The prevalence of radiographic and MRI-defined patellofemoral osteoarthritis and structural pathology: a systematic review and meta-analysis

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ABSTRACT

Background Patellofemoral osteoarthritis (PF OA) is more prevalent than previously thought and contributes to patient's suffering from knee OA. Synthesis of prevalence data can provide estimates of the burden of PF OA.

Objective This study aims to conduct a systematic review and meta-analysis on the prevalence of PF OA and structural damage based on radiography and MRI studies in different populations.

Methods We searched six electronic databases and reference lists of relevant cross-sectional and observational studies reporting the prevalence of PF OA. Two independent reviewers appraised methodological quality. Where possible, data were pooled using the following categories: radiography and MRI studies.

Results Eighty-five studies that reported the prevalence of patellofemoral OA and structural damage were included in this systematic review. Meta-analysis revealed a high prevalence of radiographic PF OA in knee pain or symptomatic knee OA (43%), radiographic knee OA or at risk of developing OA (48%) and radiographic and symptomatic knee OA (57%) cohorts. The MRI-defined structural PF damage in knee pain or symptomatic population was 32% and 52% based on bone marrow lesion and cartilage defect, respectively.

Conclusion One half of people with knee pain or radiographic OA have patellofemoral involvement. Prevalence of MRI findings was high in symptomatic and asymptomatic population. These pooled data and the variability found can provide evidence for future research addressing risk factors and treatments for PF OA.

Trial registration number PROSPERO systematic review protocol (CRD42016035649).

INTRODUCTION

Knee osteoarthritis (OA) is a leading cause of pain and disability worldwide.¹ The patellofemoral joint (PF) is commonly affected in symptomatic knee OA² and is a substantial source of symptoms associated with knee OA.³ Further to this, the PF is often affected by OA before the tibiofemoral (TF) joint and increases the risk of TF OA development and progression.^{4,5}

With a recent increase in radiography and magnetic resonance imaging (MRI) based studies focused on PF joint, the evidence on the prevalence of PF OA is expanding rapidly. A 2013 narrative literature review concluded that the prevalence of radiographic PF OA in individuals' post-ACL and/or meniscus ruptures was approximately 50%.⁶ A

recent systematic review described the prevalence of radiographic PF OA in population-based and in cohorts of people with knee pain.⁷ A large number of studies have reported PF OA in different populations (eg, post-traumatic and healthy individuals), and knowledge of population-specific prevalence is relevant for clinicians and researchers. An updated review with inclusion of different study samples (eg, post-traumatic, occupation-based, high risk of OA and healthy individuals) builds considerably on the previous systematic review⁷ and extends our current knowledge of PF OA.

MRI is the modality of choice to assess structural damage in epidemiological studies to detect early and subtle features of OA (eg, abnormal cartilage morphology and bone marrow lesions) not seen on radiography.⁸ Thus, the prevalence of PF structural damage using MRI may be higher than the prevalence determined by radiography. Including radiography and MRI-based studies in community and specific study, populations provide a comprehensive evaluation of the prevalence of PF OA and PF structural damage and extends prior reviews in this area. Thus, the objective of this study was to perform a systematic review and meta-analysis with the aim to determine the prevalence of PF OA using radiographs and MRI-defined structural PF damage in a variety of study populations.

METHODS

The study protocol was developed in consultation with guidelines provided by the Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA) statement. The protocol was prospectively registered on the PROSPERO International prospective register for systematic reviews website (<http://www.crd.york.ac.uk/PROSPERO/>) (Registration no: CRD42016035649). The reporting of this study followed the PRISMA checklist.

