## **Clinical Research Article**



# Splanchnic nerve neurolysis via the transdiscal approach under fluoroscopic guidance: a retrospective study

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<sup>1</sup>Department of Pain Management, The Fourth Affiliated Hospital of Harbin Medical University, Harbin, China <sup>2</sup>Department of Pain Management, The Second Affiliated Hospital of Harbin Medical University, Harbin, China <sup>3</sup>Department of Anesthesiology, The Fourth Affiliated Hospital of Harbin Medical University, Harbin, China

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Department of Pain Management, The Fourth Affiliated Hospital of Harbin Medical University, No. 37, Yiyuan Street, Nangang District, Harbin 150001, Heilongjiang, China Tel: +86-451-85939542 Fax: +86-451-85939730 E-mail: zhouhuacheng@163.com **Background:** Neurolytic celiac plexus block (NCPB) is a typical treatment for severe epigastric cancer pain, but the therapeutic effect is often affected by the variation of local anatomical structures induced by the tumor. Greater and lesser splanchnic nerve neurolysis (SNN) had similar effects to the NCPB, and was recently performed with a paravertebral approach under the image guidance, or with the transdiscal approach under the guidance of computed tomography. This study observed the feasibility and safety of SNN via a transdiscal approach under fluoroscopic guidance.

**Methods:** The follow-up records of 34 patients with epigastric cancer pain who underwent the splanchnic nerve block via the T11-12 transdiscal approach under fluoroscopic guidance were investigated retrospectively. The numerical rating scale (NRS), the patient satisfaction scale (PSS) and quality of life (QOL) of the patient, the dose of morphine consumed, and the occurrence and severity of adverse events were recorded preoperatively and 1 day, 1 week, 1 month, and 2 months after surgery.

**Results:** Compared with the preoperative scores, the NRS scores and daily morphine consumption decreased and the QOL and PSS scores increased at each postoperative time point (P < 0.001). No patients experienced serious complications. **Conclusions:** SNN via the transdiscal approach under flouroscopic guidance was an effective, safe, and easy operation for epigastric cancer pain, with fewer complications.

**Key Words:** Abdominal Pain; Alcohols; Autonomic Nerve Block; Cancer Pain; Celiac Plexus; Neoplasms; Nerve Block; Neurolysis; Splanchnic Nerves.

## **INTRODUCTION**

Severe abdominal pain occurs mostly in patients with upper abdominal cancers, such as pancreatic cancer, gastric cancer, and liver cancer, at the middle and terminal stages. The therapy methods include palliative tumor treatment, the World Health Organization three-step analgesic lad-

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der, and neurolytic techniques [1,2]. Most cancer pain can be relieved by drugs, but approximately 20% of patients cannot achieve satisfactory analgesia with a higher dose of opioid drugs or have a poor quality of life (QOL) due to the serious side effects of the drugs, such as nausea, constipation, and urinary retention [3].

Blocking the signal transduction of visceral pain has

Author contributions: Zhenhua Cai: Writing/manuscript preparation; Xiaolin Zhou: Writing/manuscript preparation; Mengli Wang: Investigation; Jiyu Kang: Writing/manuscript preparation; Mingshuo Zhang: Formal analysis; Huacheng Zhou: Investigation. been an effective method for treating upper abdominal pain, and the neurolytic celiac plexus block (NCPB) has been the main technique used since 1914 [4]. The NCPB has been performed with different clinical techniques according to the puncture route, the site of the needle tip, and the guidance modality (fluoroscopy, computed tomography [CT], ultrasound), including the percutaneous retrocrural, transcrural, or transaortic approaches as well as gastric endoscopic approaches [5-7]. A satisfactory therapeutic effect is determined by an ideal drug distribution, but the celiac plexus might be distorted by enlarged tumors or lymph nodes, resulting in an unsatisfactory distribution of the neurolytic agent [4,8].

