

# Association between arthralgia and imaging findings of effusion in the temporomandibular joints: a systematic review

## Abstract

**Aims:** A systematic review was conducted to determine if there is an association between joint effusion and self-reported TMJ-pain during examination and between joint effusion and arthralgia as defined by the Research Diagnostic Criteria for Temporomandibular Disorders (RDC-TMD).

**Methods:** Four reviewers separately identified observational studies evaluating a possible association between ipsilateral joint effusion, identified by MRI T2-weighted images, and TMJ pain by systematically searching three databases.

**Results:** A total of 67 articles were identified with the search strategy. However, 32 met the inclusion criteria for the systematic review.

**Conclusion:** Based on the review of 32 articles, published on this topic, body of literature was unable to provide evidence to support or refute the association between joint effusion and self-reported TMJ-pain, and between joint effusion and arthralgia as defined by RDC-TMD.

**Keywords:** temporomandibular disorders, joint effusion, MRI, arthralgia, pain

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**Abbreviations:** MRI, magnetic resonance imaging; JE, joint effusion; RDC/TMD, research diagnostic criteria for temporomandibular disorders; VGIR, visually guided TMJ irrigation; ID, internal derangement; DJD, degenerative joint disease; DDwR, disk displacement with reduction; DDwoR, disk displacement without reduction; OA, osteoarthritis; CDC/TMD, clinical diagnostic criteria for temporomandibular disorders; OR, odds ratio; VAS, visual analogue scale; ICC, interclass correlation coefficients; SSI, symptom severity index

## Introduction

The temporomandibular disorders (TMD) encompass a group of musculoskeletal and neuromuscular conditions that involve the temporomandibular joints (TMJs), the masticatory muscles, and associated tissues.<sup>1</sup> Common manifestations of TMD consist of pain of a persistent, recurring, or chronic nature in the TMJ, masticatory muscles, or in the adjacent structures; limitation or other alterations in the range of mandible motion; and TM joint noises.<sup>2</sup> It has been suggested that differential diagnosis of TMDs should be based primarily on information obtained from the patient's history, clinical examination, and when TMJ imaging procedures indicated.<sup>1</sup> Magnetic resonance imaging (MRI) is an example of such imaging procedures. It provides excellent representation of soft tissues in anatomical and semi-functional relationships. In addition, it may also be used for detection of the presence of joint effusion with acceptable levels of reliability.<sup>3</sup> Some authors have also suggested use of MRI for assessment of hard tissues; however the reliability has been shown to be poor.<sup>3-5</sup> Joint effusion is defined as a collection of fluid in the joint space. It is manifested as areas of high intensity signal in the TMJ space, on T2-weighted images.<sup>3-5</sup> It has been suggested that this

accumulation of fluid could be a surrogate of an inflammatory process that may activate or sensitize nociceptive afferent neurons within the joint.<sup>4-6</sup> Alternatively, it could result in increased intra-articular pressure, which may cause mechanical trauma, leading to hypoxia and other inflammation induced changes within the joint space, and eventually leads arthralgia.<sup>7</sup> Several studies have been conducted on the association of joint effusion and TMJ pain associated with TMD. However, the results have been inconsistent. While the majority of studies have reported a statistically significant association,<sup>4-26</sup> others have reported inconclusive findings.<sup>5-30</sup> Due to this difference in reporting, a systematic review was conducted to determine if there is an association between joint effusion and self-reported TMJ-pain during examination, as well as between joint effusion and arthralgia as defined by the Research Diagnostic Criteria for Temporomandibular Disorders (RDC-TMD).<sup>31</sup>

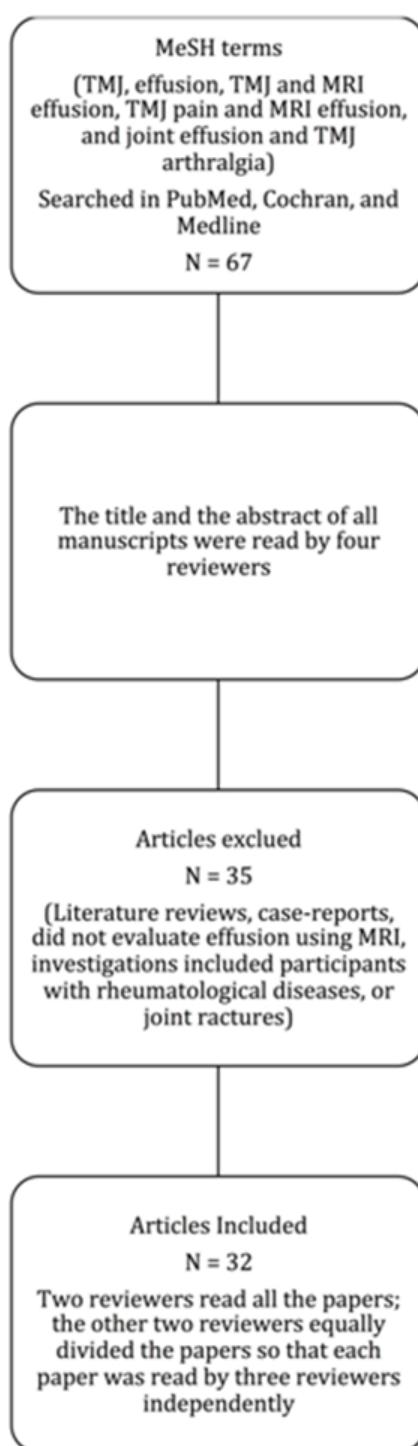
## Materials and methods

The clinical question: "Is there an association between MRI identified temporomandibular joint effusion and self-reported TMJ-pain?" was utilized to guide the review.