Literature search strategy

Using guidelines provided by the Cochrane Collaboration, a comprehensive search strategy was devised from the following electronic databases with no date restrictions: (1) MEDLINE via OVID, (2) EMBASE via OVID, (3) CINAHL via EBSCO, (4) Scopus, (5) Web of Science and (6) SPORTDiscus. The primary search strategy included search for original publications. The search strategy was deliberately simplified to ensure inclusion of all relevant papers, with all terms searched as free text and key words (where applicable): Concept 1, Patellofemoral

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(Patello-femoral, PF, PFJ, knee joint); Concept 2, Osteoarthritis (OA, arthritis, degenerative arthritis, bone marrow lesion); and Concept 3, Prevalence (prevalence, morbidity, epidemiology, diagnosis, incidence). All search terms were exploded and scope notes from each database were examined for other possible terms for modification of search strategies. The MEDLINE search strategy was adapted for other databases (online supplementary file table 1). The search strategy was limited to English language and full text. All potential references were imported into Endnote X7 (Thomson Reuters, Carlsbad, California, USA) and duplicates were removed. Two reviewers (HFH, NW) reviewed all titles returned by the database searches and retrieved suitable abstracts. Where abstracts suggested that papers were potentially suitable, the full-text versions were screened and included in the review if they fulfilled the selection criteria. Reference lists of all publications considered for inclusion were hand searched recursively and citation tracking was completed using Google Scholar until no additional eligible publications were identified. A third reviewer was consulted in case of disagreements (JJS).

Selection criteria

Cross-sectional and longitudinal studies reporting the prevalence or frequency of PF OA or PF structural damage were included. No restrictions were placed on age, sex or method of recruitment. Reviews, case reports and unpublished studies, as well as non-human studies were excluded.

Assessment of methodological quality and risk of bias

Two independent reviewers (NW and ZM), who remained blind to authors, affiliations and the publishing journal, rated the methodological quality of included studies using the Critical Appraisal tool.⁹ The Critical Appraisal tool was developed to appraise prevalence and incidence-based studies and consists of eight items (maximum score possible 8). Final study ratings for each reviewer were collated and examined for discrepancies. Any inter-rater disagreement was discussed in a consensus meeting, and unresolved items were taken to a third reviewer (HFH) for consensus. Total scores were normalised to a scale ranging from 0 to 2, for each study to assign level of methodological quality. Studies were then classified as high quality (≥ 1.4), moderate quality (1.1–1.4) or poor quality (< 1.1) based on normalised scores.¹⁰

Data management and statistical analysis

For the purposes of this systematic review, we defined prevalence as the prevalence of PF OA in community-based studies and the reported frequencies of PF OA in other populations. Data pertaining to population, sample size, sex, age, type of imaging (MRI, radiography), grading criteria, units of analysis (number of participants affected or number of knees affected) and prevalence of radiographic PF OA and MRI-defined PF structural damage (isolated PF OA/PF structural damage; combined PF OA and TF OA/PF and TF structural damage; and unclear, not clearly described whether the prevalence was isolated or combined) were independently extracted and entered into an Excel spreadsheet. If sufficient data were not reported in the published article or supplementary material provided, the corresponding author was contacted to request further data. If multiple studies presented data from one cohort, the study with the most complete data was included. PF OA and MRI-defined PF structural damage prevalence data were reported for: (1) isolated, (2) combined (PF and TF) and (3) any (isolated, combined and unclear). Meta-analysis for proportions with random effects model were performed

using MedCalc for Windows, V.16.8. Heterogeneity tests were also conducted and interpreted as follows: $I^2 \leq 25\%$, low heterogeneity; $I^2 = 25$ to $\leq 50\%$, moderate heterogeneity; and $I^2 \geq 75\%$, high heterogeneity.¹¹ Data were divided into two categories based on imaging technique used: (1) radiography and (2) MRI.

