REVIEW

Ultrasound-Guided Posterior Quadratus Lumborum Block for Acute Postoperative Analgesia in Adult Patients: A Meta-Analysis of Randomized Controlled Trials

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Objective: The quadratus lumborum block provides postoperative analgesia for patients undergoing abdominal surgery, although there are three common approaches to perform this block. The present meta-analysis investigated the effectiveness of posterior quadratus lumborum block (QLB2) after surgery.

Methods: PubMed, Embase, and the Cochrane Central Register were searched from inception to 26 August 2021 for randomized controlled trials that evaluated the analgesic efficacy of QLB2 vs control (placebo or no block). The primary outcomes were pain scores at 6 h, 12 h, and 24 h after surgery. The secondary outcomes were morphine consumption at 24 h after surgery and the postoperative complications.

Results: The present meta-analysis included 14 studies conducted with a total of 1001 patients. In comparison to control group, the QLB2 group presented significantly lower rest pain scores at 6 h (SMD -0.59; 95% CI: -1.05, -0.12; p = 0.01, I² = 84%; GRADE = moderate), 12 h (SMD: -0.83; 95% CI: -1.47, -0.19; p = 0.01; I² = 88%; GRADE = low), and 24 h (SMD: -0.37; 95% CI: -0.71, -0.03; p = 0.03; I² = 80%; GRADE = moderate) after surgery. The dynamic pain scores were significantly reduced, compared to control, in the QLB2 group at 12 h (SMD: -0.93; 95% CI: -1.52, -0.33; p = 0.002; I² = 83%; GRADE = low) and 24 h (SMD: -0.52; 95% CI: -0.93, -0.11; p = 0.01; I² = 83%; GRADE = moderate) after surgery. In addition, the QLB2 group presented reduced postoperative opioid consumption at 24 h (SMD: -0.45; 95% CI: -0.86, -0.03; p = 0.03; I² = 78%; GRADE = moderate). The subgroup analyses revealed that the analgesic benefit of QLB2 did not persist beyond 24 h when the patients were under spinal anesthesia.

Conclusion: Ultrasound-guided QLB2 could provide effective analgesia for patients under general anesthesia by decreasing the intensity of pain and opioid requirement when used within 24 h after abdominal surgery.

Keywords: quadratus lumborum block, postoperative pain, opioid, analgesia

Introduction

The analgesic effectiveness of ultrasound-guided regional anesthesia for abdominal surgeries is supported by an increasing body of evidence.¹ Regional blocks prove to be valuable and feasible tools for surgical patients, particularly in cases where neuraxial analgesia techniques cannot be performed due to certain reasons, such as coagulopathy, hypovolemia, and neurological disease. Quadratus lumborum block (QLB) is a novel regional block technique that was first reported in 2007.² In QLB, ultrasound guidance is utilized to determine the precise anatomical location of local anesthetics. Commonly, there are three approaches to QLB – lateral (termed QLB1), posterior (termed QLB2), and anterior (termed QLB3).³ Each of these approaches has a different mechanism of action, as stated in cadaveric and clinical reports.⁴ The effectiveness of postoperative analgesia of QLB has been confirmed in certain previous reports.⁵

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However, owing to the high heterogeneity and limited sample size, there was a scarcity of meta-analysis to conduct subgroup analysis of each approach of QLB. Recently, several randomized controlled trials (RCTs) found the analgesia of ultrasound-guided QLB2 remained controversial.^{6,7} Therefore, the present meta-analysis was aimed to examine the efficacy of only the ultrasound-guided single-injection QLB2 approach in adults.

Materials and Methods

Search Strategy and Selection Criteria

The present meta-analysis was conducted by following the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines.⁸ The online databases PubMed, Embase and the Cochrane Central Register were searched by two authors (CL and XMW) independently. The search was limited to RCTs on human subjects published from inception to 26 August 2021. A combination of MeSH with free-text terms was adopted as the search strategy. The search terms used were as follows: ("quadratus lumborum" [All Fields] OR ("abdominal muscles" [Mesh] AND "nerve block" [Mesh])). No language restrictions were applied during the search. In addition, the bibliographies of relevant articles were searched manually for retrieving additional studies. The objective was to search for and retrieve published reports on RCTs that investigated the effects of QLB2 compared to control (placebo or no block) in adults who had undergone abdominal surgery. The exclusion criteria were as follows: a) unavailability of full texts; b) unpublished clinical trials; c) no assessment of the analgesic outcomes; d) use of adjuvants to prolong the duration of nerve block analgesia.

Data Extraction and Quality Assessment

A standardized data collection form was developed by the authors (CL and XMW) to independently extract the following information: type of surgery, anesthesia technique, intervention and comparison, and QLB2 local anesthetic and post-operative analgesia regimen. The primary outcomes were the pain scores at 6 h, 12 h, and 24 h after surgery. The secondary outcomes were opioid consumption at 24 h after surgery and postoperative complications. Studies have demonstrated that Numerical Rating Scale may be used as a substitute for the Visual Analogue Scale.⁹ The pain score data were adjusted to a 0–10 score scale for analysis. In the case of the data that followed a non-standard distribution, the mean and standard deviation (SD) were approximated using the median and interquartile range as described previously by Wan et al¹⁰ and the Cochrane Collaboration.¹¹ Next, all the included studies were evaluated for the risk of bias, independently by the authors (XMW and CSQ), using the Cochrane Collaboration's Risk of Bias Tool. This tool included the following measurements: adequacy sequence generation, allocation of concealment, blinding throughout the study period, incomplete outcome data, selective outcome reporting, and other biases. The level of certainty was measured for the main results according to the guidelines of the Recommendation Assessment, Development, and Evaluation (GRADE) system. This system included the following measurements: risk of bias, inconsistency, indirectness, imprecision, and publication bias. The disagreements between the two evaluating authors were resolved through discussion with a third author (CL).