- Inclusion Criteria: Observational studies evaluating a possible association between ipsilateral joint effusion, identified by MRI T2-weighted images, and TMJ pain were included. Manuscripts were limited to the English language only, published from inception till January 2017.
- Exclusion Criteria: All investigations that included participants with rheumatologic diseases, or joint fractures were excluded. In addition, publications such as literature reviews and case-reports were also excluded from the review.

- c. Literature Search: Four reviewers separately identified the pertinent literature by searching PubMed, MEDLINE and Cochrane databases under the inclusion criteria of all studies that have been conducted to determine an association between joint effusion and pain in the TMJ. The MeSH terms included TMJ, effusion, MRI, pain and arthralgia.
- d. Procedure: Literature review was performed in accordance to PRISMA statement (Preferred Reporting Items for Systematic Reviews and Meta-Analyses).Four reviewers independently

read the title and the abstract of all publications that matched the MeSH search terms. The reviewers met and developed a final list of publications to be read by consensus. Two reviewers read all the papers (SKN, RFGM); the other two reviewers (HC, YG) equally divided the number papers so that each paper was evaluated independently by three reviewers (Figure 1). Articles were reviewed based on the aim of the study, demographics, methodology, diagnostic criteria, MRI protocol, results, and conclusions.



**Figure 1** Flow chart summarizing the process of literature search.

## Results

Three databases (PubMed, Cochran, and Medline) were systematically searched for articles. A total of 67 articles were identified with the search strategy. Thirty-two met the inclusion criteria, and all were observational studies. Thirty-five publications were excluded because they did not evaluate the association of TMJ effusion with TMJ pain, did not use MRI to evaluate effusion, were case-reports, or literature review studies. The summary of 32 included articles is provided in Table 1. The majority of the included studies reported a statistically significant association between joint effusion and self-reported TMJ-pain. However, during the review of these studies,

several methodological limitations were identified. For example: absence of control group;<sup>4–32</sup> lack of information about examiner or radiologist reliability;<sup>10–26</sup> failure to report details regarding the clinical examination;<sup>4–32</sup> joint pain assessed by self-report (history) only without a clinical examination;<sup>15</sup> or inappropriate statistical analysis.<sup>25</sup> Among the studies that failed to find an association between TMJ effusion and self-reported TMJ-pain, several methodological limitations were also present. These include inappropriate control groups;<sup>5–30</sup> lack of information about the evaluation of TMJ pain, or evaluation using a non-standardized examination;<sup>6–30</sup> or lack of information regarding examiner and radiologist reliability.<sup>6–30</sup>

**Table 1** Summary of the studies included in the systematic review on the association of TMJ effusion and TMJ pain

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Khawaja et al. <sup>8</sup>	Aim: To determine the clinical and radiological significance of JE in patients with TMD.	N= 158 ; 312 TMJs  59.4 % ♀  age: 31±11.1 yrs.  40.6 % ♂  age: 29.8±9.7 yrs.	Study group: Participants underwent DC/ TMD examination by calibrated examiners	T2-weighted sagittal sequences (1.5 T, 3mm thick cuts)	T-test, Pearson Chi-Square,  Generalized estimating equation analysis	No association was determined between JE and TMJ arthralgia. Statistical association was determined between JR and dis position in the coronal and in the sagittal plan.  GEE suggested that disc displacement with reduction in the sagittal plan was statistically significant contributing factor for JE.	Pain was no categorized on basis of intensity.
Oliveira et al. <sup>4</sup>	N= 74 ; 148 TMJs	  68.9 % ♀  age: 40.4±14.5 yrs.  31.1 % ♂  age: 35.9±11.2 yrs.	  Study group: Participants reported at least one sign or symptom of TMD during clinical examination	T2-weighted sagittal sequences (1.5 T, 3mm thick cuts)	Prevalence	Osteoarthritic changes were associated with JE.	No control group
	Aim: To evaluate the association between condylar bone changes and presence of TMJ effusion in symptomatic patients			JE defined: any high signal intensity in the articular space.	Fisher's exact test	Prevalence of JE in symptomatic patients was 10 %.	Clinical examination was not adequately described.
				Two radiologists, diagnosis by consensus			Reliability of clinical examiners and radiologists was not described.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Lamot U et al. <sup>14</sup>	Aim:  a. To determine correlation between MRI and clinical findings associated with TMD.  b. To assess the impact of gender and age, on this correlation.	N= 104 ; 288 TMJs  75.7 % ♀ ; 24.3 % ♂  Total sample	Study group: Participants had unilateral or bilateral presence of TMJ pain, noises/clicking, limited movement and headache. Study group was divided into 3 sub-groups based on type of ID.	T2-weighted sagittal sequences (1.5 T, 20 slices)  JE defined: any high signal intensity in anterior or posterior recess, or large amount in superior or inferior joint space.	Prevalence  Fisher's exact test	Morphological changes associated with symptoms of TMJ dysfunction. Prevalence of JE in symptomatic joints was 14.6 %  Younger age group was associated with JE	Clinical examination was not adequately described.  Reliability of clinical examiners and radiologists was not described
Santos et al. <sup>20</sup>	Aim:To assess the association between changes of the articular eminence and the condyle, articular disc changes, and the presence of TMJ effusion in symptomatic patients.	N = 71 ; 142 TMJs  69 % ♀ ; 31 % ♂  Total sample	Study group: Participants reported at least one sign or symptom of TMD during examination	T2-weighted sagittal sequences (1.5 T, 3mm thick cuts)  JE defined: any hyper-signal in the articular space.	Prevalence  Pearson's Chi-Square test	Association was present between: 1) disc form and its position; 2) condylar form and an anterior portion of disc; and 3) articular eminence form and disc form.  Prevalence of JE in symptomatic joints was 23.9 %.	No control group  Clinical examination was not adequately described.
		age: 39.4 yrs. (12 – 81 yrs.).  age: 38.7 yrs. (13 – 69 yrs.).		Two radiologist, diagnosis by consensus	Fisher's exact test		Reliability of clinical examiners and radiologists was not described

