MUSCULOSKELETAL RADIOLOGY



Efficacy of radiofrequency in lumbar facet joint pain: a systematic review and meta-analysis of placebo-controlled randomized controlled trials

Antonio Jesús Láinez Ramos-Bossini^{1,2} · Paula María Jiménez Gutiérrez^{2,3} · Fernando Ruiz Santiago^{1,2,4}

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Abstract

Background Lumbar facet joint pain (LFJP) is one of the main causes of chronic low back pain (LBP) and can be treated using radiofrequency (RF) sensory denervation. The aim of this work is to analyze the efficacy of RF in LFJP through a systematic review and meta-analysis of randomized controlled trials (RCTs) with placebo control.

Materials and methods A systematic search was conducted in the Medline (PubMed), Scopus, Web of Science databases, and the Cochrane Central Register of Controlled Trials (CENTRAL). The variables of interest were pain, functional status, quality of life (QoL), and global perceived effect (GPE) measured at different time intervals: short (<3 months), medium (>3 and <12 months), and long term (>12 months).

Results Eight RCTs with placebo control were included. RF showed significant benefits over placebo in pain relief in the short (MD – 1.01; 95% CI – 1.98 to -0.04; p = 0.04), medium (MD – 1.42; 95% CI – 2.41 to – 0.43; p = 0.005), and long term (MD – 1.12; 95% CI – 1.57 to – 0.68; p < 0.001), as well as improvement in functional disability in the short (SMD – 0.94; 95% CI – 1.73 to – 0.14; p = 0.02) and long term (SMD – 0.74; 95% CI – 1.09 to – 0.39; p < 0.001). No statistically significant differences were observed in QoL or quantitative GPE, but benefits for RF were observed in dichotomous GPE in the medium (OR 0.19; 95% CI 0.07–0.52; p = 0.001) and long term (OR 0.22; 95% CI 0.06–0.78; p = 0.02). Subgroup analyses showed more benefits for RF in LBP < 1 year in the short term and in RCTs that did not require performing an MRI for patient selection.

Conclusions RF demonstrated significant improvement in pain and functionality, but the benefits in terms of QoL and GPE are inconclusive. Future clinical trials should investigate the long-term effects of RF, its impact on quality of life, and define appropriate criteria for patient selection.

Keywords Facet joint · Pain · Radiofrequency · Placebo · Meta-analysis · Randomized controlled trial

Antonio Jesús Láinez Ramos-Bossini ajbossini@ugr.es

- ¹ Department of Radiology, Hospital Universitario Virgen de Las Nieves, Avda. Fuerzas Armadas, 18014 Granada, Spain
- ² Advanced Medical Imaging Group (TeCe22), Instituto Biosanitario de Granada (IBS.Granada), 18016 Granada, Spain
- ³ Department of Anesthesiology, Hospital Universitario Virgen de Las Nieves, 18014 Granada, Spain
- ⁴ Department of Radiology and Physical Medicine, University of Granada, Granada, Spain

Introduction

Low back pain (LBP) is a very common health problem in the adult population, with an estimated incidence of 5% and a lifetime prevalence of 60–80% [1, 2]. Chronic pain syndromes develop in 10–20% of these patients and represent a significant source of disability [2]. Among the multiple causes of chronic LBP, lumbar facet joint pain (LFJP) accounts for up to 40% of cases [3]. LFJP is characterized by the degeneration of the zygapophyseal or facet joints of two adjacent lumbar vertebrae, resulting from repetitive mechanical stress, inflammatory processes, or infections [4]. The facet joint is the only synovial joint in the spinal column, and its sensory innervation is provided by the medial branch of the dorsal root [5]. The pain associated with LFJP can radiate to the gluteal region and the posterior aspect of the leg, is exacerbated by lumbar extension and improves with slight lumbar flexion [6]. To confirm LFJP, it is often necessary to perform a diagnostic blockage test [7].

The therapeutic approach to LFJP should be multimodal, including hygienic-dietary measures, physical therapies, and pharmacological interventions [8–10]. In non-responders, second-line treatments include local anesthetic and corticosteroid infiltration, and radiofrequency (RF) of the medial branch of the dorsal root [11, 12], all of which are frequently performed by musculoskeletal and interventional radiologists. The latter procedure involves sensory denervation of the facet joint by applying an electrical field around a nerve, which alters the transmission of painful stimuli, either through direct nerve injury (continuous RF, CRF) or modulation of nerve impulses (pulsed RF, PRF) [13, 14]. According to the American Society of Interventional Pain Physicians, radiofrequency neurotomy for lumbar chronic pain patients who test positive for blocks is recommended with level II evidence and moderate strength of recommendation [15]. Similarly, current consensus practice guidelines establish that lumbar RFA may provide benefit to well-selected individuals and stress the importance of selection criteria to improve denervation outcomes [16]. These recommendations are based upon different studies of varied quality and design.

