INTERVENTIONAL RADIOLOGY



Computed tomography-guided single celiac plexus neurolysis analgesic efficacy and safety: a systematic review and meta-analysis

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Abstract

Purpose To perform a systematic review and meta-analysis of published studies to evaluate the analgesic efficacy and safety of computed tomography (CT)-guided single celiac plexus neurolysis (CPN) with the injection of a neurolytic agent into the celiac plexus in one session (CT-guided single CPN).

Methods PubMed, the Cochrane Library, and Ichushi-Web were searched for English or Japanese articles published up to February 2022, which reported findings about patients who underwent CT-guided single CPN. The outcome measures assessed in the systematic review and meta-analysis were the pain measurement scales from 0 to 10 before and after the intervention and the rate of minor and major complications.

Results The pooled pain measurement scales at pre-intervention and 1- or 2-, 7-, 30-, 60-, 90-, and 180-day post-intervention was 6.72 (95% confidence interval [CI], 4.77–9.46, $l^2 = 98\%$), 2.31 (95% CI 2.31–4.44, $l^2 = 92\%$), 2.84 (95% CI 1.39–5.79, $l^2 = 95\%$), 3.36 (95% CI 1.66–6.77, $l^2 = 98\%$), 3.19 (95% CI 1.44–7.08, $l^2 = 59\%$), 3.87 (95% CI 1.88–7.97, $l^2 = 0\%$), and 3.40 (95% CI 3.02–3.83, $l^2 =$ not applicable), respectively. The pooled minor complication rates of diarrhea, hypotension, nausea or vomiting, and pain associated with the procedure were 18% (95% CI 8–37%, $l^2 = 45\%$), 16% (95% CI 2–58%, $l^2 = 76\%$), 6% (95% CI 2–16%, $l^2 = 1\%$), and 7% (95% CI 2–21%, $l^2 = 17\%$), respectively. There was no major complication in the included studies.

Conclusion CT-guided single CPN can be performed safely and provides immediate analgesic efficacy although the amount of heterogeneity is characterized as large. Further investigation of its long-term analgesic efficacy is required.

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Graphical abstract



Keywords Celiac plexus neurolysis · Interventional radiology · Meta-analysis · Pain management · Palliative care · Systematic review

Introduction

Celiac plexus neurolysis (CPN) is an intervention for ablating the neural network of the celiac plexus with the objectives of palliating chronic abdominal pain owing to malignant and benign conditions, including pancreatic cancer, inflammatory bowel disease, and chronic pancreatitis, as well as reducing the need for narcotic analgesics [1, 2]. CPN is performed using X-ray fluoroscopy [3], endoscopic ultrasound (EUS) [4], magnetic resonance imaging [5], or computed tomography (CT) [2]. Although each of these modalities has its unique advantages and disadvantages, CT-guided CPN is adopted by interventional radiologists as it is particularly advantageous for clearly visualizing retroperitoneal structures and tumor involvement, locating the needle tip, and avoiding damage to vital organs and vessels [2, 6].

Several variants of CT-guided CPN have been reported, such as single CPN with the injection of neurolytic agents into the celiac plexus in one session (CT-guided single CPN), consecutive CPN with multiple injections of neurolytic agents into the celiac plexus through an indwelling catheter, or cryoablation of the celiac plexus [7]. Among them, CT-guided single CPN is the most widely used technique. Although the role of this technique had already been established, its analgesic efficacy and safety have not been assessed in any meta-analysis. The purpose of the present study is to perform a systematic review and meta-analysis of published studies to evaluate the analgesic efficacy and safety of CT-guided single CPN.

Materials and methods

The systematic review and meta-analysis were performed in accordance with the guidelines of the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA). No formal approval was required at our institution for this type of study.

Literature search and study selection criteria

A literature search of PubMed, the Cochrane Library, and Ichushi-Web (Igaku Chuo Zasshi; Japan Medical Abstracts Society) was systematically conducted using relevant MeSH terms and keywords among the articles published up to February 2022 (Supplement Table). The literature search was carried out with the assistance of librarians. The inclusion criteria were as follows: (1) availability of full-text articles; (2) articles reporting data from CT-guided single CPN; and (3) articles written in English or Japanese. The exclusion criteria were as follows: (1) case reports; (2) review articles; (3) letters and editorials; (4) articles with a sample size of less than 10 cases; (5) articles with no extractable data; and (6) articles with data included in subsequent articles or duplicate reports. Two authors (T.M. and R.Y.) independently conducted the literature search and article selection. If the reviewers disagreed, consensus was reached after discussion with a third reviewer (T.Y.).

Data extraction and quality assessment

For each selected article, we extracted the following data: baseline data for the article (first author; publication year; study period; countries; study design), total patient characteristics (number of patients; age; sex; malignancy or non-malignant disease), data concerning the CT-guided single CPN protocol (number of patients included in this study; type of sedation; patient position; needle gauge; local anesthetic prior to injecting neurolytic agent or not; contrast injection before CT-guided single CPN or not; local anesthetic mixed with neurolytic agent or not; neurolytic agent; amount of neurolytic agent), pain measurement scales at pre-intervention and 1- or 2-, 7-, 30-, 60-, 90-, and 180day post-intervention, and complications. The complications were classified in accordance with the classification system of the Cardiovascular and Interventional Radiological Society of Europe, i.e., from grade 1 (no complication) to grade 6 (death) [8]. Specifically, grades 1 and 2 were defined as minor complications, whereas grades 3-6 were defined as major complications.

The quality of the included studies was assessed using the Cochrane Collaboration tool for randomized clinical trials and the Risk of Bias Assessment tool for Non-randomized Studies (RoBANS) [9]. Both data extraction and quality assessment were performed independently by two reviewers (T.M. and R.Y.), with any disagreement resolved after discussion with a third reviewer (T.Y.).

