#### **REVIEW ARTICLE**



# Pedicle screw fixation of thoracolumbar fractures: conventional short segment versus short segment with intermediate screws at the fracture level—a systematic review and meta-analysis

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#### Abstract

**Objective** Posterior short-segment fixation (4-screw construct = 4S) is the conventional surgical technique for thoracolumbar fractures. The effect of adding two intermediate pedicle screws at the fractured level (6-screw construct = 6S) is still a matter of debate. This review aims to compare the results between 4 and 6S pedicle screw fixation for thoracolumbar fractures. **Methods** A systematic review and meta-analysis were performed. The databases PubMed, Embase and Google Scholar were searched until January 2020. Inclusion criteria were studies comparing 4S and 6S techniques in patients with thoracolumbar fractures. Non-comparative studies and studies without full text were excluded. Cochrane risk of bias was assessed, and the GRADE approach was used to present the quality of evidence.

**Results** Twenty-seven studies, of which 21 randomized controlled trials, with a total of 1890 patients (940 with 4S and 950 with 6S) were included. Meta-analysis showed that the 6S technique resulted in significantly lower pain scores, better short-term and long-term Cobb angles, less loss of correction and less implant failures. However, longer operation time and more blood loss were seen with the 6S technique. Length of hospital stay, Oswestry Disability Index scores and infections did not differ significantly between the 6S and 4S techniques. Quality of the evidence according to GRADE was moderate to low. **Conclusion** In the treatment of thoracolumbar fractures, adding intermediate screws at the fracture level (6S) results in less post-operative pain, better radiological outcomes and less implant failure at the cost of a longer operation time and higher blood loss.

Keywords Thoracolumbar fracture · Spine · Pedicle screw · Fracture level · Intermediate screw

# Introduction

Thoracolumbar fractures are the most common spinal injuries [1]. Although treatment remains controversial in some fracture types, posterior fixation is the most frequently used surgical technique to restore vertebral body height, correct kyphotic deformity, restore spinal stability and indirectly decompress the spinal canal [2]. The conventional surgical technique of posterior fixation is a short-segment 4-screw construct with pedicle screws one level above and one level below the fractured vertebra. However, several studies showed loss of reduction after short-segment pedicle screw fixation and undesirable rates of implant failure, especially in fractures with loss of anterior support due to comminution [3–7].

For this reason, several modifications of the surgical technique have been proposed to improve outcomes. A biomechanical study in 1994 by Dick et al. first reported the posterior short-segment fixation combined with two additional screws at the fracture level (6 screw construct) [8]. Since this method was first introduced, the technique has evolved. A series of biomechanical studies showed that the addition of screws at the fracture level could significantly increase spinal stability, provide stronger fixation and decrease the stress in the pedicle screws in the non-fractured vertebra [9–13]. However, inconsistent effects of additional screws at fracture level were reported in several

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small clinical trials, limiting the evidence for this surgical technique [14–16]. Consequently, the surgical technique with intermediate screws is still not widely used.

Therefore, this systematic review and meta-analysis were performed on all available studies and data to compare the outcomes of conventional short-segment posterior pedicle screw fixation (4-screw construct = 4S) to short-segment posterior pedicle screw fixation combined with intermediate screws at the fracture level (6-screw construct = 6S) in patients with thoracolumbar fractures.

# Methods

This systematic review and meta-analysis were performed according to the guidelines of the preferred reporting items for systematic reviews and meta-analyses statement (PRISMA) [17].

### Literature search strategy

The databases PubMed, Embase, Cochrane and Google Scholar were searched for relevant articles to answer the stated research question. The final literature search was conducted on January 15, 2020. Additional relevant studies were also manually searched. The following key search terms were used: "thoracolumbar fracture", "intermediate screw", "additional screw", "fracture level" and "fractured vertebra". The complete search consisted of the mentioned key terms and its variants.

# Selection criteria and selection of articles

Studies were included if the following inclusion criteria were met: (1) patients with thoracolumbar fractures; (2) patients undergoing posterior short-segment pedicle screw fixation; and (3) studies comparing 4S and 6S techniques. Articles were excluded based on the following criteria: (1) full text not available; (2) incorrect publication type (case reports/systematic reviews); and (3) non-comparative study design. There was no language restriction.

Titles and abstracts identified by the literature search were screened for relevancy. Irrelevant abstracts were excluded. The full text of the remaining titles and abstracts was retrieved and was assessed for eligibility. After fulltext assessment, relevant articles were included in this systematic review and meta-analysis. In addition, the reference list of each included article was screened for new and/or missing studies. If the additionally identified articles met the inclusion criteria, they were included.

#### **Data extraction**

The following variables of interest were extracted from the included studies: demographics (name of first author, publication date, study design, sample size, mean age, gender, mean follow-up period), type and level of fracture, operation techniques (open or percutaneous surgical technique, fusion status) and the outcome parameters. The outcome parameters studied in this review included operation time, hospital length of stay (LOS), intraoperative blood loss, visual analog scale (VAS), Oswestry Disability Index (ODI), short-term and long-term post-operative Cobb angle, post-operative anterior vertebral body height (AVBH), correction loss of Cobb angle, correction loss of AVBH, rate of implant failure and post-operative infection.

#### **Definitions of outcome parameters**

The VAS is a measuring instrument to quantify pain severity, with higher score indicating greater pain intensity (scale 0-10). The ODI is a questionnaire on ten domains to quantify disability due to low back pain (100% = complete disability and 0% = no disability). The Cobb angle is defined as the angle between the superior end plate of the upper vertebra and the inferior end plate of the lower vertebra. The AVBH is defined as the anterior height of the fractured vertebra.

#### Quality of the evidence assessment

The Cochrane risk of bias tool was used to determine the quality of the included RCTs. The criteria set up by the Cochrane handbook for systematic reviews of interventions [18] were used. To assess the quality of the evidence for each outcome, the Grading of Recommendations Assessment, Development and Evaluation (GRADE) was used (GRA-DEpro GDT, McMaster University, 2015). Two reviewers independently assessed the quality of each included study and the quality of the evidence. Any discrepancies were resolved through a consensus discussion.

