

Ultrasound-Guided Thoracic Paravertebral Block as a Sympathetic Blockade for Upper Extremity Neuropathic Pain: A Prospective Pilot Study

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Jeongsoo Kim¹
Ho-Jin Lee^{1,2}
Young-Ju Lee³
Chang-Soon Lee^{1,2,4}
Yongjae Yoo^{1,2,4}
Jee Youn Moon^{1,2,4}

¹Department of Anesthesiology and Pain Medicine, Seoul National University Hospital, Seoul, Republic of Korea;

²Department of Anesthesiology and Pain Medicine, Seoul National University College of Medicine, Seoul, Republic of Korea;

³Department of Anesthesiology and Pain Medicine, SMG-SNU Boramae Medical Center, Seoul, Republic of Korea;

⁴Integrated Cancer Management Center, Seoul National University Cancer Hospital, Seoul, Republic of Korea

Purpose: Ultrasound-guided thoracic paravertebral block (US-TPVB) is considered a treatment option for the management of acute pain in various pain-related conditions. We conducted a prospective pilot study to evaluate the possibility of US-TPVB as a sympathetic blockade in patients with neuropathic pain disorders in the upper extremities.

Patients and Methods: A total of 12 patients underwent US-TPVB between the T2 and T3 paravertebral space with 10 mL of 1% mepivacaine. The temperature change (°C) before and after the procedure was compared between the ipsilateral and contralateral hands. We counted the proportion of patients showing a temperature increase $\geq 1.5^{\circ}\text{C}$ and compared a change in the pain intensity before and after the procedure.

Results: The median increase in the temperature change between the ipsilateral and contralateral hands was 1.54°C (interquartile range, 1.28–2.20). There were seven patients (58.3%) who showed a temperature difference $\geq 1.5^{\circ}\text{C}$ between both hands after the US-TPVB. Eleven patients (91.7%) reported a reduction in pain according to the score on the 11-point numerical rating scale. No serious complications relevant to the procedure were reported.

Conclusion: US-TPVB could be a useful technique for sympathetic blockade in patients with upper extremity pain.

Keywords: neuropathic pain, upper extremity, paravertebral block, sympathetic block, ultrasound

Introduction

Among a plethora of various options for pain management, the sympathetic blockade has been widely used in neuropathic pain disorders, such as complex regional pain syndrome, phantom limb pain, postherpetic neuralgia, and pain due to vascular disease.^{1,2} Although the exact mechanism of pain reduction from the sympathetic block is still unknown, the hyperactive sympathetic outflow and the excitatory interactions between the sympathetic nervous system, sensory neurons, and satellite glial cells have been suggested as the associated pathophysiology of sympathetically maintained pain in response to the blockade.^{3,4}

Empirically, the stellate ganglion block (SGB) is one of the most popular sympathetic blockades to manage facial pain or pain in the upper extremities.^{5,6} SGB, however, may not always guarantee a successful outcome to control the sympathetic outflow of the upper extremity because an anatomic variant like Kuntz bypass exists,⁷ and the procedure usually targets the middle sympathetic ganglion at the C6 spinal level

Correspondence: Jee Youn Moon
Department of Anesthesiology and Pain Medicine, Seoul National University College of Medicine, Seoul 03080, Republic of Korea
Tel +82-2-2072-2462
Fax +82-2-763-9390
Email jymoon0901@gmail.com

in practice rather than the stellate ganglion.⁵ Therefore, the thoracic sympathetic ganglion block (TSGB) at the T2 or T3 spinal levels could be a confirmatory technique for managing pain in the upper extremities. Compared to the SGB, the TSGB involves a complicated procedure as the needle tip should be advanced profoundly into the anterior paravertebral space.⁸ Thus, TSGB is usually conducted under fluoroscopy (FS) or computed tomography (CT) guidance, which results in radiation exposure and inconvenient accessibility to patients and physicians at the bedside. Furthermore, due to the long needle path adjacent to the parietal pleura during TSGB, patients frequently complain of procedure-related pain accompanied by a high risk of pneumothorax.⁹