To date, various randomized controlled trials (RCTs) have been conducted comparing the efficacy of RF with placebo [17–19], intra-articular infiltration of anesthetics and corticosteroids, and different RF modalities [20–22]. However, systematic reviews and meta-analyses of these trials have shown contradictory results [23, 24], which have been attributed to factors, including small sample sizes [17, 25], significant differences in baseline variables, heterogeneous inclusion and exclusion criteria [23], presence or absence of prior diagnostic blocks [17, 26], short follow-up periods [18, 19, 27], or differences in outcome measures and RF technique used [20, 21, 28]. Additionally, various biases in different stages of the clinical trials have been noted [9, 29].

Therefore, high-quality scientific evidence is needed to update and compare the results obtained from RCTs comparing RF versus placebo. Such evidence should consider potential biases and confounding factors that may impact the analysis, shedding light on the efficacy and clinical indications of RF treatment for LFJP.

The aim of this study is to conduct a systematic review and an updated meta-analysis of placebo-controlled RCTs examining the efficacy of RF in the treatment of chronic LBP caused by LFJP.

Materials and methods

Eligibility criteria

The PRISMA (Preferred Reporting Items for Systematic Reviews and Meta-Analyses) guidelines [30] were followed to collect and report the results obtained. The design and selection criteria were based on the PICOS strategy: adult patients with chronic LBP due to LFJP (P), treated with RF-based procedures (I) compared to placebo (C), and clinical outcomes (pain, functional status, quality of life [QoL], and perceived global effect) (O). Only RCTs were included (S).

Therefore, the inclusion criteria were: RCTs on RF versus placebo for LFJP that included quantitative results for at least one of the primary outcomes, original data, and adult populations. We included all identified studies regardless the year of publication, language or study quality criteria. Efforts to obtain full-text documents (through our institutional virtual library and sending emails to authors) were conducted for studies with limited access.

The exclusion criteria were: locations or conditions other than LFJP (e.g., cervical, thoracic, disc pathology, or sacroiliac pain), RF modalities other than PRF or CRF, quasi-experimental or observational designs without a control group, letters, editorials, or conference proceedings. Figure 1 shows the flow diagram of the study.

Information sources and search strategy

A systematic search was conducted in PubMed, Scopus, Web of Science, and Cochrane Central Register of Controlled Trials (CENTRAL) databases. The search included literature published up to May 31, 2023. A search equation using the MeSH terms "facet joint," "zygapophyseal joint," "low back pain," "radiofrequency," "efficacy" was used, combining them with the boolean operators AND and OR. Only human studies with abstracts written in English or Spanish were considered, without any other restrictions. Additionally, to optimize the number of relevant results, studies of potential interest from the reference lists of the selected articles were reviewed.

The literature search was conducted by the authors (AJLRB) and (PMJG). Both evaluators are clinicians with 5-year experience in the subject area and previous experience in systematic reviews and meta-analysis methodologies. All titles and abstracts of interest were reviewed. An article that could not be unequivocally excluded based on its title and abstract was considered potentially relevant. Then, the full text of the non-excluded articles was evaluated to determine if they met all the eligibility criteria.



Fig. 1 Flow diagram of the study

Both evaluators performed the search and selection of studies independently according to PRISMA recommendations. This was performed for all steps (title and abstract, and full-text assessment). Once the evaluation was completed independently, consensus was reached for final decisions. Discrepancies between both evaluators were solved by consensus with a senior researcher (FRS).

Variables analyzed and data extraction

The treatment-related outcomes included:

- Pain relief, measured by the Visual Analog Scale (VAS) or other quantitative scales (e.g., Numeric Rating Scale, NRS).
- Improvement in functional disability measured by the Roland-Morris Disability Questionnaire (RMDQ) and the Oswestry Disability Index (ODI).
- Improvement in QoL measured by the Euro-Qol in 5 dimensions and other scales (e.g., SF-36 QoL Questionnaire, 6-item QoL scale).
- Global perceived effect (GPE) measured by the GPE scale or surrogate scores on overall subjective assessment measured as quantitative data or as data that could be

grouped as dichotomous variables (e.g., pain relief < 50% or > 50% compared to baseline, or bad/moderate vs good/ excellent overall patient satisfaction).

These variables were grouped according to the time at which they were measured as follows: short-term (< 3 months), medium-term (3–12 months), and long-term (> 12 months). If any RCT reported several measurements within one of those intervals, the data from the last one were selected.

The primary outcome measure was pain relief. The secondary outcome measures were improvement in functional status, QoL, and perceived global effect.

The data from the selected articles were extracted by the author. The data were stored in anonymized spreadsheets and the software Review Manager Web (RevMan Web) version 5.4.0 [31].

Risk of bias, heterogeneity, and publication bias

The Cochrane Risk of Bias Tool v. 2 was used to systematically address the presence of potential biases. For each RCT, the risk of bias of 10 categories was classified as low, intermediate, or high. Studies with < 5 low risk of bias items or > 2 high risk of bias items were considered as higher-riskof-bias (lower quality) studies. A subgroup analysis according to the quality of the studies was performed for each association. Publication bias was analyzed through funnel plots.