#### **Statistical analysis**

Statistical analysis for the meta-analysis was performed using Review Manager 5.3 [19]. The odds ratio (OR) and the standardized mean difference (SMD) were calculated for the dichotomous and continuous outcomes, respectively. Both the OR and MD were reported with the 95% confidence intervals (CIs). A *P* value lower than 0.05 was considered statistically significant. For the continuous outcome measures, a random effect model was used, because it cannot be assumed that all differences between effect sizes observed in our included studies are only due to sampling error [20]. For the dichotomous outcomes, the fixed effect model was used. A subgroup analysis was performed to determine difference in results for the open and percutaneous surgical technique.

# Results

## Literature search

The primary literature search showed 644 potential studies from the searched databases, and 20 additional records were identified through other sources. 620 abstracts were screened, and 77 full-text articles were evaluated after removal of irrelevant abstracts. After applying the exclusion criteria, 27 full-text articles were found to be eligible for inclusion in this systematic review and meta-analysis. See Fig. 1.

#### **Study characteristics**

The 27 selected studies included a total of 1890 patients, with 940 patients in the 4-screw construct group and 950 patients in the 6-screw construct group. Twenty studies were randomized controlled trials and 7 retrospective cohort studies. In three studies a percutaneous surgical technique was used and in 25 studies an open surgical technique.

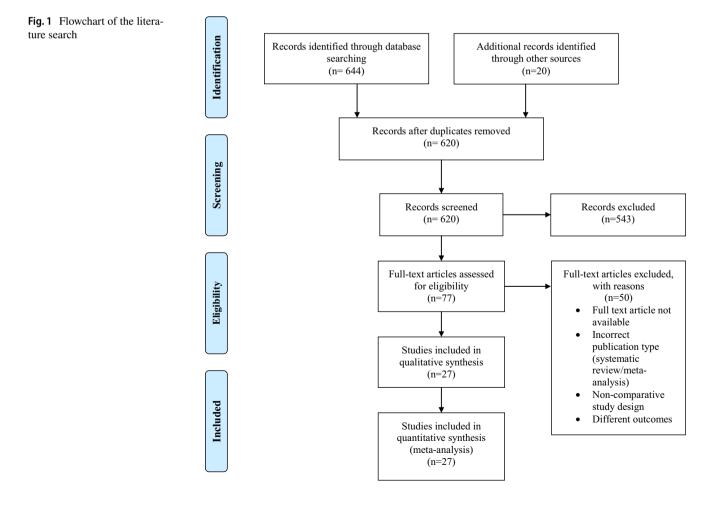
### Quality of the evidence assessment

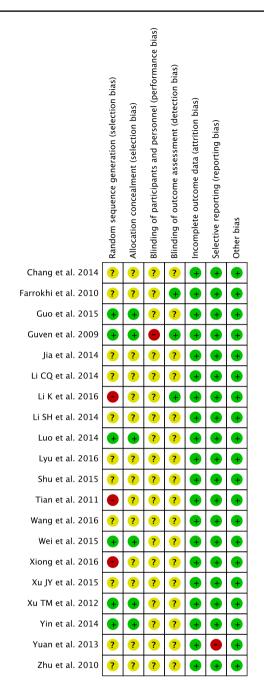
A risk of bias summary for the included RCTs is presented in Fig. 2. Quality rating of the evidence of each outcome according to GRADE is presented in Table 1.

#### **Outcome parameters**

#### **Operation time**

Fifteen studies [1, 14, 15, 21–32] provide information on the operation duration (928 patients). Combining these results shows that the placement of intermediate screws prolongs





**Fig. 2** Risk of bias summary: review of the authors' opinion on each risk of bias item for the included RCTs

the operation time significantly with a mean difference of 5.52 min (95% CI 0.79–10.26, P=0.02,  $I^2=87\%$ ) (Fig. 3).

#### Intraoperative blood loss

Sixteen studies [1, 14, 15, 21–33] including 988 patients described the amount of blood loss during surgery. The pooled analysis of these results showed a mean difference of 25.30 mL of intraoperative blood loss, favouring

the 4-screw construct significantly (95% CI 6.91–44.20,  $P < 0.01, I^2 = 96\%$ ) (Fig. 4).

#### Hospital length of stay

Information on hospital length of stay was provided in four studies [14, 15, 21, 26] (255 patients). The duration of hospital length of stay ranged from 8.7–13 days in the 6-screw group and 9–12.2 days in the 4-screw construct group. The mean difference regarding the hospital LOS was not significant (MD 0.44, 95% CI -0.68 to 1.57, P = 0.44,  $I^2 = 50\%$ ) (Fig. 5).

#### Visual analog scale

Information on the VAS is provided in nine of the included studies [1, 14, 21–23, 27, 31, 34, 35] describing 501 patients. The VAS in these studies was rated at follow-up at least three months after surgery. The pooled results showed that the 6-screw construct reduces the VAS significantly with a mean difference of 0.64 points (95% CI – 1.08 to – 0.19, P < 0.01,  $I^2 = 93\%$ ) (Fig. 6).

#### Oswestry disability index

Information about ODI was extracted from five studies including 273 patients [14, 22, 23, 26, 31]. The ODI was determined at follow-up at more than one-year post-operative. The pooled results show no significant difference between the 6-screw and 4-screw construct group (MD - 0.19, 95% CI - 1.52 to  $1.14, P = 0.78, l^2 = 41\%$ ) (Fig. 7).

#### Short-term post-operative Cobb angle

Fourteen studies [1, 15, 22–24, 27–29, 31, 32, 36–38] compared the short-term post-operative Cobb angle between the 4-screw and 6-screw construct group (713 patients). In these studies, the short-term post-operative Cobb angle was measured between 1 week and 1 month post-operative. Patients in the 6-screw construct group had significantly better shortterm post-operative Cobb angles with a mean difference of  $1.07^{\circ}$  (95% CI –1.82 to –0.32, P < 0.01,  $I^2 = 85\%$ ) (Fig. 8).

#### Long-term post-operative Cobb angle

Post-operative Cobb angle measured at follow-up after one year has been described in thirteen studies (766 patients) [1, 21–23, 26–28, 30, 31, 34, 36–38]. The 6-screw construct showed significantly better results regarding the long-term post-operative Cobb angle with a mean difference of  $3.69^{\circ}$  (95% CI – 5.20 to – 2,18, P < 0.01,  $\hat{I}^2 = 90\%$ ) (Fig. 9).