Statistical analysis

The primary outcome was the change in the 0–10 visual analog scale (VAS) or 0–10 numeric rating scale (NRS) before-and-after CT-guided single CPN to evaluate its analgesic efficacy. The secondary outcome was the rate of major and minor complications to evaluate the safety of CT-guided single CPN. The pooled pain measurement scale and the rate of major and minor complications with their 95% confidence intervals (CIs) were computed using a random-effects model based on the DerSimonian–Laird method. Heterogeneity among studies was evaluated by testing Cochran's Q statistic and the inconsistency index (I^2) statistic. For Cochran's Q test, values of q < 0.05 were considered significant. For the I^2 statistic, values of < 25% were defined as low heterogeneity, 25–50% were defined as moderate, and > 50% were defined as high heterogeneity. Egger's test was used

to analyze publication bias; values of p < 0.1 were considered significant. Meta-regression analysis was conducted to identify the source of inter-study heterogeneity. A value of p < 0.05 identified the source of heterogeneity. Egger's test and meta-regression analysis were performed if at least 4 articles were selected for each meta-analysis. The metapackage of R software (version 4.1.2; R Foundation for Statistical Computing) was used for statistical analyses.

Results

Article selection and quality assessment

From a total of 66 articles returned by the database search, 19 articles underwent a full-text review. Then, 12 reports were excluded on the basis of the eligibility criteria [10–21]. Finally, 7 articles were selected for the systematic review and meta-analysis of the analgesic efficacy and rate of major and minor complications of CT-guided single CPN (Fig. 1) [7, 22–27]. Furthermore, 5 of the included articles were case–control studies (n=2) and retrospective before-and-after studies (n=3), while 2 studies were randomized control trials (RCTs). The quality of the selected RCTs (n=2), case–control studies (n=2), and retrospective before-and-after studies (n=3) was assessed as some concerns, unclear, and high, respectively (Table 1).

Characteristics of the included studies

The 7 articles selected involved a total of 381 cases. The average-weighted mean age was 60 years, with 223 (59%) men and 158 (41%) women (Table 2). In the included articles, 79% had pancreatic cancer and 2% had non-malignant diseases, such as pancreatitis, persistent gastric ulceration, and median arcuate ligament syndrome.

In 3 of the 7 included articles, diazepam or midazolam was administered as sedation and the other articles were not clearly described (Table 3). Anterior or posterior techniques were described to access the celiac plexus with 18-to-23-gauge needles in the included articles (Table 3). Ethanol was used as a neurolytic agent in all the included articles. The average-weighted ethanol amount was 26 mL (range: 10– 40 mL). In 3 of the 7 articles, a local anesthetic (lidocaine) was injected immediately before ethanol injection and the other articles were not clearly described (Table 3). Iodinated contrast media were injected before injecting ethanol in all the included articles (Table 3). In 2 of the 7 articles, a local anesthetic (bupivacaine or lidocaine) was mixed with ethanol and the other articles were not clearly described (Table 3).



Fig. 1 Flow diagram

Table 1	Study characteristics of	of
the inclu	ided studies	

Study	ROB	Year	Study period	Study type	Location
De Cicco et al. [27]	High	1997	1989–1994	Before-after study	Italy
Lee et al. [26]	High	2000	1995-1998	Before-after study	Korea
Zhang et al. [25]	Some concerns	2008	2000-2006	RCT	China
Arai et al. [24]	Unclear	2013	2007-2008	Case-control study	Japan
Behbahani et al. [7]	Unclear	2020	2014-2019	Case-control study	US
Neuwersch-Sommereg- ger et al. [22]	High	2021	2010-2019	Before-after study	Austria
Abdelbaser et al. [23]	Some concerns	2022	2017-2019	RCT	Egypt

ROB risk of bias, RCT randomized controlled trial

Table 2 Patie	ent characteri	stics of the	included	studies

Study	Total patients	Mean age	Range	Sex (M)	Sex (F)	Primary n	nalignancy				Non- malignant disease
						Pancreas	Stomach	Colon	Liver	Other	
De Cicco et al. [27]	53	57	28–79	34	19	38	6	3	3	3	0
Lee et al. [26]	28	65	36-82	12	16	10	8	1	3	6	0
Zhang et al. [25]	56	58	38–75	35	21	56	0	0	0	0	0
Arai et al. [24]	36	68	NR	17	19	36	0	0	0	0	0
Behbahani et al. [7]	83	60	29–79	39	44	56	3	7	1	7	9
Neuwersch-Som- meregger et al. [22]	55	65	24-88	35	20	34	4	0	0	17	0
Abdelbaser et al. [23]	70	56	23–77	51	19	70	0	0	0	0	0

NR not reported

In 2 of the 7 articles, the numeric rating scale was used and in the remaining 5 articles, the visual analog scale was used for the assessment of pain. The number of articles for which the mean and standard deviation data of the pain measurement scales at pre-intervention and 1- or 2-, 7-, 30-, 60-, 90-, and 180-day post-intervention were provided were 5, 4, 4, 4, 2, 2, and 2, respectively (Table 4). Minor complications were described in detail in 5 articles (Table 5). There was no major complication in the included articles.