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Park et al. <sup>18</sup>	Aim: To investigate the value of MRI in the diagnostic process based on the RDC/TMD.	N = 100 ; 200 TMJs 68.0 % ♀ with age: 25.3±13.12 yrs.	Study group: Participants underwent RDC/TMD examination.	T2-weighted sagittal sequences (1.5 T)	Cohen's Kappa	Cohen's Kappa showed no agreement to moderate agreement between MRI and Group II diagnosis based on RDC/TMD	No information if radiologists were blind to the clinical diagnosis
		32 % ♂ with mean age: 31.9±12.3 yrs.	Study group was divided into 3 distinct Group II diagnoses based on RDC/TMD.	JE defined: any high intensity signal in articular surface or compartment.	Chi-Square test	JE and ipsilateral pain were significantly associated.	Reliability of clinical examiners and radiologists was not reported.
Bas et al. <sup>9</sup>	Aim: To evaluate the relationship between the grades of MRI depicted JE, increased capsular width (CW) measured in Ultrasonographic Imaging (USI), and joint pain.	N = 91 ; 182 TMJs 81.3 % ♀ 18.7 % ♂ age: 25 yrs.	Study group: Participants underwent a. RDC/TMD examination.	T2-weighted sagittal sequences (0.5 T, 3mm thick cuts)	Receiver Operating Characteristic curve	Cutoff value for CW was calculated to be 1.65 mm	No control group
		Total sample	b. VAS to measure pain intensity.	JE defined: area of homogenous and high signal intensity.	Friedman Test	Statistically significant association was found between VAS score and intensity of MRI- JE.	Reliability of clinical examiners and radiologists was not described
				JE was characterized into moderate and severe effusion.	Wilcoxon Test	JE absent: 2.55 VAS	
				CW was assessed by USI		Moderate JE: 2.92	
						Severe JE: 4.80	