#### Table 1 Summary of evidence graded using the GRADE approach for the included RCTs

#### Summary of findings

#### The use of the 6-screw construct compared to the 4-screw construct for patients with thoracolumbar fractures

Population: Patients with thoracolumbar fractures

Intervention: 6-screw construct

Comparison: 4-screw construct

Outcomes	Anticipated absolute effect	s (95% CI)	Relative effect (95% CI)	Number of par-	Quality of	
	Risk with 4-screw con- struct	Risk with 6-screw con- struct		ticipants (studies)	the evidence (GRADE)	
Operation time assessed with (min)	The mean operation time was 92.8 min	MD 3.88 longer [0.89, 8.65]		748 (12 studies)	⊕⊕⊕O Moderate <sup>b</sup>	
Intraoperative blood loss assessed with (mL)	The mean intraop- erative blood loss was 280.3 mL	MD 28.29 higher [5.97, 50.61]		808 (13 studies)	$ \bigoplus \bigoplus \bigoplus \bigcirc \\ Moderate^{b} $	
Hospital length of stay assessed with (days)	The mean hospital length of stay was 9.6 days	MD 0.35 longer [- 1.36, 2.07]	_	186 (3 studies)	⊕⊕OO Low <sup>a,b</sup>	
VAS	The mean VAS was 2.77	MD 0.87 lower [- 1.46, - 0.28]	_	388 (7 studies)	$\underset{Low^{a,b}}{\bigoplus} {\bigcirc} OO$	
ODI	The mean ODI was 18.8	MD 0.97 lower [- 2.46, 0.53]	_	160 (3 studies)	$ \bigoplus \bigoplus \bigoplus \bigcirc \\ Moderate^{a} $	
Short-term post-operative Cobb angle in degrees	The mean short-term post-operative Cobb angle was 5.18 degrees	MD 1.02 lower [- 2.46, 0.53]	-	607 (11 studies)	$ \bigoplus \bigoplus \bigoplus \bigcirc \\ Moderate^b $	
Long-term post-operative Cobb angle in degrees	The mean long-term post- operative Cobb angle was 11.16 degrees	MD 3.76 lower [- 5.69, - 1.82]	-	692 (11 studies)	$ \bigoplus \bigoplus \bigoplus \bigcirc \\ Moderate^{b} $	
Correction loss of Cobb angle in degrees	The mean correction loss of Cobb angle was 4.22 degrees	MD 1.78 lower [- 3.4, - 0.15]	-	254 (5 studies)	$ \bigoplus \bigoplus \bigoplus \bigcirc \\ Moderate^b $	
Correction loss of AVBH in degrees	The mean correction loss of AVBH was 8.84 degrees	MD 6.01 lower [- 7.66, - 4.36]	-	566 (8 studies)	$ \bigoplus \bigoplus \bigoplus \bigcirc \\ Moderate^b $	
Implant failure	75 per 1000	13 per 1000 [6, 27]	OR 0.16 [0.07, 0.34]	1104 (16 studies)	$ \bigoplus_{\text{Moderate}^c} \bigcirc \bigcirc$	
Post-operative infection	23 per 1000	17 per 1000) [5, 57]	OR 0.74 [0.21, 2.55]	506 (9 studies)	⊕⊕OO Low <sup>a,c</sup>	

*CI* confidence interval, *OR* odds ratio, *MD* mean difference, *AVBH* anterior vertebral body height, *VAS* visual analogue scale, *ODI* oswestry disability index, *RCTs* randomized controlled trials

GRADE working group grades of evidence

High quality: We are very confident that the true effect lies close to that of the estimate of the effect

Moderate quality: We are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different

Low quality: Our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of the effect

Very low quality: We have very little confidence in the effect estimate: The true effect is likely to be substantially different from the estimate of effect

<sup>a</sup>Risk of bias: subjective outcome and no blinding of participants/study personnel, <sup>b</sup>Inconsistency: large variations in effect, large I<sup>2</sup>, <sup>c</sup>Imprecision: low rate of events, large confidence intervals. No downgrading for indirectness or publication bias

#### **Correction loss of Cobb angle**

Seven of the included studies [1, 15, 22, 23, 25, 32, 38] provide information on the correction loss of Cobb angle

in the 6-screw and 4-screw construct group (351 patients). The 6-screw method leads to a significantly lower correction loss of Cobb angle (MD – 1.95, 95% CI – 3.10 to – 0.79, P < 0.01,  $I^2 = 87\%$ ) (Fig. 10).

	6	screw		4	screw			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
4.12.1 RCTs									
Farrokhi 2010	169	59	38	168	72	42	2.1%	1.00 [-27.74, 29.74]	
Guven 2009	128	15	18	132	15	18	6.9%	-4.00 [-13.80, 5.80]	
Jia 2013	110	5.36	30	95	4.87	28	9.5%	15.00 [12.37, 17.63]	-
Li K 2016	62.4	11.8	16	54.3	10.7	16	7.7%	8.10 [0.29, 15.91]	
Li SH 2014	110.26	26.75	45	122.64	18.62	45	7.0%	-12.38 [-21.90, -2.86]	
Lyu 2016	72.1	12.5	30	64.4	9.8	30	8.6%	7.70 [2.02, 13.38]	
Tian 2011	52	11	27	51	10	35	8.7%	1.00 [-4.31, 6.31]	- <b>-</b>
Wei 2015	126	12.4	35	119	11.8	33	8.5%	7.00 [1.25, 12.75]	<b>_</b>
Xiong 2016	121.5	17.6	33	116.4	21.9	37	7.1%	5.10 [-4.17, 14.37]	
Xu TM 2012	70	6.5	48	67	5.6	54	9.5%	3.00 [0.63, 5.37]	
Yin 2014	127.3	33.4	20	130.2	33.2	20	3.4%	-2.90 [-23.54, 17.74]	
Yuan 2013	140.6	74.3	25	125.3	67.2	25	1.3%	15.30 [-23.97, 54.57]	· · · · · · · · · · · · · · · · · · ·
Subtotal (95% CI)			365			383	80.4%	3.88 [-0.89, 8.65]	◆
Heterogeneity: Tau <sup>2</sup> =	= 46.00; 0	Chi <sup>2</sup> = 7	3.50, d	f = 11 (P	< 0.00	001); l <sup>2</sup>	= 85%		
Test for overall effect	Z = 1.60	P = 0	.11)						
4.12.2 Retrospective	cohort s	tudies							
Sun 2016	150	18	35	136	11	34	8.0%	14.00 [6.98, 21.02]	
Ye 2017	142	30	20	160.2	45.7	24	3.0%	-18.20 [-40.72, 4.32]	
Zhao 2015	115.68	12.83	35	93.08	10.98	32	8.6%	22.60 [16.90, 28.30]	
Subtotal (95% CI)			90			90	19.6%	11.06 [-2.61, 24.73]	
Heterogeneity: Tau <sup>2</sup> =	= 111.03;	Chi <sup>2</sup> =	13.66,	df = 2 (P	= 0.00	1); $I^2 =$	85%		
Test for overall effect	: Z = 1.59	P = 0	.11)						
Total (95% CI)			455			473	100.0%	5.52 [0.79, 10.25]	◆
Heterogeneity: Tau <sup>2</sup> =	= 59.67: 0	$Chi^2 = 1$	09.38.	df = 14 (	P < 0.0	0001):	$ ^2 = 87\%$	_	
Test for overall effect	,					/,	••••		-20 -10 0 10 20
Test for subgroup dif						2			Favours 6 screw Favours 4 screw