Meta-analysis

The pooled pain measurement scale at pre-intervention and 1- or 2-, 7-, 30-, 60-, 90-, and 180-day post-intervention was 6.72 (95% confidence interval [CI] 4.77–9.46, $l^2 = 98\%$, p < 0.01), 2.31 (95% CI 2.31–4.44, $l^2 = 92\%$, p < 0.01), 2.84 (95% CI 1.39–5.79, $l^2 = 95\%$, p < 0.01), 3.36 (95% CI 1.66–6.77, $l^2 = 98\%$, p < 0.01), 3.19 (95% CI 1.44–7.08, $l^2 = 59\%$, p = 0.12), 3.87 (95% CI 1.88–7.97, $l^2 = 0\%$, p = 0.35), and 3.40 (95% CI 3.02–3.83, $l^2 =$ not applicable), respectively (Figs. 2, 3). Egger's test showed a significant publication bias for pre-intervention (Fig. 4a) (p = 0.032). There was no significant publication bias for 1- or 2-, 7-, and 30-day post-intervention (Fig. 4b, c, d) (p = 0.84, 0.99, and 0.62, respectively).

The pooled rates of diarrhea, hypotension, pain associated with the procedure, and nausea or vomiting were 18% (95% CI 8–37%, $I^2 = 45\%$, p = 0.12), 16% (95% CI 2–58%, $I^2 = 76\%$, p < 0.01), 7% (95% CI 2–21%, $I^2 = 17\%$, p = 0.07), and 6% (95% CI 2–16%, $I^2 = 1\%$, p = 0.40), respectively (Fig. 5). Egger's test showed a significant publication bias for diarrhea, hypotension, pain associated with the procedure, and nausea or vomiting (Fig. 6) (p = 0.024, 0.021, 0.006, and 0.006, respectively).

Meta-regression analysis

There was no evidence that the publication year, malignancy in all patients or not, total sample size, mean age, gender, type of pain measurement scale, and amount of ethanol were associated with values of the pain measurement scales, diarrhea, and hypotension in meta-regression analysis (Table 6).

Discussion

The systematic review and meta-analysis demonstrated that CT-guided single CPN immediately reduces the pain measurement scores, and the effect seems to be sustained for at least 7 and 30 days after the intervention. The effect may be sustained for 60-, 90-, and 180-day post-intervention; however, further investigation of the long-term analgesic efficacy is required owing to the small number of studies. There was no significant difference in pain control with the ethanol