The thoracic paravertebral block (TPVB) was introduced in 1905; however, it has become popular since the 2000s along with the widespread use of ultrasonography (US).¹⁰ It was proposed as a regional anesthetic technique for the management of acute postoperative pain after thoracic and breast surgery or pain due to multiple fractured ribs instead of chronic pain control.^{11–14} A few reports have suggested that the procedure, when performed at the T2 or T3 spinal levels, could be a potential alternative to TSGB or SGB for the management of chronic pain with a low risk of pneumothorax.^{15–18} Recently, Kim et al reported that the majority of patients (80%) achieved a temperature increase ($\geq 1.5^{\circ}\text{C}$) on the palm after FS-guided TPVB at the T2 spinal level, which was superior to US-guided SGB (20.0%).¹⁶ Although the US is a more popular device with easy accessibility than FS in pain practice, it is still uncertain whether such a high proportion of patients could obtain the same temperature increase after US-guided TPVB (US-TPVB).

In this study, we hypothesized that the US-TPVB could be used as a thoracic sympathetic blockade as FS-guided TPVB was suggested in a previous study.¹⁶ We conducted a prospective case-series study to explore whether US-TPVB increases the temperature of the upper extremities and relieves neuropathic or neuralgic pain. This study was designed as a pilot study before a confirmatory trial to compare the effectiveness of US-TPVB to that of TSGB and SGB in various pain disorders in the upper extremities. We also included a review of the previous literature on TPVB in the management of chronic pain conditions.

Patients and Methods

Study Participants

This prospective case-series study was approved by the Institutional Review Board of Seoul National University

Hospital (IRB No. 1907–177-105) and conducted in accordance with the Declaration of Helsinki. From August 2019 to November 2019, 12 consecutive patients with chronic neuropathic pain in the upper extremities, aged 19–85 years, were included. Written informed consent was obtained from all patients prior to study participation. The exclusion criteria were as follows: (1) patients' refusal; (2) peripheral vascular disease in the upper extremity, such as atherosclerosis, thoracic outlet syndrome, or vasculitis; (3) a history of thoracic sympathetic ganglion neurolysis or radiofrequency ablation; (4) a history of lower cervical or upper thoracic spinal surgery; (5) an implanted spinal cord stimulator for management of upper extremity pain; (6) diaphragmatic paresis or severe respiratory disease; (7) systemic or local infection at the injection site; (8) coagulopathy or allergy to local anesthetics; (9) a deformation at the injection site; and (10) pregnancy.

Procedure and Outcome Measurement

All the US-TPVB procedures were performed by one pain physician (JY Moon). The patients were placed in the prone position and a doughnut-shaped cushion was applied to their forehead, if necessary. The vital signs of all the participants were monitored throughout the entire procedure. After sterilization, pre-scanning using a 1–5 MHz round probe (UMT-400, Mindray, Shenzhen, China) was conducted to identify the T2 paravertebral space in a paravertebral sagittal image (Figure 1A). The key anatomic structures, such as the T2 and T3 transverse processes and their relevant ribs, the superior cost-transverse ligament (SCTL), and the parietal pleura, were examined before the needle was advanced (Figure 1B). Subsequently, a 22-G spinal needle (Taechang Industry, Gongju-si, Chungcheongnam-do, Republic of Korea) was inserted at about 5 cm from the midline into the ipsilateral T2 PVS in a caudocranial orientation with a longitudinal, oblique, in-plane technique. Once the needle was advanced to the TPVS until the SCTL was breached with a pop sensation, 10 mL of 1% mepivacaine was injected after negative aspiration, which was expected to spread to two segments of the TPVS, including the sympathetic ganglia.^{19–21} The local anesthetic was delivered slowly, followed by repeated aspiration to reduce the risk of intravascular administration. The injectate was observed to push the bright hyperechoic pleura away from the needle. After it was confirmed that the injectate was completely administered, the transducer was removed, and the sliding