Fig. 3 Forest plot comparing operation time for 6S versus 4S. RCTs randomized controlled trials, CI confidence interval, IV inverse variance, SD standard deviation

	6	screw		4	screw			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
4.10.1 RCTs									
Farrokhi 2010	430	305	38	515	485	42	1.0%	-85.00 [-260.84, 90.84]	←
Guven 2009	442	50	18	430	60	18	6.7%	12.00 [-24.08, 48.08]	
Jia 2013	650	18.64	30	550	17.68	28	8.6%	100.00 [90.65, 109.35]	-
Li K 2016	50.7	12	16	46.1	9	16	8.6%	4.60 [-2.75, 11.95]	-
Li SH 2014	231.21	87.55	45	260.54	90.54	45	6.6%	-29.33 [-66.13, 7.47]	— <b></b> +
Luo 2014	1,245	221	30	873	145	30	2.8%	372.00 [277.42, 466.58]	► <b>&gt;</b>
Lyu 2016	100.7	18.9	30	89.1	17.3	30	8.6%	11.60 [2.43, 20.77]	-
Tian 2011	250	65	27	245	62	35	7.0%	5.00 [-26.98, 36.98]	
Wei 2015	355.6	84.2	35	351.3	88.7	33	6.2%	4.30 [-36.86, 45.46]	
Xiong 2016	357.6	51.3	33	325.9	87.8	37	6.9%	31.70 [-1.57, 64.97]	
Xu TM 2012	130	4.8	48	120	7.5	54	8.7%	10.00 [7.58, 12.42]	•
Yin 2014	327.6	92.7	20	325.5	87.9	20	5.0%	2.10 [-53.89, 58.09]	
Yuan 2013	140.6	74.3	25	125.3	67.2	25	6.4%	15.30 [-23.97, 54.57]	
Subtotal (95% CI)			395			413	83.2%	28.29 [5.97, 50.61]	◆
Heterogeneity: Tau <sup>2</sup> :	= 1265.37	7; Chi <sup>2</sup> =	= 405.4	3, df = 1	2 (P < )	0.0000	1); I <sup>2</sup> = 97	%	
Test for overall effect	z = 2.48	8 (P = 0)	.01)						
4.10.2 Retrospective	e cohort s	tudies							
Sun 2016	425	85	35	400	55	34	6.9%	25.00 [-8.69, 58.69]	+
Ye 2017	483.5	186.6	20	507.5	300	24	1.4%	-24.00 [-169.24, 121.24]	
Zhao 2015	229.06	28.32	35	218.88	20.39	32	8.5%	10.18 [-1.56, 21.92]	
Subtotal (95% CI)			90			90	16.8%	11.58 [0.52, 22.64]	•
Heterogeneity: Tau <sup>2</sup> :	= 0.00; Cł	ni <sup>2</sup> = 0.3	89, df =	= 2 (P = 0	).64); I <sup>2</sup>	= 0%			
Test for overall effect	z = 2.05	5 (P = 0)	.04)						
Total (95% CI)			485			503	100.0%	25.30 [6.19, 44.42]	•
Heterogeneity: Tau <sup>2</sup>	= 1086.29	): Chi <sup>2</sup> =	= 406.5	7. df = 1	5 (P < )	0.0000	1): $ ^2 = 96$	~ · -	
Test for overall effect				.,	•		_,,. 50		-200 -100 0 100 200
Test for subgroup dif		- <b>.</b>		f = 1 (P =	= 0.19).	$1^2 = 42$	.2%		Favours 6 screw Favours 4 screw
i est ist subgroup un	i ei ei ee ei		1, J, u	. 10-	5.15),	12			

Fig. 4 Forest plot comparing intraoperative blood loss for 6S versus 4S. RCTs randomized controlled trials, CI confidence interval, IV inverse variance, SD standard deviation

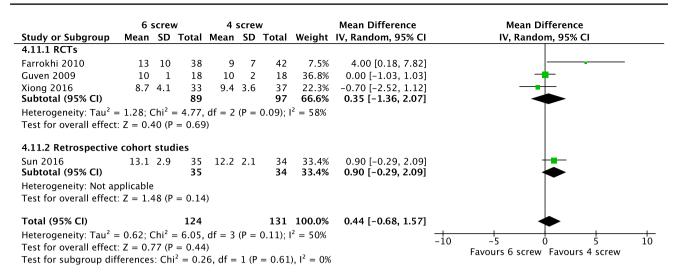


Fig. 5 Forest plot comparing hospital length of stay for 6S versus 4S. *RCTs* randomized controlled trials, *CI* confidence interval, *IV* inverse variance, *SD* standard deviation

