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REVIEW ARTICLE

Efficacy of erector spinae plane block in pain management for patients with herpes zoster: a systematic review and meta-analysis



Alexandre Yamada Fujimura Júnior (1)^a,*, Carolina Braga Moura (1)^b, Arnaldo Bastos dos Santos (1)^c

^a Faculdade de Medicina de Marília (FAMEMA), Departamento de Medicina, Marília, SP, Brazil

^b Universidade Federal Fluminense (UFF), Departamento de Neurologia, Niterói, RJ, Brazil

^c Hospital Sírio Libanês, Departamento de Anestesiologia, São Paulo, SP, Brazil

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KEYWORDSAbstNerve block;ObjeAnesthesia,Planconduction;MethHerpes zoster;izedPain;popuNeuralgia,Thepostherpetiction.outcoResueighin th(MD)95%95%-9.9;Concfit inis neis ne	ract ract ract: ract: rectives: Systematic review and meta-analysis to evaluate the efficacy of the Erector Spinae e Block (ESPB) in managing pain related to Herpes Zoster. nods: We systematically searched PubMed, Embase, Cochrane Library, and CNKI for random- trials comparing ESPB plus standard clinical treatment with clinical treatment alone. The nation included patients with acute infection and those with Postherpetic Neuralgia (PHN). primary outcome was pain intensity, and secondary outcomes included analgesic consump- Mean Difference (MD) was used for continuous outcomes, and Risk Ratio (RR) for binary omes. <i>Its:</i> Seven trials with 362 patients were included. ESPB significantly reduced pain up to t weeks (MD = -1.21; 95% CI -2.17 to -0.24; $I^2 = 89\%$). In the subgroup analysis of patients the acute stage, the benefit seemed to extend with pain reduction lasting up to 12-weeks = -1.49; 95% CI -2.61 to -0.37; $I^2 = 0\%$), and a reduction in the incidence of PHN (RR = 0.49; CI 0.28 to 0.85; I^2 : 0%). In the PHN subgroup, pain reduction was notable only at four weeks = -1.08; 95% CI -1.81 to -0.35; $I^2 = 86\%$). ESPB also reduced acetaminophen (MD = -0.6 g.day ⁻¹ ; CI -1.05 to -0.14; $I^2 = 49\%$) and pregabalin consumption (-68.58 mg.day ⁻¹ ; 95% CI -127.18 to 7; $I^2 = 41\%$) over 12 weeks. <i>Clusion:</i> ESPB seems to provide pain relief in Herpes Zoster patients, with a prolonged bene- the acute stage. Also, ESPB reduced the need for analgesics over 12 weeks. More research eded to corroborate this practice.

Abbreviations: ESPB, Erector Spinae Plane Block; PHN, Postherpetic Neuralgia; Rop, Ropivacaine; Bup, Bupivacaine; Beta, Betamethasone; Dex, Dexamethasone; Triamci, Triamcinolone; MeCbl, Methylcobalamin; MD, Mean Difference; CNKI, Chinese National Knowledge Infrastructure; CI, Confidence Interval; PRISMA, Preferred Reporting Items for Systematic Reviews and Meta-Analysis; RCTs, Randomized Clinical Trial(s); VAS, Visual Analogic Scale; NRS, Numeric Rate Scale.

* Corresponding authors.

E-mail: alexandrefujimurajr@gmail.com (A.Y. Fujimura Júnior).

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Introduction

Herpes zoster, caused by the reactivation of the varicellazoster virus in the sensory ganglia of cranial nerves and dorsal root ganglia, manifests as a painful rash along the affected dermatome. In addition to pain, this condition severely impacts quality of life, particularly in the elderly, who are more susceptible to complications such as Postherpetic Neuralgia (PHN), a chronic neuropathic condition.¹⁻⁴

Although vaccination has proven effective in reducing the incidence of herpes zoster, vaccine uptake remains low, with only 56% of eligible patients receiving it.⁵ Consequently, interventional therapies targeting different stages of the disease are still essential, especially in refractory cases where inappropriate use of analgesics, including anti-inflammatory drugs and opioids, can result in more harm than benefit.⁶

Anesthetic techniques like paravertebral and epidural blocks have demonstrated efficacy in pain relief and in reducing the incidence of PHN.^{7,8} However, these techniques require advanced expertise and carry a higher risk of complications.^{9,10}

The Erector Spinae Plane Block (ESPB) has recently emerged as a safer, easier alternative with fewer complications, such as pneumothorax and hematoma.^{11,12} Given these potential advantages, we conducted a meta-analysis to evaluate the effectiveness of ESPB in managing pain associated with herpes zoster.

Material and methods

Protocol and registration

This systematic review and meta-analysis was conducted and reported in accordance with the Cochrane Collaboration Handbook for Systematic Reviews of Interventions and the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA) guidelines.¹³ The review aimed to evaluate the efficacy of the ESPB in patients with pain related to Herpes Zoster infection. This study was prospectively registered in PROSPERO with number CRD42024566674.