Radiography studies

The Kellgren and Lawrence (KL) grading criteria¹² and Osteoarthritis Research Society International (OARSI) atlas¹³ are used to define radiographic OA in the TF compartments. There is no KL or OARSI atlas definition of PF OA based on radiographs; however, both criteria are often used to quantify the severity of radiographic OA in the PF using the skyline and/or lateral radiography views. For the purposes of this systematic review and meta-analysis, osteophytes and joint space narrowing were used to define PF OA. If prevalence for multiple radiographic OA features (eg, prevalence based on osteophytes and joint space narrowing) was reported, then prevalence based on osteophytes was chosen. Data were pooled based on the following study populations: (1) community-based (individuals randomly recruited from community), (2) knee pain/symptomatic (individuals recruited based on knee-related symptoms), (3) radiographic and symptomatic OA (individuals recruited based on symptoms and radiographic OA), (4) healthy individuals (no pain, injury or OA), (5) radiographic or high risk of OA (individuals recruited based on radiographic OA or risk of developing radiographic OA without regard to knee pain/symptoms), (6) occupational-based (individuals recruited based on their occupation/sports) and (7) post-traumatic (individuals with previous knee-related trauma, such as ACL injury or reconstruction or meniscal injury). Given that individuals recruited based on high risk of OA may or may not have had previous trauma; data from individuals with high risk of OA were not included in the post-traumatic category. The occupation-based category included different sporting and occupational activities such as long distance runners, shooters, graphic designers and monks. To determine the prevalence in individuals exposed to different activities, the data from sports and occupational activities were pooled together. Data were stratified based on intensity of activity (eg, high: soccer graphic and low: graphic designers) activities. For longitudinal studies, data from the latest time point (rather than baseline) were included. Within the eight study population categories, sensitivity analyses were conducted when > 1 study reported sufficient data for pooling based on disease severity, compartment-specific OA pattern, age and sex. Disease severity was defined as mild, presence of at least mild radiographic PF OA; and definite, presence of definite radiographic PF OA (online supplementary file table 2). Compartment-specific OA pattern was defined as: (1) isolated PF OA, (2) combined PF OA and TF OA and (3) any PF OA. Age groups for sensitivity analyses were categorised as: (1) mean age: < 50 years, (2) mean age: ≥ 50 years. These sensitivity analyses are presented in text for any PF OA and in supplementary material (online supplementary file table 3 for the isolated and combined TF OA and PF OA groups. Where possible, medial and lateral PF OA prevalence was described.

MRI studies

Currently, there is no accepted definition of MRI-defined PF OA. A definition was proposed by Hunter *et al*¹⁴ which included a definite osteophyte and partial or full thickness cartilage loss. However, this proposed definition of MRI-defined PF OA has not been further validated. Furthermore, most previous studies do not provide data on osteophytes to enable calculation of PF

OA prevalence using this definition. Therefore, for the purposes of this systematic review we will report MRI-defined structural damage. Data were pooled based on study populations described above (except for occupational-based population) as well as general population (studies that could not be categorised into one of the categories described above). Within each study population category, data were pooled based on cartilage defect and bone marrow lesions (BML) MRI features. Authors used the following terms to define cartilage defect: cartilage abnormalities, cartilage defect, full cartilage thickness loss, cartilage pathology and cartilage lesion; and the following terms were used to define BML, marrow abnormalities, marrow lesion and bone marrow oedema. To allow data pooling where possible other scoring systems were compared with the Whole-Organ MRI Score (WORMS)¹⁵ and MRI Osteoarthritis Knee Score (MOAKS)¹⁶ based on the explanation of the scoring system provided in the paper. Data were stratified based on compartment-specific OA pattern (isolated PF OA, combined PF OA and TF OA and any PF OA). Where possible, stratified analyses were conducted based on age (mean age: <50 years, ≥50 years) and sex. If possible, medial and lateral PF OA prevalence was described. Most longitudinal MRI studies provided most complete data at baseline rather than at later time points (dropouts or only ORs data for later time points); thus, this review included baseline data.

RESULTS

Search strategy, methodological quality and risk of bias

The comprehensive search strategy identified 2681 titles, with the last search conducted on 25 February 2016. Following removal of duplicate publications and conference proceedings, titles of 1105 publications were evaluated. Thirteen titles were obtained from other resources (Google Scholar and hand searching). The full texts of 144 articles were retrieved, with 117 articles meeting the selection criteria. Following removal of studies with duplicate data, 85 studies (63 radiography studies,^{2 17–78} 24 MRI studies^{39 79–101}) were included in this systematic review (tables 1 and 2, figure 1). There was one study that reported data on radiographic PF OA and MRI-defined PF structural damage.³⁵ The methodological quality scores ranged from 0 to 2 (out of 2) (online supplementary file table 4). There were 15 studies of high quality, 16 were moderate and 54 were low quality. Most studies scored negatively on items 1 (ie, study design/sampling method) and 6 (ie, response rate) and positively on items 4 (ie, measurement criteria) and 8 (ie, study subjects described) of the critical appraisal tool. A high level of heterogeneity was noted within radiography and MRI studies (I^2 range 96%–100%). The level of heterogeneity remained high (I^2 range 70%–100%) when studies were further subgrouped based on population, OA severity pattern, age and sex. Exclusion of low methodological quality studies did not decrease the heterogeneity levels.