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Kaneyama et al. <sup>13</sup>	Aim: To determine the association between JE and various soluble cytokine receptors in the synovial fluid.	N = 55 ; 55 TMJs	Study group: Participants had diagnosis of TMD based on  a. Clinical symptoms and MRI.	T2-weighted sagittal sequences (1.5 T, 3mm thick cuts) JE defined: area of high intensity signal in superior compartment. JE was characterized into grade I, II, and III based on the quantity of signal.	Mann-Whitney U test	The mean concentration of cytokine receptors in the synovial fluid were significantly higher in joints with JE than in joints without JE.	Gender and age demographics not provided.
			b.VAS used to measure ipsilateral joint pain intensity.	Three radiologist, diagnosis by consensus.	Spearman's correlation	JE was reported in 55.5 % joints.	Clinical examination was not adequately described.
Nakaoka et al. <sup>32</sup>	Aim: To investigate the changes of JE on the MRI and pathology observed in arthroscopy after VGR.	N = 56 ; 56 TMJs	Study group: Participants underwent clinical examination.	T2-weighted sagittal sequences (0.3 T)	Wilcoxon signed-rank sum test	No statistically significant difference was observed between severity of JE and VAS.	Reliability of clinical examiners and radiologists was not described.
	a. To correlate these findings with clinical variables.	40 participants completed the study	Study group had a clinical diagnosis of unilateral chronic closed lock.	JE defined: area of high signal intensity.	Mann-Whitney U test	JE was significantly higher in the group with poor surgical outcome.	No control group.
		Total sample		JE was characterized into grade I, II, and III.	Spearman's correlation	No statistical correlation between JE and synovial lining score.	Clinical examination was not adequately described.
		age 43 yrs.		Three radiologist, diagnosis by consensus			Reliability of clinical examiners and radiologists was not described.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Costa et al. <sup>10</sup>	Aim: To assess the correlation of TM joint ID in patients with the presence of headache, bruxism, and joint pain using MRI.	N = 58 ; 58 TMJs	Study group: Participants underwent a. RDC/TMD examination by a calibrated examiner.	T2-weighted sagittal sequences (2 T, 1.5 mm thick cuts)	Chi-square test	Participants with TMD and headaches had significantly more ID than the control group.	High signal in the inferior TM joint space was not considered as indicative of JE. Reliability of clinical examiners and radiologists was not described.
		TMD group: 42 participants, 83.3 % ♀, 16.7 % ♂. Age range of 16-83 yrs.	b. Headache assessment by a trained neurologist.	JE defined: area of high signal intensity in the region of upper and lower joint spaces.	Fisher's exact test	JE was statistically more prevalent in group with TMD and headaches	Tables and text do not match.
Farina et al. <sup>28</sup>	Aim: To evaluate the association between conventional and enhanced MRI findings, and TMJ pain.	N = 38 ; 78 TMJs	Study group: Participants had TMD pain and dysfunction.	T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts)	Chi-square	33.3 % joints with pain had JE.	Age information of the participants not provided.
		81.6 % ♀, 18.4 % ♂	Control group: Participants did not have any TMD pain or dysfunction.	JE defined: presence of high signal intensity within joint space in open- or closed-mouth images.	T-test	21.7 % of joints without pain had JE.	Clinical examination was not adequately described.
					Multivariate logistic regression analysis.	Statistical correlation between TMJ pain and JE was not observed.	Reliability of clinical examiners and radiologists was not described.
						Odds ratio that painful TMJ had JE was 1.2	34.2 % joints had been on NSAID before MRI.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Limchaichana et al. <sup>5</sup>	Aim: To evaluate the association between MRI findings and clinical symptoms of TMD.	N = 60 ; 120 TMJs 78.3 % ♀, 21.7 % ♂	Study group: Participants underwent RDC/TMD examination by a calibrated examiner.	T2-weighted sagittal sequences (1.5 T)  JE defined: area of more than a line or dots of high signal intensity in the region of superior or inferior joint space.	T-test  Fisher's exact test	18 % of joints with myofascial pain only, had JE.  33 % of joints with myofascial pain and arthralgia/OA had JE.	Overlapping of diagnosis – both groups had masticatory muscle pain.
Guler et al. <sup>29</sup>	Aim: To determine the association between JE, protein concentration in synovial fluid, and TMJ pain.	N = 31 ; 31 TMJs  Study group: 81.3 % ♀, 18.7 % ♂.  age: 31 yrs. (17-57 yrs.)  Control group: 66.7 % ♀, 33.3 % ♂.  age: 28 yrs. (17-42 yrs.)	Study group: Participants underwent RDC/TMD examination.  a. VAS to measure pain intensity.  Participants were divided into:  a. DDwoR, with arthralgia and OA.  b. DDwoR without any pain-related TMD diagnosis.	Radiologist had inter-observer agreement of 0.69 for JE  JE defined: high-signal area in the upper and lower joint spaces.  JE was characterized into 4 levels depending on quantity of joint fluid.	Chi-square test  Kappa statistic	No statistical difference between two groups in prevalence of JE.  7.7 % joints in control group had JE.  75 % joints in study group had JE.  No significant association was observed between level of pain and JE.	Reliability of clinical examiners and radiologists was not described.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Yano et al. <sup>26</sup>	Aim: To evaluate the association between joint fluid and various disk pathological conditions.	N = 17; 34 TMJs  87.5 % ♀, 12.5 % ♂  age of total sample was 20.5 yrs. (12 – 31 yrs.)	Study group: Participants were diagnosed as having ID of either one or both TMJs.	T2-weighted sagittal sequences (1 T/ 1.5 T, 3 mm thick cuts)  JE defined: high-signal area in the upper and lower joint spaces.	Chi-square test	85.3 % of joints had JE  No significant difference observed between quantity of joint fluid and grade of ID.	Small sample size.  Mean age of the participants is younger than the mean age of patients with TMD.
Emshoff et al. <sup>33</sup>	Aim: To evaluate the association between TMD sub-groups and MRI diagnosis of TMJ ID, osteoarthritis, JE, and bone marrow edema.	N = 164; 164 TMJs  TMJ pain group: 86.4 % ♀, 13.6 % ♂.  age: 36.9 yrs. (12-69 yrs.)  Control group: 67.4 % ♀, 32.6 % ♂.  age: 38.3 yrs. (15-64 yrs.)	Study group: Participants underwent (CDC/TMD) examination.	T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts)  JE was characterized into 4 levels depending on quantity of joint fluid.	Chi-square test	Significant difference was observed between presence of pain and the quantity of joint fluid  Significant association was present between TMJ pain and the MRI diagnosis of TMJ ID for the ID type I group, Capsulitis/Synovitis group, the ID type III group and DJD group.	Clinical examination was not adequately described.  Reliability of clinical examiners and radiologists was not described.  DJD diagnosis was diagnosed using a clinical examination.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Emshoff et al. <sup>34</sup>	Aim: To evaluate whether the MRI variables of TMJ ID, OA, and/or JE may predict the presence of TMJ pain.	N = 42 ; 84 TMJs  95.2 % ♀, 4.8 % ♂.  age of the sample: 38.8 yrs. (16 -77 yrs.)	Study group: Participants underwent CDC/TMD examination.	T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts)	Chi-square test	TMJ ID type-III were significantly associated with high rate of TMJ ID.  JE defined: area of more than one high signal intensity line in the region of joint space, evident in at least two consecutive sections.	Primarily clinical symptoms used without confirming the diagnosis of arthralgia.  A statistically significant association was found between JE and TMJ pain.  JE was observed in 45.2 % of TMJs with pain, while 14.3 % of TMJs without pain had JE.
Guler et al. <sup>11</sup>	Aim: To correlate MRI findings of JE, disc displacement, condylar bone changes and disc form with clinical findings of pain and sounds in patients with bruxing behavior.	N = 94 ; 188 TMJs  Study (bruxing) group: 86.7 % ♀ and 13.3 % ♂.  age: 29 yrs. (13 – 63 yrs.)  Control group: 86.7 % ♀, 13.3 % ♂.  age: 26 yrs. (14 – 50 yrs.)	Study group: Participants underwent examination for	T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts)	Chi-Square	Significantly higher prevalence of condylar bony changes present in joints with displaced disks in the study group.  JE defined: area of high signal intensity in the region of the upper and lower joint spaces.	Data of the subjects included in the analysis were meeting the exclusion criteria.  Reliability of clinical examiners was not described.  Significant association was present between JE and TMJ pain.  30 % of painful joints in the study group, and 59 % of the painful joints in the control group had JE.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Yamamoto et al. <sup>25</sup>	Aim: To evaluate the association between quantity of JE and TMD associated pain and dysfunction.	N = 293; 577 TMJs  82.6 % ♀, 17.4 % ♂.	Study group: Participants had TMD pain and dysfunction.	T2-weighted sagittal sequences (1 T/ 1.5 T)	Kruskal-Wallis test	Statistically significant difference was present for JE, between painful and non-painful joints in the DDwR group.	Clinical examination was not adequately described.
		age: 31.4 yrs. (10-78 yrs.)		JE defined: high-signal area in the upper and lower joint spaces.	Scheffe test	No statistically significant difference was present in painful and non-painful joints of normal, DDwR, or OA groups.	Radiologists were not blind to the clinical diagnosis
				JE was characterized into 4 levels.	Wilcoxon rank sum		Reliability for the quantification criteria for JE not provided.
Emshoff et al. <sup>35</sup>	Aim: To determine if MRI findings are predictive of TMJ pain.	N = 169; 338 TMJs.  85.2 % ♀, 14.8 % ♂.	Study group: Participants underwent CDC/TMD examination.	T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts).	Chi-square test	In multiple logistic regression analysis, ID was significantly associated with TMJ pain; however, OA, JE, and bone marrow edema were statistically not significant.	Radiologists were not blind to the clinical diagnosis.
		age of the sample: 36.9 yrs. (15 -86 yrs.)	Intra-examiner reliability was between K ≥ .75 and K = 1.0.	Intra observer reliability was determined (K ≥ .75 – K = 1.0).		JE was observed in 44.2 % of painful TMJs, and 28.3 % non-painful TMJs	According to multiple logistic model, JE is not associated with TMJ pain, while according to Chi-square test it is.
				JE defined: area of more than one high signal intensity line in the region of joint space, evident in at least two consecutive sections.	Multiple logistic regression analysis	A statistically significant correlation was present between TMJ pain and JE.	