The splanchnic nerve is the main origin of the celiac plexus, which consists of the nociceptive afferent fibers of the upper abdominal parenchymatous organ and is situated posterior to the diaphragmatic crura, piercing the crura of the diaphragm at the T11-12 levels to join the celiac ganglion [1]. The retrocrural space is a potential closed triangular space, and the effect of splanchnic nerve neurolysis (SNN) is rarely influenced by the surrounding structures [8]. Several trials have indicated that SNN results in better pain relief, less opioid consumption, fewer complications, and a better QOL compared with the NCPB [1,9,10]. Furthermore, patients have longer pain relief and more satisfaction after SNN compared with NCPB [11]. The primary puncture method for SNN is the retrocrural or paravertebral approach, guided by fluoroscopy or CT. The paravertebral pathway under fluoroscopic guidance has a risk of pneumothorax and vascular injury [5]. Thus, SNN under CT guidance has more advantages, and the transdiscal approach could further avoid injury to the paraspinal organs [12]. However, patients receive a larger dose of radiation under CT guidance, and not all hospitals are qualified to use CT guidance.

Therefore, the authors aimed to perform SNN via a transdiscal approach under fluoroscopic guidance and observe the feasibility and safety of this technique for treating epigastric cancer pain.

## **MATERIALS AND METHODS**

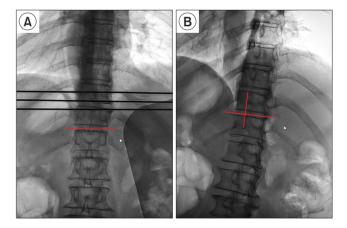
#### 1. Participants

This was a retrospective observational study of patients with upper abdominal cancer pain who underwent the splanchnic nerve block or the SNN via the T11-12 transdiscal approach under fluoroscopic guidance. Patients admitted from October 2016 to October 2019 at the Fourth Affiliated Hospital of Harbin Medical University were enrolled. Inclusion criteria included (1) diagnosis of abdominal pain and referred pain of the back that was secondary to cancer; (2) severe pain with a numeric rating scale (NRS) score  $\geq$  7 after pharmacotherapy, intolerable side effects from opioid therapy, or a desire for further treatment; and (3) more than 50% relief in pain intensity with diagnostic splanchnic nerve block. Exclusion criteria included coagulation dysfunction, local or systemic infection, uncontrolled hypotension, poor cardiac or poor pulmonary function, intestinal obstruction, and patients' refusals to participate. The study protocol was approved by the Ethics Committee of the Fourth Affiliated Hospital of Harbin Medical University (Protocol No: 2020-WZYSLLSC-05) and the study was conducted in accordance with the Declaration of Helsinki and its later amendments.

### 2. Procedure

Each patient had fasted for 6 hours and was water deprived for 2 hours, and the narcotics were stopped on the procedure day. The patient was placed in a prone position with a pillow under the upper abdomen, and a peripheral intravenous line was established. The patient was given inhaled oxygen at 3 L/min with a nasal cannula and received a rapid infusion of 500 mL of 0.9% saline. Vital signs were monitored, including electrocardiogram, blood pressure, and oxygen saturation. Preoperatively, 0.02 mg/kg midazolam was intravenously injected for sedation.

The vertebral bodies of T11 and T12 were identified in the anteroposterior (AP) view, and the image intensifier was adjusted to keep the inferior and superior endplates



**Fig. 1.** The location of puncture site. (A) The inferior and superior endplate of the T11-12 intervertebral disc kept in a line (red line) in the anteroposterior view. The 3 black lines were Kirschner wires for preliminary positioning assistance. (B) The X-ray tube ball was then rotated to an oblique position ipsilateral, made the tip of the superiorarticular process of T12 point to the midpoint of the T11 vertebral body in the oblique view. The intersection of the 2 red lines was the puncture point on the skin.