	Study No. of												
$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	study	patients ed in this	Type of sedation	Patient po	sition	Needle gaug	e Local : prior to neuroly	anesthetic o injecting ytic agent	Contrast injection CPN	media before	Local anesthet mixed with neu lytic agent	c Neurolytic agent tro-	Amount of ethanol (mL)
	De Cicco et al. [27] 53		Diazepam	Anterior		20, 22	Lidoca (Imme befor	uine diately re CPN)	Yes		Bupivacaine	Ethanol	30
	Lee et al. [26] 28		NR	NR		18, 21, 22	Lidoca (Imme befor	uine diately re CPN)	Yes		NR	Ethanol	30
Arai e al. [24]19NRPosterior23NRYesNREthanol14Behchaun e al. [23]34NRAnterior23NRYesEthanol40Newessers (al. [23]34MidacolamPosterior18NRYesEthanol20Newessers (al. [23]34MidacolamPosterior18NRYesEthanol20McMeser et al. [23]34MidacolamPosterior18NRYesEthanol20McMoser et al. [24]Anterior18NRYesYesEthanol20McMoser et al. [25]34NRYesYesNRNRPosteriorMc nerger et al. [26]9NRNRNRNRNRNRMc nerger et al. [26]9NRNRNRNRNRNRMc nerger et al. [26]9NRNRNRNRNRNRMc nerger et al. [26]9NRNRNRNRNRNRMot setter et al. [26]9NRNRNRNRNRNRMot setter al. [27]810271127123413006Mot setter al. [27]9NRNRNRNRNRNRNRMot setter al. [26]910112711271234130.6006Mot setter al. [27]93127 <td< td=""><td>Zhang et al. [25] 29</td><td></td><td>Diazepam</td><td>Posterior</td><td></td><td>23</td><td>Lidoca (Immed befor</td><td>uine diately re CPN)</td><td>Yes</td><td></td><td>NR</td><td>Ethanol</td><td>20</td></td<>	Zhang et al. [25] 29		Diazepam	Posterior		23	Lidoca (Immed befor	uine diately re CPN)	Yes		NR	Ethanol	20
Betholmain et al. [1] 33 NR NR Zeamoly of the constraints of the constraint of the constraints of the constraint of the constraints	Arai et al. [24] 19		NR	Posterior		23	NR		Yes		NR	Ethanol	14
Neuversch-Som- meregeret al. [22] NR Anterior 23 NR Ethanol 10 Modekseret al. [23] Midazolan Restrict 18 N Yes Idocatine Ethanol 20 Mothesseret al. [23] Midazolan Restrict 18 N Yes Idocatine Ethanol 20 Mr na reported File Di N N N Yes Indocatine Ethanol 20 Mr an reported Fain Sub Pain N N N N N N Mr and reported Fain Sub Day 1 or 2 SD Day 3 Day 3 Day 3 Day 3 Motion 10 27 N NR NR NR NR NR NR Material 1 21 11 27 12 34 1 N N N Locate al. [26] 9 NR NR NR NR NR NR NR Material 1 21 1 27 62 NR NR NR Material 1 21 1 23 31 NR NR NR Material<	Behbahani et al. [7] 33		NR	NR		22	NR		Yes		NR	Ethanol	40
Motolescental_23 34 Midazolam Posterior 18 NR Yss Lidocaine Ethanol 20 Minor reported Amon 20 No 1 No 1	Neuwersch-Som- 47 meregger et al. [22]		NR	Anterior		23	NR		Yes		NR	Ethanol	10
$\label IT and reported the reported for the reported state and after CT-guided single cellac plexus neurolysis (CPN) of the included state scale from 0 to 10) To 10 and the resonand after CT-guided single cellac plexus neurolysis (CPN) of the included state scale from 0 to 10) To 2.7 1.7 4.4 2.7 6.2 2.2 NR $	Abdelbaser et al. [23] 34		Midazolam	Posterior		18	NR		Yes		Lidocaine	Ethanol	20
Pre SD Day 1 or 2 SD Day 3 SD Day 90 SD Day 180 SD De Cicco et al. [27] 8 1.0 2.7 1.7 4.4 2.7 6.2 2.2 NR S0 S0 <td< th=""><th>Study Pair</th><th>1 measurer</th><th>nent scale (from 0</th><th>to 10)</th><th></th><th></th><th></th><th></th><th></th><th></th><th></th><th>Remarks</th><th></th></td<>	Study Pair	1 measurer	nent scale (from 0	to 10)								Remarks	
$ \begin{array}{cccccc} De Cicco et al. [27] & 8 & 1.0 & 2.7 & 1.7 & 4.4 & 2.7 & 6.2 & 2.2 & NR & VAS \\ Lee et al. [26] & 9 & NR $	Pre	SD	Day 1 or 2 SI	Day 7	SD	Day 30 Si	D Day 60	SD	Day 90 S	SD Da	iy 180 SD		
Lee et al. [26]9NRNRNRNRNRNRNRNRVASZhang et al. [25]9.40.71.30.81.71.12.71.23.413.91.2NRNRNR90 ($n=27$), Day 90 ($n=25$)Arai et al. [24]NRNRNRNRNRNRNRNRNRNRNRArai et al. [24]NRNRNRNRNRNRNRNRNRNRBehbahani et al. [7]52.33.72.73.93.12.51.9NRNRNRNRNRBehbahani et al. [7]52.33.72.73.93.12.51.9NRNRNRNRNRNeuversch-Sommereg-5.11.92.42.0NRNRNRNRNRNRNR90 ($n=5$)Neuversch-Sommereg-5.11.92.42.0NRNRNRNRNRNRNR90 ($n=5$)Ser et al. [22]4.91.3NR2.31.02.70.93.01.0NRNR3.030.90 ($n=5$)Abdelbaser et al. [23]6.91.3NR2.31.02.70.93.01.0NRNR3.03.03.0Abdelbaser et al. [23]6.91.3NR2.41.02.70.93.01.0NRNR3.03.01.0NR	De Cicco et al. [27] 8	1.0	2.7 1.5	7 4.4	2.7	6.2 2.	.2 NR	NR	NR	NR NF	k NR	VAS	
Zhang et al. [25] 9.4 0.7 1.3 0.8 1.7 1.1 2.7 1.2 3.4 1 3.9 1.2 NR NR NR 90, 60 (n=27), Day 90 (n=25) Arai et al. [24] NR 90, 60 (n=27), Day 90 (n=25) Arai et al. [24] NR 90, 60 (n=27), Day 90 (n=25) Behbahani et al. [7] 5 2.3 3.7 2.7 3.9 3.1 2.5 NR 30, 50 (n=25), Day 90 (n=25) 30, 50 (n=27), Day 90 (n=25) 30, 50 (n=26) 30, 50 (n=26) 30, 50 (n=5) NN 30, Day 90 (n=5) NR 30, Day 90 (n=5) 30, Day 90 (n=5) 30, Day 90 (n=5) NR 30, Day 90 (n=5) NR </td <td>Lee et al. [26] 9</td> <td>NR</td> <td>NR NI</td> <td>R NR</td> <td>NR</td> <td>NR N</td> <td>'R NR</td> <td>NR</td> <td>NR</td> <td>NR NF</td> <td>R NR</td> <td>VAS</td> <td></td>	Lee et al. [26] 9	NR	NR NI	R NR	NR	NR N	'R NR	NR	NR	NR NF	R NR	VAS	
Arai et al. [24] NR S0, Day 90 (n=5) 30, Day 90 (n=5) 30, Day 90 (n=5) S0, Day 90 (n=2) S0, Day 9	Zhang et al. [25] 9.4	0.7	1.3 0.8	3 1.7	1.1	2.7 1.	.2 3.4	1	3.9 1	I.2 NF	k NR	Day 30, 60 $(n = 27)$, Da	/ 90 (<i>n</i> =25)
Behbahani et al. [7] 5 2.3 3.7 2.7 3.9 3.1 2.5 NR NR Day 2 (n=33), Day 7 (n=14), Day 30 Neuwersch-Sommereg- 5.1 1.9 2.4 2.0 NR NR NR NR NR 90 (n=5) Neuwersch-Sommereg- 5.1 1.9 2.4 2.0 NR MS 90 (n=5) 0.0 49 (n=5) 0.0 49 (n=2) 0.0 49 (n=2) 0.0 1.0 NR NR NR NR NR NR NR NR NR 1.2 VAS ger et al. [22] 6.9 1.3 NR NR 2.0 0.0 0.0 0.0 NR 3.4 1.2 VAS	Arai et al. [24] NR	NR	4.0 NI	R NR	NR	NR N	IR NR	NR	NR	NR NF	k NR	NRS	
Neuwersch-Sommereg- 5.1 1.9 2.4 2.0 NR MA	Behbahani et al. [7] 5	2.3	3.7 2.5	7 3.9	3.1	2.5 1.	.9 NR	NR	2.6 2	2.5 NF	R NR	Day 2 $(n=33)$, Day 7 $(130, 20, 20)$	t=14), Day
Abdelbaser et al. [23] 6.9 1.3 NR NR 2.3 1.0 2.7 0.9 3.0 1.0 NR NR 3.4 1.2 VAS	Neuwersch-Sommereg- 5.1 ger et al. [22]	1.9	2.4 2.() NR	NR	NR	IR NR	NR	NR	AR NF	k NR	NRS	
	Abdelbaser et al. [23] 6.9	1.3	NR NI	R 2.3	1.0	2.7 0.	.9 3.0	1.0	NR	NR 3.4	t 1.2	VAS	