Eligibility criteria

Studies were included in this review if they met the following criteria: 1) Randomized Clinical Trials (RCTs); 2) Enrolled patients with pain related to herpes zoster in two contexts: acute herpes zoster (reactivation of a viral infection with severe pain in a specific dermatome associated with vesicular erythema) and PHN (persistent pain lasting more than three months after the resolution of acute infection and dermatological lesions); 3) Compared clinical treatment combined with the application of ESPB to clinical treatment alone; and 4) Assessed at least one of the targeted outcomes of this meta-analysis. Studies without a control group or with overlapping patient populations were excluded. No restrictions were applied regarding the date or language of publication, as part of an effort to broaden the search.

Search strategy and study selection

We conducted a systematic search of the MEDLINE, Cochrane, Embase, and China National Knowledge Infrastructure (CNKI) databases from their inception until July to September 2024. The search strategy used the following terms: "erector spinae plane block"; "erector spinae block"; "herpes-zoster"; "herpes zoster"; "postherpetic"; "varicella zoster virus"; "Varicella-Zoster Virus" and "Chickenpox". The complete search strategy, using Boolean operators and specific models adapted for each database searched, is available in Supplementary Table 1.

All identified articles were systematically evaluated according to the inclusion and exclusion criteria. Article selection was performed independently by two authors (A.Y and A.B), with any disagreements resolved by consensus. To enhance the search for studies, the authors reviewed the references and related studies of the included articles to locate additional studies that met the inclusion criteria.

Data extraction

Two reviewers (A.Y and C.B) independently read the included studies and extracted data on 1) Pain scale over each available week or month; 2) Consumption of any analgesics, whether opioids or anti-inflammatory drugs, or any neuroleptics used by patients during clinical follow-up; 3) Number of patients who developed PHN; 4) Adverse effects, including minor ones such as nausea, vomiting, dizziness, headache, or transient hypotension, and major ones such as bleeding and pneumothorax. Additionally, data were collected regarding the type of anesthetic drug used for ESPB, the number of applications performed during the study, the clinical treatment protocol, and key epidemiological characteristics, including the age and sex of the patients. All extracted data were cross verified for accuracy by a third reviewer (A.B).

Regarding the pain score, we used the Visual Analog Scale (VAS), graded from 0 to 10, as the standard for representing our forest plots. We considered, based on a previous study, the equivalence between VAS and the Numeric Rating Scale (NRS).¹⁴ For continuous outcomes, we extracted the mean and standard deviation for each group. When only the median and quartiles were available, we converted them into mean and standard deviation.^{15,16}

To calculate the daily mean intake of analgesics used over 12 to 11 weeks of follow-up, we used the weighted mean,¹⁷ in the same way, that each daily mean reported over different time intervals was adjusted according to the duration of each interval. Thus, longer time intervals had more weight in calculating the average over 12 weeks. In the same way, we calculated the combined average standard deviation over the 12-week period using a more complex equation.¹⁸

Quality assessment

Quality assessment of RCTs was performed using the Cochrane Collaboration's tool¹⁹ for assessing risk of bias in randomized trials, in which studies are scored as high, low, or some concerns of risk of bias in 5 domains: selection, performance, detection, attrition, and reporting biases. This risk of bias evaluation was performed independently by two authors (A.Y and C.B) with disagreements resolved by consensus. To create the risk of bias figure, we used the Robvis tool.²⁰

Data analysis

The effects of ESPB on continuous outcomes were evaluated using the Mean Difference (MD) with a 95% Confidence Interval (95% CI), while binary outcomes were assessed using the Risk Ratio (RR) with a 95% CI.

For binary outcomes, we employed the Mantel-Haenszel method, and for continuous outcomes, we used the inversevariance method. Heterogeneity was assessed through Cochran's Q test, I² statistics, and Tau-squared, utilizing the restricted maximum likelihood estimator. Heterogeneity was categorized as low (I² = 0%-25%), moderate (I² = 26% -50%), or high (I² > 50%). The fixed-effects model was used for outcomes with low heterogeneity (I² < 25%) and the random-effects model for studies with moderate to high heterogeneity (I² > 25%). All statistical analyses were performed using Review Manager version 5.4 (Cochrane Center, The Cochrane Collaboration, Denmark).

We also conducted subgroup analysis and sensitivity analysis to evaluate the effect of ESPB in the different clinical phases of herpes zoster. Due to the limited number of studies for each outcome, we did not perform publication bias analysis.

Results

Study selection and characteristics

As shown in Figure 1, the initial search identified 100 studies. After removing duplicates and ineligible studies, 24 remained and were fully reviewed based on the inclusion criteria. Of these, 7 RCTs²¹⁻²⁷ were included, comprising 362 individuals with Herpes Zoster-related pain. Participants



Figure 1 Flow diagram illustrating the study selection process according to the PRISMA guidelines.

were equally divided between those receiving ESPB combined with clinical therapy and those receiving clinical therapy alone. Of the total, 192 had acute Herpes Zoster, while the remaining 170 patients had PHN. Medication therapy for those with acute infection included antiviral drugs (acyclovir or valacyclovir), either alone or in combination with pregabalin, gabapentin or methylcobalamin. For those with PHN, the primary medication used was pregabalin or gabapentin, either alone or combined with an anti-inflammatory. Further details on the study characteristics are reported in Table 1.