Statistical Analysis

Meta-analysis techniques (Revman 5.3, The Cochrane Collaboration 2014) were employed to combine the data wherever possible. In the case of continuous data, the mean difference (MD) was calculated when the outcome in the studies was obtained using the same scale; otherwise, standardized mean difference (SMD) was calculated. Dichotomous data were summarized as the risk ratio (RR) with the associated 95% confidence interval (CI). In the case of significant heterogeneity ($I^2 > 50\%$) among the included trials, random-effects modeling was adopted to calculate the pooled effect size; otherwise, fixed effects modeling was selected. When $I^2 > 50\%$, one study was omitted sequentially to identify the potential sources of heterogeneity. A p-value of <0.05 was considered statistically significant.

Results

A total of 2065 citations were retrieved in the initial database search, from which only those trials that fulfilled the inclusion criteria were retained (Figure 1). Finally, 14 trials conducted with a total of 1001 patients were included in the

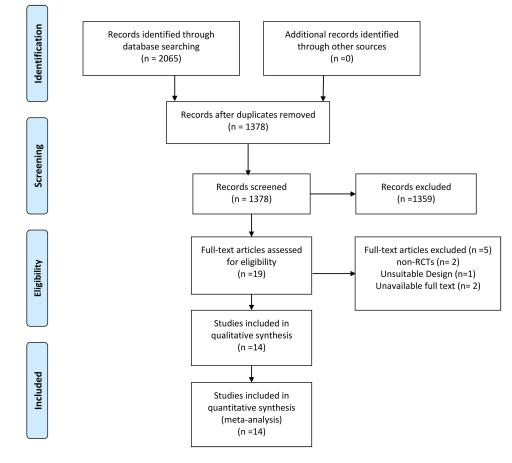


Figure I Flow diagram.

present meta-analysis. Table 1 presents the characteristics of the included RCTs. All included RCTs were published between the years 2015 and 2021.

Study Characteristics

Six studies^{12–17} involved patients who underwent elective cesarean section, six studies^{7,18–22} involved patients who underwent laparoscopic surgery (cholecystectomy, radical gastrectomy, colorectal resection, renal surgery and gynecologic surgery), and the remaining two studies^{6,23} involved patients who underwent open inguinal hernia repair surgery and full abdominoplasty. Seven RCTs^{12–17,23} were performed under spinal anesthesia, while the remaining seven RCTs^{6,7,18–22} employed general anesthesia. In three RCTs,^{12,15,16} QLB2 and control groups were compared and both groups received intrathecal morphine (ITM). The local anesthetic of QLB2 was ropivacaine in seven studies,^{6,7,13,16,18,21,22} bupivacaine in five studies,^{12,14,17,19,23} and levobupivacaine in two studies.^{15,20} One RCT¹⁶ included four groups and involved two comparisons between the QLB2 and control groups, with or without ITM (spinal anesthesia with 100 µg of intrathecal morphine). Each of the two comparisons was considered a separate trial in the present analysis.

Quality Assessment of the Selected Studies

According to the Cochrane Collaboration's tool, most of the trials included in the present meta-analysis presented a low risk of bias (Figure 2). Table 2 presents the level of certainty for the main results.

Table	I	Trial	Characteristics
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Author/Year	Type of Surgery	Anesthetic Technique	Intervention and Comparison (Number of Participants)	Timing of Block	QLB Block Local Anesthetic	Postoperative Analgesia
Bagbanci 2020 ²³	Open inguinal hernia repair surgery	0.5% isobaric bupivacaine (SA)	I.QLB (19) 2.No block (20)	Preoperative	0.25% bupivacaine 20mL	Fentanyl PCIA, bolus 25 μg, lockout 10 mins, loading dose 50 μg, no continuous infusion; IV paracetamol 1 g Q6h
Blanco 2015 ¹⁷	Caesarean section	15 mg bupivacaine and 20 μg fentanyl (SA)	I.QLB (25) 2.Placebo (23)	Postoperative	0.125% bupivacaine 0.2 mL/kg per side	Morphine PCIA, bolus I mg, lockout 5 mins, no continuous infusion; oral paracetamol I g Q6h and oral diclofenac 50 mg Q8h
Bjelland 2019 ⁶	Full abdominoplasty	GA	I.QLB (23) 2.Placebo (23)	Preoperative	0.375% ropivacaine 20 mL per side	Morphine PCIA, bolus 1 mg, lockout 5 mins, no continuous infusion; oral paracetamol 1 g Q6h
Boulianne 2020 ⁷	Laparoscopic colorectal resection	GA	I.QLB (31) 2.Placebo (31)	Preoperative	0.375% ropivacaine 20mL per side	Opioids (with or without patient-controlled devices) and acetaminophen
Fujimoto 2019 ²⁰	Elective laparoscopic gynecological surgery	GA	I.QLB (29) 2.No block (27)	Postoperative	0.25% levobupivacaine 25~30mL per side	Fentanyl PCIA, bolus 10 μ g, lockout 10 mins, 10 μ g/h continuous infusior or non-steroidal anti-inflammatory drugs as needed if pain scores >5
Irwin 2020 ¹⁵	Caesarean section	10–11.5 mg bupivacaine, 20 μg fentanyl+ 100 μg morphine (SA)	I.QLB+ ITM (44) 2.No block+ ITM (42)	Postoperative	0.25% levobupivacaine 20mL per side	Morphine PCIA, bolus I mg, lockout 5 mins, no continuous infusion; ora paracetamol I g Q6h and diclofenac 75 mg Q12h
Ishio 2017 ¹⁸	Laparoscopic gynecological surgery	GA	I.QLB (35) 2.No block (35)	Postoperative	0.375% ropivacaine 20 mL per side	Acetaminophen or diclofenac IV Q4h as needed
Ökmen 2018 ¹⁹	Laparoscopic cholecystectomy	GA	I.QLB (30) 2.Placebo (29)	Preoperative	0.25% bupivacaine 0.3 mL/kg per side	Tramadol PCIA, bolus 0.3 mg/kg, lockout 20 mins, a demand dose of 10 mg; 1 g paracetamol Q8h if pain scores >5
Salama 2020 ¹⁴	Caesarean section	12.5 mg of 0.5% bupivacaine and 10 μg fentanyl (SA)	I.QLB (30) 2.Placebo (30)	Postoperative	0.375% bupivacaine 24 mL per side	Morphine PCIA, bolus I mg, lockout 5 min, max dose 48 mg/4 h, no continuous infusion; IV paracetamol I g if pain scores >3
Tamura 2019 ¹⁶	Caesarean section	II–I3 mg bupivacaine and I0 μg fentanyl ± 100 μg morphine (SA)	I.QLB+ ITM (34) 2.Placebo+ ITM (38) 3.QLB (36) 4.Placebo (38)	Postoperative	0.3% ropivacaine 0.45 mL/kg per side	Oral diclofenac sodium 50 mg as needed if pain scores <3; IV pentazocine 15 mg with pain scores 3–5.9; IV acetaminophen 15mg/kg and pentazocine 15 mg if pain scores >6

