

A comparison of the postoperative analgesic efficacy between epidural and intravenous analgesia in major spine surgery: a meta-analysis

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Abstract: Postoperative analgesia remains a challenge for orthopedic surgeons. The aim of this meta-analysis is to compare the efficacy of epidural analgesia (EA) and intravenous patient-controlled analgesia (IV-PCA) following major spine surgery. We searched electronic databases, including the PubMed, EMBASE, Ovid and Cochrane databases, for randomized controlled trials (RCTs) published before June 2016. The quality of the included trials was assessed using the Cochrane risk-of-bias tool. Random effects models were used to estimate the standardized mean differences (SMDs) and relative risks (RRs), with the corresponding 95% confidence intervals (CI). Subgroup analyses stratified by the type of epidural-infused medication and epidural delivery were also performed. A total of 17 trials matched the inclusion criteria and were chosen for the following meta-analysis. Overall, EA provided significantly superior analgesia, higher patient satisfaction and decreased overall opioid consumption compared with IV-PCA following major spine surgery. Additionally, no differences were found in the side effects associated with these two methods of analgesia. Egger's and Begg's tests showed no significant publication bias. We suggest that EA is superior to IV-PCA for pain management after major spine surgery. More large-scale, high-quality trials are needed to verify these findings.

Keywords: adolescent idiopathic scoliosis, lumbar fusion, epidural analgesia, intravenous application, perioperative pain

Introduction

Postoperative pain is not only a matter of worsened patient-oriented outcomes, such as quality of life, it also substantially affects recovery, increases postoperative morbidities and prolongs the length of hospital stay. ^{1–3} For patients undergoing major spine surgeries, postoperative pain typically lasts for at least 3 days. ⁴ Although multiple analgesic strategies are used, no consensus has been established. Additionally, postoperative analgesia remains a challenging problem for orthopedic surgeons. In recent years, epidural analgesia (EA) and intravenous patient-controlled analgesia (IV-PCA) have both been identified as effective methods of postoperative pain control following spinal surgery, and the efficacies of pain control of these two methods have been compared by several randomized controlled trials (RCTs). However, despite some individual studies demonstrating the analgesic benefits of EA, no published systematic review has examined which method is better. We searched all RCTs comparing the analgesic effects of EA and IV-PCA following spinal surgeries and conducted this meta-analysis to compare EA and IV-PCA and to clarify which method is better in terms of pain control and adverse effects.

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Methods

Search strategy

This systematic review and meta-analysis was based on the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines.⁵ We searched MEDLINE, EMBASE, Cochrane Library and Ovid to retrieve related studies published before June 2016 with combinations of the following keywords: "epidural", "intravenous", "spine surgery (lumbar fusion/lumbar laminectomy/scoliosis correction/spinal fracture fixation)", "postoperative", "pain", "efficacy", "analgesia" and "randomized controlled trial". The language was restricted to English and Chinese. A manual search was also conducted to identify relevant publications from the citation lists of the included articles.

Inclusion and exclusion criteria

Trials were included if they met the following criteria: 1) the study was an RCT; 2) patients received major spine surgeries, including lumbar fusion, lumbar laminectomy, scoliosis correction and spinal fracture fixation; 3) EA was defined as a medication delivered into the epidural space by infusion, a patient-controlled device or repeated bolus dosing; 4) the relative risk (RR) with 95% confidence interval (95% CI; or crude data that allowed their calculation) were reported; and 5) the language was restricted to English and Chinese.

The following types of studies were excluded: 1) case series, case reports, reviews and observational studies; 2) studies incorporating the use of nonpharmacologic interventions as a part of the analgesic regimen; 3) studies for which the crude data were not given or the odds ratio (OR) or RR could not be calculated; and 4) publications that were not written in English or Chinese.

Data extraction and study quality assessment

Two authors independently read the full texts and extracted relevant data from each publication. The following data were extracted: first author, year of publication, country, research design, number of participants, male/female ratio, analgesic regimen used and the postoperative pain scores. Data regarding analgesia-related complications, including nausea, vomiting, pruritus and paresthesia, were also extracted. Graphical data were converted to numerical data independently by the two authors, and disagreements were resolved by consensus.