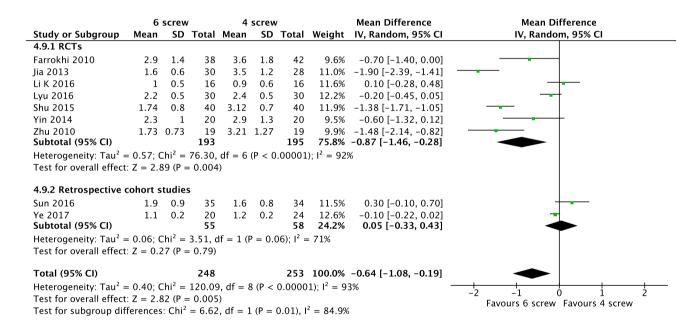


Fig. 6 Forest plot comparing visual analog scale for 6S versus 4S. RCTs randomized controlled trials, CI confidence interval, IV inverse variance, SD standard deviation

#### Correction loss of anterior vertebral body height

# Twelve studies [14, 22, 23, 25, 26, 32, 33, 38–42] describe values for the correction loss of anterior vertebral height of the 4- and 6-screw construct groups (862 patients). A meta-analysis of these values shows that the 6-screw construct leads to a significantly lower correction loss of the AVBH than the 4-screw method with a mean difference of 4.36 mm (95% CI – 6.56 to – 2.16, P < 0.01, $I^2 = 98\%$ ) (Fig. 11).

#### Implant failure

Twenty studies [1, 14, 15, 21–30, 32–34, 39, 40, 42, 43] describe rates of post-operative implant failure (1514 patients). The pooled results show that a significantly lower rate of implant failure was seen in the 6-screw construct group with an OR of 0.26 (95% CI 0.15–0.47, P < 0.01,  $l^2 = 0\%$ ) (Fig. 12).

	6 screw			4	screw	,		Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
4.8.1 RCTs									
Jia 2013	19.8	3.6	30	21.2	3.1	28	28.1%	-1.40 [-3.13, 0.33]	
Li K 2016	10	4.7	16	9.4	4.6	16	12.9%	0.60 [-2.62, 3.82]	
Xiong 2016 <b>Subtotal (95% CI)</b>	24.4	17.6	33 <b>79</b>	25.8	17.6	37 <b>81</b>		-1.40 [-9.66, 6.86] - <b>0.97 [-2.46, 0.53]</b>	
Heterogeneity: Tau <sup>2</sup> =	= 0.00; 0	Chi <sup>2</sup> =	1.16, c	df = 2 (	P = 0.5	56); I <sup>2</sup> =	= 0%		
Test for overall effect	Z = 1.2	27 (P =	= 0.20)						
4.8.2 Retrospective	cohort s	tudie	s						
Sun 2016	16.7	1.46	35	15.5	3.8	34	34.0%	1.20 [-0.17, 2.57]	+ <b>e</b> -
Ye 2017 <b>Subtotal (95% CI)</b>	12.9	3.5	20 55	14	3.7	24 <b>58</b>	22.5% <b>56.5%</b>	,	
Heterogeneity: Tau <sup>2</sup> =	= 1.81; C	Chi <sup>2</sup> =	3.17, c	df = 1 (	P = 0.0	)7); I <sup>2</sup> =	= 68%		Ī
Test for overall effect	: Z = 0.1	l8 (P =	= 0.86)						
Total (95% CI)			134			139	100.0%	-0.19 [-1.52, 1.14]	•
Heterogeneity: Tau <sup>2</sup> = Test for overall effect Test for subgroup dif	Z = 0.2	28 (P =	= 0.78)	,					-10 -5 0 5 10 Favours 6 screw Favours 4 screw

Fig. 7 Forest plot comparing Oswestry Disability Index for 6S versus 4S. RCTs randomized controlled trials, Cl confidence interval, IV inverse variance, SD standard deviation

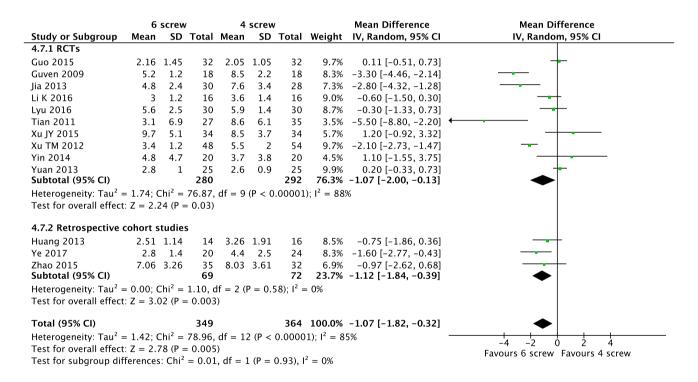


Fig. 8 Forest plot comparing short-term post-operative Cobb angle for 6S versus 4S. *RCTs* randomized controlled trials, *CI* confidence interval, *IV* inverse variance, *SD* standard deviation

#### Post-operative infection

Information on post-operative infections is provided in twelve studies [1, 14, 15, 21–27, 31, 43] (765 patients). Fixed-effect analysis showed no significant difference between the groups concerning infections post-operatively (OR 0.66, 95% CI 0.21 to 2.07, P = 0.47,  $I^2 = 0\%$ ) (Fig. 13).

#### Subgroup analysis

Three of the included studies used a percutaneous technique for the pedicle screw fixation and 24 studies applied an open surgical technique, including 226 and 1654 patients, respectively. A subgroup analysis was performed to analyse the outcomes for the open and percutaneous surgical technique separately. For the open surgical technique, the pooled