Pangthipampai	Caesarean	0.5% hyperbaric	I.QLB+ ITM (18)	Postoperative	0.25% bupivacaine	Morphine PCIA, bolus I mg, lockout 5 mins, no continuous infusion; oral
202112	section	bupivacaine 2–2.2 mL +	2.Placebo+ ITM (20)		25 mL per side	acetaminophen I g Q6h and ibuprofen 400 mg Q8h
		200 µg morphine (SA)				
Li 2021 ²¹	Laparoscopic	GA	I.QLB (32)	Preoperative	0.4%	Sufentanil PCIA, bolus 5 µg, lockout 10 mins, 0.625 µg/h continuous
	renal surgery		2.No block (32)		ropivacaine	infusion
					30 mL	
Borys 2021 ¹³	Caesarean	0.5% hyperbaric	I.QLB (35)	Postoperative	0.375%	Morphine PCIA, bolus I mg, lockout 5 mins, no continuous infusion; IV
	section	bupivacaine 1.8–2.6 mL	2.Placebo (33)		ropivacaine	paracetamol Q6h; IV morphine 5 mg if pain scores >4
		(SA)			0.2 mL/kg per	
					side (up to	
					20 mL)	
Zhu 2021 ²²	Laparoscopic	GA	I.QLB (30)	Preoperative	0.3%	Sufentanil PCIA
	radical		2.Placebo (29)		ropivacaine	
	gastrectomy				20 mL per side	

Abbreviations: QLB, quadratus lumborum block; ITM, intrathecal morphine; PCIA, patient-controlled intravenous analgesia; IV, intravenous; GA, general anesthesia; SA, spinal anesthesia.

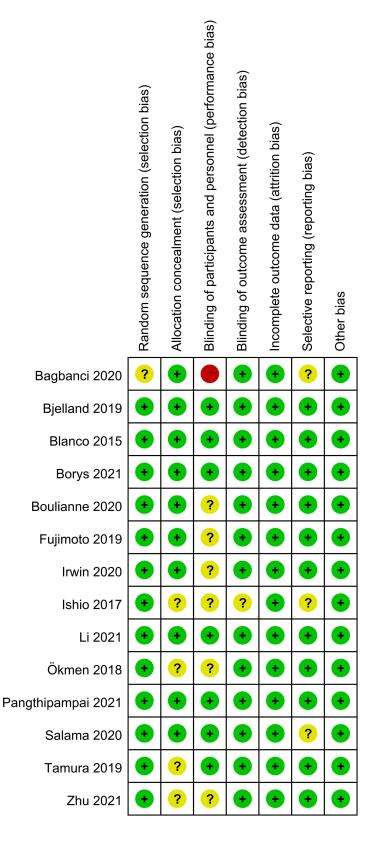


Figure 2 Risk-of-bias summary.

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Abbreviations: "+", low risk of bias; "?", unclear risk of bias; "-", high risk of bias.

