds ratio		
Study or subgroup	Events	Total	Events	Total	Weight	M-H, fixed, 95% CI		M-H	, fixed, 95% (	2	
Jutley 2005	2	21	15	22	22.4%	0.05 [0.01, 0.27]	+	-			
Kang 2013	9	28	18	23	22.7%	0.13 [0.04, 0.47]			-		
Kang 2014	3	33	8	19	15.6%	0.14 [0.03, 0.61]	-		—		
Mao 2014	9	27	10	13	15.2%	0.15 [0.03, 0.68]		-			
Salati 2008	3	22	9	18	14.5%	0.16 [0.03, 0.73]			—		
Yang 2013	2	16	7	19	9.5%	0.24 [0.04, 1.41]					
Total (95% Cl)		147		114	100.0%	0.13 [0.07, 0.24]		•			
Total events	28		67								
Heterogeneity: Chi <sup>2</sup> =	1.84, df=5 (P	=0.87); I <sup>2</sup>	=0%								
Test for overall effect:	Z =6.47 (P<0	).00001)					0.02	0.1 Favours [experi	1 imental] Fav	10 ours [control]	50

Figure 6 Forest plot of OR and its 95% CI of paresthesia between uniportal and three-port VATS groups. OR, odds ratios; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

#### Qin et al. Unportal vs. three-port VATS for spontaneous pneumothorax

	Expe	erimenta	al		Contro	I		Std. mean difference		Std. mean diff	ierence	
Study or subgroup	Mean	SD	Total	Mean	SD	Total	Weight	V. fixed, 95% CI		IV. fixed, 95	% CI	
Chen, P. R. 2012	4.5	0.7	10	4.95	0.39	20	6.1%	-0.86 [-1.65, -0.06]				
Huang 2014	3.1	1.2	99	4.2	1	101	44.1%	-0.99 [-1.29, -0.70]				
Kang 2014	4.06	1.2	33	5	1.83	19	11.4%	-0.63 [-1.21, -0.06]				
Ocakcioglu 2015	3.42	0.94	37	4.23	0.92	40	17.4%	-0.86 [-1.33, -0.39]				
Song 2015	3.9	1.2	37	5.2	1.3	23	12.4%	–1.04 [–1.59, 0.48]				
Yang 2013	4.1	1.7	27	4.8	2.2	13	8.6%	-0.37 [-1.03, 0.30]			-	
Total (95% CI)			243			216	100.0%	-0.87 [-1.07, -0.68]		•		
Heterogeneity: Chi <sup>2</sup> =	=3.84, df=	5 (P=0.	57); l <sup>2</sup> =	0%					-2			<u> </u>
Test for overall effect	: Z =8.75	(P<0.00	0001)						-2	Favours [experimental]	Favours [con	itrol]

Figure 7 Forest plot of SMD and its 95% CI of VAS at 24 h between uniportal and three-port VATS groups. SMD, standardized mean differences; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

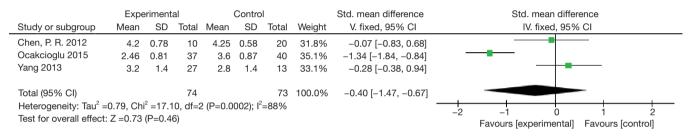


Figure 8 Forest plot of SMD and its 95% CI of VAS at 72 h between uniportal and three-port VATS groups. SMD, standardized mean differences; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

	Expe	rimenta	al		Control			Std. mean difference	Std. mean difference
Study or subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV. fixed, 95% CI	IV. fixed, 95% CI
Chen, P. R. 2012	3.3	0.48	10	3.55	0.6	20	6.1%	-0.43 [-1.20, 0.34]	
Huang 2014	2.5	1	99	2.9	1.3	101	46.1%	-0.34 [-0.48, -0.08]	
Liang 2013	1.91	0.59	23	2.29	0.61	27	11.0%	-0.62 [-1.19, -0.05]	
Ocakcioglu 2015	1.96	0.59	37	2.55	0.78	40	16.5%	-0.84 [-1.31, -0.37]	
Song 2015	2.5	1.5	37	3.9	1.8	23	12.1%	-0.85 [-1.40, -0.31]	
Yang 2013	2.7	1	27	2.6	1.1	13	8.2%	0.09 [-0.57, 0.76]	
Total (95% CI)			233			224	100.0%	-0.49 [-0.68, -0.30]	◆
Heterogeneity: Chi <sup>2</sup> =	8.15, df=	5 (P=0.	15); l <sup>2</sup> =	39%					
Test for overall effect:	: Z =5.04	(P<0.00	0001)						–1 –0.5 0 0.5 1 Favours [experimental] Favours [control]