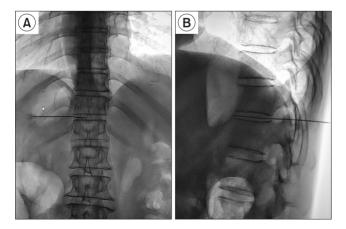
of the T11-12 intervertebral disc in a line (Fig. 1A). The fluoroscopy tube was then rotated to an oblique ipsilateral position, making the tip of the superior articular process of T12 point to the midpoint of the T11 vertebral body (Fig. 1B). The puncture point on the skin was the projection of the intersection of the middle point of the lateral margin of the superior articular process of the T12 and the T11-12 intervertebral disc (the first position point). The ideal final position of the needle tip would be at the middle of the disc in the AP view and at the front edge of the vertebral body in the lateral view.

Before the puncture, the needle (22 gauge, 150 mm) was curved approximately 10 degrees at the first proximal cm from the needle tip. Using coaxial technology, after reaching the first landing point, the needle tip was slipped over the upper joint and entered into the disc. When the tip of the needle approached the front edge of the vertebral body, the stylet was pulled out, and a 1 mL syringe with 0.2 mL saline was connected. Then, the needle was pushed forward carefully and stopped immediately when injection resistance in the syringe disappeared. The fluoroscopy was then adjusted to confirm whether the needle tip was in the correct position (Fig. 2). After the needle tip reached the correct position, a 5 mL mixture of contrast agent and local anesthetic, which contained iohexol 2.5 mL (concentration 300 mgI/mL) and the mixture of 1% lidocaine + 0.5% ropivacaine 2.5 mL, was injected with an increment of 1 mL. The ideal spread would be seen as a "honeycomb" covering the bilateral vertebral body in the AP view and confined to the prevertebral tissue plane covering the bilateral T11-12 vertebral body in the lateral view (Fig. 3). If the contrast agent was distributed on a single side, the needle was withdrawn and used to perform a second puncture to supplement the distribution of the agent, till the drug distribution was satisfactory. In rare cases, a second needle from the ipsilateral or contralateral side was used to supplement the distribution of the agent. The patient was returned to the ward if the vital signs were stable and no complaints of discomfort were made after 10 minutes of observation. The patient was ordered to maintain the prone position for 2 hours and then moved to a supine position for 24 hours.

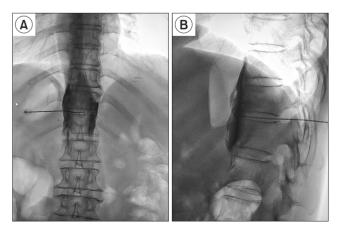
A diagnostic block was performed the day before with a total of 5 mL 1% lidocaine and 0.5% ropivacaine. If the effect of the diagnostic block was satisfactory, the SNN was carried out with an identical procedure the next day. After the contrast agent was distributed satisfactorily, a total 5 mL 99.7% alcohol (Tianjin Tianli Chemical Reagent Co., Ltd., Tianjin, China) was injected within 2 minutes, and then 0.5 mL of 0.9% normal saline was injected before the withdrawal of the needle to avoid alcohol flow into the path of the puncture.

#### 3. Outcome measures

The degree of pain was evaluated by the NRS (0-10), where a score of 0 means no pain and a score of 10 indicates the maximum level of intolerable pain, and the QOL, in which a score of 10 was the best and a score of 0 was the worst [13], prior to the procedure and at 1 day, 1 week, 1 month, and 2 months postprocedure. The daily morphine equivalent was recorded during the follow-up. The incidences of side effects or complications related to the operation were recorded, including pneumothorax, back pain, hypotension, diarrhea, discitis, and neurological effects, such as nerve injury and paresthesia. Patient satisfaction was evaluated using the patient satisfaction scale (PSS), which recorded the improvement in symptoms, such as pain relief, som-



**Fig. 2.** Ideal position of the needle. (A) Anteroposterior view, the tip of the needle arrived at the midline, which was bilateral vertebral arch inner range. (B) Lateral view, with the needle tip at the anterior border of the vertebral body.



**Fig. 3.** Ideal spread of the contrast agent. (A) Anteroposterior view, the ideal spread would be seen as a "honeycomb" covering the bilateral vertebral body. (B) Lateral view, contrast agent was confined to the prevertebral tissue plane covering the bilateral T11-12 vertebral body.