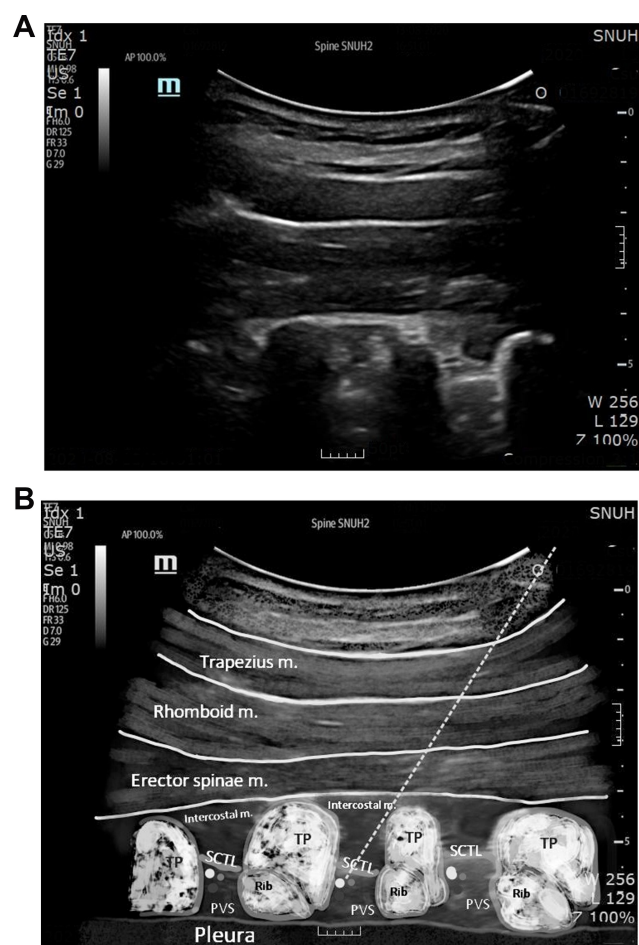


Figure 1 Ultrasound-guided thoracic paravertebral block in a sagittal image. Ultrasound image (A) and schematic image (B) are shown.

Notes: The dotted line indicates the entry path of the needle. After passing the SCTL, 10 mL of 1% mepivacaine was injected.

Abbreviations: PVS, paravertebral space; SCTL, superior costotransverse ligament; TP, transverse process; m, muscle.

movement of the pleura was assured. After a 30-min observation with the confirmation of no severe adverse events, the patients were allowed to leave the pain center.

The participants' temperatures were measured in both palms with an infrared imaging thermometer (FLIR® TG165TM, The World's Sixth Sense®, Wilsonville, OR, USA). The measurements were obtained immediately before and 20 min after the procedure by a research nurse, unaware of which side each patient received TPVB on. To ensure consistency between participants, a point 3 cm below the palmar aspect of the third metacarpophalangeal joint (hand) was used as a landmark. The temperature measurements were recorded at a perpendicular angle and a fixed distance of 30 cm with a recently calibrated infrared imaging thermometer. The difference in temperature change (°C) between the ipsilateral and contralateral

hands at 20 min after the US-TPVB was calculated using the formula [Post-treatment temperature of the block side – Pretreatment temperature of the block side] – [Post-treatment temperature of the other side – Pretreatment temperature of the other side]. If the difference was $\geq 1.5^{\circ}\text{C}$, we defined the procedure to be successful.²² The proportion of patients reaching $\geq 1.5^{\circ}\text{C}$ increase in the temperature in the ipsilateral hand compared to the contralateral hand was also recorded.

Pain intensity in each patient in the upper extremity before and 20 min after the US-TPVB was compared using an 11-point numerical rating scale (NRS) pain score. Other information, such as the duration of pain (months), diagnosis relevant to upper extremity pain, prescribed analgesics, and other comorbidities, were also collected. The possible complications (vascular puncture, hypotension, pleural puncture, pneumothorax, seizure, ipsilateral or bilateral Horner syndrome, ipsilateral brachial plexus block, or hemidiaphragmatic paresis)^{10,23} were evaluated throughout the observation.

Continuous data were expressed as median (interquartile range, IQR) or mean (standard deviation, SD) and categorical data as proportions (%).

Review of Previous Studies

The previous literature was searched to investigate TPVB in patients with neuralgic or neuropathic pain disorders. We excluded studies conducted for the purpose of preemptively controlling pain after surgery. The keywords {"paravertebral" AND ("complex regional" OR "neuralgia" OR "neuropathic")} were inserted in PubMed, and the relevant literature was identified. Our review was restricted to clinical studies conducted in humans and only adults aged ≥ 18 years. Accessible articles in English and all types of studies, such as case reports, observational studies, retrospective studies, and randomized control studies, were considered. After the search, two board-certified pain physicians (J Kim and Y-J Lee) independently screened the abstracts and the titles of articles that could be eligible for this review and excluded the articles that did not match our goals.