Statistical analysis

For variables measured on different scales (e.g., functional status), standardized mean differences (SMDs) with a 95% confidence interval (CI) were calculated. In the case of different scales with the same range (e.g., pain), mean differences (MDs) were applied, as in Shih et al. [11]. When standard deviations (SDs) were not available for a given variable, they were calculated using the standard error (SE) through the formula $SD = SE \cdot \sqrt{N}$. For studies with sample sizes greater than 70 patients, SD was estimated from 95% confidence intervals using the formula $SD = \sqrt{N} \cdot \frac{(U_L - L_L)}{3.92}$, and for studies with smaller sample sizes, SD was estimated using the formula $SD = \sqrt{N} \cdot \frac{(U_L - L_L)}{4.13}$. If SDs or 95% CIs were not available, SD values were imputed based on the median of SDs from all studies in the same group [32–34].

The inverse variance-weighted method with a randomeffects model was applied to quantitative outcomes, and the Mantel–Haenszel method was applied for dichotomous GPE variables. The I² statistic was used to analyze heterogeneity among studies (non-relevant, moderate, or substantial, with cutoff values of I² < 40%, 40% < I² < 75%, and I² > 75%, respectively) [35]. Sensitivity analyses were performed in cases of significant heterogeneity (I² > 40%) by sequentially removing each study to estimate its contribution to the overall analysis. Two-tailed tests were conducted, with significance set at p < 0.05. Statistical analyses were performed using RevMan web [31].

Results

Baseline characteristics of patients

The RCTs included in the meta-analysis encompassed data from 472 patients, with 249 in the RF group and 223 in the placebo group. The smallest RCT included 30 patients (Gallagher et al. 1994), while the largest included 150 patients (Moussa et al. 2020). Of the eight included studies, seven compared CRF with placebo, and one (Tekin et al. 2007) compared CRF, PRF, and placebo. Therefore, data from the latter were analyzed based on the corresponding subgroups, following the example of Maas et al. [29]. Table 1 summarizes the baseline characteristics of the participants in each study.

Risk of bias

A high risk of bias was detected in several RCTs, specifically for performance (n=1), detection (n=1), attrition (n=3), information (n=1), and other biases (n=6). Figure 2 summarizes the analysis of risk of bias in the RCTs included in the meta-analysis. Globally, 4 studies [19, 21, 27, 36] were considered of lower quality, mainly due to combinations of selection, blinding and attrition biases. The rest of RCTs were considered of higher quality (i.e., lower risk of bias) according to the criteria detailed in the methodology.

Pain relief

Pain relief was measured using the VAS in all studies except Van Tilburg et al. (2016), who used the 11-NRS. Statistically significant differences were found favoring RF over placebo in the short (MD – 1.01; 95% CI – 1.98 to – 0.04; p = 0.04), medium (MD – 1.42; 95% CI – 2.41 to – 0.43; p = 0.005), and long term (MD – 1.12; 95% CI – 1.57 to – 0.68; p < 0.001). Heterogeneity among the studies was high, particularly in the short and medium-term analyses (I²=90 and 91%, respectively). The results of the analysis are shown in Fig. 3.

Improvement in functional status

Statistically significant benefits of RF over placebo were observed in the short (SMD - 0.94; 95% CI - 1.73 to - 0.14; p = 0.02) and long term (SMD - 0.74; 95% CI - 1.09 to - 0.39; p < 0.0001). For medium term outcomes, a trend toward significance favoring RF was observed (SMD - 1.43;

Main author of the trial (year)	No. of com- parison groups	Treatment group	Z	Age		Evaluation time (months)	Total follow- up (months)	Initial VAS X (s.d.)	Initial ODI X (s.d.)	Quality of life
Leclaire et al. (2001)	2	CRF	36	46.7 (9.3)	12/24	0, 1, 3	3	51.9 (26.7)	38.3 (14.7)	
		Sham	34	46.4 (9.8)	13/21			51.5 (20.8)	36.4 (14.6)	
Nath et al. (2008)	2	CRF	20	56	6/14	0, 6	9	6.03		
		Sham	20	53	9/11			4.35		
Van Tilburg et al. (2016)	2	CRF	30	65	14/16	1, 3	12	7.2		
		Sham	30	58	12/18			7.4		
Tekin et al. (2007)	Э	CRF	20	60.5 (8.5)	9/11	0, 6, 12	12	6.5 (1.5)	39.2	
		RFP	20	59.6 (7.7)	8/12			6.6 (1.6)	39.4	
		Sham	20	57.9 (9.3)	9/11			6.8 (1.6)	40.1	
Van Wijk et al. (2005)	2	CRF	40	46.9 (11.5)	10/30	3	3	5.8 (1.8)		
		Sham	41	48.1 (12.6)	13/28			6.5 (1.8)		
Van Kleef et al. (1999)	2	CRF	15	46.6(7.4)	5/10	2	2	5.2(1.7)	20.3(3.8)	21.6(3.6)
		Sham	16	41.4(7.5)	6/10			5.2(1.6)	21.6(3.6)	20.3(3.8)
Moussa et al. (2020)	ę	CRF	50	57.3	16/34	3, 6, 12, 24, 36	36	8.25(1.56)*	31^{*}	
		Sham	50	56.9	17/33			7.73(1.41)*	50*	
Gallagher et al. (1994)	2	CRF	18	ND	QN	0, 1, 6	9	5.8		
		Sham	12	ND	QN			7.2		