Prevalence of patellofemoral OA based on radiography

Community-based population

In community-based populations, the overall prevalence of isolated PF OA from four studies^{19 24 26 28} was (mean proportion: (95% CI)) 7% (5 to 10), combined PF OA and TF OA from four studies^{19 24 26 28} was 17% (10 to 26), and any PF OA based on nine studies^{18 19 24 26 28 39 46 54 75} was 38% (28 to 50) (figure 2A–C). In the any PF OA group, the prevalence of mild OA severity was 33% (17 to 51) from three studies^{28 46 75} and definite OA severity was 40% (28 to 53) from six studies.^{18 19 24 26 39 54} The prevalence of any PF OA in community-based population was 32% (24 to 42) in those aged 50 years or over from

eight studies.^{19 24 26 28 39 46 54 75} Only one study described prevalence of isolated compartment-specific PF OA,²⁶ with prevalence of medial PF OA at 0.3% in women and 0.7% in men, and the prevalence of lateral PF OA at 1.6% in women and 3.7% in men. Sensitivity analyses based on sex revealed that the prevalence of any PF OA in women was 41% (31 to 51) from six studies^{18 19 24 26 28 75} and 47% (23 to 71) in men from four studies.^{18 19 24 26}

Knee pain or symptomatic population

Overall prevalence of isolated PF OA was 19% (11 to 29) from eight studies,^{2 21 32 42 52 53 59 69} combined PF OA and TF OA was 34% (25 to 43) from seven studies^{2 32 42 52 53 59 69} and any PF OA was 43% (32 to 55) from 12 studies^{2 21 22 32 42 52 53 59 60 64 69 77} (figure 2D–F). For any PF OA, the prevalence of mild and definite OA severity was 37% (24 to 51) from seven studies^{2 21 22 32 52 60 69} and 49% (30 to 67) from six studies,^{2 42 53 59 64 77} respectively. Age-based prevalence of any PF OA in individuals under 50 years was 54% (16 to 90) from two studies^{69 77} and in those 50 years or over was 43% (31 to 56) from eight studies.^{2 21 22 42 53 59 60 64} Sex-based prevalence of any PF OA was 46% (23 to 70) in women^{53 60} and 58% (27 to 86) in men.^{53 60 69}

Radiographic and symptomatic knee osteoarthritis

Overall prevalence of isolated PF OA was 20% (11 to 32) from four studies,^{25 30 33 55} combined PF OA and TF OA was 43% (8 to 83) from two studies^{25 55} and any PF OA was 57% (43 to 70) from 13 studies^{25 27 30 33 35 37 41 45 50 55 61 74 78} (figure 2G–I). In the any PF OA group, the prevalence of mild severity was 56% (41 to 70) from 12 studies.^{25 30 33 35 37 41 45 50 55 61 74 78} The prevalence in individuals 50 years or over was 58% (42 to 72) from 12 studies^{25 27 30 33 35 37 41 45 55 61 74 78} and the prevalence of any PF OA in women was 36% (33 to 38)^{27 45} and men was 35% (16 to 58) from two studies.^{27 45}

Healthy individuals

Data from four studies were included in meta-analyses to determine the prevalence of PF OA in healthy individuals.^{34 60 63 72} Overall prevalence of any PF OA in healthy individuals (no pain, injury or OA) was 17% (6 to 33) (figure 3A). Sensitivity analyses based on sex could only be performed in women revealing the prevalence of PF OA in healthy women at 15% (1 to 43) from two studies.^{60 72}

Radiographic knee OA or at risk of developing OA

Overall prevalence of any PF OA in individuals with radiographic OA or at risk of OA was 48% (35 to 61) from four studies^{36 38 43 48} (figure 3B), with prevalence based on mild and definite OA severity as follows: 54% (17 to 89) from two studies^{36 48} and 45% (30 to 60) from two studies,^{38 43} respectively. In this group, the prevalence of any PF OA in women was 41% (8 to 80) from two studies.^{38 43}