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 Table 5
 Complications of

 CT-guided single celiac
 plexus neurolysis (CPN) of the

 included studies

Study	Minor complic	cation				Major	
	Hypotension	Diarrhea	Vomiting or nausea	Pain	Other	complica- tion	
De Cicco et al. [27]	NR	NR	NR	NR	NR	0	
Lee et al. [26]	5	6	0	0	0	0	
Zhang et al. [25]	13	6	0	4	Drunkenness (2)	0	
Arai et al. [24]	6	0	0	0	0	0	
Behbahani et al. [7]	0	9	4	0	0	0	
Neuwersch-Sommereg- ger et al. [22]	NR	NR	NR	NR	NR	0	
Abdelbaser et al. [23]	1	2	2	3	0	0	

NR not reported

injection volume in this meta-regression analysis. Moreover, the demographics (age, gender), disease characteristics (malignancy in all patients or not), publication year, and total sample size were not associated with the values of pain measurement scales in this meta-analysis.

In 372 cases (98%) of a total of 381 cases in the 7 included articles, CT-guided single CPN has been performed for chronic abdominal pain associated with malignancy. In particular, chronic abdominal pain associated with pancreatic cancer accounted for 79% of all the patients in the included studies. This result is consistent with the fact that pancreatic cancer is recognized as one of the most painful malignancies with substantial suffering and is often unresponsive to typical medical management [1, 28].

The procedure of CT-guided single CPN shows that anterior or posterior techniques have been described to access the celiac plexus with 18-to-23-gauge needles in the included studies. Among them, 20 gauge or smaller needles were used in most of the included articles, which seems to be sufficient for CT-guided single CPN. Although they have not been mentioned in the included articles, lateral decubitus, posterior intradiscal, and transaortic approaches have been reported [2]. Currently, the choice of these techniques should be individualized on the basis of the operator's preference, patient's anatomy and comorbidities, and extent of the disease. However, because the anterior approach nearly always involves passage through the visceral organs (especially the liver and the stomach), care should be taken to minimize damage to these structures by choosing the shortest route through them, avoiding large vessels and dilated biliary ducts, and minimizing needle repositioning [6]. After iodinate contrast media were injected to determine the region of opacification and ensure the extravascular needle position, ethanol was injected as a neurolytic agent in all the included articles. These results indicate that ethanol is generally the first-choice agent for CT-guided single CPN.

Diarrhea (18% [95% CI 8-37%]) and hypotension (16% [95% CI 2–58%]) were found to be relatively frequent minor complications in this meta-analysis. These expected minor complications are due to the destruction of sympathetic signals, which causes the parasympathetic nervous system to remain unopposed. Diarrhea was transient in the included articles. However, diarrhea is rarely persistent and refractory [29]. Hypotension usually only requires an adequate intravenous bolus of normal saline during or after the procedure [30]. Other common minor complications included pain associated with the procedure (7% [95% CI 2-21%]) and nausea/vomiting (6% [95% CI 2–16%]), which was transient in the included articles. The included articles indicate that it is necessary to consider administering a local anesthetic immediately before ethanol injection or a mixture of ethanol and a local anesthetic because ethanol injection may cause severe temporary pain. There was no major complication in the included articles. Bleeding complications that require

Fig. 2 Forest plot of the pooled pain measurement scales at preintervention and post-intervention. a Forest plot of the overall pooled pain measurement scales at pre-intervention. **b** Forest plot of the overall pooled pain measurement scales at 1- or 2-day post-intervention. c Forest plot of the overall pooled pain measurement scales at 7-day post-intervention. d Forest plot of the overall pooled pain measurement scales at 30-day post-intervention. e Forest plot of the overall pooled pain measurement scales at 60-day post-intervention. f Forest plot of the overall pooled pain measurement scales at 90-day post-intervention. g Forest plot of the overall pooled pain measurement scales at 180-day post-intervention