The quality of the included trials was assessed by two independent reviewers using the Cochrane risk-of-bias tool for RCTs.⁷ The bias domains that were evaluated with this

tool included selection, performance, detection, attrition and reporting bias.

Statistical analysis

Statistical analyses were performed using Stata version 12.0 (Stata Corporation, College Station, TX, USA). We analyzed the RRs with 95% CIs for dichotomous variables and the standardized mean difference (SMD) with 95% CIs for continuous variables. Adjusted risk estimates were also abstracted. The unadjusted risk estimates were calculated using original data when the adjusted risk estimates were unavailable. The I-squared (I2) statistic was used to assess statistical heterogeneity among the studies. A value of $l^2 > 50\%$ reflected significant heterogeneity.8 Both fixed and random effects models were used to pool the data in our study. However, the random effects model is preferable for surgical research because patients who are operated on in different centers have varying risk profiles and selection criteria for each surgical technique. Considering the possible strong heterogeneity resulting from potential differences across the trials, we performed a subgroup analysis stratified by medication type (opioids alone, local anesthetic alone and local anesthetic with an opioid) and epidural delivery type (patient-controlled analgesia and continuous infusion analgesia) for outcomes that were reported by at least two trials. The robustness of the combined results was evaluated by a sensitivity analysis; namely, the analysis was repeated after the exclusion of one study at a time. 10 Begg's funnel plot and Egger's test for possible publication bias were used in analyses that included >10 studies.

Results

A total of 206 articles were identified after the initial search. After the exclusion of 128 articles that were irrelevant to our research, 78 articles remained. In total, 28 articles were retrieved for full-text review, and 20 studies were identified. Finally, three studies were excluded because the crude data could not be extracted. Therefore, 17 trials matched the inclusion criteria and were chosen for the following metaanalysis (Figure 1). These trials were dated from 1989 to 2016, had prospective randomized designs and included a total of 938 participants, of which 508 (54.2%) received EA and 430 (45.8%) underwent IV-PCA. The characteristics of the retrieved studies are presented in Table 1. Of the 17 included trials, eight trials enrolled patients undergoing lumbar fusion, 11-18 six enrolled patients undergoing adolescent idiopathic scoliosis (AIS) correction^{19–24} and three enrolled patients undergoing other major spine surgeries.^{25–27}

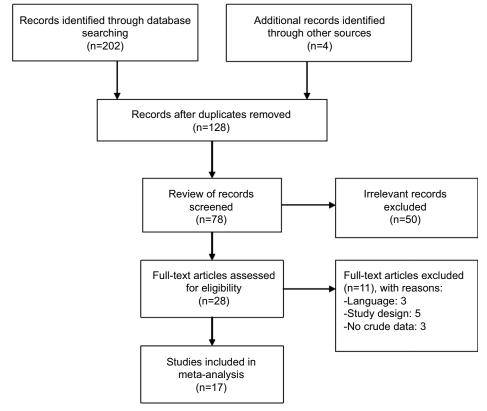


Figure 1 Flow chart of the process of screening of studies.

All IV-PCA patients received opioids for pain control. For EA regimens, 12.2% of the epidural patients received opioids alone, whereas 14.8% received a local anesthetic alone and 73% received a local anesthetic and opioid. A study by Prasartritha et al²⁷ compared three regimens (epidural opioids, epidural opioids and a local anesthetic, and intravenous opioids), and we divided this trial into two comparisons (epidural opioids versus intravenous opioids; and epidural opioids and a local anesthetic versus intravenous opioids) in the following analysis. For the application of epidural analgesics, six studies utilized patient-controlled epidural analgesia (PCEA), and 11 studies utilized continuous epidural infusion analgesia (CEIA). Blumenthal et al^{19,20} compared the efficacy of two types of epidural catheter infusion with intravenous infusion, while the other studies compared the efficacy of a single type of epidural catheter infusion with intravenous infusion. Klatt et al²³ compared three delivery techniques (a single catheter, double catheter and intravenous), and we divided this trial into two comparisons (single catheter versus intravenous delivery, and double catheter versus intravenous delivery). More than half of the included trials described the randomization methods (a computerized list or random number table). All studies reported detailed information about withdrawal and dropouts. Blinding is not a necessity in our analysis because the placement of a sham epidural catheter that is not used for pain management is unequivocally not considered to be ethical.²⁸ Therefore, the methodology of all included studies could be considered to be of moderate-to-high quality. The results of the quality assessment with the Cochrane risk-of-bias checklist are summarized in Figures S1 and S2.