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Manfredini et al. <sup>16</sup>	Aim: To evaluate the predictive value of clinical symptoms for MRI findings of JE.	N = 61	Study group: Participants underwent a. RDC/TMD examination.	T2-weighted sagittal sequences (0.5 T, 3 mm thick cuts) JE defined: area of high signal intensity inside the joint space.	Multiple logistic analysis Goodness-of-fit test	Except for pain during joint play and pain in the TM joint during dynamic tests, rest of the clinical variables were included in the logistic regression analysis.	Gender demographics were not provided.
Emshoff et al. <sup>36</sup>	Aim: To evaluate the association between MRI findings of OA, and/or JE, and TMJ pain.	N = 112 ; 224 TMJs.  86.6 % ♀, 13.4 % ♂  age: 38.2 yrs. (15-78 yrs.)	Study group: Participants underwent CDC/TMD examination.	T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts)  JE defined: area of more than one high signal intensity line in the region of joint space, evident in at least two consecutive sections.	Chi-square  Model Chi-square test	Among single parameters, presence of pain in the TMJ with lateral palpation provided good diagnostic accuracy of 76.2 %.	No control group.
						Reliability of radiologists was not described.	Reliability of clinical examiners and radiologists was not described.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Segami et al. <sup>30</sup>	Aim: To evaluate the association between JE and synovial fluid constituents in patients with ID and OA.	N = 108; 108 TMJs  Study group: 90 % ♀, 10 % ♂.	Study group: Participant underwent  a. MRI	T2-weighted sagittal sequences (1.5 T, 3 mm thick slices).  JE defined: high-signal area in the upper and lower joint spaces. It was further characterized into 4 levels depending on quantity of joint fluid.	Mann-Whitney U test	No statistically significant difference was observed in pain intensity among study groups.  Joints with JE had significantly higher total protein concentration, and Interleukin – 6 levels than joints without JE.	Control group only had ♂.  Clinical examination was not adequately described.
		age: 34.9 yrs. (13-73 yrs.)	b. VAS to measure pain intensity.	Grade 0 and 1 were excluded.		30 % participants in study group were on NSAID.	
Tanaka et al. <sup>23</sup>	Aim:	Control group: All ♂.  age 30 yrs.	Study group had ID or OA, with or without JE.  Control group were symptom free.	FS T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts).	Chi-square test	Detection rate of JE by FS was significantly greater than by T2-weighted MRI.	Clinical examination was not adequately described.
	a. To compare JE evaluation between T2-weighted MRI and Fat saturation (FS) MRI.	76.5 % ♀, 23.5 % ♂.		T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts).	Kappa statistic test	Statistically significant association between JE and TMJ pain.	Reliability of radiologists was not described.
	b. To determine association between JE and MRI findings.	age of case group: 29.3 yrs. (17-52 yrs.)  20 participants were in control group.  65 % were ♀, 35 % ♂.  age of control group: 23.3 yrs. (18-35 yrs.)		JE defined: area of high signal intensity greater than a moderate amount of fluid.		Kappa test indicates good agreement between pain and JE ( $K = 0.66$ ).  FS indicates 70 % of painful joints and 3 % of pain-free joints having JE.	

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Haley et al. <sup>12</sup>	Aim: To investigate the association between TMJ-Pain and clinical and MRI findings.	N = 85; 170 TMJs.	Study group: Participants underwent standardized clinical examination.	T2-weighted sagittal sequences (1.5 T).	Chi-square test	69 % of joints with pain had JE ( $p = .001$ ). OR for this association was 3.8.	All participants were ♀.
	All participants were female.	Age: 16-49 yrs.	Examiners had acceptable levels of reliability.	JE defined: area of high signal intensity area within either joint space in open and closed position.	OR	80 % of joints with pain had ID ( $p = .332$ ). OR for this association was 1.8.	
Rudisch et al. <sup>19</sup>	Aim: To investigate the association between TMJ-Pain and MRI findings of ID and JE.	N = 41 ; 82 TMJs.	a. Study group: Participants underwent CDC/TMD examination.	T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts)	Chi-square	84 % of joints with pain had reported experiencing pain at the onset of study ( $p = .001$ ). OR for this association was 49.	Reliability of clinical examiners and radiologists was not described.
	78 % ♀, 22 % ♂.	age: 39.1 yrs. (17-78 yrs.)		JE defined: area of more than one high signal intensity line in the region of joint space, evident in at least two consecutive sections.	Kappa statistical test	58.5 % of joints with pain and 26.8 % of joints without pain had JE.	
						ID and JE were statistically associated with TMJ-Pain.	
						TMJ-Pain and ID had a K value of 0.34 (poor).	
						TMJ-Pain and JE had a K value of 0.32 (poor).	