а	Study	Total	Mean	SD Me	an I	Mean 95	%-CI Weight
	Behbahani et al (2020) Neuwersch-Sommeregger et al (20 Abdelbaser et al (2022) De Cicco et al (1997) Zhang et al (2008)	33 21) 47 34 53 29	5.00 2.30 5.10 1.90 6.90 1.30 8.00 1.00 9.40 0.70	000 000 000	÷	5.00 [4.27; 5.10 [4.58; 6.90 [6.48; 8.00 [7.74; 9.40 [9.15;	5.85] 18.9% 5.67] 19.8% 7.35] 20.3% 8.27] 20.5% 9.66] 20.5%
	Random effects model Heterogeneity: $l^2 = 98\%$, $\tau^2 = 0.0739$,	196 p < 0.01		5 6 7	789	6.72 [4.77;	9.46] 100.0%
b	Study	Total	Mean	SD Me	an I	Mean 95	%-CI Weight
	Zhang et al (2008) Neuwersch-Sommeregger et al (20 De Cicco et al (1997) Behbahani et al (2020)	29 21) 47 53 33	1.30 0.80 2.40 2.00 2.70 1.70 3.40 2.40			1.30 [1.04; 2.40 [1.89; 2.70 [2.28; 3.40 [2.67;	1.63]24.9%3.05]24.7%3.20]25.8%4.33]24.6%
	Random effects model Heterogeneity: $I^2 = 92\%$, $\tau^2 = 0.1544$,	162 p < 0.01		1.5 2 2.5	3 3.5 4	2.31 [1.21;	4.44] 100.0%
С	Study To	tal Mean	SD	Mean	Mean	95%-CI V	Veight
	Zhang et al (2008) Abdelbaser et al (2022) Behbahani et al (2020) De Cicco et al (1997)	29 1.70 34 2.30 14 3.90 53 4.40	1.1000	*	1.70 2.30 3.90 4.40	[1.34; 2.15] [1.99; 2.66] [2.57; 5.91] [3.73; 5.19]	25.3% 26.5% 22.0% 26.3%
	Random effects model 1 Heterogeneity: $I^2 = 95\%$, $\tau^2 = 0$	30).1864, <i>p</i> <	0.01	234	2.84 [1.39; 5.79] 1	00.0%
d	Study To	tal Mean	SD	Mean	Mean	95%-CI V	Veight
	Behbahani et al (2020) Zhang et al (2008) Abdelbaser et al (2022) De Cicco et al (1997)	52.50272.70342.70536.20	1.9000 — 1.2000 0.9000 2.2000		2.50 2.70 2.70 	[1.28; 4.87] [2.28; 3.19] [2.41; 3.02] [5.64; 6.82]	17.4% 27.1% 27.7% 27.8%
	Random effects model 1 Heterogeneity: $I^2 = 98\%$, $\tau^2 = 0$	19).1859, <i>p</i> <	0.01	2 3 4 5	3.36 [[1.66; 6.77] 1	00.0%
е	Study To	tal Mean	SD	Mean	Mean	95%-CI V	Weight
	Abdelbaser et al (2022) Zhang et al (2008)	34 3.00 27 3.40	1.0000 1.0000		3.00 3.40	[2.68; 3.36] [3.04; 3.80]	49.8% 50.2%
	Random effects model Heterogeneity: $I^2 = 59\%$, $\tau^2 = 0$	61 0.0046, <i>p</i> =	0.12 2	1 I I 3 4 5	3.19	[1.44; 7.08] 1	00.0%
f	Study To	tal Mean	SD	Mean	Mean	95%-Cl \	Veight
	Behbahani et al (2022) Zhang et al (2008)	5 2.60 25 3.90	2.5000 — 1.2000		2.60 3.90	[1.12; 6.04] [3.46; 4.40]	2.0% 98.0%
	Random effects model Heterogeneity: $I^2 = 0\%$, $\tau^2 = 0$,	30 <i>p</i> = 0.35	1	2 3 4 5 6	3.87 [7 8	(1.88; 7.97] 1	00.0%
g	Church	T-4-1 PT					
	Study) 34 3	an SL	wiean	Me:	40 [3.02:3.8	31

 $I^2 = not applicable$

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blood transfusion following fluoroscopic-guided CPN [31], lower-extremity paralysis following EUS-guided CPN [32], and thrombosis of the celiac trunk leading to hepatic, splenic, gastric, or bowel infarction following EUS-guided CPN [33] have been reported as rare major complications. Moore et al. strongly recommended CT guidance, which precisely locates the position of the needle's point and bevel immediately before injection of the neurolytic agent, thereby avoiding complications [34]. However, publication bias was found with respect to minor complications in the meta-analysis. In other words, the minor complications reported in some studies may be underestimated. Thus, our understanding of the risks associated with CT-guided single CPN may be further limited by the underreporting of minor and severe complications.

This study has some limitations. First, the number of included studies was limited and only 7 studies that satisfied the inclusion criteria were included. In particular, the meta-analysis of the pain measurement scales 60- and 90-day post-intervention was performed in only two studies. Further, the mid-term and long-term results of CT-guided single CPN must be confirmed. Second, we could not perform subgroup analysis of the needle size, needle tip position, and access route owing to the small number of included studies. Future research should focus on these factors. Third, comparisons with other modalities, especially EUS-guided CPN, could not be made. Multicenter, large-sample, highquality cohort studies and RCTs for CT-guided single CPN should be included in future. Despite these limitations, this systematic review and meta-analysis can provide useful information for the current clinical practice of CT-guided single CPN.

In conclusion, CT-guided single CPN can be performed safely and provides immediate analgesic efficacy although the amount of heterogeneity is characterized as large.

Fig. 4 Funnel plot of the pooled pain measurement scales at pre-intervention and post-intervention. a Funnel plot of the overall pooled pain measurement scales at pre-intervention. b Funnel plot of the overall pooled pain measurement scales at 1- or 2-day post-intervention. c Funnel plot of the overall pooled pain measurement scales at 7-day post-intervention. d Funnel plot of the overall pooled pain measurement scales at 30-day post-intervention



Fig. 5 Forest plot of the pooled minor complication rate. **a** Forest plot of the overall pooled rate of diarrhea. **b** Forest plot of the overall pooled rate of hypotension. **c** Forest plot of the overall pooled rate of pain associated with the procedure. **d** Forest plot of the overall pooled rate of nausea or vomiting