Occupation-based population

Four studies reported occupation-based prevalence of PF OA.^{51 68 72 76} Overall prevalence of any PF OA in individuals in occupations or sports such as long distance running, soccer, shooting, floor layers, graphic designers and monks was 21% (9 to 37) (figure 3C). For any PF OA, the prevalence based on mild OA severity was 29% (10 to 52) from three studies.^{68 72 76} The prevalence of any PF OA in individuals 50 years and over was 18% (9 to 28) from three studies.^{51 68 72} Sensitivity analyses based on sex revealed the prevalence of any PF OA in men was 14% (9 to 20) from two studies.^{51 68} Analysis could not be

Table 1 Details of included radiography studies													
Author, year	Additional information	Sex	Age	N (knees)	Diagnostic criteria	Isolated PF OA (%)		Lateral	Combined (%)		Unclear PF OA (%)	Severity	
						Overall	Medial		Overall				
Community-based population													
Al-Arfaj and Al-Boukai ¹⁸ 2002	First two patients visiting a clinical invited everyday	F	40±15	133	KL				81			Definite	
Baker et al. ¹⁹ 2004	Randomly from community over the age of 60 years	M	40±15	167	KL				88			Definite	
		F	68±6	1475	KL	8			26			Definite	
		M	69±6	997	KL	9			12			Definite	
		F	62±11	283	KL					53		Definite	
Braga ²⁴ 2009	African-Americans (>45 years) recruited by probability sampling	M	60±11	147	KL				60			Definite	
		F	62±11	799	KL					42		Definite	
		M	61±10	728	KL					54		Definite	
		F	72±5	383		3	0	2	28	29	2	15	Definite
Cho ²⁶ 2016	Random sample of individuals (>65 years) selected from a longitudinal study	M	71±5	298		5	1	4	8	13	1	7	Definite
		F	59±7	325	Burnett, 1994 (JSN and osteophyte)	13			15				Mild
Cicuttini ²⁸ 1997	Unrelated women selected from a group participating in twin study of OA	F											
Gross ³⁹ 2012	Offspring (and their spouses) of participants from original Framingham Heart Study and population-based sample of Framingham community	FM	63±9	985 (1159)	OARSI				8			Definite	
Hunter Zhang ⁴⁶ 2007	Random sample from Health ABC study	FM	74±3	595	OARSI				49			Mild	
Lanyon ⁵⁴ 1998	Random sampling, stratified by age from the lists of two general practitioner	FM	62 (40–80)	452	Altman				13			Definite	
Szoeke ⁷⁵ 2006	Postmenopausal women selected from another population-based study	F	60±3	224	OARSI				22			Mild	
Tangtrakulwanich ⁷⁶ 2012	Individuals (>40 years) without rheumatic diseases using primary care	FM	>40	576	0–3 scale				38			Mild	
Healthy individuals													
Englund ³⁴ 2005		FM	56±21	68	OARSI				9			Mild	
McAlindon ⁶⁰ 1992		M	>55	78					5			Mild	
		F	>55	162					5			Mild	
Naredo ⁶⁵ 2005		FM	68±8	10	KL				70			Mild	
Spector ⁷² 1996		F	54±6	215	KL				28			Mild	
Knee pain or symptomatic population													
Barret ²¹ 1990		FM	78 (51–93)	1894 (2197)	Alhback				18			Mild	
Bennett ²⁷ 2007		FM	54±13	39	ACR				62			Mild	
Davies ³² 2002		FM	>40	174 (206)	Ahlback	9			13			Mild	
Duncan ² 2006		FM	66 (50–93)	777	KL	24			40			Definite	
Hinman ⁴⁷ 2014		FM	54±10	224		25	3	11	44 ⁴⁷			Definite	
Kumm ⁵² 2012	6-year follow-up	FM	51±6	128	Line drawing atlas	19			38			Mild	
Lacey ⁵³ 2008		F	50–64	200	KL	19			27			Definite	
		M	50–64	158		35			25			Definite	
		F	>65	207		21			47			Definite	
		M	>65	180		24			62			Definite	
McAlindon ⁶⁰ 1992		M	>55	86	KL				12			Mild	
		F	>55	187					26			Mild	
McAlindon ⁵⁹ 1996		FM	81±5	608	Osteophyte/JSN	5			20			Definite	
Neame ⁶⁴ 2004	Right knee	FM	64±9	1729 (1718)					14			Definite	
Continued													