Results

The demographic data and clinical parameters of the 12 patients in this study are shown in Table 1. Among the 12 patients, complex regional pain syndrome was the most common diagnosis ($n = 5$), followed by postherpetic neuralgia ($n = 4$), peripheral nerve injury ($n = 2$), and

Table 1 Demographics and Clinical Parameters of the Study Participants

No.	Age (Years)	Sex	BMI (kg/m ²)	Smoking (Yes/No)	Diagnosis	Duration of Pain (Months)	Block Site (Right/Left)	Pre-NRS	Post-NRS
1	60	Male	25.1	No	PHN	60	Right	3	1
2	64	Female	24.2	No	PHN	74	Right	7	7
3	66	Female	18.8	No	PSSS	68	Left	8	5
4	63	Male	23.7	No	PNI	60	Right	7	3
5	19	Female	18.1	No	PHN	48	Left	9	0
6	60	Female	26.8	No	PNI	70	Right	4	1
7	52	Female	30.5	No	CRPS	21	Left	6	2
8	72	Female	24.6	No	PHN	13	Right	7	1
9	41	Male	19.4	Yes	CRPS	78	Right	8	6
10	58	Female	26.3	No	CRPS	43	Left	7	4
11	42	Male	21.4	Yes	CRPS	40	Right	5	2
12	49	Male	22.8	Yes	CRPS	28	Right	7	5

Note: Pre-NRS refers to the pain intensity before the procedure, and post-NRS means the pain intensity 20 minutes after the procedure.

Abbreviations: BMI, body mass index; CRPS, complex regional pain syndrome; NRS, numerical rating scale; PHN, post-herpetic neuralgia; PNI, peripheral nerve injury; PSSS, post-spinal surgery syndrome.

postspinal surgery syndrome ($n = 1$). The median difference in the temperature change between the ipsilateral arm and the contralateral arm was 1.54°C (IQR, $1.28\text{--}2.20^{\circ}\text{C}$). A difference in temperature change $\geq 1.5^{\circ}\text{C}$ between the ipsilateral and contralateral hands was detected in seven patients (58.3%) (Figure 2). The median 11-point NRS pain score was 7 (IQR, $5.5\text{--}7.2$) at baseline and decreased to 2.5 (IQR, $1.0\text{--}5.0$) at 20 min after the US-TPVB. Overall, the patients ($n = 11$, 92%) reported pain reduction followed by the procedure, except for one patient who reported the same pain intensity after the procedure. Ipsilateral ptosis was observed in two patients; however, there were no cases of pneumothorax, hypotension,

vascular puncture, seizure, or other complications during and after the procedure.

Among a total of 92 articles searched by the keywords in PubMed, we eliminated 62 pieces of literature due to subjects irrelevant to our study and duplicate articles (Figure 3). Out of the remaining articles ($n = 30$), 22 were excluded because the majority of them focused on preemptive or postoperative surgical pain control ($n = 18$), two articles were not conducted in humans, one article was review literature, and one was not in English. Finally, eight articles regarding TPVB in patients with neuropathic or nerve-related pain disorders were included in our review (Table 2), which comprised a randomized cross-over study