Pooled analysis of all studies and subanalysis

This meta-analysis of RCTs demonstrates that combining ESPB with clinical treatment is more effective than medication alone in controlling pain. One week after therapy, the MD in the VAS scale between the ESPB and control group was -0.98 (95% CI -1.29 to -0.67; $I^2 = 0\%$; Fig. 2A). Two weeks after, it was -1.43 (95% CI -2.10 to -0.75; $I^2 = 83\%$; Fig. 2B); three weeks after, it was -0.88 (95% CI -1.24 to -0.53; $I^2 = 0\%$; Fig. 2C); four weeks after, it was -1.47 (95% CI -2.01 to -0.93; $I^2 = 76\%$; Fig. 2D). At eight weeks, the MD was -1.21 (95% CI: -2.17 to -0.24; $I^2 = 89\%$; Fig. 2E). However, after this period, no statistical significance was observed. At 12 weeks, the effect size was -0.81 (95% CI -1.93 to 0.31; $I^2 = 53\%$; Fig. 2F).

In the subanalysis of the acute phase of Herpes Zoster, four studies^{21-23,25} reported pain scale data. One week after the follow-up began, the MD between the groups was -0.92 (95% CI -1.63 to -0.21; I² = 0%; Fig. 3A). Two weeks later, this value was -2.18 (95% CI -3.5 to -0.85; I² = 71%; Fig. 3B). In four weeks, this value was -1.99 (95% CI -2.59 to -1.40: I^2 = 23%; Fig. 3C), reinforcing the benefit of ESPB in pain management. Unlike the overall pooled result mentioned above, the subgroup analysis of patients with acute-phase herpes zoster showed pain score reduction even after 12 weeks, with a MD of -1.49 (95% CI -2.61 to -0.37; $I^2 = 0\%$; Fig. 3D). This finding is further supported by the combination of three studies^{22,23,25} that evaluated the number of patients that developed PHN at the third month of followup. The risk ratio of 0.49 (95% CI 0.28 to 0.85; $I^2 = 0\%$; Fig. 3E) indicates a significant reduction PHN in patients undergoing ESPB.

In the subgroup analysis of patients with PHN, the MD one week after ESPB was -0.99 (95% CI -1.34 to -0.64; $I^2 = 0\%$; Fig. 4A). At four weeks, the MD was -1.08 (95% CI -1.81 to -0.35; $I^2 = 86\%$; Fig. 4B). However, by eight weeks, the MD was -1.08 (95% CI -2.19 to 0.02; $I^2 = 93\%$; Fig. 4C), demonstrating no clear conclusions regarding its benefit at this time point. Beyond this period, only one study²⁴ assessed pain scores in patients with PHN and demonstrated pain score reduction only up to 1 week.

Regarding analgesic medication, three studies^{22,24,25} reported the daily dose of acetaminophen over 12 weeks. Our meta-analysis revealed that patients undergoing ESPB required a lower amount of this analgesic, with an MD of -0.60 g per day (95% CI -1.05 to -0.14; $I^2 = 49\%$; Fig. 5A). Additionally, one study²³ evaluated the total dose of acetaminophen used over 24 weeks of follow-up, showing that the ESPB group had a significantly lower total dose of this analgesic (78.53g vs. 153.83g, p = 0.024). Only one study²² evaluated opioids, demonstrating a significant difference in

daily tramadol consumption over 12 weeks as well (36 mg vs. 245 mg, p = 0.001).

Additionally, two studies^{24,25} assessed the daily use of pregabalin over 12 weeks and, through analysis of the combined data, indicated that patients undergoing ESPB required a lower dose of this anticonvulsant, with a reduction of -68.58 mg per day (95% CI -127.18 to -9.97; $l^2 = 41\%$; Fig. 5B). Two other studies^{23,27} evaluated the total dose of pregabalin. In the study conducted by Cao²⁷ patients in the intervention group required a significantly lower total amount of pregabalin over the 4-week evaluation period (1.8 g vs. 7.28 g; p < 0.005). In the study conducted by Abdelwahab,²³ the same medication was also significantly lower in the intervention group over a 24-week period (24,550.8 mg vs. 35,575.0 mg, p = 0.041).

Regarding minor adverse effects such as dizziness, drowsiness, and nausea, three studies^{24,26,27} reported these complications, with the combined statistical analysis showing no significant difference between ESPB and placebo (RR = 0.71; 95% CI 0.36 to 1.38; $I^2 = 0\%$; Supplementary Fig. 1). Finally, no major complications, such as pneumothorax or bleeding, were reported in the studies.