Indexes	Preoperative	Postoperative			
	(n = 31)	1 d (n = 31)	1 wk (n = 31)	1 mo (n = 30)	2 mo (n = 29)
QOL scores	1.2 ± 1.0	$5.3 \pm 0.8^{a}$	$6.8 \pm 1.2^{\circ}$	$7.2 \pm 0.9^{a}$	7.3 ± 1.1ª
PSS scores	$1.3 \pm 0.5$	6.5 ± 1.1°	$7.2 \pm 1.0^{\circ}$	$6.8 \pm 2.5^{\circ}$	7.2 ± 2.1 <sup>a</sup>
NRS scores	7.6 ± 2.1	$3.7 \pm 1.0^{\circ}$	$2.6 \pm 0.7^{\circ}$	$2.8 \pm 0.6^{\circ}$	$2.9 \pm 0.8^{\circ}$
Daily consumption of morphine (mg/d)	182 ± 36	$90 \pm 14^{a}$	$52 \pm 7^{a}$	60 ± 11ª	$58 \pm 9^{a}$

Values are presented as mean ± standard deviation.

QOL: quality of life, PSS: patient satisfaction scale, NRS: numeric rating scale.

 $^{a}P < 0.001$ , compared with the patients in the preoperative time.

nolence, operation time, intestinal function, recovery of appetite, and weight gain, and was assessed by the patient with a linear analog scale (with 0 indicating very satisfied and 10 indicating very dissatisfied) [12].

#### 4. Statistical analysis

Statistical analysis was performed with the Statistical Package for Social Sciences (SPSS) (ver. 20.0; IBM Co., Armonk, NY). Numerical data are expressed as the means  $\pm$  standard deviations. Repeated measures analysis of variance was performed for repeated measurements. The level of significance was set at P < 0.05.

## RESULTS

A total of 40 patients were assessed for eligibility. Four patients did not meet the inclusion criteria (mainly due to laboratory test abnormalities or changes in the characteristics of the pain, such as neuropathic and/or somatic pain), and 2 patients declined to take part in SNN due to the potential surgical risks. Thirty-four patients were finally enrolled in the trial. The mean age of the included patients (22 males and 12 females) was 60 ± 12 years (range 48-74 years). The procedures were performed for pain secondary to pancreatic cancer (18 patients), gastrointestinal tract cancer (6 patients), and hepatobiliary cancer (10 patients). Three of the patients had an unsatisfactory effect from the diagnostic splanchnic nerve block, and they only underwent the splanchnic nerve block. Thus, 31 patients underwent both the diagnostic splanchnic nerve block and the SNN, and the number of transdiscal operations was 65. The average operating time was  $15.8 \pm 5.2$  minutes. Seventeen (26.2%) patients had a nonideal distribution of contrast agent, 14 (21.5%) of which were satisfactorily resolved through adjustment of the needle direction, and the other 3 through a second needle puncture, resulting in a single needle success rate of 95.4%.

Compared with the preoperative NRS score, the post-

 Table 2. The complications in patients who underwent splanchnic nerve

 neurolysis (n = 31)

Complication	Number of cases (%)
Burning pain in the abdomen	3 (9.7)
Diarrhea	4 (12.9)
Hypotension	3 (9.7)
Transient backache	5 (16.1)
Pneumothorax	0
Discitis	0
Paresthesia	0

operative NRS score decreased significantly (Table 1). The NRS score at 1 week after the operation was  $2.6 \pm 0.7$ , which was lower than the preoperative score ( $7.6 \pm 2.1$ ) (P < 0.001), and was maintained at a low level by the final 2 months follow-up. Simultaneously, the postprocedure daily morphine equivalent (mg/d) (1 day, 1 week, 1 month, 2 months) decreased significantly after SNN from 182  $\pm$  36 mg to 52  $\pm$  7 mg (at 1 week) (Table 1). Two patients even stopped taking morphine 1 week after the SNN until the end of follow-up. Unfortunately, one patient died at 1 month and another at 7 weeks. The QOL and PSS scores increased after SNN, especially after 1 week (Table 1).