Continued

Table 1 Continued

Author, year	Additional information	Sex	Age	N (knees)	Diagnostic criteria	Isolated PF OA (%)	Combined (%)	Unclear PF OA (%)	Severity
						Overall	Medial	Lateral	
Left knee									
Sadat Ali ⁵⁹ 1996		FM		(1723)				14	Definite
		M	41±7	103 (126)	JSN, osteophytes and varus deformity	45		30	Mild
Thorstenson ⁷⁷ 2009			45 (35–54)	125	JS width <5mm			33	Mild
Radiographic knee OA or high risk of knee OA									
Et ³⁶ 1998	Knee OA	FM	56±11	240 (369)	Altman			34	Unclear
Glass ³⁸ 2014	Knee OA or High risk	F	62±8	1618 (3236)	KL			22	Definite
		M	62±8	1094 (2188)	KL			18	Definite
Huang ⁴³ 2000	Hand, hip or knee OA	F	64 (29–87)	270	KL			62	Definite
Jones ⁴⁸ 1993	Knee OA	FM	62 (18–91)	30 (60)	Osteophyte			75	Mild
Radiographic and symptomatic OA									
Ciuttini ²⁷ 2002		M	63±10	44	Osteophyte and JSN			48	Definite
		F	63±10	66	Osteophyte and JSN			38	Definite
Cooper ³⁰ 1994		FM	>55	109	Radiograph atlas of knee OA	31			Mild
Elah ³³ 2000		FM	66±11	292	OARSI	11			Mild
Ersöz ³⁵ 2003		FM	62±9	20 (40)	KL			88	Mild
Farrokh ³⁷ 2013		FM	65±9	167	KL			80	Mild
Hinman ⁴¹ 2002		FM	68±7	41	ACR			81	Mild
Hunter Niu ⁴⁵ 2005		F	68±6	1500	KL			36	Mild
		M	74±3	595	KL			26	Mild
Kerna ⁵⁰ 2013		FM	45±6	438	Line drawing atlas			48	Mild
Ledingham ⁵⁵ 1993		FM	65 (34–91)	252	Modified Thomas 1995	24		64	Definite
Messier ⁶¹ 2005		FM	74±1	10	KL			90	Mild
Szebenyi ⁷⁴ 2006			66±10	167 (334)	OARSI			65	Mild
Van der Esch ⁷⁸ 2014		FM	62±8	298	Osteophyte			86	Mild
Post-traumatic population									
Ahn ¹⁷ 2012	Postoperative ACLR	FM	29±9	117	IKDC			60	Mild
Barenius ²⁰ 2014	ACLR with patellar graft	FM	39±6	69	KL			32	Definite
	Uninjured contralateral	FM			KL			12	Definite
	Semitendinosus ACLR graft	FM	42±7	65	KL			36	Definite
Barenius ²⁰ 2014	Uninjured contralateral	FM	42±7	65	KL			12	Definite
Bourke ²³ 2012	Isolated ACL injury with ACLR	FM	27 (15–46)	118	IKDC			12	Mild
Cohen ²⁹ 2007	ACLR	FM	45±10	62	Fairbank			74	Mild
Culvenor ³¹ 2014	ACLR	FM	54±11	250	OARSI	20		48	Mild
Englund ³⁴ 2005	Medial meniscectomy	FM	54±11	67	OARSI			19	Mild
	Lateral meniscectomy	FM	42 (22–62)	67	IKDC			27	Mild
Hertel ⁴⁰ 2005	ACLR—patellar	FM	57±12	89	IKDC			19	Mild
Hulet ⁴⁴ 2015	Arthroscopic partial lateral meniscectomy	FM	31 (15–61)	100	IKDC			33	Mild
Jarvela ⁴⁷ 2001	ACLR with patellar graft	FM	27 (18–38)	56	KL			47	Mild
Keays ⁴⁹ 2007	ACLR with patellar/semiotendinosus and gracilis graft	FM	26±10	249	KL			25	Mild
Li ⁵⁶ 2011	ACLR	FM	26±10	249	KL			11	Mild
Li ⁵⁶ 2011	Uninjured contralateral	FM	26±10	249	KL			3	Mild