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Segami et al. <sup>6</sup>	Aim: To characterize JE with arthroscopic findings	N = 47 ; 47 TMJs  93.6 % ♀, 6.4 % ♂  age: 39 yrs. (13-76 yrs.)	Study group: Participants underwent  I. Clinical examination.  2. VAS to measure pain intensity.	T2-weighted sagittal sequences (1.5T, 3 mm thick slices).  JE defined: high-signal area in the upper and lower joint spaces.  JE was characterized into 4 levels.  Grade 0 and I were excluded.	Mann-Whitney U test  Spearman's correlation co-efficient	68.1 % of joints had JE.  VAS score had no statistical correlation with the presence of JE.	Control group only had ♂.  Clinical examination was not adequately described.  Reliability of clinical examiners and radiologists was not described.
Shaefer et al. <sup>7</sup>	Aim:  a. To evaluate the association between TMJ arthralgia and JE.  b. To evaluate the validity of 1 pound of palpating pressure for diagnosing arthralgia.	N = 30 ; 30 TMJs  100 % ♀  age of study group: 31.3±9 yrs.  age of control group: 31.6±6 yrs.	Study group: Participants underwent  a. RDC/TMD examination, Modified SSI for severity of symptoms.  b. Examiners were calibrated.	T2-weighted sagittal sequences  JE defined: presence of any high intensity signal within either joint space.  Reliability of the radiologist to determine JE was K = 0.85.	Sensitivity and Specificity  T-test  Paired t-test	85 % of subjects with TMJ arthralgia had JE.  72 % of subjects without any TMJ pain had JE.  MRI effusion sensitivity was 85 %, and specificity was 28 %.	All subjects were ♀.  The criterion for TMJ arthralgia was not consistent with RDC/TMD specifications.  Use of the most painful joint instead of both or an average.
							Sensitivity and specificity of 1 pound of palpating pressure was 27 % and 100 %, respectively.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Suenaga et al. <sup>21</sup>	Aim: To evaluate the association between JE, contrast enhancement of JE, nitric oxide present in the joint, and TMD symptoms.	N = 77 ; 154 TMJs  84.6 % ♀, 15.4 % ♂.	Study group: Participants underwent  a. Clinical examination.	T2-weighted sagittal sequences (1.5 T, 5 mm thick cut).	Chi-square test	56 % of symptomatic TMJs and 9 % of non-symptomatic TMJs had JE.  JE was significantly associated with pain, and joint sounds.	Reliability of clinical examiners and radiologists was not described.  Clinical diagnosis was based on non-standardized examination.
Larheim et al. <sup>15</sup>	Aim: To evaluate the association between JE and MRI diagnosis of TMJ ID, bone marrow abnormalities, and pain.	N = 523	Study group: Participants underwent evaluation by a questionnaire.	Post-contrast T1-weighted oblique sagittal sequence (1.5 T, 5 mm thick cut).  JE defined: area of high signal intensity in the region of the superior space.	Spearman's rank correlation	Contrast enhancement of JE was significantly higher in the symptomatic groups than T2-weighted images.	High signal in the inferior TM joint space was not considered as indicative of JE.
				T2-weighted sagittal sequences (1.5 T, 3 mm thick cuts).	T-test	13.4 % TMJs were diagnosed as having JE.	Clinical evaluation of TMJs was carried out by a questionnaire.
				JE defined: area of high signal intensity greater than a moderate amount of fluid.	Fisher's exact test	Compared to pain-free TMJs, joints with pain had statistically more cases of JE, ID, and cortical bone abnormalities.	Reliability of questionnaire and radiologists was not described.
						Regression analysis indicated that TMJ-Pain side difference was positively dependent on JE and condyle marrow abnormalities.	Prevalence of pain in joints with JE was not provided.
						Regression analysis	

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
Takahashi et al. <sup>22</sup>	Aim: To investigate the association between JE, joint pain, and protein levels in joint lavage fluid of patients with ID and OA.	N = 26 ; 38 TMJs  84.2 % ♀, 15.8 % ♂  age: 43.2 yrs. (16 – 67 yrs.)	Study group: Participants underwent clinical examination.	T2-weighted sagittal sequences (1.5 T, 5 mm thick cut).	Fisher's exact test	80 % of TMJs with pain and 38.5 % of pain-free TMJs had JE ( $p < .05$ ).	Reliability of clinical examiners and radiologists was not described.
Adame et al. <sup>27</sup>	Aim: To correlate JE with clinical and MRI findings.	N = 142 ; 169 TMJs  88.9 % ♀, 11.1 % ♂  age: 25.4 yrs. (14-74 yrs.)	Study group: Participants underwent clinical examination.	T2-weighted sagittal sequences (0.5T, 5 mm thick slices).	Pearson's Chi-square test	65 % of participants with JE had TM joint pain.	Clinical examination was not adequately described.  Definition of the control group was not clear.
Murakami et al. <sup>17</sup>	Aim: To investigate the association between TMJ pain levels and JE.	N = 19 ; 19 joints  All participants were ♀ and were diagnosed as having disk displacement without reduction.	Study group: Participants underwent a. Clinical examination.	T2-weighted sagittal sequences (1.5 T).  JE defined: area of more than a line of high signal intensity in the region of superior or inferior joint space.	Paired t-test	52.6 % of participants had JE.  No statistically significant difference was observed between the groups for VAS pain score, pain questionnaire scores, and total pain score.	All participants were ♀.  Clinical examination was not adequately described.

Table Continued...