	6	screw			screw			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
4.6.1 RCTs									
Farrokhi 2010	14	8	38	19	10	42	5.8%	-5.00 [-8.95, -1.05]	
Guo 2015	3.57	1.94	32	5.64	2.06	32	9.2%	-2.07 [-3.05, -1.09]	
Jia 2013	6.7	3.1	30	12.1	6.5	28	7.4%	-5.40 [-8.05, -2.75]	
Li K 2016	5.5	2.1	16	6.4	1.9	16	8.8%	-0.90 [-2.29, 0.49]	+
Li SH 2014	8.35	5.08	45	9.67	6.44	45	7.7%	-1.32 [-3.72, 1.08]	
Lyu 2016	8.6	2.1	30	9.6	2.2	30	9.1%	-1.00 [-2.09, 0.09]	
Shu 2015	10.4	6.7	40	20.8	8.3	40	6.6%	-10.40 [-13.71, -7.09]	
Xiong 2016	5.8	3.4	33	10.7	9.2	37	6.7%	-4.90 [-8.08, -1.72]	
Xu JY 2015	9.8	4.5	34	9.7	2.9	34	8.4%	0.10 [-1.70, 1.90]	_ <b>_</b>
Yin 2014	8.3	7.7	20	9.8	6.1	20	5.4%	-1.50 [-5.81, 2.81]	
Yuan 2013	2.7	2	25	13.1	4.7	25	8.2%	-10.40 [-12.40, -8.40]	_ <b>.</b>
Subtotal (95% CI)			343			349	83.2%	-3.76 [-5.69, -1.82]	◆
Heterogeneity: Tau <sup>2</sup>	= 9.04; (	Chi <sup>2</sup> =	112.35	, df = 1	LO (P <	0.000	01); $I^2 = 9$	91%	
Test for overall effect	C: Z = 3.8	80 (P =	= 0.000	1)					
4 C 2 Detre en estive	cohort s	tudie	2						
4.6.2 Retrospective			,						
Huang 2013		1.25	, 14	5.12	1.07	16	9.3%	-2.61 [-3.45, -1.77]	
•		1.25	14 20	5.12 9.1	1.07 6	16 24	9.3% 7.6%	-2.61 [-3.45, -1.77] -4.80 [-7.29, -2.31]	
Huang 2013	2.51	1.25	14					. , .	
Huang 2013 Ye 2017	2.51 4.3	1.25 1.5	14 20 <b>34</b>	9.1	6	24 <b>40</b>	7.6% <b>16.8%</b>	-4.80 [-7.29, -2.31]	
Huang 2013 Ye 2017 <b>Subtotal (95% CI)</b>	2.51 4.3 = 1.50; 0	1.25 1.5 Chi <sup>2</sup> =	14 20 <b>34</b> 2.67, d	9.1 f = 1 (F	6	24 <b>40</b>	7.6% <b>16.8%</b>	-4.80 [-7.29, -2.31]	
Huang 2013 Ye 2017 <b>Subtotal (95% CI)</b> Heterogeneity: Tau <sup>2</sup> Test for overall effect	2.51 4.3 = 1.50; 0	1.25 1.5 Chi <sup>2</sup> =	14 20 <b>34</b> 2.67, d	9.1 f = 1 (F	6	24 <b>40</b> .0); I <sup>2</sup> =	7.6% <b>16.8%</b> 63%	-4.80 [-7.29, -2.31] -3.38 [-5.43, -1.33]	
Huang 2013 Ye 2017 <b>Subtotal (95% CI)</b> Heterogeneity: Tau <sup>2</sup>	2.51 4.3 = 1.50; 0	1.25 1.5 Chi <sup>2</sup> =	14 20 <b>34</b> 2.67, d	9.1 f = 1 (F	6	24 <b>40</b> .0); I <sup>2</sup> =	7.6% <b>16.8%</b>	-4.80 [-7.29, -2.31]	
Huang 2013 Ye 2017 <b>Subtotal (95% CI)</b> Heterogeneity: Tau <sup>2</sup> Test for overall effect	2.51 4.3 = 1.50; 0 t: Z = 3.2	1.25 1.5 Chi <sup>2</sup> = 23 (P =	14 20 <b>34</b> 2.67, d = 0.001 <b>377</b>	9.1  f = 1 (F )	6 P = 0.1	24 <b>40</b> .0); I <sup>2</sup> = <b>389</b>	7.6% 16.8% 63% 100.0%	-4.80 [-7.29, -2.31] -3.38 [-5.43, -1.33] -3.69 [-5.20, -2.18]	
Huang 2013 Ye 2017 <b>Subtotal (95% CI)</b> Heterogeneity: Tau <sup>2</sup> Test for overall effect <b>Total (95% CI)</b>	2.51 4.3 = 1.50; ( t: Z = 3.2 = 6.23; (	1.25 1.5 Chi <sup>2</sup> = 23 (P = Chi <sup>2</sup> =	14 20 <b>34</b> 2.67, c = 0.001 <b>377</b> 115.58	9.1  f = 1 (F ) ;, df = 1	6 P = 0.1	24 <b>40</b> .0); I <sup>2</sup> = <b>389</b>	7.6% 16.8% 63% 100.0%	-4.80 [-7.29, -2.31] -3.38 [-5.43, -1.33] -3.69 [-5.20, -2.18]	-10 -5 0 5 10 Favours 6 screw Favours 4 screw

Fig. 9 Forest plot comparing long-term post-operative Cobb angle for 6S versus 4S. *RCTs* randomized controlled trials, *CI* confidence interval, *IV* inverse variance, *SD* standard deviation

	6	screw	,	4	screw	1		Mean Difference	Mean Difference		
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI		
4.4.1 RCTs											
Guven 2009	2.8	3.2	18	3.7	3.1	18	11.1%	-0.90 [-2.96, 1.16]			
Jia 2013	2.7	1.6	30	6.3	4.2	28	12.7%	-3.60 [-5.26, -1.94]	<b>_</b>		
Li K 2016	2.6	1.5	16	2.7	1.2	16	15.4%	-0.10 [-1.04, 0.84]			
Lyu 2016	2.9	1.4	30	3.7	2.2	30	15.4%	-0.80 [-1.73, 0.13]			
Wei 2015 <b>Subtotal (95% CI)</b>	1.38	0.95	35 <b>129</b>	4.86	1.75	33 125		-3.48 [-4.15, -2.81] -1.78 [-3.40, -0.15]			
Heterogeneity: Tau <sup>2</sup> = Test for overall effect <b>4.4.2 Retrospective</b>	: Z = 2.1	L4 (P =	= 0.03)	dt = 4	(۲ < 0	.00001	); I <sup>2</sup> = 919	%			
Huang 2013	0	1.2	14	1.86	1.66	16	15.1%	-1.86 [-2.89, -0.83]	_ <b>_</b>		
Zhao 2015 Subtotal (95% CI)	2.63	1.95	35 <b>49</b>	5.51	3.28	32 <b>48</b>		-2.88 [-4.19, -1.57] -2.29 [-3.27, -1.30]	<b>→</b>		
5 /	Heterogeneity: $Tau^2 = 0.16$ ; $Chi^2 = 1.45$ , $df = 1$ (P = 0.23); $I^2 = 31\%$ Test for overall effect: Z = 4.54 (P < 0.00001)										
Total (95% CI)			178					-1.95 [-3.10, -0.79]	•		
Heterogeneity: Tau <sup>2</sup> = Test for overall effect Test for subgroup dif	: Z = 3.3	30 (P =	-4 -2 0 2 4 Favours 6 screw Favours 4 screw								