Continued

Table 1 Continued

Author, year	Additional information	Sex	Age	N (knees)	Diagnostic criteria	Isolated PF OA (%)	Combined (%)	PF + TF OA	Undeclared PF OA (%)	Medial	Lateral	Severity
Liden ⁵⁷ 2008	ACLR with patellar graft	FM	30 (17–52)	72	Ahlaback and Fairbank				14			Mild
Lohmander ⁵⁸ 2004	ACLR with hamstring graft	FM	29 (15–59)	41					5			Mild
	ACL injury with surgery	F	31 (26–40)	41	OARSI				20			Mild
	ACL injury without surgery	F	31 (26–40)	26					4			Mild
	ACLR with patellar graft	FM		83	IKDC				76			Mild
Murray ⁶² 2012	Uninjured contralateral	FM		42					59			Mild
Neuman ⁶⁵ 2009	ACL injury with meniscal injury	FM	43±8	22	OARSI				18			Mild
	ACL injury without meniscal injury	FM	43±8	38	OARSI				3			Mild
Neuman ⁶⁵ 2009	ACLR with meniscal injury	FM	43±8	11	OARSI				55			Mild
	ACLR without meniscal injury	FM	43±8	4	OARSI				25			Mild
Overall (ACL with or with meniscal or reconstruction)		FM	43±8	75	OARSI			9				Mild
Oiestad ⁶⁶ 2013	ACLR	FM	71±8	181					26			Mild
Rofth ⁶⁷ 1985	ACLR (non-augmented)	FM		38	Osteophyte size				13		40	Mild
	ACLR (augmented)	FM		43	Osteophyte size				4		14	Mild
Sajovic ⁷⁰ 2006	ACLR with semitendinosus and gracilis	FM	24 (14–42)	28	IKDC				7			Mild
	ACLR patellar tendon	FM	27 (16–46)	26	IKDC				12			Mild
Salmon ⁷¹ 2006	ACLR—13-year follow-up	FM	27 (25–28)	43	IKDC				26			Mild
Sward ⁷³ 2013	ACL injured (with or without ACLR) with varus alignment	FM	42±7	36	OARSI			22				Mild
	ACL injured (with or without ACLR) with valgus/neutral	FM	39±6	29	OARSI			7				Mild
Occupation-based population												
Kujala ⁵¹ 1995	Long distance runners	M	60±5	28	KL				11			Definite
	Soccer players	M	57±5	31	KL				16			Definite
	Weight lifters	M	60±5	29	KL				28			Definite
	Shooters	M	61±5	29	KL				3			Definite
	Floor layers	M	53 (39–68)	134	Modified Ahlback Scale				9			Mild
Rytter ⁶⁸ 2009	Graphic designers	M	58 (40–70)	120	Modified Ahlback Scale				18			Mild
Spector ⁷² 1996	Ex-athletes	F	52±6	81	KL				42			Mild
Tangtrakulwanich ⁷⁶ 2006	Monk		44±18	261	KL			33				Mild

Where possible, age is presented as mean±SD or mean (range).

Mild severity indicates at least mild OA severity and definite indicates definite OA severity (equivalent to KL≥2).

ACLR, ACL reconstruction; AC/R, American College Rheumatology; F, female; FM, both female and male; IKDC, International Knee Documentation Committee; JSN, joint space narrowing; M, male; OA, osteoarthritis; OARSI, Osteoarthritis Research Society International; PF OA, patellofemoral osteoarthritis.