Quality assessment

Individual RCT appraisal is reported in Supplementary Figure 2. Two^{22,24} of the six RCTs included in this systematic review were assessed as having a low risk of bias. These trials were double-blinded and clearly described an adequate randomization process. Another three studies^{21,23,25} were rated as having some concerns, primarily because they did not provide sufficient information regarding the blinding of outcome assessors and patients, raising uncertainty about the potential for bias in this domain. The remaining trials^{26,27} were assessed as having a high risk of bias because, in addition to not mentioning the blinding of assessors and patients, they also did not provide a detailed explanation of how the randomization process was conducted, and they did not have a study protocol before the initiation of RCT registered in Clinical Trials.

Discussion

Our article is the first systematic review and meta-analysis conducted to evaluate whether ESPB therapy can improve clinical symptoms in patients with herpes zoster-related pain. This approach highlights the potential of ESPB as an alternative management strategy, particularly for cases unresponsive to conventional pharmacological treatments. The key findings demonstrate that ESPB therapy resulted in substantial pain reduction, improved patient-reported outcomes, and significantly decreased the need for analgesics, notably pregabalin, acetaminophen, and tramadol. Additionally, no adverse effects were reported, reinforcing the safety profile of this intervention.

In the combined analysis of all included studies, significant heterogeneity was identified in pain outcomes, particularly at the second, fourth, and eighth weeks of clinical follow-up as evidenced in Figure 2. This variability can be attributed to differences in the studied populations, as the pooled analysis included patients in the acute phase of

Study	Anesthetic Composition for ESPB and Application Frequency	Standard oral medicines	Patients ESPB / Control	Male (%) ESPB / Control	Phase of pain in study	Age ^a (years) ESPB / Control	Baseline pain ^b ESPB / Control	Follow-up, weeks	Country	Outcomes available
Patil ²¹ 2024	20 mL Bup 0.25% + 8 mg Dex, applied only once	No specified	20 / 20	75 / 55	Acute	56.10 / 57.60	7.7 / 7.5	8	India	Incidence of PHN, pain score (2, 4 and 8 w), and rescue analgesic require- ment.
Lin ²² 2021	25 mL Rop. 0.4%, applied daily for 3 days	Valacyclovir + MeCbl	26 / 26	46.2 / 50	Acute	68.2 / 65.2	6 / 6.8	12	China	Incidence of PHN, pain score (1, 4, and 12 w), acetaminophen and trama- dol consumption
Abdelwahab ²³ 2022	Epinephrine + 2.5 mL Bup 0.5% + 8 mg Dex, applied only once	Acyclovir + Pregabalin	30 / 30	40 / 43.3	Acute	59.47 / 61.3	7 / 7	24	Egypt	Incidence of PHN, pain score (1, 3, 4, 12, and 24 w), pregabalin and acet- aminophen consumption
Ahmed ²⁴ 2022	20 mL Bup 0.25%, applied only once	Pregabalin + Acetaminophen	25 / 25	56 / 52	PHN	56.16 /54.36	7 / 7	12	Egypt	Pain score (each week until 12 weeks), pregabalin and acetaminophen consumption
El-Sayed ²⁵ 2021	20 mL Bup 0.25%, applied once: after 2 w, second application, if VAS > 6	Acyclovir + Pregabalin	20 / 20	N/A	Acute	N/A	8.8 / 9	12	Egypt	Incidence of PHN, pain score (2, 4, and 12 w), pre- gabalin and acetamino- phen consumption
Xiang ²⁶ 2018	20 mL Rop 0.15% + 0.3 mL Beta + 0.5 g MeCbl Applied once; repeat at 2 w or 4 w if needed	Gabapentin	30 / 30	N/A	PHN	71.2/71.5	7.59 / 7.54	10	China	Pain score (1, 4, 6, 8, and 10 w)
Cao ²⁷ 2019	20 mL Rop 0.5% + 20 mg Tri- amci + 0.5 mg MeCbl Applied weekly for 4 w	Pregabalin	30 / 30	46.67 / 50	PHN	65/65	7.3 / 7.4	8	China	Pain score (1, 2, 3, 4, 5, 6 and 8 w), pregabalin consumption

ESPB, Erector Spinae Plane Block; PHN, Postherpetic Neuralgia; Rop, Ropivacaine; Bup, Bupivacaine; Dex, Dexamethasone; Beta, Betamethasone; Triamci, Triamcinolone; MeCbl, Methylcobalamin; N/A, Not Available.

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^a Mean age; ^b Pain using VAS (0–10) and reporting mean.