Fig. 10 Forest plot comparing correction loss of Cobb angle for 6S versus 4S. RCTs randomized controlled trials, CI confidence interval, IV inverse variance, SD standard deviation

data on the VAS, short-term and long-term post-operative Cobb angle, correction loss of Cobb angle and AVBH and implant failure all showed significant results favouring the 6-screw construct group. A significantly higher blood loss, but no significantly longer operation time was seen in the 6-screw construct with the open technique. For the percutaneous technique, a significantly longer operation time and significantly better results of long-term Cobb angle and correction loss of AVBH were seen in the 6-screw group compared to the 4-screw group. For the percutaneous technique, the 6-screw construct did not result in an increase in blood loss compared to the 4-screw construct. For the outcomes ODI and post-operative infection, the pooled analysis showed no significant difference between the 6-screw and

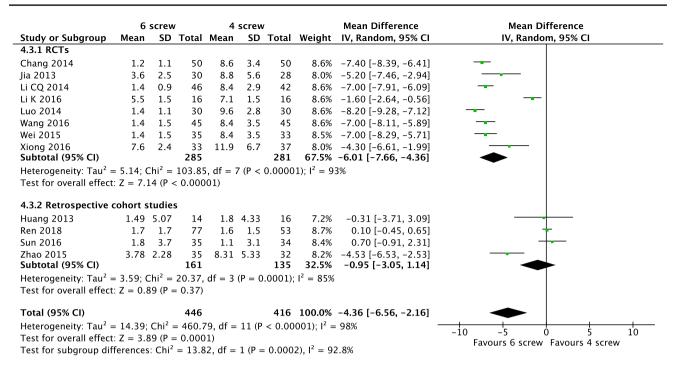


Fig. 11 Forest plot comparing correction loss of anterior vertebral body height for 6S versus 4S. *RCTs* randomized controlled trials, *CI* confidence interval, *IV* inverse variance, *SD* standard deviation

4-screw construct group for both percutaneous and open techniques. No studies describing the hospital length of stay made use of the percutaneous surgical technique.

# Discussion

In this systematic review, short-segment fixation (4S construct) was compared to short-segment fixation with additional intermediate screws at the fracture level (6S construct) in the treatment of thoracolumbar fractures.

The results from our meta-analysis reveal that a 6S construct significantly improves outcomes concerning postoperative pain, short-term and long-term Cobb angles, correction loss of Cobb angle and AVBH and implant failure. Although these results were significantly better in 6S compared to 4S patients, questions remain about the clinical relevance of these differences. The mean difference in VAS of 0.64 and the mean gain in short-term Cobb angle of 1 degree were statistically significant in favour of the 6S. The clinical impact of these small differences, however, remains questionable. The mean difference in long-term Cobb angle of almost 4 degrees and an odds ratio of 0.26 for implant failure seems to be more clinically relevant. This, however, did not result in a significant difference in the measured ODI scores after one year, which may be explained by the fact that the ODI was only studied in 3 RCTs and 2 retrospective studies with a total of 273 patients. The clinical impact of a prolongation of the operation duration with a mean of approximately 6 min and a higher blood loss of 25.30 mL also remains questionable. Although 6S constructs seem to have small but significantly better results on many outcomes, future studies should focus more on patient-reported outcomes and quality of life measurements, in which the benefits and risks should be weighted in a cost-effectiveness analysis [44].

Our meta-analysis shows significantly better results concerning radiological outcomes and implant failure in the 6S construct. These results can be explained by the fact that placement of intermediate screws at the fracture level improves the construct rigidity and stiffness, as described in several biomechanical studies [8–12, 45]. A study by Dobran et al. showed that short-segment fixation with intermediate screws provides radiological results similar to long-segment posterior fixation in which 2 levels above and 2 levels below the fracture are fixated [46]. However, a short-segment 6-screw construct could decrease the size of the immobilized segment and therefore retains more spinal motion compared to a long-segment fixation. Moreover, 6 instead of 8 screws are needed in a short-segment 6S construct and when an open technique is used, a smaller incision size will be necessary [47].

Klezl et al. showed that the addition of vertebral body stenting (VBS) in posterior stabilization could also lead to improvement in pain and functional outcomes [48]. The study by Pflugmacher et al. presented similar improved

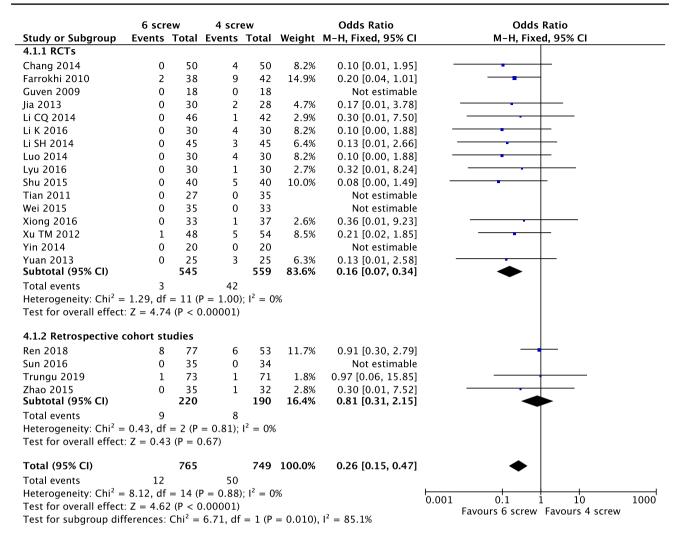


Fig. 12 Forest plot comparing implant failure for 6S versus 4S. *RCTs* randomized controlled trials, *CI* confidence interval, *IV* inverse variance, *SD* standard deviation

results for balloon kyphoplasty combined with posterior instrumentation [49]. Bornemann et al. demonstrated that radiofrequency kyphoplasty in combination with posterior fixation also leads to better results on VAS and ODI scores than with posterior fixation alone [50]. Posterior short-segment fixation with intermediate screws was compared to short-segment fixation with balloon kyphoplasty by Zhang et al. Their study showed peri-operative results favouring intermediate screws but clinical and radiological outcomes favouring balloon kyphoplasty [51]. Future studies on a combination of posterior short-segment fixation with kyphoplasty and intermediate screws through the kyphoplasty cement might be interesting. However, the relation of VBS, balloon kyphoplasty and radiofrequency kyphoplasty to short-segment posterior fixation with intermediate screws in terms of peri-operative, clinical and radiological outcomes requires further elucidation to draw definite conclusions.