 Table 1
 Baseline characteristics of the patients included in the meta-analysis



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•	••	Ð	•	Ð	Ð	••	Ð	••	Random sequence generation (selection bias)
•	•	••	••	••	••	Ð	••	••	Allocation concealment (selection bias)
•	•	•	•	Ð	•	Ŧ	<mark>.</mark> ي	•	Blinding (performance bias and detection bias): Self-reported outcomes
•	•	••	•	•	•	•	••	Ŧ	Blinding (performance bias and detection bias): Objective outcomes
•	•	•	•	•	•	••	Ŧ	Ŧ	Blinding of participants and personnel (performance bias):
•	•	•	•	•	••	••	•	••	Blinding of outcome assessment (detection bias): Self-reported outcomes
•	•	•	Ð	•	•	•	Ŧ	••	Blinding of outcome assessment (detection bias): Objective measures
•	•	•	••	•	•	•	•	••	Incomplete outcome data (attrition bias): All outcomes
->	•	••	••	-0	•	••	••	••	Selective reporting (reporting bias)
•	•	Ð	•	•	•	•	•	Ŧ	Other blas

Fig. 2 Risk of bias assessment for the trials included in the meta-analysis. Each evaluated item is indicated as "+" for low risk of bias, "?" for unclear risk, and "-" for high risk

95% CI – 3.24 to – 0.37; p = 0.12). Overall, significant differences favoring RF were found (SMD – 1.04; 95% CI – 1.65 to – 0.44; p < 0.0007). Heterogeneity was high, particularly in the short and medium term (I² = 87 and 90%, respectively). The results are shown in Fig. 4.

Improvement in QoL

Only two RCTs (Nath et al., 2008; Van Kleef et al., 1999) included results on QoL, using different questionnaires. QoL was analyzed for short and medium term outcomes combined, and no statistically significant differences were observed (SMD – 0.28; 95% CI – 0.75 to 0.18; p = 0.23).

Heterogeneity among the studies was low ($I^2 = 0\%$). The results of the analysis are shown in Fig. 5.

Global Perceived Effect

GPE was analyzed in six RCTs [17, 18, 20, 21, 25, 27]. No statistically significant differences were found in GPE measured as a continuous variable (SMD 0.04; 95% CI – 0.55 to 0.63; p = 0.90). The heterogeneity among studies was moderate (I² = 64%).

Moussa et al. (2020), Van Wijk et al. (2005), and Tekin et al. (2007) measured GPE as variables that could be grouped as dichotomous. In the short term, the results