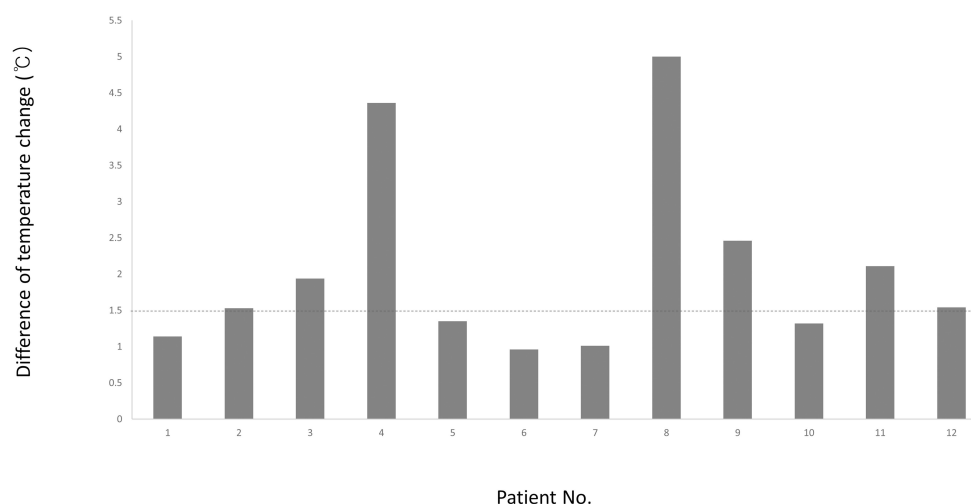


Figure 2 The difference of temperature change between the ipsilateral and contralateral hands at 20 minutes after the ultrasound-guided thoracic paravertebral block.
Note: The horizontal line drawn at 1.5°C indicates the success criterion.

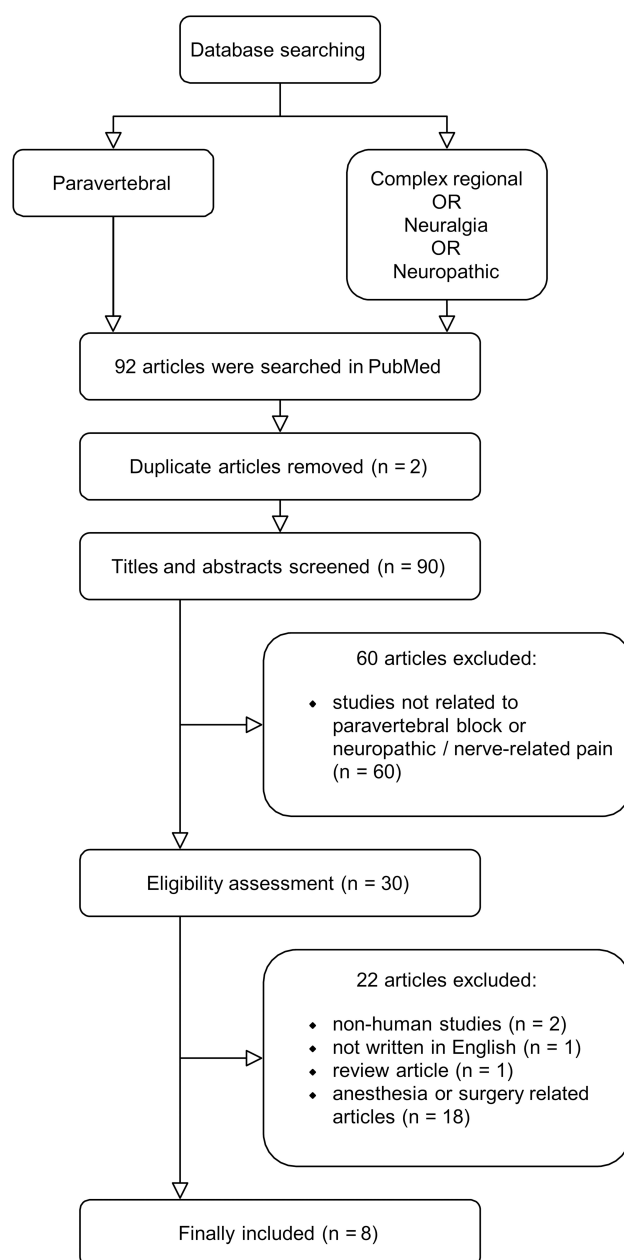


Figure 3 Flow diagram of included previous studies.

(n = 1),¹⁶ randomized controlled trials (n = 4),^{24–27} a case-series study (n = 1),²⁸ and case reports (n = 2).^{18,29} Among them, three were performed on herpes zoster,^{25–27} two on postherpetic neuralgia,^{28,29} two on complex regional pain syndrome,^{16,18} and one on intercostal neuralgia.²⁴ All the studies reported a reduction in the intensity of pain after the procedure. There were only two studies that used the US device during the TPVB at the affected PVS in patients with pain from postherpetic neuralgia or herpes zoster.^{27,28} Two studies were performed to block the thoracic sympathetic ganglia;^{16,18} one was conducted using an FS with

a small volume of local anesthetics (5 mL),¹⁸ and the other was performed using an FS-guided TPVB to compare their effectiveness to US-guided SGB.¹⁶ The latter described that FS-guided TPVB at the T2 spinal level was more effective in blocking thoracic sympathetic ganglia with greater pain relief.¹⁶

Discussion

This prospective case-series study investigated TPVB under US guidance, with 10 mL of local anesthetics, as an alternative to blocking the thoracic sympathetic ganglia in chronic neuropathic pain of the upper extremities. In our study, over half of all patients (n = 7, 58.3%) showed a significant increase in temperature ($\geq 1.5^{\circ}\text{C}$) after the procedure. Almost all patients (n = 11, 92%) experienced pain reduction in their upper extremities 20 min after the procedure.