Table 2 The GRADE Level of Certainty for Main Outcome

Outcomes	No of	No of			Quality Assessr	nent		Effect SMD (95% CI)	Quality	Importance
	Studies	Patients	Risk of Bias	Inconsistency	Indirectness	Imprecision	Other Considerations			
Rest pain scores at 6 h	8	509	No serious	Serious ^a	No serious	No serious	None	SMD -0.59 (-1.05, -0.12)	Moderate	Critical
Rest pain scores at 12 h	6	355	No serious	Serious ^a	No serious	Serious ^b	None	SMD -0.83 (-1.47, -0.19)	Low	Critical
Rest pain scores at 24 h	11	686	No serious	Serious ^a	No serious	No serious	None	SMD -0.37 (-0.71, -0.03)	Moderate	Critical
Dynamic pain scores at 6 h	4	291	No serious	Serious ^a	No serious	Serious ^b	None	SMD -0.33 (-0.79, 0.12)	Low	Critical
Dynamic pain scores at 12 h	5	300	No serious	Serious ^a	No serious	Serious ^b	None	SMD -0.93 (-1.52, -0.33)	Low	Critical
Dynamic pain scores at 24 h	9	571	No serious	Serious ^a	No serious	No serious	None	SMD -0.52 (-0.93, -0.11)	Moderate	Critical
Opioid consumption at 6 h	3	190	No serious	Serious ^a	No serious	Serious ^b	None	SMD -0.43 (-0.87,0.01)	Low	Critical
Opioid consumption at 12 h	2	134	No serious	Serious ^a	No serious	Serious ^b	None	SMD -0.13 (-1.08,0.82)	Low	Critical
Opioid consumption at 24 h	8	453	No serious	Serious ^a	No serious	No serious	None	SMD -0.45 (-0.86,-0.03)	Moderate	Critical

Notes: ^aQuality was rated down for inconsistency because I2>50%. ^bQuality was rated down for imprecision due to total population size is less than 400. Abbreviations: SMD, standard mean difference; CI, confidence interval.

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	QLB2 Control						:	Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
Blanco 2015	0.33	0.79	25	3.67	2.37	23	11.2%	-1.89 [-2.58, -1.20]	
Fujimoto 2019	29.17	17.54	29	33.33	23.48	27	12.5%	-0.20 [-0.72, 0.33]	
Irwin 2020	0.67	1.07	44	1.33	1.54	42	13.2%	-0.50 [-0.92, -0.07]	
Pangthipampai 2021	0.33	0.78	28	0.33	0.78	27	12.5%	0.00 [-0.53, 0.53]	
Tamura(1) 2019	2.32	0.91	34	2.25	1.28	38	13.0%	0.06 [-0.40, 0.52]	_ _
Tamura(2) 2019	4.21	2.33	36	4.39	2.37	38	13.1%	-0.08 [-0.53, 0.38]	
Zhu 2021	3	1.56	30	4	1.56	29	12.5%	-0.63 [-1.16, -0.11]	
Ökmen 2018	1.25	0.88	30	3	1.15	29	11.9%	-1.69 [-2.29, -1.09]	
Total (95% CI)			256			253	100.0%	-0.59 [-1.05, -0.12]	•
Heterogeneity: Tau ² =	0.37; Ch	i ² = 44.	58, df =	: 7 (P <	0.00001	l); ² = 8	34%	_	
Test for overall effect:	Z = 2.49	(P = 0.	01)						-2 -1 0 1 2 Favours [QLB2] Favours [control]

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	C	QLB2		С	ontrol		5	Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Bagbanci 2020	3.1	1.24	19	4.1	1.91	20	16.1%	-0.61 [-1.25, 0.04]	
Blanco 2015	0.67	1.57	25	3	1.58	23	16.1%	-1.46 [-2.10, -0.81]	_
Borys 2021	2.33	2.32	35	3.67	2.32	33	17.3%	-0.57 [-1.06, -0.09]	
Irwin 2020	1.12	1.53	44	1.48	2.3	42	17.7%	-0.18 [-0.61, 0.24]	
Pangthipampai 2021	0.33	0.78	28	0.33	0.78	27	17.0%	0.00 [-0.53, 0.53]	+
Ökmen 2018	1.25	0.88	30	3	0.57	29	15.9%	-2.32 [-2.99, -1.65]	
Total (95% CI)			181			174	100.0%	-0.83 [-1.47, -0.19]	
Heterogeneity: Tau ² =	0.56; Ch	i ² = 40).52, df	= 5 (P <	< 0.000	001); l²	= 88%		
Test for overall effect:	Z = 2.54	(P = 0	0.01)						-2 -1 0 1 2 Favours [QLB2] Favours [control]

	C	QLB2		C	ontrol		:	Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
Bagbanci 2020	2.26	1.28	19	2.9	1.65	20	8.2%	-0.42 [-1.06, 0.21]	+
Blanco 2015	1.67	2.36	25	2	1.58	23	8.7%	-0.16 [-0.73, 0.41]	
Borys 2021	2	3.09	35	2.83	3.1	33	9.4%	-0.27 [-0.74, 0.21]	
Fujimoto 2019	2.17	1.95	29	2	1.57	27	9.1%	0.09 [-0.43, 0.62]	
Irwin 2020	1.46	1.88	44	1.82	2.38	42	9.8%	-0.17 [-0.59, 0.26]	
Ishio 2017	1.5	1.8	35	4.4	2.1	35	9.0%	-1.47 [-2.00, -0.94]	
Pangthipampai 2021	1.67	2.34	28	1.33	2.35	27	9.0%	0.14 [-0.39, 0.67]	
Tamura(1) 2019	1.95	0.99	34	1.97	1.17	38	9.5%	-0.02 [-0.48, 0.44]	-+-
Tamura(2) 2019	2.67	1.6	36	2.42	1.12	38	9.6%	0.18 [-0.28, 0.64]	- -
Zhu 2021	2	1.56	30	2.67	0.78	29	9.1%	-0.53 [-1.05, -0.01]	
Ökmen 2018	1	0.57	30	2.5	1.19	29	8.6%	-1.60 [-2.19, -1.00]	
Total (95% CI)			345			341	100.0%	-0.37 [-0.71, -0.03]	•
Heterogeneity: Tau ² =	0.27; Ch	i² = 48	8.96, df	= 10 (P	< 0.00	0001); I	² = 80%		
Test for overall effect:	,		,	- (-		,,			-2 -1 0 1 2 Favours [QLB2] Favours [control]

Figure 3 Forest plot of postoperative rest pain score. (A) Rest pain scores at 6 h after surgery. (B) Rest pain scores at 12 h after surgery. (C) Rest pain scores at 24 h after surgery.