	Radi	ofrequer	ncy	Sham				Mean difference	Mean difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	IV, Random, 95% Cl
1.1.1 Short term									
Gallagher et al.	3.4	0.69	18	6	0.98	12	6.9%	-2.60 [-3.24 , -1.96]	
Leclaire et al.	5.23	2.7	36	4.44	1.8	34	5.9%	0.79 [-0.28 , 1.86]	
Moussa et al.	3.05	1	50	2.63	1.3	50	7.3%	0.42 [-0.03 , 0.87]	
Tekin et al. (CRF)	2.3	1.4	20	4.3	1	10	6.4%	-2.00 [-2.87 , -1.13]	
Tekin et al. (PRF)	2.8	1.5	20	4.3	1	10	6.3%	-1.50 [-2.40 , -0.60]	
Van Kleef et al.	2.83	2.4	15	4.77	2.5	16	4.2%	-1.94 [-3.67 , -0.21]	
Van Tilburg et al.	5.3	1.8	30	5.5	1.9	30	6.2%	-0.20 [-1.14 , 0.74]	
van Wijk et al.	3.7	1.8	40	4.9	1.8	41	6.6%	-1.20 [-1.98 , -0.42]	
Subtotal (95% CI)			229			203	49.9%	-1.01 [-1.98 , -0.04]	
Heterogeneity: Tau ² =	1.72; Chi2 =	= 80.41, 0	df = 7 (P <	< 0.00001)	; l² = 91%				•
Test for overall effect:	Z = 2.04 (P	= 0.04)							
1.1.2 Medium term									
Gallagher et al.	4.4	0.72	18	7	0.85	12	7.1%	-2.60 [-3.18 , -2.02]	
Moussa et al.	3.25	1.1	50	5.66	0.4	50	7.5%	-2.41 [-2.73 , -2.09]	-
Nath et al.	3.9	4.8	20	3.7	4.8	20	2.2%	0.20 [-2.78, 3.18]	
Tekin et al. (CRF)	2.3	1.3	20	3.1	0.8	10	6.7%	-0.80 [-1.56 , -0.04]	
Tekin et al. (PRF)	2.9	1.6	20	3.1	0.8	10	6.4%	-0.20 [-1.06 , 0.66]	
Subtotal (95% CI)			128			102	29.9%	-1.42 [-2.41 , -0.43]	
Heterogeneity: Tau ² =	1.00; Chi ² =	= 38.67, 0	df = 4 (P <	< 0.00001)	; I² = 90%				•
Test for overall effect:	Z = 2.80 (P	= 0.005)							
1.1.3 Long term									
Moussa et al.	6.25	0.6	41	7.46	0.2	41	7.7%	-1.21 [-1.40 , -1.02]	-
Tekin et al. (CRF)	2.4	1.1	20	3.9	1.2	10	6.3%	-1.50 [-2.39 , -0.61]	
Tekin et al. (PRF)	3.5	1.3	20	3.9	1.2	10	6.2%	-0.40 [-1.34 , 0.54]	
Subtotal (95% CI)			81			61	20.2%	-1.12 [-1.57 , -0.68]	•
Heterogeneity: Tau ² =	0.07; Chi2 =	= 3.24, df	= 2 (P =	0.20); l ² =	38%				•
Test for overall effect:	Z = 4.94 (P	< 0.0000	01)						
Total (95% CI)			438			366	100.0%	-1.14 [-1.66 , -0.61]	•
Heterogeneity: Tau ² =	0.93; Chi ² :	= 171.39,	df = 15 (P < 0.0000	01); I² = 91	1%			•
Test for overall effect:	Z = 4.25 (P	< 0.000	1)						-4 -2 0 2 4
Test for subgroup diffe	erences: Ch	i² = 0.37,	df = 2 (P	= 0.83), l ²	= 0%			Favours	Radiofrequency Favours Place

Fig. 3 Forest plot comparing radiofrequency versus sham for pain relief at different time intervals after treatment or sham intervention

showed a trend toward significance favoring RF (OR 0.55; 95% CI 0.31 to 1; p = 0.05). Statistically significant differences favoring RF were observed in the medium (OR 0.19; 95% CI 0.07–0.52; p = 0.001) and long term (OR 0.22; 95% CI 0.06 to 0.78; p = 0.02). Overall, significant benefits in favor of RF over placebo were found (OR 0.38; 95% CI 0.24–0.6; p < 0.0001). Heterogeneity was low (I²=0–3%). The results of the analysis are shown in Fig. 6.

Subgroup analysis

Low back pain duration prior to patient inclusion

Regarding pain relief in the short term, the RCTs that established LBP duration > 1 year in the inclusion criteria [17, 21, 25] showed no significant differences, while significant differences favoring RF were found in the group of LBP < 1 year [18–20, 27, 36] (MD – 1.16; 95% CI – 2.11

to -0.20). In the medium and long term there were similar results between both groups.

For functional status, significant differences favoring RF were found in both groups in the medium and long term. Although no significant differences were observed in the short term in either group, a trend toward significance (p=0.11) was observed in the group of LBP < 1 year. Supplementary File 1 presents the forest plots for the subgroup analyses.

MRI prior to patient inclusion

Regarding pain relief, the RCTs that included the performance of MRI as an inclusion criterion [17, 21, 27] showed significant differences favoring RF in the long term (MD: -1.21; 95% CI -1.40 to -1.02), but not in the short or medium term. In the RCTs where the previous performance of MRI was not established as inclusion