The thoracic paravertebral space is a potential wedge-shaped compartment adjacent to the posterolateral vertebral body. It is continuous with the intercostal space laterally and the epidural space medially. The dorsal and ventral branches of the spinal nerve and intercostal nerve traverse posteriorly to the paravertebral space. Anteriorly, it contains the rami communicantes, hemi-azygos vein, and sympathetic trunk.^{10,30,31} Hence, the TPVB could produce ipsilateral, segmental, somatic, and sympathetic nerve blockade. In addition, the space extends cranially and caudally,^{19,32} which was the case in at least two patients of our study, resulting in Horner syndrome with ptosis after US-TPVB. Previous cadaver studies demonstrated that US-guided injection with 10–20 mL of contrast dye was visible in the intercostal space to the sympathetic chain with cranial and caudal extensions.^{19,32} In those studies, the paravertebral spreads of contrast media were highly variable, which might contribute significantly to analgesic effects in each patient. Besides, the approach may affect the spread of the injectates in US-TPVB, as reported in a recent cadaveric study.³³ In the study, an intercostal approach covered the intercostal area rather than reaching the sympathetic chain anteriorly.³³ So far, previous clinical and cadaveric studies have focused on the intercostal and epidural spreading for managing post-operative or acute pain rather than blocking the sympathetic chain.^{19,30,32} Therefore, it was necessary to investigate whether TPVB could be used as the thoracic sympathetic blockades for the management of chronic pain. Our result suggests the possibility of US-TPVB at the T2 and T3 spinal levels to be used for the TSGB.

Table 2 Previous Studies of Paravertebral Block for Neuropathic or Nerve-Related Pain

	Dx	Study Design	Groups (N)	Device	Technique	Dose	Outcome
Kim et al ¹⁶	CRPS	Randomized cross-over study	G1: SGB (15) G2: TPVB (15)	US for SGB, FS for TPVB	SGB at the C6 level TPVB at the T2 level	SGB: 5 mL of 1% LDC TPVB: 10 mL of 1% LDC	The % of successful outcome ($\Delta T \geq 1.5^\circ\text{C}$) was higher in G2 (80%) than in G1 (20%, $p < 0.05$). The 11-pointed NRS pain scores after the procedure were lower in G2 (G1, 5.3 ± 1.2 vs G2, 3.9 ± 1.3 ; $p < 0.05$).
Xiao et al ²⁴	Intercostal neuralgia	RCT	G1: PG (30) G2: TPVB (30) G3: PG + TPVB (30)	NSt	TPVB at the affected PVS	PG: 150–600 mg/day TPVB: 20 mL (NS + 0.75% RPV 5 mL + betamethasone 1 mL + Vitamin B12 1 mg)	After 6 weeks, The VAS pain score in G3 was lower than those in G2 and G1 (G1, 26.5 ± 0.7 vs G2, 34.5 ± 0.7 vs G3, 12.4 ± 0.9 ; $p < 0.005$)
Gungor et al ¹⁸	CRPS	Case report	N = 1	FS	TPVB at the T2 PVS	5 mL of 0.125% BPV	The successful sympathetic blockade was achieved ($\Delta T > 2.0^\circ\text{C}$), and pain and tingling senses were reduced by 50% immediately after the procedure.
Zhao et al ²⁸	PHN	Case series	N = 27	US	TPVB at the affected PVS	10 mL (NS + 0.2% methylene blue 2 mL + 0.75% RPV 5 mL)	The VAS pain score was decreased from 7.6 ± 1.5 to 1.6 ± 1.3 at 2 weeks ($p < 0.05$)
Naja et al ²⁹	PHN	Case report	N = 1	NSt	TPVB at the T1 PVS (a bolus) and T3 PVS (via a catheter)	20 mL (0.5% BPV 19 mL + 150 μg clonidine 1 mL) injected through a catheter every 48 h for 3 weeks	The patients with 7–8/10 on the VAS pain score at baseline achieved pain-free after the procedure for an 8-month follow-up period.
Makharita et al ²⁵	HZ	RCT	G1: P (68) G2: TPVB (70)	FS	TPVB at the affected PVS	P: 10mL NS TPVB: 10 mL (NS + 8mg dexamethasone + 0.5% BPV 5 mL)	The VAS pain score was lower in the G2 at 3 weeks (G1, 1.5 ± 1.9 vs G2, 0.5 ± 1.4 ; $p < 0.001$). Total PG and AAP consumptions were reduced in G2 vs G1 (both $p < 0.001$).
Ji et al ²⁶	HZ	RCT	G1: M (68) G2: M + PVB (64)	NSt (PVB)	PVB: Affected PVS	M: 800 mg acyclovir 5 times/d for 7 days + diclofenac 50 mg up to 4 times/d PVB: 10 mL (0.25% BPV + 40 mg methyl-prednisolone) injected every 48 h for a week	The % of patients with pain decreased in G2 than G1 at 1 month (G1, 45% vs G2, 13%; $p < 0.001$). The incidence of PHN also decreased, and at 12 months, the % of patients with pain was different (G1, 16% vs G2, 2%; $p < 0.05$).