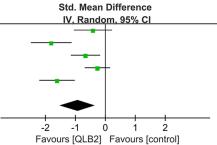
Effect of Interventions

Pain Scores at 6 h, 12 h, and 24 h After Surgery

The pain scores at 6 h, 12 h, and 24 h after surgery were determined. A random effects model was adopted to calculate the pooled effects size ($I^2 > 50\%$). In comparison to control, the QLB2 group exhibited significantly lower pain scores at 6 h (SMD -0.59; 95% CI: -1.05, -0.12, p =0.01, $I^2 = 84\%$; GRADE = moderate; Figure 3A), 12 h (SMD: -0.83; 95% CI: -1.47, -0.19; p = 0.01; $I^2 = 88\%$; GRADE = low; Figure 3B), and 24 h (SMD: -0.37; 95% CI: -0.71, -0.03; p = 0.03; $I^2 = 80\%$; GRADE = moderate; Figure 3C) after the surgery and at rest. The dynamic pain scores at 6 h after surgery did not differ significantly between the two groups (SMD: -0.33; 95% CI: -0.79, 0.12; p = 0.15; $I^2 = 73\%$; GRADE = low; Figure 4A). However, the dynamic pain scores were significantly reduced

	QLB2				ontrol		5	Std. Mean Difference	Std. Mean Difference		
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl		
Irwin 2020	2.8	3.14	44	4.4	2.61	42	26.2%	-0.55 [-0.98, -0.12]			
Tamura(1) 2019	2.11	1.34	34	2.13	1.81	38	25.3%	-0.01 [-0.48, 0.45]	_ + _		
Tamura(2) 2019	4.79	2.39	36	4.5	2.54	38	25.5%	0.12 [-0.34, 0.57]			
Ökmen 2018	1.5	1.19	30	2.75	1.45	29	23.1%	-0.93 [-1.47, -0.39]			
Total (95% CI)			144			147	100.0%	-0.33 [-0.79, 0.12]	•		
Heterogeneity: Tau ² =	0.16; Cł	ni² = 1′	1.22, df	= 3 (P =	= 0.01); l ² = 7	3%				
Test for overall effect:	Z = 1.43	(P = (0.15)						-2 -1 0 1 2 Favours [QLB2] Favours [control]		

Study or Subgroup	Mean	00	-					
	Mean	SD	lotal	Mean	SD	Total	Weight	IV, Random, 95% CI
Bagbanci 2020	4.89	1.82	19	5.65	1.78	20	19.1%	-0.41 [-1.05, 0.22]
Blanco 2015	0.67	1.57	25	4.33	2.37	23	18.5%	-1.81 [-2.49, -1.13]
Borys 2021	3.67	2.32	35	5	1.55	33	21.0%	-0.66 [-1.15, -0.17]
Irwin 2020	3.5	3.07	44	4.27	2.69	42	21.7%	-0.26 [-0.69, 0.16]
Ökmen 2018	1	1.29	30	3	1.15	29	19.7%	-1.61 [-2.21, -1.02]
Total (95% CI)			153			147	100.0%	-0.93 [-1.52, -0.33]



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		QLB2		-	ontrol			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
Bagbanci 2020	3.36	1.6	19	4	1.94	20	10.3%	-0.35 [-0.98, 0.28]	
Blanco 2015	1.67	2.36	25	3.67	2.37	23	10.6%	-0.83 [-1.42, -0.24]	
Borys 2021	3.75	2.51	35	5	1.55	33	11.4%	-0.59 [-1.07, -0.10]	
Irwin 2020	3.73	2.29	44	4.55	3.57	42	11.8%	-0.27 [-0.70, 0.15]	+
Ishio 2017	2.9	1.9	35	6.3	2.1	35	10.9%	-1.68 [-2.23, -1.13]	_ _
Pangthipampai 2021	4.33	2.34	28	4	3.13	27	11.1%	0.12 [-0.41, 0.65]	
Tamura(1) 2019	2.95	1.78	34	3.25	1.73	38	11.6%	-0.17 [-0.63, 0.29]	
Tamura(2) 2019	4.14	1.35	36	3.81	1.13	38	11.6%	0.26 [-0.19, 0.72]	+
Ökmen 2018	1.5	1.19	30	3.25	1.45	29	10.8%	-1.30 [-1.87, -0.74]	
Total (95% CI)			286			285	100.0%	-0.52 [-0.93, -0.11]	◆
Heterogeneity: Tau ² =	0.32; Ch	i² = 45	5.81, df	= 8 (P <	< 0.000	001); l²	= 83%		
Test for overall effect:									-2 -1 0 1 2 Favours [QLB2] Favours [control]

Figure 4 Forest plot of postoperative dynamic pain score. (A) Dynamic pain scores at 6 h after surgery. (B) Dynamic pain scores at 12 h after surgery. (C) Dynamic pain scores at 24 h after surgery.

in the QLB2 group at 12 h (SMD: -0.93; 95% CI: -1.52, -0.33; p = 0.002; I² = 83%; GRADE = low; Figure 4B) and 24 h (SMD: -0.52; 95% CI: -0.93, -0.11; p = 0.01; I² = 83%; GRADE = moderate; Figure 4C) after surgery.