	Radi	ofrequer	icy	Sham				Std. mean difference	Std. mean difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
1.2.1 Short term									
Leclaire et al.	33.6	6.1	36	33.7	5.7	34	9.8%	-0.02 [-0.49 , 0.45]	_
Moussa et al.	-34.9	6.1	50	-33.6	5.7	50	10.0%	-0.22 [-0.61, 0.17]	-
Tekin et al. (CRF)	25.6	6.5	20	30.5	5.7	10	8.9%	-0.76 [-1.55 , 0.02]	
Tekin et al. (PRF)	24.4	5.7	20	30.5	5.7	10	8.8%	-1.04 [-1.85 , -0.23]	
Van Kleef et al.	19.93	6.1	15	39.69	5.7	16	7.7%	-3.26 [-4.38 , -2.14]	
Subtotal (95% CI)			141			120	45.1%	-0.94 [-1.73 , -0.14]	•
Heterogeneity: Tau ² =	0.69; Chi ² :	= 31.54, 0	df = 4 (P <	(0.00001)	; l² = 87%				•
Test for overall effect:	Z = 2.30 (P	= 0.02)							
1.2.2 Medium term									
Moussa et al.	-30.3	6.65	50	-10.8	5.7	50	9.5%	-3.12 [-3.72 , -2.53]	-
Tekin et al. (CRF)	25.1	6.4	20	28.9	5.7	10	8.9%	-0.60 [-1.37, 0.18]	
Tekin et al. (PRF)	25.3	6.9	20	28.9	5.7	10	8.9%	-0.54 [-1.31, 0.24]	
Subtotal (95% CI)			90			70	27.3%	-1.43 [-3.24 , 0.37]	
Heterogeneity: Tau ² =	2.40; Chi2 =	= 38.75, 0	tf = 2 (P <	(0.00001)	; l² = 95%				
Test for overall effect:	Z = 1.56 (P	= 0.12)							
1.2.3 Long term									
Moussa et al.	-6.3	6.6	41	-2	5.7	41	9.9%	-0.69 [-1.14 , -0.24]	-
Tekin et al. (CRF)	28	7.1	20	33.6	5.7	10	8.8%	-0.82 [-1.61 , -0.02]	
Tekin et al. (PRF)	28.5	6.1	20	33.6	5.7	10	8.8%	-0.83 [-1.62 , -0.04]	
Subtotal (95% CI)			81			61	27.6%	-0.74 [-1.09 , -0.39]	•
Heterogeneity: Tau ² =	0.00; Chi ² :	= 0.13, df	= 2 (P =	0.94); l² =	0%				
Test for overall effect:	Z = 4.17 (P	< 0.000	1)						
Total (95% CI)			312			251	100.0%	-1.04 [-1.65 , -0.44]	•
Heterogeneity: Tau ² =	0.90; Chi ² =	= 98.52, 0	df = 10 (P	< 0.0000	1); I ² = 909	%			•
Test for overall effect:	Z = 3.40 (P	= 0.0007	7)						-4 -2 0 2 4
Test for subgroup diffe	erences: Ch	i² = 0.69,	df = 2 (P	= 0.71), l ²	= 0%			Favours	Radiofrequency Favours Sham

Fig. 4 Forest plot comparing radiofrequency versus placebo for improvement in functional disability at different time intervals after treatment or sham intervention

	Radi	ofrequen	су		Sham			Std. mean difference	Std. mean difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% CI	IV, Fixed, 95% CI
1.3.1 Short and Medi	um term								
Nath et al.	2.57	0.49	20	2.73	0.69	20	56.5%	-0.26 [-0.88 , 0.36]	
Van Kleef et al.	17.07	9.1	15	19.98	9.1	16	43.5%	-0.31 [-1.02, 0.40]	_
Subtotal (95% CI)			35			36	100.0%	-0.28 [-0.75, 0.18]	
Heterogeneity: Chi2 =	0.01, df = 1	(P = 0.92	2); I ² = 0%	D					•
Test for overall effect:	Z = 1.19 (P	= 0.23)							
Total (95% CI)			35			36	100.0%	-0.28 [-0.75 , 0.18]	
Heterogeneity: Chi2 =	0.01, df = 1	(P = 0.92	2); I ² = 0%	D					-
Test for overall effect:	Z = 1.19 (P	= 0.23)						_	1 + 1 + 1 + 1 + 1 + 1 + 1 + 1 + 1 + 1 +
Test for subgroup diffe	rences: No	t applicat	ole					Favours R	adiofrequency Favours Sham

Fig. 5 Forest plot comparing RF treatment with placebo for improvement in quality of life at different time intervals after treatment or sham intervention

criterion [18–20, 25, 26], significant differences favoring RF were found in the short term (MD – 1.42; 95% CI – 2.35 to – 0.50), with a trend toward significance in the medium (p = 0.11) and long (p = 0.08) term. For functional status, no significant differences were found in the group with prior MRI in the short term, but significant differences favoring RF were found in the medium (SMD: -3.12; 95% CI: -3.72 to -2.53) and long