Although 28 studies were performed on the use of intermediate screws, no literature exists on the ideal characteristics of these intermediate screws. In our included literature, different types and sizes of screws were used as intermediate screws. Guven et al. and Chang et al. used shorter screws in the fractured vertebra compared to the screws in the non-fractured vertebra [15, 39]. The 6-screw construct by Farrokhi et al. used all same size screws [21]. Poly-axial intermediate screws were used by Sun et al., whereas monoaxial intermediate screws were used by Guven et al. [14, 15]. Hence, there is no consensus on what size and type of intermediate screw are the most favourable. Future (biomechanical) studies are needed on this topic.

The results of our systematic review and meta-analysis are in line with previous and smaller systematic reviews

	6 scr	ew	4 scr	ew		Odds Ratio	Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M–H, Fixed, 95% Cl
4.2.1 RCTs							
Farrokhi 2010	2	38	4	42	48.7%	0.53 [0.09, 3.06]	
Guven 2009	0	18	0	18		Not estimable	
Jia 2013	0	30	0	28		Not estimable	
Li K 2016	0	16	0	16		Not estimable	
Lyu 2016	1	30	0	30	6.4%	3.10 [0.12, 79.23]	
Tian 2011	0	27	0	35		Not estimable	
Wei 2015	0	35	0	33		Not estimable	
Xiong 2016	1	33	2	37	24.7%	0.55 [0.05, 6.32]	
Yin 2014	0	20	0	20		Not estimable	
Subtotal (95% CI)		247		259	79.9%	0.74 [0.21, 2.55]	
Total events	4		6				
Heterogeneity: Chi <sup>2</sup> =	0.95, df	= 2 (P	= 0.62);	$I^2 = 0\%$			
Test for overall effect	Z = 0.47	7 (P = (	0.64)				
4.2.2 Retrospective of	cohort st	udies					
Sun 2016	0	35	0	34		Not estimable	
Trungu 2019	0	73	1	73	20.1%	0.33 [0.01, 8.20]	
Ye 2017	0	20	0	24		Not estimable	
Subtotal (95% CI)		128		131	20.1%	0.33 [0.01, 8.20]	
Total events	0		1				
Heterogeneity: Not ap	plicable						
Test for overall effect	Z = 0.68	8 (P = 0	).50)				
Total (95% CI)		375		390	100.0%	0.66 [0.21, 2.07]	•
Total events	4		7				
Heterogeneity: Chi <sup>2</sup> =	1.14, df	= 3 (P	= 0.77;	$l^2 = 0\%$			
Test for overall effect							0.001 0.1 1 10 1000 Favours 6 screw Favours 4 screw
Test for subgroup dif	ferences:	Chi <sup>2</sup> =	0.21, df	= 1 (P	= 0.64),	$l^2 = 0\%$	Favours o screw Favours 4 screw

Fig. 13 Forest plot comparing post-operative infection for 6S versus 4S. RCTs randomized controlled trials, CI confidence interval, IV inverse variance, SD standard deviation

comparing 4S construct with the 6S construct [52–54]. However, these previous reviews were no longer up to date (included studies until 2017) or did not include all available data due to a language restriction. To date, this is the first and largest review in which all available and up to date literature on this topic is included. Both RCTs and retrospective cohort studies were included in this systematic review, but a subgroup meta-analysis was performed for RCTs and retrospective studies. For most outcomes, the meta-analysis for the RCT and retrospective studies corresponded well; therefore, results from the retrospective studies seem to be reliable and valuable for the overall meta-analysis.

This review still has some limitations. Although most studies were randomized controlled trials, sample sizes were small and in most studies, the type of randomization was not clearly stated or mentioned at all. This potential risk of bias together with the non-blinded fashion both devaluated the quality of evidence in the GRADE, especially for the outcomes that were subjective, or patient reported. Together with the statistical heterogeneity ( $I^2$ ), this led to the devaluation of the quality of evidence for most outcomes to "moderate" or "low".

Moreover, the included studies seem to have heterogeneous populations with different indications for posterior fixation. Although all studies included fractures from level T10 to L5, different fracture types with different classification systems (type A, B or C according to AO/Magerl and type A, B, C or D according to Denis) and patients different neurological status (Frankel A to E) were included in the different studies. Some studies even did not explicitly describe the number of patients in each fracture type or neurological status at all. Mahar et al. showed that the placement of screws at fracture level improves construct rigidity and shields the fractured vertebra from anterior loads [45]. Therefore, especially comminuted fractures that are less stable due to lack of anterior support might benefit more from rigid constructions with intermediate screws. Future studies should direct at which type of fractures benefit the most from 6-screw constructs.

The use of open and percutaneous techniques also implied heterogeneity and poses another limitation of this review. The majority of the included studies used an open technique, while only 3 used a percutaneous technique. Previous studies have shown that percutaneous techniques might result in better outcomes than open techniques, regardless of the placement of additional screws [55]. Besides, a 6S construct might have different impact in percutaneous surgery than in open surgery. Our subgroup analysis group showed less differences between the 4S and 6S constructs in percutaneous surgery, but this might be the result of smaller sample sizes and therefore statistically insignificance.

# Conclusion

This systematic review and meta-analysis show that in the treatment of thoracolumbar fractures, the addition of intermediate screws at the fractured level results in significantly less post-operative pain, better radiological outcomes and less implant failure at the cost of a longer operation time and higher blood loss. However, most differences in outcome were small and the overall quality of the evidence was moderate to low. More studies on this topic are needed and should be directed more at patient reported outcomes and at which specific fracture types benefit the most from intermediate screws.

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#### **Compliance with ethical standards**

**Conflict of interest** The authors declare that they have no conflict of interest.

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