No serious complications occurred in any patient, including shock, intractable hypotension, pneumothorax, discitis, transient paresthesia, and paraplegia. Three patients (9.7%) suffered from burning pain in the abdomen, which diminished after 24 hours. Four patients (12.9%) suffered from mild diarrhea, which was alleviated in 2–3 days with treatment, including aggressive hydration (oral or parenteral) and antidiarrheal agents. Three patients (9.7%) had postural hypotension, which resolved within 24–48 hours. Five patients (16.1%) suffered from transient backache, which was relieved after taking nonsteroidal anti-inflammatory drugs (**Table 2**). No discomfort was found in three patients who underwent the diagnostic block.

## DISCUSSION

In this trial, SNN was performed successfully via a novel transdiscal approach under fluoroscopic guidance. The SNN procedure was performed over a shorter operating time, and the patients received less radiation. Meanwhile, the patients felt satisfied, as indicated by the decreased morphine intake and increased PSS and QOL scores. A few patients had transient complications, such as postural hypotension, transient backache and mild diarrhea, from which they recovered quickly. Most importantly, no serious complications occurred, including nerve injury and pneumothorax.

NCPB is an effective therapeutic option for epigastric cancer pain [14,15]. Edelstein et al. [16] reported that among 87 patients who had undergone an NCPB, 35 (40%) patients received a major or complete sustained reduction in pain. However, in advanced malignancies, the celiac plexus anatomy may be distorted by the underlying malignancy or the celiac lymph nodes enlarged; thus, the target of nerve damage can be hard to reach or the spread of neurolytic agents may be limited [8,17]. Our data confirmed that the spread of the agent was confined to the front edge of the T11-12 vertebral body and was not affected by the abdominal tumors. Additionally, the performance of an NCPB through a transaortic, retrocrural pathway might lead to complications, including paraplegia, abdominal aortic dissection, renal injury, hematuria, or intravascular injection. Sometimes, NCPB cannot be performed in patients with accompanying severe systemic diseases [4,10].

The thoracic splanchnic nerve lies in a small triangular space with well-defined landmarks and boundaries, and has a less variable anatomical relationship with surrounding structures; thus, SNN is easier to perform compared to a conventional NCPB. Süleyman Ozyalçin et al. [10] found that SNN was an alternative to the NCPB. Ahmed and Arora [8] also confirmed that SNN was an effective alternative to the NCPB in patients whose celiac anatomy was distorted by tumors. Kapural et al. [11] performed the NCPB and SNN at different time intervals on the same patient and found that SNN could deliver local anesthetic to the paravertebral compartment medial to the pleural cavity, which was in close proximity to the greater and lesser splanchnic nerves; thus, SNN provided much longer pain relief than the NCPB. Plancarte et al. [12] proved that SNN could provide analgesia and decrease morphine consumption in patients with upper abdominal malignancies.

SNN is usually performed via a paravertebral approach under fluoroscopic and CT guidance or a transdiscal approach under CT guidance [12]. The final position of the needle tip in the paravertebral approach lies in the anterior third of the vertebral body on the lateral fluoroscopic

view [1]. The blocking solution is thus more easily spread into the intervertebral foramen, leading to paresthesia and paraplegia and even a high risk of pricking the pleura and paravertebral vessels. Plancarte et al. [12] proved that the SNN under CT guidance through a transdiscal approach with a unilateral puncture resulted in a bilateral block. However, patients usually receive a large dose of radiation under CT guidance, and not all hospitals are qualified to use CT guidance. Therefore, we performed SNN through the T11-12 intervertebral disc under fluoroscopic guidance. In addition, the curved needle technique made the puncture direction easier to adjust if the drug distribution was unsatisfactory. In the early stage, we used another needle to remedy deficiencies in drug distribution according to previous reports [8,11]. After the curved needle was applied with expertise, one needle could complete the operation.