	Radi	ofrequer	nev.		Sham			Std mean difference	Std mean difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	IV, Random, 95% CI
1.4.1 Short term									
Van Kleef et al.	1.33	1.25	15	0.37	7 1.25	16	29.1%	0.75 [0.02 , 1.4	81
Van Tilburg et al.	3.4	1	30	3.7	7 1.3	30	37.8%	-0.26 [-0.76 , 0.2	51
Subtotal (95% CI)			45			46	66.8%	0.21 [-0.77 . 1.15	91
Heterogeneity: Tau ² =	0.40: Chi ² :	= 4.87. df	f = 1 (P =	0.03); l ²	= 79%			•	
Test for overall effect:	Z = 0.42 (P	= 0.67)		,					
1.4.2 Medium term									
Nath et al.	2.75	1.2	20	3.05	5 1.2	20	33.2%	-0.25 [-0.87 , 0.3	81
Subtotal (95% CI)			20			20	33.2%	-0.25 [-0.87 . 0.34	B1
Heterogeneity: Not ap	plicable							• • • • • • • • • • • • • • • • • • • •	
Test for overall effect:	Z = 0.77 (P	= 0.44)							
Total (95% CI)			65			66	100.0%	0.04 [-0.55 , 0.65	3]
Heterogeneity: Tau ² =	0.17; Chi ² :	= 5.55, dt	f = 2 (P =	0.06); l ²	= 64%			-	
Test for overall effect:	Z = 0.13 (P	= 0.90)							
Test for subgroup diffe	rences: Ch	i² = 0.59,	df = 1 (P	= 0.44),	l² = 0%			Favou	rs Radiofrequency Favours Sham
	Radio	freque	ncy	Shar	n		Odds ra	tio (Non-event)	Odds ratio (Non-event)
Study or Subgroup	e Event	s To	tal Ev	ents	Total	Weight	М-Н, І	Fixed, 95% Cl	M-H, Fixed, 95% Cl
1.5.1 Short term									
Moussa et al.		30	50	25	50	25.7%		0.67 [0.30 , 1.47]	
van Wijk et al.		24	39	16	39	24.3%		0.43 [0.18 , 1.08]	
Subtotal (95% CI)			89		89	50.0%	9	0.55 [0.31 , 1.00]	•
Total events:		54		41					
Heterogeneity: Chi ²	= 0.48, df	= 1 (P =	= 0.49); I	² = 0%					
Test for overall effect	ct: Z = 1.94	4 (P = 0	.05)						
1.5.2 Medium term									
Moussa et al.		21	50	6	50	31.7%		0.19 [0.07 , 0.52]	_ _
Subtotal (95% CI)			50		50	31.7%	-	0.19 [0.07 , 0.52]	◆
Total events:		21		6					•
Heterogeneity: Not a	applicable								
Test for overall effect	ct: Z = 3.20	0 (P = 0.	.001)						
1.5.3 Long Term									
Moussa et al.		3	41	0	41	5.9%		0.13 [0.01 , 2.65]	
Tekin et al. (CRF)		19	20	7	10	6.5%		0.12 [0.01 , 1.39]	
Tekin et al. (PRF)		17	20	7	10	5.8%		0.41 [0.07 , 2.56]	
Subtotal (95% CI)			81		61	18.3%		0.22 [0.06 , 0.78]	
Total events:		39		14					
Heterogeneity: Chi2	= 0.79, df	= 2 (P =	= 0.67); I	² = 0%					
Test for overall effect	ct: Z = 2.34	4 (P = 0.	.02)						
Total (95% CI)			220		200	100.0%		0.38 [0.24 , 0.60]	•
Total events:	1	14		61					
Heterogeneity: Chi ²	= 5.16, df	= 5 (P =	= 0.40); I	² = 3%					0.01 0.1 1 10 100
Test for overall effect	ct: Z = 4.09	9 (P < 0.	.0001)					Favours R	adiofrequency Favours Sham
Test for subgroup di	ifferences:	Chi ² = 4	4.09, df =	= 2 (P =	0.13), I ² =	= 51.1%			

Fig. 6 Forest plot comparing radiofrequency versus placebo for global perceive effect (GPE) at different time intervals after treatment or sham intervention. Top, GPE measured as a continuous variable. Bottom, GPE measured as a dichotomous variable

(SMD: -0.69; 95% CI: -1.14 to -0.24) term. In the other group, significant differences were observed in the short, medium and long term. Supplementary File 2 presents the forest plots for the subgroup analysis.

Heterogeneity and publication bias

A subgroup analysis of all associations including 3 or more studies was performed according to their quality (Supplementary File 3). The heterogeneity of most associations (5 out of 7) disappeared in the highest-quality subgroup, suggesting that the low quality of certain studies might represent a relevant source of heterogeneity. Shortterm associations presented more favorable estimates for RF in the higher-quality subgroup.

Regarding publication bias, funnel plots were obtained only for pain and functional status due to the low number of studies in the other analyzed variables [37]. The funnel plots did not suggest publication bias (Supplementary File 4).

Sensitivity analysis

The sensitivity analysis showed differences in pain and functional status when excluding certain studies in different time intervals. Specifically, for pain, the exclusion of the studies by Gallagher et al. (1994) in the medium term and Moussa et al. (2020) in the long term led to significant modifications (from "favorable to RF" to "no significant differences"). In the short term, no significant variations in the overall effect size were observed.

Regarding functional status, the exclusion of Van Kleef et al. (1997) in the short term led to significant modifications (from "significant in favor of RF" to "not significant"), and a reduction in heterogeneity (I^2 from 87 to 51%). Similarly, the exclusion of Moussa et al. (2020) in the medium term led to significant modifications (from "not significant" to "favorable to RF"), associated with significant changes in heterogeneity (from 95 to 0%). No sensitivity analysis was conducted for the remaining comparisons due to the low number of studies.