Author and Year	Study aim	Demographics	TMJ pain assessment methodology	Imaging assessment methodology	Statistics	Conclusion	Limitations
		age: 39.1 yrs. (14-61 yrs.)	b.VAS and Pain level questionnaire to measure intensity of pain.				Reliability of clinical examiners and radiologists was not described
Westesson et al. <sup>24</sup>	Aim: To investigate the association between JE and clinical symptoms of TMJ pain, and MRI diagnosis of ID and OA.	N = 390 ; 780 TMJs	Study group: Participants had TMD pain and dysfunction.	T2-weighted sagittal sequences (1.5 T).	No report on statistical analysis.	7 % of joints with normal disk position had JE.	No demographic details provided.
			Control group: Participants did not have any TMD pain or dysfunction.	JE defined: more than a line of high signal intensity in the region of superior or inferior joint space.	40 % of joints with DDwR had JE.		Clinical examination was not adequately described.
					50 % of joints with DDwoR had JE.		Details for blindness of the radiologist not provided.
					27 % of joints with OA had JE.		Reliability of clinical examiners and radiologists was not described.
						Statistically significant association was present between JE and joints with severe pain.	

## Discussion

A systematic review was conducted to determine if there is an association between joint effusion, self-reported TMJ pain and arthralgia of the TMJ. Based on the review of 32 articles published on this topic, the body of literature was unable to provide evidence to support or refute the association between joint effusion and self-reported pain TMJ pain during examination or between joint effusion and arthralgia, as defined by RDC-TMD. Systematic review of the studies indicated that the majority of investigations had reported a statistically significant association between joint effusion and TMJ self-reported pain during examination and joint effusion and arthralgia as defined by RDC-TMD. However, these results need to be interpreted with caution due to the presence of several crucial methodological limitations. Some of these studies failed to apply a standardized clinical examination.<sup>4-36</sup> This may result in failure to identify the true source of pain associated with TMD, or may result in using indeterminate classifications, such as the inclusion of participants with “tenderness”

to palpation instead of “pain”, or inclusion of participants with pain of muscular origin rather than of the TMJ. Furthermore, some investigations failed to report the reliability of the clinical examiners and radiologists.<sup>10-26</sup> Reliability is the overall consistency of a measure and is considered an important factor in identification of TMD and MRI-based findings since the prevalence of both conditions in the normal population range relatively low,<sup>31</sup> and overlooking reliability may influence the results in favour of a false positive association and inconsistent reporting. Another reason for disparity in reporting may be due to poor consensus among the authors for the operational definition of joint effusion. Some of the investigators defined effusion as presence of a high intensity signal on T2 images only in the superior joint space, while others considered presence of a high intensity signal in the superior or inferior joint space as indicative of joint effusion. Similarly, some authors quantified the joint effusion into multiple relative categories. However, none of these investigations reported the reliability and validity of radiologist to quantify effusion, which limits

the utility of this measure. Nonetheless, among the studies reporting quantification of effusion, the results were not consistent.

## Conclusion

In summary, based on the findings of this systematic review, the association between MRI diagnosis of joint effusion and self-reported TMJ-pain as well as between joint effusion and arthralgia, as defined by RDC-TMD, was not determined. This may be attributed to several methodological limitations, and heterogeneity in the operational definition of joint effusion. Investigations using validated and reliable clinical and radiologic criteria with appropriate case and control groups that have been adequately characterized are required. This will help determine, if any, association is present between MRI diagnosis of joint effusion and joint pain, and joint effusion and arthralgia.

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## Conflicts of interest

The authors declare no potential conflicts of interest with respect to the authorship and/or publication of this article.