A. VAS in 1 week

	ESPB Control							Mean Difference		Mean Difference			
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% CI	1	IV, Fixed, 95% CI			
Abdelwahab 2022	4.2674	2.1407	30	5	2.7245	30	6.3%	-0.73 [-1.97, 0.51]					
Ahmed 2022	2.15722	2	28	2.7186	3.1445	25	4.7%	-0.56 [-2.00, 0.88]					
Cao 2019	5.2674	0.973	30	6.15568	1.5588	30	22.5%	-0.89 [-1.55, -0.23]					
Lin 2021	3.307	1.9608	26	4.3213	1.1765	28	12.8%	-1.01 [-1.88, -0.14]					
Xiang 2018	4.01	0.84	30	5.08	0.84	30	53.7%	-1.07 [-1.50, -0.64]		-			
Total (95% CI)			144			143	100.0%	-0.98 [-1.29, -0.67]		•			
Heterogeneity: Chi2 = 1	4 (P = 0.9	(5); ² =	0%					4		-			
Test for overall effect:	Z = 6.14 (F	> < 0.000	01)							Favours [ESPB] Favours [control]	4		

B.VAS in 2 weeks

	ESPB			(Control			Mean Difference		Mean Dif	ference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total Weight IV, Random, 95% C		IV, Random, 95% C	1	IV, Rando	m, 95% CI	
Ahmed 2022	2.3578	2.3584	25	2.7186	3.1445	26	11.7%	-0.36 [-1.88, 1.16]				
Cao 2019	4.6435	0.7784	30	5.3565	0.7784	30	26.1%	-0.71 [-1.11, -0.32]		-		
El-Sayed 2021	5.8201	1.9946	20	7.3201	1.1968	20	17.3%	-1.50 [-2.52, -0.48]				
Patil 2024	3.3	2.2	20	6.15	0.7	20	17.4%	-2.85 [-3.86, -1.84]	_	-		
Xiang 2018	3.1	0.55	30	4.71	0.48	30	27.5%	-1.61 [-1.87, -1.35]		-		
Total (95% CI)			125			126	100.0%	-1.43 [-2.10, -0.75]		•		
Patil 2024 3.3 2.2 Giang 2018 3.1 0.55 Total (95% CI) 1 feterogeneity: Tau² = 0.42; Chi² = 23.87, df fest for overall effect: Z = 4.14 (P < 0.0001)				(P < 0.0	001); l ² =	83%			-	1 1	2	1
Study or Subgroup Mean SD Tr. Ahmed 2022 2.3578 2.3584 Cao 2019 4.6435 0.7784 B-Sayed 2021 5.8301 1.9946 B-Sayed 2021 5.8301 3.22 Xiang 2018 3.1 0.55 Total (95% CI) - - Hetarogeneity: Tau ² = 0.42; Chi ² = 23.87; df - -									-4	Favours [ESPB]	Favours [control]	4

C. VAS in 3 weeks

		ESPB		c	Control			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% C	I IV, Fixed, 95% CI
Abdelwahab 2022	3.6435	2.3363	30	4	1.5568	30	12.6%	-0.36 [-1.36, 0.65]	
Ahmed 2022	2.3578	2.3584	25	2.7186	3.1445	25	5.4%	-0.36 [-1.90, 1.18]	
Cao 2019	1.3586	0.7784	30	2.3565	0.7784	30	82.0%	-1.00 [-1.39, -0.60]	-
Total (95% CI)			85			85	100.0%	-0.88 [-1.24, -0.53]	•
Heterogeneity: Chi ² =	1.82, df =	2 (P = 0	.40); 12	= 0%					-4 -2 0 2 4
lest for overall effect:						Favours [ESPB] Favours [control]			

D.VAS in 4 weeks

	1	ESPB		(Control			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean SD Total			Weight	IV, Random, 95% C	CI IV, Random, 95% CI
Abdelwahab 2022	2.5	2.22	30	4	1.48	30	14.5%	-1.50 [-2.45, -0.55]	·
Ahmed 2022	2.15722	2	25	2.3673	2.3588	25	11.3%	-0.21 [-1.42, 1.00]	
Cao 2019	2.3565	0.7784	30	3.25	0.74	30	23.5%	-0.89 [-1.28, -0.51]	
EI-Sayed 2021	3	1.5957	20	5	1.5957	20	14.0%	-2.00 [-2.99, -1.01]	· · · ·
Patil 2024	3.4	2.6	20	6.15	0.74	20	11.6%	-2.75 [-3.93, -1.57]	·
Xiang 2018	2.33	0.44	30	4	0.56	30	25.1%	-1.67 [-1.92, -1.42]	· •
Total (95% CI)			155			155	100.0%	-1.47 [-2.01, -0.93]	•
Heterogeneity: Tau ² =	0.28; Chi ²	= 20.74,	df = 5	(P = 0.00	09); I ² = 1	76%			
Test for overall effect:	Z = 5.36 (F	< 0.000	01)						Favours [ESPB] Favours [control]

E. VAS in 8 weeks

		ESPB				ontrol			Mean Difference		Mean Differer	ice	
_	Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI		IV, Random, 95	5% CI	
	Ahmed 2022	1.6422	0.7861	25	2	1.5722	25	26.6%	-0.36 [-1.05, 0.33]				
	Cao 2019	2.8565	1.5568	30	3.6435	0.7784	30	27.3%	-0.79 [-1.41, -0.16]				
	Patil 2024	2.45	3.05	20	4.3	2.27	20	16.0%	-1.85 [-3.52, -0.18]	_	-		
	Xiang 2018	1	0.59	30	3	0.42	30	30.1%	-2.00 [-2.26, -1.74]		-		
	Total (95% CI)			105			105	100.0%	-1.21 [-2.17, -0.24]		-		
	Heterogeneity: Tau ² =	0.79; Chi	² = 27.93	, df = 3	(P < 0.0	0001); l ²	= 89%			-4	-2 0	2	4
	Test for overall effect:	Z = 2.46	(P = 0.01))						-4	Favours [ESPB] Favo	urs [control]	4