Opioid Consumption at 6 h, 12 h, and 24 h After Surgery

The opioid consumption at 6 h (SMD: -0.43; 95% CI: -0.87, 0.01; p = 0.05; I² = 55%; GRADE = low; Figure 5A) and 12 h (SMD: -0.13; 95% CI: -1.08, 0.82; p = 0.78; I² = 86%; GRADE = low; Figure 5B) did not differ significantly between the two groups. The opioid consumption at 24 h after the surgery was assessed in eight trials, $^{6,12,15,17,20-23}$ which included a total of 453 patients (230 patients in the QLB2 group and 223 patients in the control group). In comparison to control group, the QLB2 group presented reduced postoperative opioid consumption at 24 h (SMD: -0.45; 95% CI: -0.86, -0.03; p = 0.03; I² = 78%; GRADE = moderate; Figure 5C).

Side-Effects

Among the included studies, six trials reported the incidence of postoperative nausea and vomiting (PONV).^{6,12,19,21–23} A fixed-effects model was adopted as there was no significant heterogeneity among these studies ($I^2 = 0\%$). No significant difference in PONV was observed between the two groups (RR: 0.67; 95% CI: 0.45, 1.01; p = 0.06;

4									
	C	QLB2		C	ontrol		:	Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
Blanco 2015	2.83	5.11	25	10.33	11.06	23	28.8%	-0.87 [-1.46, -0.27]	_ _
Fujimoto 2019	83	12	29	94	32	27	32.2%	-0.46 [-0.99, 0.08]	
Irwin 2020	3	3.07	44	3.33	3.84	42	38.9%	-0.09 [-0.52, 0.33]	
Total (95% CI)			98			92	100.0%	-0.43 [-0.87, 0.01]	•
Heterogeneity: Tau ² =	0.08; Cł	ni² = 4.	.42, df =	= 2 (P =	0.11); I	² = 55%	,		
Test for overall effect:	7 = 1.93	(P = 0)	0.05)						-2 -1 0 1 2
		. (.	0.00)						Favours [QLB2] Favours [control]

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QLB2				c	ontrol		:	Std. Mean Difference	Std. Mean Difference		
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI		
Blanco 2015	8	5.93	25	14	11.85	23	47.8%	-0.64 [-1.22, -0.06]			
Irwin 2020	7	6.67	44	5	5.19	42	52.2%	0.33 [-0.10, 0.76]	+=-		
Total (95% CI)			69			65	100.0%	-0.13 [-1.08, 0.82]			
Heterogeneity: Tau ² = 0.40; Chi ² = 6.94, df = 1 (P = 0.008); l ² = 86%											
Test for overall effect:	Z = 0.27		-2 -1 0 1 2 Eavours [OL B2] Eavours [control]								

Favours [QLB2] Favours [control]

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	(QLB2		Control			:	Std. Mean Difference	Std. Mean Difference		
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI		
Bagbanci 2020	184.21	52.84	19	401.25	116.26	20	9.8%	-2.33 [-3.17, -1.50]			
Bjelland 2019	9.3	5.4	23	11.3	5.4	23	12.3%	-0.36 [-0.95, 0.22]			
Blanco 2015	11	11.01	25	22	19.75	23	12.3%	-0.68 [-1.27, -0.10]	_ _		
Fujimoto 2019	304	58	29	301	50	27	12.9%	0.05 [-0.47, 0.58]	_ _		
Irwin 2020	16.33	16.09	44	14.67	15.35	42	13.9%	0.10 [-0.32, 0.53]			
Li 2021	119	40	32	118	36	32	13.2%	0.03 [-0.46, 0.52]	_ _		
Pangthipampai 2021	11.5	11.05	28	16.5	17.48	27	12.8%	-0.34 [-0.87, 0.19]			
Zhu 2021	35.77	10.2	30	41.72	11	29	12.9%	-0.55 [-1.07, -0.03]			
Total (95% CI)			230			223	100.0%	-0.45 [-0.86, -0.03]	•		
Heterogeneity: Tau ² = 0.28; Chi ² = 32.53, df = 7 (P < 0.0001); l ² = 78%									-2 -1 0 1 2		
Test for overall effect:	Z = 2.11 (-2 -1 0 1 2 Favours [QLB2] Favours [control]								

Figure 5 Forest plot of opioid consumption. (A) Opioid consumption at 6 h after surgery. (B) Opioid consumption at 12 h after surgery. (C) Opioid consumption at 24 h after surgery.

	QLB	2	Control		Control			Risk Ratio		Risk Ratio			
Study or Subgroup	Events	<u>nts Total Events Total</u>		Weight	M-H, Fixed, 95% C	CI M-H, Fixed, 95% CI							
Bagbanci 2020	0	19	5	20	13.2%	0.10 [0.01, 1.62]	←		<u> </u>				
Bjelland 2019	8	23	9	23	22.2%	0.89 [0.42, 1.89]		-	-				
Li 2021	9	32	12	32	29.6%	0.75 [0.37, 1.53]		_					
Pangthipampai 2021	0	28	1	27	3.8%	0.32 [0.01, 7.57]		•					
Zhu 2021	9	30	11	29	27.6%	0.79 [0.39, 1.62]		_					
Ökmen 2018	0	30	1	29	3.8%	0.32 [0.01, 7.61]							
Total (95% Cl)		162		160	100.0%	0.67 [0.45, 1.01]		•					
Total events	26		39										
Heterogeneity: Chi² = 3.05, df = 5 (P = 0.69); l² = 0%							⊢ 0.01			10	100		
Test for overall effect: $Z = 1.90$ (P = 0.06)								0.1 Favours [QLE	32] Favou	10 Irs [control]	100		

Figure 6 Forest plot of postoperative nausea and vomiting.