The dose of nerve neurolysis varies greatly in the clinical practice and trials, and the ultimate goal is to achieve complete drug coverage of the involved nerve while avoiding excessive dosage and serious complications. The alcohol dose (approximately 12-20 mL) and concentration (50%-100%) resulting in SNN in clinical studies are often mentioned [3,18]. Ahmed and Arora [8] used 6 mL of 50% alcohol on both sides (a total of 12 mL), while Shwita et al. [1] and Amr et al. [19] applied 10 mL of 70% and 100% alcohol, respectively, on each side (a total of 20 mL) to perform the neurolysis. The overdose of alcohol could spread to adjacent structures, and lead to more serious complications, including paresthesia or paraplegia [20,21]. In this study, 5 mL contrast agent did not spread to the first half of the vertebral body, and was able to meet the clinical requirement. Usually, higher concentrations of alcohol have a more destructive effect. Higher concentrations and lower doses of alcohol were used in this study to enhance the nerve block effect and reduce complications.

The agents currently used for chemical neurolysis are alcohol, phenol, adriamycin, and methylene blue. Adriamycin mainly acts on the dorsal root ganglia and has a slow onset [22], while methylene blue has a short acting time. Alcohol acts by denaturing proteins, extracting fatty substances, and precipitating lipoproteins and mucoproteins, damaging both Schwann and nerve cells, and resulting in Wallerian degeneration. Phenol is primarily a local anesthetic at lower concentrations and becomes more neurolytic at higher concentrations [23], which could produce a block lasting 3-6 months [3]. Koyyalagunta et al. [4] found that alcohol and phenol had no difference in complications and pain improvement. In the present study, 99.7% alcohol was chosen, which is safe according to previous literature and can provide a more effective block than other concentrations.

The incidence of mild diarrhea (12.9%), postural hypotension (9.7%), and transient backache (16.1%) in this trial were less than those in a previous study, which was 23%–30%, 19%–50%, and 27%, respectively [1]. Plancarte et al. [12] found that SNN via the transdiscal approach had a minor risk of complications (*e.g.*, pneumothorax) compared with traditional approaches, including the retrocrural and paravertebral approaches, because of the use of a single needle through the disk and a lower volume of neurolytic agent. Simultaneously, the transdiscal approach was a safer route for the needle pathway for avoiding traumatic injury to the arteries branching to the Adamkiewicz artery [12]. In addition, no opioid withdrawal syndrome was found during this trial, which might be related to a reduced dose of opioids was applied to assist analgesia.

In brief, the main advantages of the transdiscal approach were as follows: (1) it avoids alcohol flow into the intervertebral foramen, which might result in paresthesia and nerve root neurolysis; (2) it avoids injury to the lumbar arteries (which can lead to paraplegia) and other nearby organs, such as the liver, kidney, intestine, pancreas, *etc.*; and (3) the puncture needle is relatively fixed, and the operation is easy, effective, and safe.

Discitis is the main concern for the transdiscal approach. Recent studies have proven that the risk of discitis is very low, and could be further reduced by the use of prophylactic antibiotics [20,23,24]. Furthermore, no discitis was previously found during SNN and superior hypogastric plexus neurolysis through the intervertebral disc approach [12,25]. Although the probability of discitis is low, some measures should be taken, such as decreased puncture times, improved operation skills, and prophylactic antibiotics, to further reduce the risk of this complication.

There are some limitations in our study: (1) Although the punctures were 100% successful, we still required the use of a second needle (3 out of 65) at the early stage of the procedure when the drug distribution was not ideal. (2) This trial had no prospective design, and the sample size was small; thus, the success rate of the puncture and the complication ratio require further observations. Simultaneously, the different effects among different types of cancer were not observed. (3) There was no comparative study between the SNN and traditional NCPB, and the effects of different drugs and different doses of alcohol were not observed in this trial. (4) Although there were fewer complications in this puncture approach, the ultrasoundassisted puncture might be safer, including avoiding the possibility of pleural puncture and vascular injury.

In conclusion, SNN performed through the transdiscal approach under fluoroscopic guidance is a safe and easy operation that has many advantages, including accurate positioning, being a more targeted operation, and producing minimal complications. This method could be popularized to treat abdominal cancer pain.

## **CONFLICT OF INTEREST**

No potential conflict of interest relevant to this article was reported.

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