F. VAS in 12 weeks

		ESPB		0	Control			Mean Difference		Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% C	i	IV, Random, 95% CI	
Abdelwahab 2022	1.2674	2.1407	30	2.713	3.1137	30	33.0%	-1.45 [-2.80, -0.09]		_	
Ahmed 2022	1.7156	1.5722	25	1.7156	1.5722	25	45.8%	0.00 [-0.87, 0.87]			
El-Sayed 2021	0.6799	1.1968	20	2.2591	4.3882	20	21.1%	-1.58 [-3.57, 0.41]	_		
Total (95% CI)			75			75	100.0%	-0.81 [-1.93, 0.31]		-	
Heterogeneity: Tau ² =	0.52; Chi	2 = 4.24,	df = 2 (P = 0.12); l ² = 53	%			-		_
Test for overall effect:	Z = 1.42	(P = 0.16	5)						-4	-2 0 2 Eavours (ESPB) Eavours (control)	4

Figure 2 Analysis of ESPB therapy in reducing pain in patients with herpes zoster-related pain across several weeks of follow-up.

herpes zoster and those with PHN. In the acute phase, pain is primarily inflammatory, driven by varicella-zoster virus reactivation and local immune responses, resulting in peripheral sensitization.^{28,29} In contrast, postherpetic neuralgia is characterized by chronic neuropathic pain, stemming from nerve damage, demyelination, and central nervous system remodeling.³⁰⁻³³ These distinct mechanisms likely influenced treatment efficacy and contributed to the observed heterogeneity, highlighting the need to consider disease stage in interpreting pooled outcomes.

Another factor that justifies the observed heterogeneity is the varying methods by which ESPB was performed in the intervention group across the included studies. For example, three studies^{21,23,24} analyzed a single injection of ESPB. On the other hand, other studies adopted repeated applications of ESPB, ranging from a second application in the fourth week, depending on the patient's pain level, to weekly applications during the first four weeks. This introduces significant variability in pain control and management across studies, increasing heterogeneity, particularly when analyzed over longer follow-up periods, especially after 4 weeks. Another reason for high heterogeneity is the composition of ESPB. Some studies utilized adjuvant medications alongside local anesthetics during ESPB. For instance, Patil et al.²¹ performed the block using bupivacaine combined with 8 mg of dexamethasone, while Xiang et al.²⁶ employed ropivacaine with 0.3 mL of betamethasone. According to previous meta-analyses^{34,35} and randomized trials³⁵ in other clinical contexts, perineural corticosteroids can prolong the duration of the block and enhance the efficacy of local

A.VAS in 1 week

			ESPB		0	Control		Mean Difference			Mear	nce		
_	Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% CI		IV, F	xed, 95	% CI	
	Abdelwahab 2022	4.2674	2.1407	30	5	2.7245	30	33.0%	-0.73 [-1.97, 0.51]			-		
	Lin 2021	3.307	1.9608	26	4.3213	1.1765	28	67.0%	-1.01 [-1.88, -0.14]			-		
											-			
	Total (95% CI)			56			58	100.0%	-0.92 [-1.63, -0.21]		-	•		
	Heterogeneity: Chi ² = 0	0.13, df =	1 (P = 0	.72); l ²	= 0%					-4	-2		2	4
	Test for overall effect:	Z = 2.53	(P = 0.01)							Favours [ESF	B] Fave	ours [control]	-

B.VAS in 2 weeks

		ESPB		0	Control			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
El-Sayed 2021	5.8201	1.9946	20	7.3201	1.1968	20	49.9%	-1.50 [-2.52, -0.48]	_
Patil 2024	3.3	2.2	20	6.15	0.7	20	50.1%	-2.85 [-3.86, -1.84]	
Total (95% CI)			40			40	100.0%	-2.18 [-3.50, -0.85]	
Heterogeneity: Tau ² = Test for overall effect:	0.64; Chi Z = 3.22	² = 3.39, (P = 0.00	df = 1 1)	(P = 0.07); l ² = 71	%			-4 -2 0 2 4 Favours [ESPB] Favours [control]

C. VAS in 4 weeks

		ESPB			Control		Mean Difference			Mean Difference			
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% C	0	IV, Fixe	d, 95% CI		
Abdelwahab 2022	2.5	2.22	30	4	1.48	30	38.7%	-1.50 [-2.45, -0.55]					
El-Sayed 2021	3	1.5957	20	5	1.5957	20	36.1%	-2.00 [-2.99, -1.01]					
Patil 2024	3.4	2.6	20	6.15	0.74	20	25.2%	-2.75 [-3.93, -1.57]					
Total (95% CI)			70			70	100.0%	-1.99 [-2.59, -1.40]		+			
Heterogeneity: Chi ² =	2.59, df	= 2 (P =)	0.27); 1	² = 23%					-4	-2	0	2	4
Test for overall effect:	2 = 0.50	(P < 0.0	0001)							Favours [ESPB]	Favours [co	ontrol]	