Pain

Postoperative pain scores were recorded using a 0-10 cm visual analog scale (VAS), on which a score of zero indicates no pain and a score of 10 indicates the worst conceivable pain.²⁹ Fourteen studies reported postoperative VAS scores. 11,12,14-22,24-26 The meta-analysis results demonstrated that, overall, EA provided significantly superior analgesic effects compared with IV-PCA at all time points.

On postoperative day 1, epidural patients in the lumbar surgery group had a lower VAS score than IV-PCA patients (p=0.029, SMD = -1.15, 95% CI = -2.19 to -0.12). For patients undergoing scoliosis correction or other spine surgeries, no significant difference was noted between EA and IV-PCA (Figure 2).

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First author,	Country Design	Design	Operations	¥4						
year	•	1		N (male/female)	Mean age, years (±SD)	Type of delivery	Medication	N (male/ female)	Mean age, years (±SD)	Medication
Johnson, 1989 ¹³	S	UB, RCT	Lumbar fusion	12 (7/5)	14	Continuous, single-catheter	Morphine, 0.5 mg/h (1 mg/ mL)	13 (8/5)	39	Morphine sulfate
Joshi, 1995 ¹⁴	Ireland	UB, RCT	Lumbar laminectomy 10 (6/4)	10 (6/4)	45 (3)	Continuous, single-catheter	Fentanyl, 4–10 mL/h	10 (8/2)	49 (4)	Morphine
Cohen, 1997"	SN	DB, RCT	Lumbar fusion	21	45 (2.6)	Continuous, single-catheter	(2 µg/ mL) Morphine sulfate (0.004%)	21	45 (2.5)	Morphine sulfate
Cassady, 2000 ²¹	Sn	UB, RCT	AIS elective spinal	17 (2/15)	14.6 (1.8)	Continuous, single-catheter	and bupivacaine (0.0625%) Bupivacaine (0.125%),	16 (3/13)	14.4 (1.6)	(1 mg/mL) Morphine sulfate
Fisher. 2003 ¹²	Canada	DB. RCT	fusion Lumbar fusion	36 (17/19)	20	Patient-controlled, single-	epinephrine and fentanyl (0.0025 mg/mL) Fentanyl (5 ug/ mL).	38 (15/23)	15	Fentanyl (10 us/mL)
		i i			1	catheter	bupivacaine (I mg/mL) and epinephrine (I ug/ mL)		į	0
O'Hara, 2004²4	SN	DB, RCT	AIS elective spinal	12	Z. R.	Continuous, single-catheter	Fentanyl (5 µg/mL) and	6	Z Z	Morphine sulfate
Blumenthal,	SN	UB, RCT	AlS correction	15 (4/11)	17 (5)	Continuous, double-catheter	Ropivacaine (0.3%)	15 (3/12)	17 (5)	Morphine
Blumenthal, 2006 ¹⁹	Sn	UB, RCT	AIS correction	15 (4/11)	14 (1)	Continuous, double-catheter	Ropivacaine (0.3%)	15 (5/10)	(I)	Morphine
Schenk, 2006 ¹⁸	Germany	DB, RCT	Lumbar fusion	28 (15/13)	42	Patient-controlled, single- catheter	Ropivacaine (0.125%) and sufentanil (1.0µg/ mL)	30 (21/9)	50	Morphine
Hu, 2008 ²⁶	China	UB, RCT	Internal fixation of thoracolumbar verrehral fractures	30	<u>ح</u> ح	Patient-controlled, single- catheter	Bupivacaine (0.125%) and morphine (0.04 mg/mL)	30	۳ Z	Morphine
Fu, 2009 ²⁵	China	UB, RCT	Spine surgery	35	۲ ۲	Patient-controlled, single-catheter	Fentanyl (5 µg/mL) and ropivacaine (0.2%)	35	Z Z	Fentanyl (12 µg/mL)
Gauger, 2009^{22}	SN	UB, RCT	AIS elective spinal fusion	(61/0) 61	15.1 (1.5)	Patient-controlled, single-catheter	Hydromorphone (10 µg/mL) 19 (4/15) and bupivacaine (0.1%)	19 (4/15)	14.7 (1.8)	Hydromorphone
Kluba, 2010 ¹⁵	Germany	UB, RCT	Lumbar fusion	29 (14/15)	57	Continuous, single-catheter	Ropivacaine (0.2%) and sufentanil (5 µg/mL)	23 (14/9)	62	Piritramide
Prasartritha, 2010 ²⁷	Thailand	UB, RCT	Spine surgery	40 (morphine 18/22), 40 (combined 14/26)	55 (morphine), 52 (combined)	Continuous, single-catheter	Morphine (0.1 mg/mL); morphine (0.1 mg/mL) and bupivacaine (0.0625%)	39 (16/23)	50	Morphine
Klatt, 2013 ²³	S	UB, RCT	AIS elective spinal fusion	21 (single), 18 (double)	Z Z	Continuous, single-catheter and double-catheter	Bupivacaine (0.1%) and fentanyl (2 µg/mL)	21	Z Z	Morphine
Li, 2015¹6	China	UB, RCT	Lumbar fusion	65 (16/49)	63.2 (10.2)	Patient-controlled, single- catheter	Sufentanil (75 µg), ropivacaine (150 mg), and droneridol (25 mg)	55 (15/40) 65.3 (9.6)	65.3 (9.6)	Sufentanil (150 µg), droperidol (2.5 mg)
Park, 2016 ¹⁷	Korea	UB, RCT	UB, RCT Lumbar fusion	45 (20/25)	65.8 (9.1)	Continuous, single-catheter	Ropivacaine (0.2%)	41 (22/19) 67.1 (7.9)	(67.1 (7.9)	Fentanyl (10 µg/mL)