## References

1. Greene CS. Diagnosis and treatment of temporomandibular disorders: emergence of a new care guidelines statement. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2010;110(2):137–139.
2. Dworkin SF, LeResche L. Research diagnostic criteria for temporomandibular disorders: review, criteria, examinations and specifications, critique. *J Craniomandib Disord.* 1992;6(4):301–355.
3. Ahmad M, Hollender L, Anderson Q, et al. Research diagnostic criteria for temporomandibular disorders (RDC/TMD): development of image analysis criteria and examiner reliability for image analysis. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2009;107(6):844–860.
4. Oliveira JX, Rosa JA, Dutra ME, et al. Assessing joint effusion and bone changes of the head of the mandible in MR images of symptomatic patients. *Braz Oral Res.* 2013;27(1):37–41.
5. Limchaichana N, Nilsson H, Ekberg EC, et al. Clinical diagnoses and MRI findings in patients with TMD pain. *J Oral Rehabil.* 2007;34(4):237–245.
6. Segami N, Nishimura M, Kaneyama K, et al. Does joint effusion on T2 magnetic resonance images reflect synovitis? Comparison of arthroscopic findings in internal derangements of the temporomandibular joint. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2001;92(3):341–345.
7. Shaefer JR, Jackson DL, Schiffman EL, et al. Pressure-pain thresholds and MRI effusions in TMJ arthralgia. *J Dent Res.* 2001;80(10):1935–1939.
8. Khawaja SN, Crow H, Mahmoud RF, et al. Is There an Association Between Temporomandibular Joint Effusion and Arthralgia? *J Oral Maxillofac Surg.* 2017;75(2):268–275.
9. Bas B, Yilmaz N, Gokce E, et al. Ultrasound assessment of increased capsular width in temporomandibular joint internal derangements: relationship with joint pain and magnetic resonance grading of joint effusion. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2011;112(1):112–117.
10. Costa AL, D'Abreu A, Cendes F. Temporomandibular joint internal derangement: association with headache, joint effusion, bruxism, and joint pain. *J Contemp Dent Pract.* 2008;9(6):9–16.
11. Guler N, Yatmaz PI, Ataoglu H, et al. Temporomandibular internal derangement: correlation of MRI findings with clinical symptoms of pain and joint sounds in patients with bruxing behaviour. *Dentomaxillofac Radiol.* 2003;32(5):304–310.
12. Haley DP, Schiffman EL, Lindgren BR, et al. The relationship between clinical and MRI findings in patients with unilateral temporomandibular joint pain. *J Am Dent Assoc.* 2001;132(4):476–481.
13. Kaneyama K, Segami N, Yoshimura H, et al. Increased levels of soluble cytokine receptors in the synovial fluid of temporomandibular joint disorders in relation to joint effusion on magnetic resonance images. *J Oral Maxillofac Surg.* 2010;68(5):1088–1093.
14. Lamot U, Strojan P, Surlan Popovic K. Magnetic resonance imaging of temporomandibular joint dysfunction-correlation with clinical symptoms, age, and gender. *Oral Surg Oral Med Oral Pathol Oral Radiol.* 2013;116(2):258–263.
15. Larheim TA, Westesson PL, Sano T. MR grading of temporomandibular joint fluid: association with disk displacement categories, condyle marrow abnormalities and pain. *Int J Oral Maxillofac Surg.* 2001;30(2):104–112.
16. Manfredini D, Tognini F, Zampa V, et al. Predictive value of clinical findings for temporomandibular joint effusion. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2003;96(5):521–526.
17. Murakami K, Nishida M, Bessho K, et al. MRI evidence of high signal intensity and temporomandibular arthralgia and relating pain. Does the high signal correlate to the pain? *Br J Oral Maxillofac Surg.* 1996;34(3):220–224.
18. Park JW, Song HH, Roh HS, et al. Correlation between clinical diagnosis based on RDC/TMD and MRI findings of TMJ internal derangement. *Int J Oral Maxillofac Surg.* 2012;41(1):103–108.
19. Rudisch A, Innerhofer K, Bertram S, et al. Magnetic resonance imaging findings of internal derangement and effusion in patients with unilateral temporomandibular joint pain. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2001;92(5):566–571.
20. Santos KC, Dutra ME, Warmling LV, et al. Correlation among the changes observed in temporomandibular joint internal derangements assessed by magnetic resonance in symptomatic patients. *J Oral Maxillofac Surg.* 2013;71(9):1504–1512.
21. Suenaga S, Abeyama K, Hamasaki A, et al. Temporomandibular disorders: relationship between joint pain and effusion and nitric oxide concentration in the joint fluid. *Dentomaxillofac Radiol.* 2001;30(4):214–218.
22. Takahashi T, Nagai H, Seki H, et al. Relationship between joint effusion, joint pain, and protein levels in joint lavage fluid of patients with internal derangement and osteoarthritis of the temporomandibular joint. *J Oral Maxillofac Surg.* 1999;57(10):1187–1193; discussion 1193–1184.
23. Tanaka T, Morimoto Y, Masumi S, et al. Utility of frequency-selective fat saturation T2-weighted MR images for the detection of joint effusion in the temporomandibular joint. *Dentomaxillofac Radiol.* 2002;31(5):305–312.
24. Westesson PL, Brooks SL. Temporomandibular joint: relationship between MR evidence of effusion and the presence of pain and disk displacement. *AJR Am J Roentgenol.* 1992;159(3):559–563.
25. Yamamoto M, Sano T, Okano T. Magnetic resonance evidence of joint fluid with temporomandibular joint disorders. *J Comput Assist Tomogr.* 2003;27(5):694–698.
26. Yano K, Sano T, Okano T. A longitudinal study of magnetic resonance (MR) evidence of temporomandibular joint (TMJ) fluid in patients with TMJ disorders. *Cranio.* 2004;22(1):64–71.
27. Adame CG, Monje F, Offnoz M, et al. Effusion in magnetic resonance imaging of the temporomandibular joint: a study of 123 joints. *J Oral Maxillofac Surg.* 1998;56(3):314–318.

28. Farina D, Bodin C, Gandolfi S, et al. TMJ disorders and pain: assessment by contrast-enhanced MRI. *Eur J Radiol.* 2009;70(1):25–30.
29. Guler N, Uckan S, Imirzalioglu P, et al. Temporomandibular joint internal derangement: relationship between joint pain and MR grading of effusion and total protein concentration in the joint fluid. *Dentomaxillofac Radiol.* 2005;34(3):175–181.
30. Segami N, Miyamaru M, Nishimura M, et al. Does joint effusion on T2 magnetic resonance images reflect synovitis? Part 2. Comparison of concentration levels of proinflammatory cytokines and total protein in synovial fluid of the temporomandibular joint with internal derangements and osteoarthritis. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2002;94(4):515–521.
31. Schiffman E, Ohrbach R, Truelove E, et al. Diagnostic Criteria for Temporomandibular Disorders (DC/TMD) for Clinical and Research Applications: Recommendations of the International RDC/TMD Consortium Network\* and Orofacial Pain Special Interest Groupdagger. *J Oral Facial Pain Headache.* 2014;28(1):6–27.
32. Nakaoka K, Hamada Y, Holmlund AB, et al. The changes of joint effusion on MRI and arthroscopic findings after visually guided TMJ irrigation correlated to the clinical outcome. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2009;108(1):99–104.
33. Emshoff R, Brandlmaier I, Gerhard S, et al. Magnetic resonance imaging predictors of temporomandibular joint pain. *J Am Dent Assoc.* 2003;134(6):705–714.
34. Emshoff R, Brandlmaier I, Bertram S, et al. Risk factors for temporomandibular joint pain in patients with disc displacement without reduction - a magnetic resonance imaging study. *J Oral Rehabil.* 2003;30(5):537–543.
35. Emshoff R, Brandlmaier I, Bertram S, et al. Relative odds of temporomandibular joint pain as a function of magnetic resonance imaging findings of internal derangement, osteoarthritis, effusion, and bone marrow edema. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod.* 2003;95(4):437–445.
36. Emshoff R, Brandlmaier I, Bertram S, et al. Magnetic resonance imaging findings of osteoarthritis and effusion in patients with unilateral temporomandibular joint pain. *Int J Oral Maxillofac Surg.* 2002;31(6):598–602.