0.06 [0.02; 0.16] 100.0%

author	Events	Total		Proportion	95%-CI	Weight
Arai et al (2013)	0	19 ⊫–		0.00	[0.00; 0.18]	4.1%
Abdelbaser et al (2022)	2	34 —		0.06	[0.01; 0.20]	13.5%
Zhang et al (2008)	6	29		0.21	[0.08; 0.40]	25.8%
Lee et al (2000)	6	28		0.21	[0.08; 0.41]	25.7%
Behbahani et al (2020)	9	33		0.27	[0.13; 0.46]	30.8%
Random effects model Heterogeneity: $I^2 = 45\%$, τ^2	² = 0.1423	143 5, p = 0.12		0.18	[0.08; 0.37]	100.0%
h		· ·				
D						
author	Events	Total		Proportion	95%-CI	Weight
Behbahani et al (2020)	0	33 -	<u> </u>	0.00	[0.00; 0.11]	12.2%
Abdelbaser et al (2022)	1	34 -+	<u> </u>	0.03	[0.00; 0.15]	16.7%
Lee et al (2000)	5	28		0.18	[0.06; 0.37]	23.3%
Arai et al (2013)	6	19		0.32	[0.13; 0.57]	23.3%
Zhang et al (2008)	13	29		0.45	[0.26; 0.64]	24.6%
Random effects model Heterogeneity: $I^2 = 76\%$, τ^2	2 = 1.7406	143 - 6, <i>p</i> < 0.01 0		0.16	[0.02; 0.58]	100.0%
С						
C author	Events	Total		Proportion	95%-CI	Weight
C author Behbahani et al (2020)	Events 0	Total 33 ≖–	<u> </u>	Proportion	95%-CI [0.00: 0.11]	Weight 8.6%
C author Behbahani et al (2020) Lee et al (2000)	Events 0 0	Total 33 ा— 28 ा—		Proportion 0.00 0.00	95%-CI [0.00; 0.11] [0.00: 0.12]	Weight 8.6% 8.6%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013)	Events 0 0 0	Total 33 ⊷ 28 ⊷ 19 ⊷		Proportion 0.00 0.00 0.00	95%-CI [0.00; 0.11] [0.00; 0.12] [0.00: 0.18]	Weight 8.6% 8.6% 8.6%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022)	Events 0 0 0 3	Total 33 ा— 28 ा— 19 ा— 34 →		Proportion 0.00 0.00 0.00 0.09	95%-CI [0.00; 0.11] [0.00; 0.12] [0.00; 0.18] [0.02; 0.24]	Weight 8.6% 8.6% 8.6% 34.4%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008)	Events 0 0 0 3 4	Total 33 ा−− 28 ा−− 19 1−− 34 −− 29		Proportion 0.00 0.00 0.00 0.09 0.14	95%-CI [0.00; 0.11] [0.00; 0.12] [0.00; 0.18] [0.02; 0.24] [0.04; 0.32]	Weight 8.6% 8.6% 34.4% 39.8%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008) Random effects model Heterogeneity: $l^2 = 17\%$, τ^2	Events 0 0 3 4 2 = 0.1925	Total 33 28 19 34 29 143 0, p = 0.31 0		Proportion 0.00 0.00 0.09 0.14 0.07	95%-Cl [0.00; 0.11] [0.00; 0.12] [0.00; 0.18] [0.02; 0.24] [0.04; 0.32] [0.02; 0.21]	Weight 8.6% 8.6% 34.4% 39.8% 100.0%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008) Random effects model Heterogeneity: $I^2 = 17\%$, τ^2 d	Events 0 0 3 4 2 = 0.1925	Total 33 ा−− 28 п−− 19 п−− 34 −− 29 143 0, p = 0.51 0		Proportion 0.00 0.00 0.09 0.14 0.07	95%-CI [0.00; 0.11] [0.00; 0.12] [0.00; 0.18] [0.02; 0.24] [0.04; 0.32] [0.02; 0.21]	Weight 8.6% 8.6% 34.4% 39.8% 100.0%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008) Random effects model Heterogeneity: /² = 17%, τ ²	Events 0 0 3 4 2 = 0.1929 Events	Total 33 ⊫− 28 ⊫− 19 ⊫− 34 − 29 143 0, <i>p</i> = 0.31 0		Proportion 0.00 0.00 0.09 0.14 0.07 Proportion	95%-CI [0.00; 0.11] [0.00; 0.12] [0.02; 0.24] [0.04; 0.32] [0.02; 0.21]	Weight 8.6% 8.6% 34.4% 39.8% 100.0% Weight
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008) Random effects model Heterogeneity: / ² = 17%, τ ² d author Zhang et al (2008)	Events 0 0 3 4 2 = 0.1925 Events 0	Total 33 28 19 34 29 143 0 Total 29		Proportion 0.00 0.00 0.09 0.14 0.07 Proportion 0.00	95%-CI [0.00; 0.11] [0.00; 0.12] [0.02; 0.24] [0.04; 0.32] [0.02; 0.21] [0.02; 0.21]	Weight 8.6% 8.6% 34.4% 39.8% 100.0% Weight 9.5%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008) Random effects model Heterogeneity: $I^2 = 17\%$, τ^2 d author Zhang et al (2008) Lee et al (2000)	Events $0 \\ 0 \\ 0 \\ 3 \\ 4$ $2^{2} = 0.1929$ Events $0 \\ 0$	Total 33 28 19 34 29 143 0 Total 29 28 0		Proportion 0.00 0.00 0.09 0.14 0.07 Proportion 0.00 0.00	95%-CI [0.00; 0.11] [0.00; 0.12] [0.02; 0.24] [0.04; 0.32] [0.02; 0.21] [0.02; 0.21] [0.00; 0.12] [0.00; 0.12] [0.00; 0.12]	Weight 8.6% 8.6% 34.4% 39.8% 100.0% Weight 9.5% 9.4%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008) Random effects model Heterogeneity: $I^2 = 17\%, \tau^2$ d author Zhang et al (2008) Lee et al (2000) Arai et al (2013)	Events $0 \\ 0 \\ 0 \\ 3 \\ 4$ $2^{2} = 0.1925$ Events $0 \\ 0 \\ 0 \\ 0$	Total 33 28 19 34 29 143 0 Total 29 28 19 29 19 29 19 19		Proportion 0.00 0.00 0.09 0.14 0.07 Proportion 0.00 0.00 0.00 0.00 0.00	95%-CI [0.00; 0.11] [0.00; 0.12] [0.02; 0.24] [0.04; 0.32] [0.02; 0.21] [0.02; 0.21] [0.00; 0.12] [0.00; 0.12] [0.00; 0.12]	Weight 8.6% 8.6% 34.4% 39.8% 100.0% Weight 9.5% 9.4% 9.4%
C author Behbahani et al (2020) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022) Zhang et al (2008) Random effects model Heterogeneity: $I^2 = 17\%, \tau^2$ d author Zhang et al (2008) Lee et al (2000) Arai et al (2013) Abdelbaser et al (2022)	Events $0 \\ 0 \\ 0 \\ 3 \\ 4$ $2^{2} = 0.1925$ Events $0 \\ 0 \\ 0 \\ 2$	Total 33 = 28 = 19 = 34 19 = 34 29 = 143 p = 0.31 0 Total 29 = -28 = -28 19 = -28 19 = -28 = -28 19 = -28 10 = -2		Proportion 0.00 0.00 0.09 0.14 0.07 Proportion 0.00 0.00 0.00 0.00 0.00 0.00	95%-CI [0.00; 0.11] [0.00; 0.12] [0.02; 0.24] [0.04; 0.32] [0.02; 0.21] [0.02; 0.21] [0.00; 0.12] [0.00; 0.12] [0.00; 0.18] [0.01: 0.20]	Weight 8.6% 8.6% 34.4% 39.8% 100.0% Weight 9.5% 9.4% 28.7%