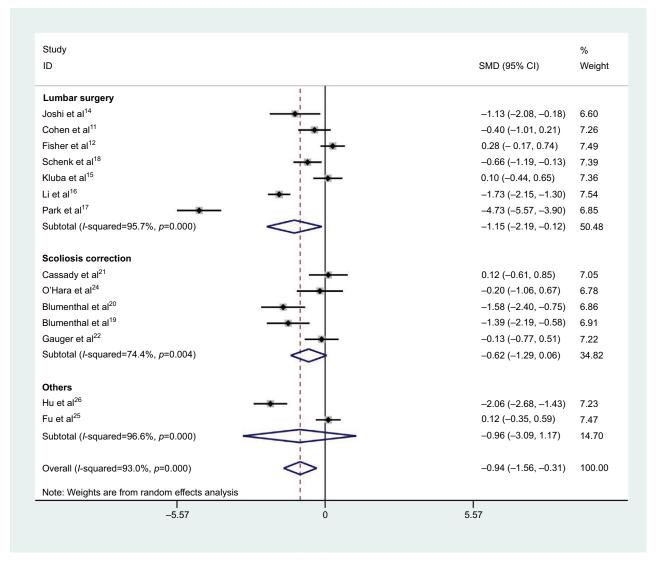


Figure 2 Summary effect of postoperative pain scores 24 hours after surgery.

Notes: Diamonds represent pooled estimates, and width of the diamonds represents 95% Cls. The p-value was calculated using chi-squared test.