Figure 6). The differences in the methods used for reporting outcomes in different studies and the lack of relevant data rendered it impossible to conduct the meta-analysis of postoperative pruritus and sedation. The main adverse event associated with QLB2 was lower limb numbness or weakness. With the use of QLB2, Fujimoto et al reported five cases

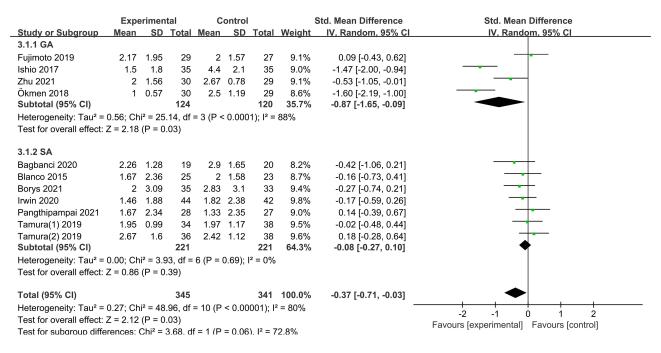


Figure 7 Forest plot of rest pain scores at 24 h after surgery by subgroup. Abbreviations: GA, general anesthesia; SA, spinal anesthesia.

of lower limb numbness in 29 patients who underwent laparoscopic gynecological surgery,²⁰ while Ökmen et al reported 2 cases of lower limb sensory loss and weakness in 30 patients who underwent laparoscopic cholecystectomy.¹⁹

Subgroup Analysis

The studies were divided into two subgroups –general anesthesia and spinal anesthesia. The results of the subgroup analysis revealed that the general anesthesia group presented significantly reduced rest pain scores (SMD: -0.87; 95% CI: -1.65, -0.09; p = 0.03; $I^2 = 88\%$) and dynamic pain scores (SMD: -1.50; 95% CI: -1.89, -1.10; p < 0.00001; $I^2 = 0\%$) at 24 h after the surgery compared to the spinal anesthesia group (rest pain scores: SMD: -0.08; 95% CI: -0.27, 0.01; p = 0.39; $I^2 = 0\%$; Figure 7; dynamic pain scores: SMD: -0.24; 95% CI: -0.52, 0.03; p = 0.08; $I^2 = 52\%$; Figure 8). The resting pain scores at 6 h after surgery and the opioid consumption at 24 h after surgery did not present any notable differences between the two subgroups (Table 3). The risk of PONV also did not differ significantly between the two groups (Table 3). The subgroup analysis could not be performed for the other primary outcomes as only one trial was available for analysis in each subgroup.

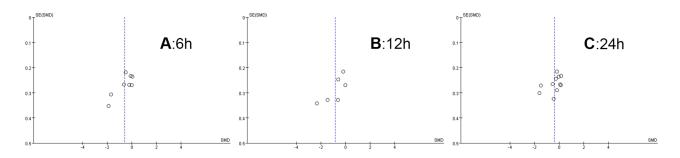


Figure 8 Begg's funnel plot of rest pain scores.

Notes: (A) Funnel plot of rest pain scores at 6 h after surgery; (B) Funnel plot of rest pain scores at 12 h after surgery; (C) Funnel plot of rest pain scores at 24 h after surgery.

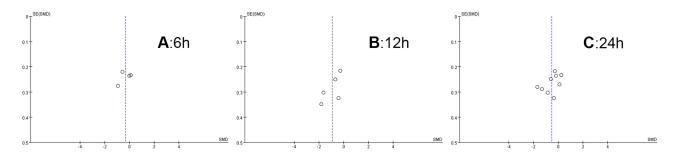


Figure 9 Begg's funnel plot of dynamic pain scores.

Notes: (A) Funnel plot of rest pain scores at 6 h after surgery; (B) Funnel plot of rest pain scores at 12 h after surgery; (C) Funnel plot of rest pain scores at 24 h after surgery.

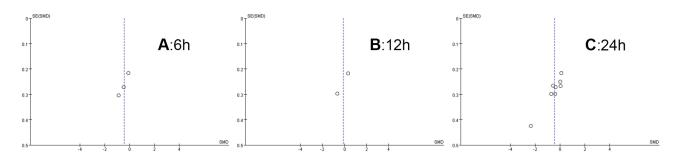


Figure 10 Begg's funnel plot of opioid consumption.

Notes: (A) Funnel plot of rest pain scores at 6 h after surgery; (B) Funnel plot of rest pain scores at 12 h after surgery; (C) Funnel plot of rest pain scores at 24 h after surgery.

Publication Bias

According to the funnel plots (Figures 8-10), no evident publication bias was observed in the present meta-analysis.