Figure 9 Forest plot of SMD and its 95% CI of VAS at 48 h between uniportal and three-port VATS groups. SMD, standardized mean differences; CI, confidence interval; VATS, video-assisted thoracoscopic surgery.

of PSS at 24 h, 48 h and 72 h in the uniportal vs. threeport VATS groups was 1.90 vs. 2.55, 2.36 vs. 2.83, 2.33 vs. 2.44, respectively. There was no heterogeneity between the uniportal and three-port VATS groups (PSS at 24 h: P=0.96, I<sup>2</sup>=0%; at 48 h: P=0.73, I<sup>2</sup>=0%; at 72 h: P=0.91, I<sup>2</sup>=0%), so a fixed effects model was selected. This metaanalysis suggested that the uniportal VATS groups had a significantly lower PSS at 24 h and 48 h compared to the three-port VATS groups (at 24 h: SMD =-0.81; 95% CI: -1.21 to -0.41; P<0.0001; at 48 h: SMD =-0.69; 95% CI: -1.08 to -0.29; P=0.0007). However, there was no significant differ ence between the uniportal and three-port VATS groups in terms of PSS at 72 h (SMD= -0.13; 95% CI: -0.52 to -0.25; P=0.50). The results were summarized in *Table 3*.

#### Sensitivity analysis

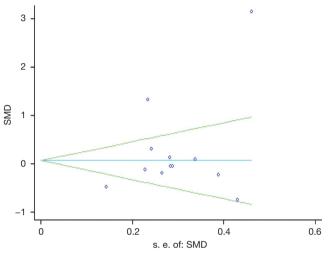
To evaluate the influence of single studies and analyze the effects of heterogeneity on the pooled SMDs of operative

#### Journal of Thoracic Disease, Vol 7, No 12 December 2015

	-		-				
Endpoint analysis	No. of studies	Mean	SMD (95% CI)	P value	I <sup>2</sup> <sub>het</sub>	P <sub>het</sub>	Model
At 24 h	2	1.90 <i>vs.</i> 2.55	-0.81 (-1.21, -0.41)	<0.0001	0%	0.96	Fixed
At 48 h	2	2.36 vs. 2.83	-0.69 (-1.08, -0.29)	0.0007	0%	0.73	Fixed
At 72 h	2	2.33 vs. 2.44	-0.13 (-0.52, 0.25)	0.50	0%	0.91	Fixed

Table 3 Summary of the PSS in the uniportal and three-port VATS groups

VATS, video-assisted thoracoscopic surgery; No, number; SMD, standardized mean differences; 95% CI, 95% confidence interval; I<sup>2</sup><sub>het</sub>, I<sup>2</sup> test for heterogeneity; P<sub>het</sub>, P for heterogeneity; h, hour; Fixed, fixed effect model.



Begg's funnel plot with pseudo 95% confidence limits

Figure 10 Funnel plots for the recurrence in studies included in the meta-analysis.

time, length of postoperative drainage and hospital stay, we performed a sensitivity analysis by estimating the average OR in the absence of each study.

For the operative time, two studies (23,28) influenced the comparable outcome in the sensitivity analysis of each study. Heterogeneity was acceptable (P=0.21,  $I^2$ =25%). Thus, a fixed effects model was selected. There was a significant difference between these two groups (SMD= -0.17; 95% CI: -0.32 to -0.01; P=0.04). In both studies excluded after sensitivity analysis, the operative time in the uniportal VATS groups was longer than that in the three-port VAST groups (uniportal vs. three-port: 1.8±0.1 h vs. 1.3±0.2 h, and  $55.2\pm15.5$  min vs.  $35.9\pm14.0$  min). The most likely reason for the high heterogeneity of operative time in one study was that the suture was used in the uniportal VATS groups, while endoscopic staplers were used in the three-port VATS groups (28). The cause of high heterogeneity in another excluded study may have been a change to multi-DOF forceps by operators and assistants.

Regarding length of postoperative drainage, after excluding two studies (23,28) based on sensitivity analysis, heterogeneity was non-existent among the remaining 6 studies (P=0.76,  $I^2$ =0%), and a significant difference was found (SMD= -0.23; 95% CI: -0.41 to -0.05; P=0.01).