D.VAS in 12 weeks

	ES	Control			Mean Difference			Mean Difference				
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% CI		IV, Fixe	d, 95% CI	
Abdelwahab 2022	1.2674 2	2.1407	30	2.713	3.1137	30	68.5%	-1.45 [-2.80, -0.09]		-		
El-Sayed 2021	0.6799 1	.1968	20	2.2591	4.3882	20	31.5%	-1.58 [-3.57, 0.41]	_		_	
Total (95% CI)			50			50	100.0%	-1.49 [-2.61, -0.37]		-		
Heterogeneity: Chi ² = 0.01, df = 1 (P = 0.91); l ² = 0%										-2 (4
Test for overall effect: Z = 2.61 (P = 0.009)										Favours [ESPB]	Favours [control]	

E. Risk Ratio of PHN in 12 weeks

	ESPB		Control		Risk Ratio		Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	M-H, Fixed, 95% CI
Abdelwahab 2022	8	30	12	30	42.3%	0.67 [0.32, 1.39]	
El-Sayed 2021	2	20	6	20	21.1%	0.33 [0.08, 1.46]	
Lin 2021	4	26	10	24	36.6%	0.37 [0.13, 1.02]	
Total (95% CI)		76		74	100.0%	0.49 [0.28, 0.85]	•
Total events	14		28				
Heterogeneity: Chi ² = 1	1.23, df =	2 (P = (1 01 1 10 100			
Test for overall effect:	Z = 2.55 (P = 0.0	0.0	Favours [ESPB] Favours [control]			

Figure 3 Subgroup analysis of ESPB therapy in patients in the acute phase of herpes zoster across several weeks of follow-up. (A–D) Analysis of pain reduction. (E) Risk ratio for the development of Postherpetic Neuralgia (PHN).

anesthetics. Given the inflammatory nature of herpes zoster-associated pain, the use of corticosteroids in ESPB may further potentiate its effect. Therefore, another factor that could influence and contribute to the observed heterogeneity is the inclusion of studies^{27,26} that utilized perineural administration of methylcobalamin alongside the anesthetic, which may introduce variability in outcomes. The perineural use of methylcobalamin in combination with local anesthetics has been investigated in studies^{36,37} addressing herpes zoster-related pain, demonstrating significant pain relief. Methylcobalamin, the active form of vitamin B12, exhibits neuroprotective effects, facilitates neuronal regeneration, and reduces nerve hyperexcitability, thereby alleviating neuropathic pain. To explore heterogeneity, ESPB was evaluated in the acute phase of the disease as well as in patients with PHN. In the acute stage, the results demonstrated greater homogeneity of outcomes and more prolonged pain control, lasting up to 12 weeks, as assessed by the combined analysis of studies in this phase. This can be attributed to patients in the acute phase who underwent the block experiencing a lower rate of pain chronification, defined as PHN over 12 weeks. In other words, fewer patients in the ESPB group reported persistent pain, leading to a reduction in pain scale scores. This finding is supported by the observed RR of 0.49, as depicted in Figure 2E, which corroborates this assertion. A brief review of the literature analyzing other types of nerve blocks reveals similar findings. A meta-analysis conducted by Kim et al.⁷ reported comparable results regarding



Figure 4 Subgroup analysis of ESPB therapy in patients with Postherpetic Neuralgia (PHN).

the risk of PHN in patients in the acute phase who underwent paravertebral block, a technique that shares a similar mechanism of action with ESPB. In this study, an RR of 0.37 was observed, with a p-value of 0.01.

Regarding the subgroup of patients with PHN, only three studies^{24,26,27} were included, and heterogeneity remained high, primarily due to differences in block application frequency. Ahmed et al.²⁴ performed a single application and observed pain scale improvements only during the first week of follow-up, with outcomes analyzed at 12 weeks. In contrast, Cao et al.²⁷ administered weekly blocks during the first 4 weeks, and Xiang et al.²⁶ applied the block at 2 or 4 weeks based on pain levels, resulting in benefits lasting up to 8 weeks. These variations significantly influenced the combined results, as shown in the forest plot. Notably, heterogeneity was zero at the 1 week follow-up, when all

intervention groups received a single block. However, heterogeneity increased substantially at 4 and 8 weeks, likely due to additional applications in two studies.^{26,27} Despite these differences, the studies demonstrated pain relief, particularly during the first week of treatment. An interesting observation is that repeated applications resulted in prolonged pain control, as indicated by two studies,^{27,26} lasting up to 8 weeks of follow-up, even though the last application was administered in the fourth week. In other words, this approach provided an additional month of pain relief without any further interventions.