(Continued)

Table 2 (Continued).

	Dx	Study Design	Groups (N)	Device	Technique	Dose	Outcome
Zhao et al ²⁷	HZ	RCT	G1: M (44) G2: M + TPVB (43)	US	TPVB at the affected PVS	M: antiviral drug and nutritional therapy TPVB: 10 mL (0.2% methylene blue 2 mL + 0.75% RPV 5 mL + NS)	The VAS pain score decreased at 1 week in G2 than G1 (G1, 5.9 ± 1.5 vs G2, 2.9 ± 1.6 ; $p < 0.05$).

Note: Values are presented as mean \pm standard deviation or a number or proportion (%).

Abbreviations: AAP, acetaminophen; BPV, bupivacaine; CRPS, complex regional pain syndrome; FS, fluoroscopy; G, group; HZ, herpes zoster; LDC, lidocaine; M, medication; NRS, numerical rating scale; NS, normal saline; NST, nerve stimulator; P, placebo; PG, pregabalin; PHN, postherpetic neuralgia; PVB, paravertebral block; PVS, paravertebral space; RCT, randomized controlled trial; RPV, ropivacaine; SGB, stellate ganglion block; TPVB, thoracic paravertebral block; US, ultrasound; VAS, visual analogue scale.

In order to perform a sympathetic blockade of the upper extremity, US-guided SGB or FS-guided TSGB has been mainly implemented.^{34,35} However, SGB is incomplete due to Kuntz bypass^{36,37} and TSGB has a higher risk of pneumothorax because the needle should reach the anterior paravertebral space (Figure 4).^{10,20} For this reason, TSGB usually requires FS guidance. Otherwise, US-TPVB has an advantage for physicians and patients as it is conducted at the bedside without radiation exposure. In addition, TPVB targets the posterior paravertebral space, which lowers the risk of complications, such as pneumothorax. However, because the thoracic sympathetic ganglia are located anteriorly in the paravertebral space, blockade of these ganglia after US-TPVB is uncertain. In our study, after US-TPVB

using 10 mL of local anesthetics, only seven participants (58.3%) showed an increase $\geq 1.5^\circ\text{C}$ on their ipsilateral hands compared to the contralateral hands, which was considered as a meaningful increase after sympathetic blockade in previous studies.^{5,16,22} Presumably, 10 mL of local anesthetics would not be enough to reach the target sympathetic ganglia in some cases. Therefore, future studies could be conducted to investigate the adequate volume of injectates enough to soothe thoracic sympathetic ganglia during the US-TPVB because higher volumes of local anesthetics are commonly used during the US-guided procedure in the thoracic paravertebral areas.^{30,38}

Our brief review noted a few studies reporting the effectiveness of TPVB in treating several neuropathic or