Abbreviations: Cl, confidence interval; SMD, standardized mean difference.

On postoperative day 2, the epidural regimens were better than IV-PCA for both lumbar surgery (p=0.045, SMD =-0.93, 95% CI =-1.84 to -0.02) and scoliosis correction (p=0.01, SMD =-0.77, 95% CI =-1.35 to -0.18) groups (Figure 3).

On postoperative day 3, no significant difference was observed between EA and IV-PCA in patients undergoing lumbar surgery (p=0.608, SMD =-0.21, 95% CI =-1.03 to 0.60). For patients undergoing scoliosis correction, EA provided better pain control (p=0.005, SMD =-0.71, 95% CI =-1.22 to -0.21) (Figure 4).

When the subgroup analyses were performed, epidural-infused local anesthetic alone and CEIA provided better pain control following scoliosis correction at all time points. The detailed data are presented in <u>Tables S1 and S2</u>.

Nausea and vomiting

Thirteen trials, including nine lumbar fusion trials, three scoliosis correction trials and one other spinal surgery trial, reported the rate of nausea and vomiting following postoperative analgesia. $^{11-13,15-18,21,23,26,27}$ The meta-analysis results showed no significant difference between the EA and IV-PCA groups (p=0.56, RR =0.88, 95% CI =0.57 to 1.35 for subcategory lumbar fusion; p=0.71, RR =0.97, 95% CI =0.82 to 1.15 for the scoliosis correction subcategory) (Figure S3). When the nine lumbar fusion trials were further divided by the type of epidural medication infused and the epidural delivery type, no significant differences were found in the incidence of nausea and vomiting. The details are shown in Figures S4 and S5, respectively.

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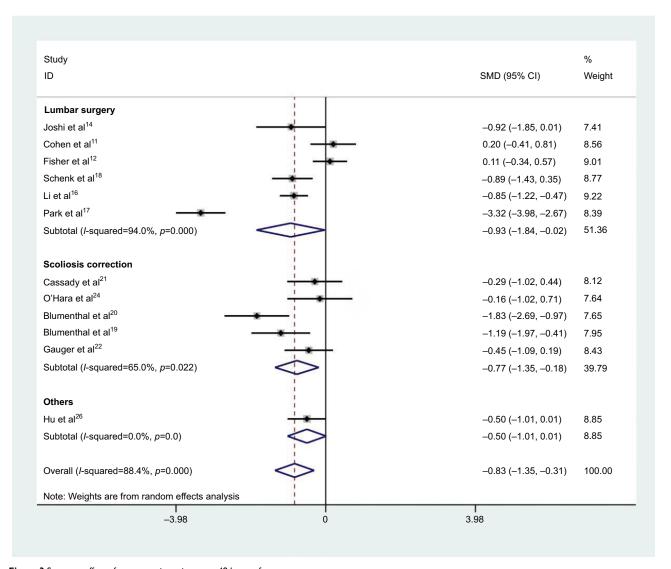


Figure 3 Summary effect of postoperative pain scores 48 hours after surgery.

Notes: Diamonds represent pooled estimates, and width of the diamonds represents 95% Cls. The p-value was calculated using chi-squared test.

Abbreviations: Cl, confidence interval; SMD, standardized mean difference.

Pruritus

Nine trials, including 11 comparisons, reported the incidence of pruritus, $^{11,12,16-18,21,23,26,27}$ with two showing a statistically significant reduction in the EA group compared with the IV-PCA group. 17,26 The meta-analysis of all trials did not reveal any significant difference between the groups (p=0.23, RR =1.52, 95% CI =0.77 to 2.99 for subcategory lumbar fusion; p=0.27, RR =1.26, 95% CI =0.84 to 1.91 for the scoliosis correction subcategory) (Figure S6). When the lumbar fusion trials were stratified by the epidural delivery type, no significant difference in pruritus was observed (Figure S7). Furthermore, in the subgroup stratified by the type of epidural medication (Figure S8), epidural-infused local anesthetic with an opioid had a borderline elevated risk for the incidence of pruritus (p=0.051, RR =1.67, 95% CI =1.00 to 2.78).