The ESPB significantly reduced the use of analgesic medications. The available data from the included studies primarily evaluated pregabalin, acetaminophen, and tramadol, highlighting a substantial reduction in analgesic dependence when ESPB was employed. This reduction in analgesic use

A. Acetamino	phen ι	use d	over	12	weeks
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	ESPB			Control				Mean Difference	Mean Difference	
Study or Subgroup	Mean SD Total		Mean SD Total		Weight	IV, Random, 95% CI	IV, Random, 95% CI			
Ahmed 2022	163	65.03	25	211.86	77.55	25	69.7%	-48.86 [-88.53, -9.19]		
EI-Sayed 2021	61.709	105.709	20	175.635	174.286	20	30.3%	-113.93 [-203.26, -24.59]		
Total (95% CI)			45			45	100.0%	-68.58 [-127.18, -9.97]	•	
Heterogeneity: Tau ² = 873.18; Chi ² = 1.70, df = 1 (P = 0.19); l ² = 41% Test for overall effect: Z = 2.29 (P = 0.02)								-	-200 -100 0 100 200 Favours [ESPB] Favours [control]	

Figure 5 Mean Difference (MD) in the consumption of analgesics and neuroleptic medications.

was tracked in studies for up to 12 weeks. Notably, ESPB in neuropathic conditions, as presented in this review, underscores the need for further research. This is particularly important as avoiding the excessive use of systemic medications with potential adverse effects is critical in clinical practice, especially in elderly patients, often burdened with additional comorbidities and medication. Analgesics such as acetaminophen and tramadol can cause serious side effects when taken in excess.³⁸ Excessive consumption of acetaminophen can be particularly associated with severe liver damage, including acute liver failure, as well as allergic reactions and, in rare cases, severe skin conditions like Stevens-Johnson syndrome.^{39,40} Opioids, on the other hand, are linked to both dependence and addiction, alongside notable side effects such as nausea and constipation.⁴¹

The ESPB, a recently developed therapy by Forero et al.,⁴² has been gaining traction and is based on the injection of anesthetic between the erector spinae muscle and the transverse process of the vertebra, guided by an ultrasound device.⁴³ This technique differs from other types of blocks, such as epidural and paravertebral blocks, due to its ease of execution and relative safety. One feared complication common to any type of anesthetic block is the risk related to the systemic absorption of local anesthetics. which can result in respiratory depression and central nervous system effects. These risks can be mitigated by adhering to the maximum allowable dose of the anesthetic and employing simple measures, such as aspirating the needle before injecting. This review did not identify any major adverse effects, including pneumothorax and hematoma. Minor side effects, such as nausea and dizziness, were comparable between the intervention and control groups.

This study has several limitations that need to be highlighted. Firstly, significant heterogeneity was identified in the combined analysis, possibly reflecting disparities in how the intervention was performed across the studies. To address this issue, a subgroup analysis was conducted; however, due to the limited number of studies and, consequently, the small number of patients analyzed, drawing precise conclusions remains challenging. Another factor that contributed to the heterogeneity was the varying follow-up periods across the studies, with some evaluating the intervention only at 8 weeks while others extended it to a longer period. The frequency of clinical follow-ups also varied, with some studies reporting outcomes weekly, while others used more spaced intervals. Consequently, when the outcomes were combined, this generated a publication bias.

Another notable issue is the imprecision of the statistical calculations. Some studies required the conversion of medians and interquartile ranges into means and standard deviations. Although this was done using the Cochrane-recommended method, such conversions often distort results due to skewed data. Therefore, the findings presented in this study should be interpreted with caution.

The fact that all studies were conducted in only two countries, China and India, introduces a potential bias that limits the generalizability and applicability of the findings. Indeed, research restricted to specific geographic regions may introduce publication biases related to the healthcare systems, demographic characteristics, and genetic factors of the analyzed populations. In terms of study quality, it is important to highlight that two out of the seven studies included were categorized as having a high risk of bias. This stems, in part, from the inclusion of peer-reviewed trials and studies present in the gray literature. While gray literature can provide valuable insights and broaden the scope of the review, its inclusion introduces severe limitations. Specifically, gray literature often contains studies with inconsistent methodological rigor and may not meet the highest standards of quality.

Finally, the outcomes related to analgesic consumption were also notably limited. This is because the studies evaluated only the use of pregabalin, acetaminophen, and tramadol, without considering other medications such as gabapentin and other opioids.

Conclusion

In this review, ESPB appears to be effective in reducing pain and the use of analgesic medications in patients with herpes zoster-related pain. The potential benefits seem to be more prolonged during the acute phase of the disease. However, further studies with standardized ESPB application protocols are necessary to validate and better understand these findings.

Conflicts of interest

The authors declare no conflicts of interest.

Disclosures

All authors report no relationships that could be construed as a conflict of interest. All authors take responsibility for all aspects of the reliability and freedom from bias of the data presented and their discussed interpretation.

Supplementary materials

Supplementary material associated with this article can be found in the online version at doi:10.1016/j.bjane.2025. 844598.

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