Discussion

This meta-analysis included eight placebo-controlled RCTs with a total of 472 patients (249 in the experimental group and 223 in the sham group). The results indicate that RF provides significant benefits in terms of pain relief and improvement in functional disability in the short, medium, and long term compared to placebo, which is consistent with previous studies [17, 21, 25]. However, the benefits in terms of QoL and perceived global effect are inconclusive, mainly due to the low number of RCTs that evaluated these variables in a comparable manner, although there are cues suggesting favorable benefits in GPE, consistent with previous studies [38]. Overall, the analysis of risk of bias indicates that the quality of the studies is adequate, although there may be information biases, as reported elsewhere [29, 38]. Although no clear signs of publication bias were found, its assessment is limited due to the low number of RCTs. A high heterogeneity was found among studies, with the sensitivity analysis showing a mild influence of some studies (e.g., Gallagher et al., Moussa et al.).

The subgroup analysis according to the quality of the studies showed interesting results. First, the heterogeneity disappeared in most of the associations in the subgroup of higher-quality studies. This suggests that lower-quality studies represent a relevant source of heterogeneity and, therefore, future RCTs should focus on avoiding these biases. Second, in the higher-quality subgroup, the pooled estimates of short-term outcomes were much higher than in the lower-quality (less reliable) group (-1.57 vs. - 0.42 for pain relief, -1.63 vs. - 0.14 for improvement in functional disability). This fact reinforces our results, showing that higher-quality studies showed even more favorable outcomes for RF in the short term.

Several systematic reviews and meta-analyses on this topic have been published. For example, a systematic review of RCTs conducted by Leggett et al. (2014) reported shortterm benefits in favor of RF [39], while Manchikanti et al. (2020) found long-term pain relief benefits with level II evidence [15]. These findings contradict a Cochrane review published by Maas et al. (2015), which reported the absence of high-quality studies suggesting benefits of RF in chronic LBP [29]. Lee et al. (2017) published a meta-analysis with 7 RCTs and a total of 454 patients [40] which found benefits favoring RF compared to the control group in pain relief for up to 12 months, in line with our findings. Similarly, Chen et al. (2019) evaluated the efficacy of RF in the treatment of LFJP and sacroiliac joint pain [38], and reported favorable results for RF in pain relief, functionality, and QoL. Very recently, the meta-analysis conducted by Janapala et al. (2021) concluded that there is level II evidence in favor of RF efficacy [41]. These results are consistent with our findings, which includes the largest number of placebocontrolled RCTs published to date, and focuses exclusively on traditional RF modalities (continuous and pulsed).

A noteworthy aspect of our study is the evaluation of QoL, which has only been assessed in the previous metaanalysis by Chen et al. (2019). We found no significant differences between groups, although the number of RCTs included is very low. In addition, we conducted an analysis on GPE and found significant differences in favor of RF. However, the number of studies is limited, warranting a more comprehensive and standardized approach when assessing QoL and GPE in future research.

The subgroup analysis based on the duration of LBP suggests an influence on the response to RF, being more favorable in less chronic cases. These findings could be explained by differences in structural spine changes or central sensitization phenomena [42]. The other subgroup analysis shows that patients who did not undergo an MRI before inclusion in the study reported more pain relief in the short term. This could be explained by a selection bias,

as patients with different conditions that could show a more favorable response to RF might have been included, or by the influence of MRI on patients' expectations of treatment [43, 44]. It should be noted that these subgroup analyses were predicated on certain biologically plausible hypotheses that might introduce bias, such as temporal variations in painrelated neuromodulation and macroscopic edema resulting from local inflammation. Other potential sources of bias, such as heterogeneity in sham-related procedures, a wellknown controversial topic in spinal pain-related procedures [45], have been discussed elsewhere [15, 31] and therefore were not specifically explored in this meta-analysis.

This meta-analysis has several strengths and some limitations. Remarkable strengths include the quantitative analysis of QoL and GPE, which have been poorly analyzed in previous studies [38], and the subgroup analyses conducted, which allowed to suggest hypotheses to consider in future research. Regarding the limitations, there is a relatively low number of RCTs, which limits the statistical power of the meta-analysis [46], high heterogeneity in study design, inclusion criteria, and outcome measures. In addition, the cutoff points chosen for short, medium, and long-term follow-up times lack universal consensus and thus could entail a potential source of bias, although they are comparable to previous meta-analyses (e.g., [29]). These limitations should be taken into account in future studies.

Conclusion

Radiofrequency treatment for Lumbar facet joint pain provides significant benefits compared to placebo in terms of pain relief in the short, medium, and long term, as well as improved functionality in the short and long term. However, the evidence for benefits in quality of life and perceived global effect is inconclusive. The duration of low back pain and performing an MRI before treatment may influence therapeutic response. Future clinical trials should investigate the long-term effects of RF, its impact on quality of life, and define appropriate criteria for patient selection.

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Declarations

Conflict of interests The authors have no relevant financial or non-financial interests to disclose.

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Consent to participate Not applicable.

Consent to publish Not applicable.

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