Paresthesia

Four studies reported the incidence of paresthesia following postoperative analgesia, 12,15,22,23 with none individually showing a statistically significant reduction in the EA versus IV-PCA groups. The meta-analysis of all studies also demonstrated no difference between these two analgesic methods for different surgeries (p=0.054, RR =7.56, 95% CI =0.97 to 59.17 for lumbar fusion; p=0.45, RR =1.60, 95% CI =0.48 to 5.34 for scoliosis correction) (Figure S9).

The amount of opioids used

The amount of perioperative opioids used was reported in seven studies. 11,12,14,17,19,20,24 O'Hara et al²⁴ reported no difference in morphine consumption between the groups but provided no detailed data, and therefore this study was excluded from the analysis. Pooled data from the other six studies

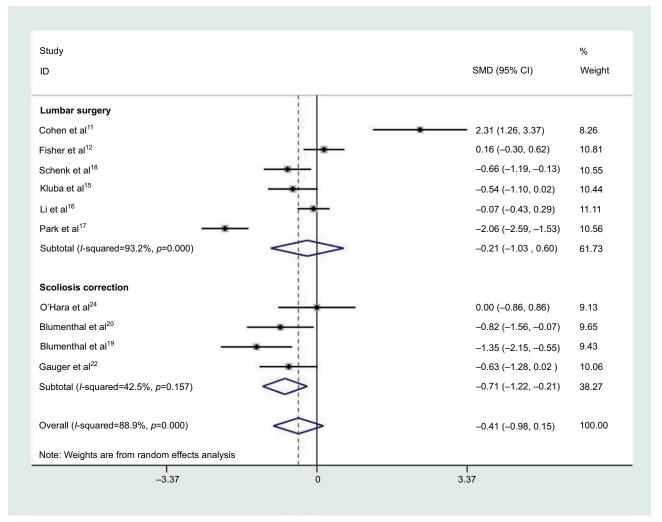


Figure 4 Summary effect of postoperative pain scores 72 hours after surgery. Notes: Diamonds represent pooled estimates, and width of the diamonds represents 95% Cls. The p-value was calculated using chi-squared test. Abbreviations: CI, confidence interval; SMD, standardized mean difference.

revealed a significantly reduced amount of opioids in the EA group. The results were as follows: p=0.012, SMD =-3.12, 95% CI = -5.56 to -0.68 for the lumbar fusion subcategory; and p < 0.01, SMD = -1.45, 95% CI = -2.02 to -0.87 for the scoliosis correction subcategory (Figure S10).

Patient satisfaction

Five of 17 studies assessed patient satisfaction with postoperative pain management, and all of them applied a numeric scale, with a higher score indicating higher satisfaction. 12,15,17,19,20 The results showed that patients undergoing scoliosis correction were significantly more satisfied in the EA group (p<0.01, SMD =2.25, 95% CI =1.59 to 2.90), whereas the satisfaction of patients undergoing lumbar fusion did not significantly differ between treatment types (p=0.09, SMD =0.39, 95% CI =-0.06 to 0.84) (Figure S11).

Motor block

Incidence of motor block was reported by two studies. 17,18 Both studies compared the efficacy of EA and IV-PCA following lumbar fusion. The results indicated that patients in the EA group had a 15 times risk of experiencing motor block (p=0.008, RR =15.07, 95% CI =2.04 to 111.34) (Figure S12).