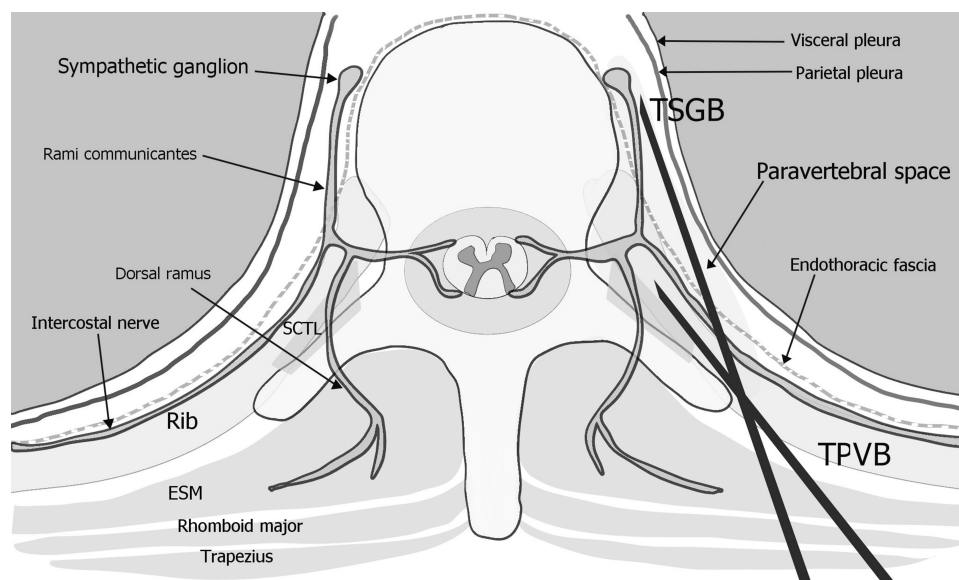


Figure 4 Schematic axial image of the thoracic paravertebral space.

Note: The needle of the TPVB towards the posterior paravertebral space, and the needle of the TSGB towards the sympathetic ganglia at the anterior paravertebral space.

Abbreviations: ESM, erector spinae muscle; SCTL, superior costotransverse ligament; TPVB, thoracic paravertebral block; TSGB, thoracic sympathetic ganglion block.

nerve-related pain (Table 2). Among those pieces of literature, a recent study by Kim et al described that FS-guided TPVB using 10 mL of 1% lidocaine achieved a temperature increase $\geq 1.5^{\circ}\text{C}$ in 80% of subjects ($n = 12/15$).¹⁶ They used the same volume (10 mL) as that of our study. However, technically, all injectates were administered after confirming needle tips using contrast media under FS guidance, which might result in higher success rates in their study. Otherwise, using the US, real-time imaging helps physicians detect the location of the needle tip. However, during the procedure, an initial volume of the injectates may quickly spread outside the superior costotransverse ligament, out of the paravertebral space. In this case, the physicians need to advance the needle a little further, and subsequently, the remaining volume should be administered. We suggest that US-TPVB should be performed in two steps, such as initial hydro-dissection with normal saline (<1 mL) to verify the needle position followed by the second step of injection with local anesthetics, which could improve the accuracy of the procedure.

This study has several limitations. First, it was a case series report with a narrative review to show the possibility of US-TPVB as an alternative to TSGB. Therefore, the head-to-head studies to compare US-TPVB and TSGB or SGB are necessary with reasonable sample size. Second, among several sympathetic functions, we applied only a difference in temperature changes using infrared thermography. The other measurements using the changes in skin conductance or the amount of sudomotor activity could help confirm the complete sympathetic blockade after the procedure.³⁹ Third, in terms of the technical issue, we advanced the needle using a longitudinal parasagittal image in a caudocranial orientation. The other technique, such as using a lateral-to-medial approach in a transverse image, should be investigated to achieve the same result. Although almost all the patients in this study, who were diagnosed with neuropathic or nerve-related pain in their upper extremities, reported pain reduction at 20 min after the block, the clinical and long-term effectiveness of US-TPVB should be investigated in various chronic pain conditions.

Conclusion

In conclusion, although further confirmatory studies are warranted, this case-series study suggested that US-TPVB might be a useful method for managing patients with neuropathic pain in the upper extremity.

Acknowledgments

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Disclosure

The authors declare no conflicts of interest.

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