Random effects model 143						
Heterogeneity: $I^2 = 1\%$, $\tau^2 = 0.2078$, $p = 0.4$	0					
()	0.05	0.1	0.15	0.2	0.25

Fig. 6 Funnel plot of the pooled minor complication rate. **a** Funnel plot of the overall pooled rate of diarrhea. **b** Funnel plot of the overall pooled rate of hypotension. **c** Funnel plot of the overall pooled rate of pain associated with the procedure. **d** Funnel plot of the overall pooled rate or vomiting



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Table 6 Exploration ofheterogeneity within the studiesvia meta-regression analysis

	R^2	95% CI	p value
Pre-intervention			-
Publication year	31.93%	$\left(-0.0506 \text{ to } 0.0154\right)$	0.188
Malignancy in all patients or not	11 79%	$\begin{bmatrix} -0.5852 \text{ to } 1.3140 \end{bmatrix}$	0.309
Total sample size	4.36%	$\begin{bmatrix} -0.0455 \text{ to } 0.0224 \end{bmatrix}$	0.359
Mean age	30.54%	$\begin{bmatrix} -0.1570 \text{ to } 0.0505 \end{bmatrix}$	0.201
Gender (male)	20.26%	$\begin{bmatrix} -2.5373 \text{ to } 6.7358 \end{bmatrix}$	0.245
VAS vs NRS	9.90%	$\begin{bmatrix} -0.6052 \text{ to } 1.2929 \end{bmatrix}$	0.332
Day 1 or 2 post-intervention			0.000
Publication year	0%	$\left(-0.0980 \text{ to } 0.1151\right)$	0.763
Malignancy in all patients or not	7.82%	$\begin{bmatrix} -2.4761 \text{ to } 1.4550 \end{bmatrix}$	0.38
Total sample size	0.00%	$\begin{bmatrix} -0.0568 \text{ to } 0.0891 \end{bmatrix}$	0.441
Mean age	0%	(-0.3211 to 0.3664)	0.804
Gender (male)	0%	$\left(-20.4548 \text{ to } 12.7661\right)$	0.424
VAS vs NRS	0%	$\left(-2.5459 \text{ to } 2.4483\right)$	0.941
Amount of ethanol	0%	$\left(-0.0627 \text{ to } 0.0983\right)$	0.441
Day 7 post-intervention			
Publication year	0%	[-0.1226 to 0.1000]	0.705
Malignancy in all patients or not	0%	[-8.8991 to 6.6454]	0.596
Total sample size	0%	(-0.0927 to 0.1048)	0.817
Mean age	0%	[-0.7207 to 0.8548]	0.749
Gender (male)	0%	[-2.9541 to 2.1370]	0.561
Amount of ethanol	55.43%	(-0.0471 to 0.1283)	0.185
Day 30 post-intervention			
Publication year	54.43%	[-0.0918 to 0.0278]	0.148
Malignancy in all patients or not	0%	[-2.4293 to 3.1420]	0.637
Total sample size	15.26%	(-0.1034 to 0.0586)	0.356
Mean age	0%	(- 0.8796 to 0.7105)	0.692
Gender (male)	0%	[-15.9297 to 16.7487]	0.924
Amount of ethanol	0%	(-0.1358 to 0.1560)	0.794
Diarrhea			
Publication year	0%	(-0.1959 to 0.1421)	0.648
Malignancy in all patients or not	0%	(- 3.5523 to 1.8914)	0.403
Total sample size	0%	(-0.0655 to 0.0771)	0.812
Mean age	0%	(-0.4036 to 0.3978)	0.983
Gender (male)	77.78%	(-13.0196 to 6.4167)	0.359
Amount of ethanol	25.82%	(-0.0527 to 0.1599)	0.207
Hypotension			
Publication year	8.34%	(-0.3755 to 0.1444)	0.252
Malignancy in all patients or not	39.47%	(- 3.4912 to 9.3659)	0.242
Total sample size	5.05%	(-0.1535 to 0.0570)	0.241
Mean age	0%	【 − 0.4243 to 0.6678 】	0.529
Gender (male)	0%	(-25.4775 to 21.8367)	0.822
Amount of ethanol	14.72%	[-0.3576 to 0.1547]	0.297

CI confidence interval, NRS numeric rating scale, VAS visual analog scale

Further investigation of its long-term analgesic efficacy is required.

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Declarations

Conflict of interest On behalf of all the authors, the corresponding author states that there are no conflicts of interest.

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