For postoperative hospital stay, sensitivity analysis of each study showed that, there was no heterogeneity (P=0.32,  $I^2$ =14%) and a significantly difference was found (SMD= -027; 95% CI: -0.42 to -0.11; P=0.0009) between the uniportal and three-port VATS groups in summarized data from the 10 studies, except for one study (23).

#### **Publication bias**

Publication bias was assessed using Egger's and Begg's test in STATA version 12.0. No publication bias in the comparison of recurrence (Egger P=0.67; Begg P=0.72), operative time (Egger P=0.24; Begg P=0.84), or length of postoperative drainage (Egger P=0.72; Begg P=1.00) was detected. And Egger's and Begg's test for the other endpoints, which were evaluated by fewer than 10 studies, were not performed. The results are summarized in *Figures 10–12*.

#### Discussion

Uniportal VATS was described as inserting the video thoracoscope and two forceps through a single incision. Uniportal VATS was performed under general anesthesia using one-lung ventilation with the patient in lateral recumbency. The size of the skin incision should be associated with the size of the forceps, thoracoscope and trocar. Most publications reported that the skin incisions made was 1.5-3.0 cm in length through the  $4^{th}-7^{th}$ intercostal space on the mid-axillary, anterior or posterior axillary line. Most surgeons preferred a wound protector or a trocar to fix on the incision window. We suggested that a rubber wound protector should be a better choice (31), due to the provision of an adequate space for operating,

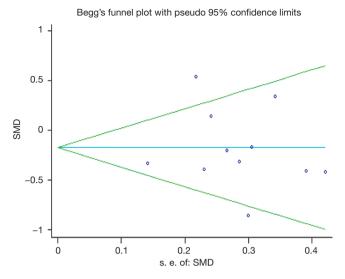
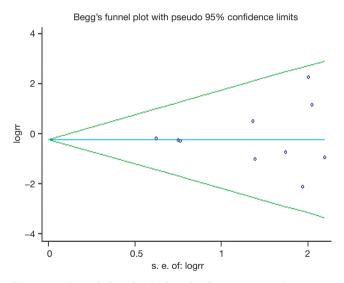


Figure 11 Funnel plots for the operative time in studies included in the meta-analysis.



**Figure 12** Funnel plots for the length of postoperative drainage in studies included in the meta-analysis.

protection of intercostal tissue and prevention of nerve entrapment. Next, a 5- or 10-mm 30° or 0° thoracoscope and straight shaft forceps or multi-DOF forceps were required to perform lobectomy. Multi-DOF forceps, which allowed the surgeon to manipulate bullae or blebs on a different axis, could diminish this interference, which was caused by inadequate space for operating through the same incision. After checking for air leak and bleeding, one drainage tube was inserted through the incision. We concluded form these studies that, uniportal VATS was performed better with a wound protector to fix on the incision window, a multi-DOF forceps to operate, and staplers for lobectomy during the surgical procedure.

We conducted this meta-analysis to evaluate the evidence comparing uniportal to three-port VATS for spontaneous pneumothorax and to assess the advantages of uniportal VATS. This meta-analysis evaluated the mortality, recurrence, complications, postoperative stay, operative time, length of postoperative drainage, paresthesia, VAS, and PSS of in uniportal and three-port VATS in patients with spontaneous pneumothorax using the best evidence available to date. Compared to the three-port VATS groups, there was no significant difference in the uniportal VATS groups in terms of mortality, recurrence, postoperative stay, operative time, and length of postoperative drainage. This meta-analysis indicated that the uniportal VATS groups were significantly associated with lower paresthesia, postoperative pain, and higher patients satisfaction in the short-term.

The mortality rate was 0% as reported in both groups. Complications were reported in only 13/320 (4%) patients (six patients in the uniportal VATS groups and seven patients in the three-port VATS groups) in six studies(17,20,21,29-31), including two patients with wound infection, nine patients with air leak, and two patients with pleural effusion. No serious complications occurred. This meta-analysis indicated that using the uniportal VATS technique did not caused more deaths, complications or recurrences during the perioperative period. This uniportal VATS technique was as clinically safe as the three-port VATS technique and did not lend to a worse outcome.

This meta-analysis showed that uniportal VATS did not increase operative time, or prolong postoperative drainage or postoperative hospital stay comparing with three-port VATS. Theoretically, a smaller incision makes operation more difficulty. Based on the surgeons' experiences, the operative time may be longer when they first begin using the technique. After adapting the technique and gaining more practice, the speed of the procedure accelerated quickly. For postoperative drainage, most studies reported bleeding and the flux of postoperative drainage could be ignored both in uniportal VATS groups and three-port VATS groups. This meant that uniportal VATS did not increase the risk of injury of to any vessels in the thoracic cavity, which may occur due to the difficulty of the uniportal VATS technique. These results showed that uniportal VATS was a safe and effective surgical approach for spontaneous pneumothorax, and did not pose an additional burden on

hospital resources.