Discussion

The main findings of the present meta-analysis, which had a moderate-to-low level of certainty, are presented ahead. In comparison to control (placebo or no block), the ultrasound-guided QLB2 group presented significant reductions in the rest pain scores at 6 h,12 h, and 24 h after surgery, dynamic pain scores at 12 h and 24 h after surgery, and opioid consumption in the first 24 h after abdominal surgery in adults. However, the subgroup analysis showed that the spinal anesthesia group did not reduce the postoperative rest and dynamic pain scores at 24 h in the presence of QLB2. The

Table 3 The Result of Subgroup Analysis

Outcomes	Subgroup	No of Trails	No of Patients	Effect SMD/RR (95% CI)	Р	l ²
Rest pain scores at 6 h	General anesthesia	3	174	SMD -0.83 (-1.66, 0.00)	0.05	86%
	Spinal anesthesia	5	335	SMD -0.44 (-1.01, 0.13)	0.13	84%
Rest pain scores at 24 h	General anesthesia	4	244	SMD -0.87 (-1.65, -0.09)	0.03	88%
	Spinal anesthesia	7	442	SMD -0.08 (-0.27, 0.01)	0.39	0%
Dynamic pain scores at 24 h	General anesthesia	2	129	SMD -1.50 (-1.89, -1.10)	<0.00001	0%
	Spinal anesthesia	7	442	SMD -0.24 (-0.52, 0.03)	0.08	52%
Opioid consumption at 24 h	General anesthesia	4	225	SMD -0.20 (-0.49, 0.10)	0.19	21%
	Spinal anesthesia	4	228	SMD -0.76 (-1.61, 0.10)	0.08	89%
Postoperative nausea and vomiting	General anesthesia	4	228	RR 0.78 (0.51, 1.19)	0.25	0%
	Spinal anesthesia	2	94	RR 0.15 (0.02, 1.15)	0.07	0%

Abbreviations: SMD, standard mean difference; CI, confidence interval; RR, risk ratio.

patients who were under general anesthesia could get better 24 h pain-related outcomes compared to those under spinal anesthesia. This could be attributed to the anesthetic effect of spinal anesthesia or the intrathecal opioid use, which covering up the effect of QLB2. In addition, perioperative QLB2 could not reduce the risk of PONV and presented no major complications.

Recently, several systematic reviews on QLB have indicated the analgesic efficacy of QLB, although such findings should be interpreted cautiously depending on different approaches of QLB. Two meta-analyses reported that the opioid consumption at 24 h after surgery was reduced in the QLB (using different QLB approaches) group.^{5,24} Nevertheless, the subgroup analyses reported by Korgvee et al⁵ demonstrated that QLB2 did not reduce opioid consumption at 24 h after surgery, compared with placebo, no block or other peripheral block. Their results are not consistent with those observed in the present meta-analysis. This difference could be attributed to the lack of sample size and the differences in the selection method used for control groups between the two analyses. Another reason could be related to the observation of the subgroup analysis conducted in the present study, which revealed that QLB2 did not reduce the rest pain scores and dynamic pain scores at 24 h after surgery when the patients were under spinal anesthesia.

The quadratus lumborum muscle is surrounded by thoracolumbar fascia, which encases the dorsal muscles extending from the thoracic region to the lumbar spine. It is generally accepted that local anesthetics spread to the paravertebral space through the thoracolumbar fascia to exert their analgesic effects. Moreover, it is reported that the blockade of the sympathetic fibers in the thoracolumbar fascia and visceral analgesia could contribute to the analgesic efficacy of QLB.^{4,17} According to the cadaveric reports, each approach of QLB has a different mechanism of action. In QLB1 or QLB2, the injected local anesthetics are reportedly confined to the thoracolumbar fascia or transversus abdominis plane (TAP). In QLB3, the injected local anesthetics may spread to the mid to lower thoracic paravertebral space and the lumbar nerve roots.^{25,26} The three-dimensional computed tomography images of patients have revealed that QLB1 spread in the TAP, QLB2 spread in the TAP and posterior region of the quadratus lumborum muscle, and transverse oblique paramedian QLB3 spread to the lumbar and thoracic paravertebral regions. Tamura et al²⁷ reported that in magnetic resonance imaging, QLB2 appeared to have a wider dye spread compared to QLB1. Furthermore, QLB3 is a deep approach, which is reported to result in a greater motor blockade and the risk of needle trauma of the pleura and kidney.^{26,28} Consequently, in clinical practice, a higher number of people select QLB2. Since there is a lack of studies comparing the different approaches of OLB, whether the other approaches are better than OLB2 remains unclear so far. One small-sample-size study reported that the analgesic effect of OLB3 was a superior to that of OLB2 in cesarean delivery.²⁹ However, Brixel, et al³⁰ recently reported that 30 mL of the QLB2 solution could reach multiple locations around the quadratus lumborum muscle when sonographic localization was used. Therefore, future studies should consider investigating the effects of the different approaches of QLB using varying dosages of the local anesthetic.

Our meta-analysis has several limitations. Firstly, when sensitivity analysis was performed by omitting one study sequentially, high heterogeneity was still observed. This heterogeneity could be attributed to the different types of surgery, different types of anesthesia, and different durations of the QLB2 procedure adopted in the studies. Moreover, the differences in the study designs, the dosages and types of the local anesthetic, and the postoperative analgesia regimen could also have contributed to the increased heterogeneity. Second, as non-opioid drugs were used for post-operative pain management in certain studies, it was not possible to extract relevant data from all the included trials. Third, the success rate of sensory blocking could not be evaluated adequately as a few patients were under general anesthesia.

Conclusion

Ultrasound-guided QLB2 could provide effective analgesia for adult patients by decreasing the intensity of pain and opioid requirements within 24 h after abdominal surgery. However, when used in conjunction with spinal anesthesia, QLB2 appeared to lose its superior efficacy. Nonetheless, the use of QLB2 is recommended in the setting of general anesthesia.

Abbreviations

QL block, quadratus lumborum block; QLB2, posterior quadratus lumborum block; RCT, randomized controlled trial; SMD, standardized mean difference; CI, confidence interval; RR, relative risk; PONV, postoperative nausea and vomiting; SD, standard deviation; IQR, interquartile range; ITM, intrathecal morphine; TAP, transversus abdominis plane.

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Disclosure

The authors declare that they have no competing interests in this work.

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