Publication bias and sensitivity analysis

A funnel plot of the logarithm of RR (logRR) against the logarithm of standard error was constructed for the analysis of the incidence of nausea and vomiting, in which 13 comparisons were included (Figure 5). Visual inspection of the funnel plots did not show any remarkable asymmetry. In addition, Egger's and Begg's tests showed no significant publication bias (p=0.179 for Egger's test and p=0.502

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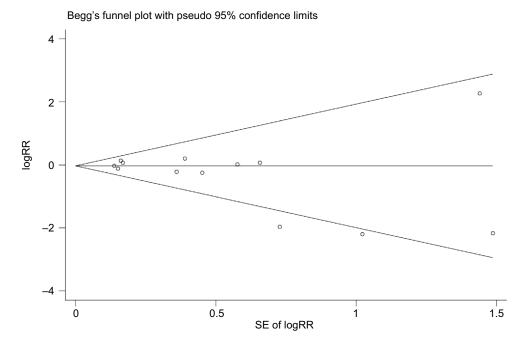


Figure 5 Funnel plot for the meta-analysis on the incidence of nausea and vomiting.

Notes: Each point represents a separate study for the indicated association. The horizontal line indicates the effect size.

Abbreviations: SE, standard error; RR, risk ratio; logRR, natural logarithm of risk ratio.

for Begg's test). A sensitivity analysis was conducted by excluding each study one by one, and the pooled results were not significantly attenuated, which confirmed the robustness of the data (Figure S13).

Discussion

Severe pain is encountered by most patients undergoing major spine surgery. EA and IV-PCA are two common options for treating postoperative pain, and previous systematic reviews have shown that EA provides superior pain management. 6,30 However, both systematic reviews enrolled small numbers of patients undergoing spine surgery, and no universally agreed-upon consensus exists regarding which analgesic approach performs better in spine surgery patients.²² In the literature, Joshi et al¹⁴ compared the efficacy and safety of EA with IV-PCA in the management of postoperative pain after lumbar laminectomy and found that EA was superior for controlling pain. However, in a prospective randomized double-blind clinical trial conducted by Cohen et al,11 the data suggested no clinical advantage of EA over IV-PCA for spinal fusion patients. A trial by Cassady et al²¹ demonstrated that EA and IV-PCA were comparably effective and safe for scoliosis correction procedures in adolescents, which was consistent with the findings reported by O'Hara et al,24 but was in contrast to the findings reported by Gauger et al,²² who found EA to be more effective than IV-PCA for pain relief. These discrepancies are likely due to the relatively small sample sizes, which provided limited statistical power.

Therefore, we performed this meta-analysis to comprehensively evaluate the ability of EA and IV-PCA to improve pain control following major spine surgeries.

We conducted an analysis stratified by surgery type, and for patients undergoing scoliosis correction, no difference was found for the use of EA and IV-PCA for pain control on postoperative day 1. However, on postoperative days 2 and 3, EA significantly alleviated pain compared to IV-PCA. In contrast, for patients undergoing lumbar fusion, EA provided better pain management than IV-PCA on postoperative days 1 and 2 but failed to have a superior analgesic effect on day 3. We believe this difference was due to two reasons. First, patients undergoing scoliosis correction were all children and adolescents, while patients undergoing lumbar fusion were mostly adults with degenerative spinal diseases. Additionally, adolescents have been reported to have more pain than adults after surgery.31 Second, the researchers adjusted their analgesic doses daily to facilitate patient comfort, which may influence the results.²² When we further investigated the effects of the type of epidural delivery on our results, we found that CEIA provided better pain control following scoliosis correction at all time points. For patients undergoing lumbar fusion, neither CEIA nor PCEA had better analgesic effects than IV-PCA. This result was not consistent with the research presented by Lu et al, 32 which confirmed that PCEA was more effective in relieving pain than IV-PCA for spinal fusion patients. The explanation for this discrepancy may be as follows. First, Lu et al³² grouped patients undergoing