Residual neurological complications in the uniportal VATS groups were considered significantly lower than in the three-port VATS groups. Compared to the threeport VATS groups, the uniportal VATS groups showed significantly lower in incidences of paresthesia. Patients in the uniportal VATS groups suffered from less postoperative pain at 24 h and 72 h, but equal pain at 48 h. There was no significant difference between the uniportal and threeport VATS groups in the analysis for VAS at 48 h. One reason may be the limited amount of data which came from only three studies with a total of 147 patients. As threeport incisions caused more injuries to the intercostal nerves than uniportal incisions did, more residual neurological symptoms, such as paresthesia, post-operative pain, even sexual dysfunction (16), can be unrelenting and difficult to treat with analgesics. The included studies reported that the patients in the three-port VATS groups required more postoperative epidural or intravenous patient-controlled analgesia (PCA) treatment than those in the uniportal VATS groups. Short-term outcomes showed that residual paresthesia and pain were lower and patients satisfaction was higher in the uniportal VATS groups. These advantages of uniportal VATS would lead to being the preferred choice of surgeons' and patients' choices. With its popularity and application, uniportal VATS may be an alternative technique to three-port VATS. However a scientific, randomize trial with a large sample size is needed to investigate long-term outcomes.

The most common patient complaints were paresthesia and wound pain, which may be caused by injury to the intercostal nerves when the operative approach was chosen-either VATS or open thoracotomy-penetrates the intercostal space. To prevent these problems, we summarized the following strategies based on surgeons' feedback: (I) single incision laparoscopic surgery (SILS) system combined with single-port technique was used as a novel method in uniportal VATS for treatment of pneumothorax (33). This SILS system, which was made of elastic elastomeric material, can diminish compression of the intercostal space. (II) A wound protector, which was used to fix on the incision window, may have a greater advantage in preventing postoperative pain than multi-incision trocar (31). (III) The 2-mm video-assisted endoscope, which was reported to be used in the two-port VATS procedure (34), is smaller than the 5-mm video-assisted thoracoscope, and could therefore be used with a single smaller incision,

reducing injury to the intercostal nerves. (IV) Pre-emptive wound infiltration with a local anesthetic, which was reported to reduce postoperative pain in needlescopic VATS procedures for the treatment of palmar hyperhidrosis in a randomized trial (35), may be applied in uniportal VATS procedure. Blocking of the central sensitization may reduce wound pain. (V) Another uniportal VATS procedure, known as the subxiphoid technique (36), may not cause injury to the intercostal nerves because the incision is made on the subxiphoid area. This subxiphoid uniportal VATS technique may be an innovation in the single-incision approach.

In this meta-analysis, only two studies reported a conversion from uniportal VATS to three-port VATS. Chen *et al.* (21) reported only that the conversion from uniportal VATS to three-port VATS for hemostasis is difficult, but the number of conversions is not. Igai *et al.* (23) reported that only one patient required conversion to a three-port approach because of severe adhesion. To overcome these problems for surgeons who are new to uniportal VATS, we suggest that more practice for surgeons may lesd to fewer injuries, and well-selected patients should be considered.

# Limitation

Despite having 17 case-control studies that compare uniportal VATS and three-port VATS, randomized controlled trials had not been conducted. Small sample size was another limitation in most studies. A multi-center study with large-sample-size has not yet been reported. For these reasons, heterogeneity existed in recurrence, operative time, length of postoperative drainage and length of postoperative hospital stay.

#### Conclusions

This paper is the first meta-analysis that compared uniportal with three-port VATS for spontaneous pneumothorax. The uniportal VATS technique neither increased mortality or recurrence rate nor prolonged the operative time, length of postoperative drainage or postoperative hospital stay. However, this new technique could reduce patients' postoperative pain and paresthesia, and improve patients' satisfaction. This meta-analysis indicates that the uniportal VATS is a safe, feasible and effective treatment for spontaneous pneumothorax. A reduction in paresthesia and post-operative pain were among the advantages of uniportal VATS.

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None.

2286

# Footnote

*Conflicts of Interest:* The authors have no conflicts of interest to declare.

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