scoliosis correction and patients undergoing lumbar fusion together, which led to exaggerated results. Second, they calculated the OR for dichotomous variables and used a fixedeffect model in their analysis. In contrast, we considered it more appropriate to calculate the RR and to use a random effects model with RCTs, which is the more conservative approach.³³ Furthermore, we evaluated the effects of epidural medications on our results. Surprisingly, EA with opioids and local anesthetics did not provide a significant benefit for different types of surgery. However, local anesthetic-based EA significantly relieved pain following scoliosis correction surgery. Both related trials used ropivacaine as the epiduralinfused drug. 19,20 However, another local anesthetic, bupivacaine, failed to support the better analgesic effect of epidurals compared with IV-PCA.²¹ This lack of an analgesic effect may have been related to the lower dosage of bupivacaine used due to safety concerns.³⁴ Compared to bupivacaine, ropivacaine has been suggested to significantly improve safety profiles and to provide effective pain relief at lower concentrations.³⁵ Recently, Park et al¹⁷ conducted a prospective comparative trial including 94 patients harboring lumbar spinal stenosis. The patients were randomized to receive either EA with ropivacaine (51 patients) or conventional IV-PCA (43 patients). During the first postoperative 3 days, significantly lower pain scores were observed in the EA group. However, the benefits of ropivacaine-based EA should be interpreted with caution due to the limited number and small sample sizes of relevant trials. The number of epidural catheters has also been postulated to influence the analgesic effects. Blumenthal et al^{19,20} compared the analgesic effects of two intraoperatively placed epidural catheters with IV-PCA, and the results indicated that the double epidural catheter technique provided better postoperative analgesia. In another trial, Klatt et al compared single and double epidural catheters with IV-PCA. The results demonstrated that while single epidural analgesia and IV-PCA provided good pain control, double epidural catheter analgesia provided consistently better pain control than either the single epidural catheter technique or IV-PCA.²³ Both trials were conducted in adolescent patients, and future studies are needed to confirm whether this double-catheter technique would also be beneficial in adult patients.

Regarding the analgesia-related side effects, we did not find any significant difference between EA and IV-PCA. The overall side effect rates reported were comparable to those observed by other sources. The cumulative incidence of nausea and vomiting from EA was reported to be 27.6%, and the incidence of pruritus was 32.8%. In our analysis, the percentages of nausea and vomiting and pruritus were 28.1% and 23.7%, respectively.

Motor block after spine surgery, which may confound the diagnosis of neurodeficits due to the surgery itself, was regarded as one of most important issues in several trials.^{17,18} Motor block was not reported after administration of 10 mL of ropivacaine (0.1%) by Gottschalk et al.³⁷ In a trial by Schenk et al,¹⁸ motor block occurred after the initial bolus of 14 mL of 0.125% ropivacaine in five of 28 patients (17.9%) in the EA group. In a study by Park et al,¹⁷ an epidural infusion of ropivacaine (0.2%) caused a transient motor block in 10 of 47 patients (22%); however, reducing the infusion rate led to the return of motor function. Thus, the optimal concentration and infusion rate of local anesthetics should be considered before beginning epidural pain management.

This analysis has some limitations. First, publication bias cannot be excluded because only publications in English and Chinese were included in our analysis.³⁸ A possibility exists that we missed certain eligible studies. Second, separate evaluations of pain at rest and with activity have become the more accepted method for evaluating pain.²² Most of the included trials failed to use this method, which may attenuate the pain levels. Third, one must consider the validity and difficulty in measuring outcomes such as pain levels among adolescent patients, which was the main reason that we divided the trials according to the type of surgery and chose the SMD as the estimate. Fourth, the sample sizes of the included studies were relatively small, which led to a possible type II error and therefore limited the power of our results. Finally, 13 of 17 included trials used an unblended design in the allocation of treatment and the assessment of outcomes.

Conclusion

Despite these limitations, the results of this meta-analysis suggest that EA has several advantages over IV-PCA with respect to pain management, patient satisfaction and overall opioid consumption after major spine surgery. No difference was observed in the side effects associated with these two analgesic methods. Continuous EA is superior for controlling pain in adolescents undergoing scoliosis correction. Local anesthetics seem to be candidate drugs for epidural regimens in the future. More large-scale, high-quality trials that consider multiple variables (ie, types of medication infused, the optimal infusion rate, optimal medication concentration and the number of catheters) are warranted to verify these findings.

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Disclosure

The authors report no conflicts of interest in this work.

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