Table 2 Details of included MRI studies

Author, year	Additional information	Sex	Age	N (knees)	Diagnostic criteria	Isolated PF OA (%)			Combined (%)			Unclear PF OA (%)			Feature	Case definition
						Overall	Medial	Lateral	PF + TF OA	Overall	Medial	Lateral				
Community-based population																
Ding ⁸⁵ 2005	<45 years	FM	<45	167	0–4 scale					16				Cartilage defect	≥2	
	>45 years	FM	≥45	205						38				Cartilage defect	≥2	
Gross ³⁹ 2012*	FOA cohort	FM	64±9	985 (1159)	WORMS		50	18						Cartilage damage	≥2	
Stefanik ⁹⁵ 2013	FOA cohort	FM	64±8	970	WORMS	20			44					Cartilage damage	≥2	
Wang ⁹⁹ 2015	TASOAC cohort	FM	62±7	904	WORMS					19				BML	≥1	
										40				Cartilage defect	≥2	
Healthy individuals																
Sharma ⁹² 2014	OAI cohort	M	60±9	375	MOAKS	25			33					Cartilage damage	>0	
					WORMS	33			18					BML	>0	
		F	60±9	474	MOAKS	34			30					Cartilage damage	>0	
					WORMS	35			17					BML	>0	
Wang ¹⁰⁰ 2012		FM	42±7	38	0–4 scale					3				Cartilage defect	≥2	
Knee pain or symptomatic population																
Amin Baker ⁷⁹ 2009*	BOKS cohort	M	68±9	154	WORMS						81	68		Maximal cartilage morphology	≥2	
		F	64±9	111	WORMS						91	84		Maximal cartilage morphology	≥2	
Amin Goggins ⁸⁰ 2008*	Heavy lifting (BOKS cohort)	M	69±9	40	WORMS					60				Cartilage morphology	≥2	
	Occupational exposures	M	64±9	47	WORMS					72				Cartilage morphology	≥2	
	No heavy lifting/occupational exposures	M	70±9	98	WORMS					47				Cartilage morphology	≥2	
Amin Guermazi ⁸¹ 2008	BOKS cohort	FM	67±9	265	WORMS					91				Cartilage morphology	≥2	
Cai ⁸² 2015	Anhui osteoarthritis cohort	FM	56±12	174	0–4 scale					82				Cartilage damage	≥2	
		FM			0–3 scale					52				BML	≥1	

Continued

Table 2 Continued

Author, year	Additional information	Sex	Age	N (knees)	Diagnostic criteria	Isolated PF OA (%)	Combined (%)	Unclear PF OA (%)	Feature	Case definition
Crema ⁸⁴ 2014		FM	58±10	163	0–3 scale			13	BML	≥1
Peterfy ⁹⁰ 2004		FM	61±8	19	WORMS			94	Cartilage abnormality	≥1
								16	BML	≥1
								81	Osteophytes	≥1
Tsavlas ⁸⁷ 2012	Various knee-related clinical conditions	FM	≤50	315	ICRS			5	Cartilage lesion	≥2
		FM	>50	200	ICRS			33	Cartilage lesion	≥2
Radiographic knee OA or high risk of knee OA										
Gross 2012 ³⁹	MOST cohort	FM	62±8	1381 (1621)	WORMS	69	36		Cartilage damage	≥2
Stefanik Gross 2015 ^{95*} (O&C)	MOST cohort	FM	69±8	1137	WORMS	22	13	63	BML	≥1
Stefanik Gross ⁹⁴ 2015 [†] (ACR)	MOST cohort	F	66±8	653 (2594)	WORMS			51	Cartilage damage	≥2
								29	BML	≥1
		M	66±8	400 (1486)	WORMS			43	Cartilage damage	≥2
								23	BML	≥1
Runhaar ⁹¹ 2014		F	56±3	348 (529)	MOAKS	44	35	18	BML	≥1
				348 (467)		47	42	18	Cartilage defect	≥1
				348 (408)		33	27	25	Osteophytes	≥1
Radiographic and symptomatic OA										
Chan ²⁵ 1991		FM	58	20	0–3 scale	0		75	Cartilage loss	≥1
						0		75	Osteophytes	≥1
Post-traumatic population										
Culvenor ⁸⁵ 2015	ACLR with or without meniscus repair	FM	30±8	111	MOAKS	12		2	BML	≥1
Van Meer ⁹⁸ 2016	ACL rupture	FM	25±33	143	MOAKS			29	Osteophytes	≥1
Wang ¹⁰⁰ 2012	Arthroscopic partial medial meniscectomy (2 years post)	FM	43±5	63	0–4 scale	19			Cartilage defect	≥2

Continued

Table 2 Continued

Author, year	Additional information	Sex	Age	N (knees)	Diagnostic criteria	Isolated PF OA (%)	Combined (%)	Unclear PF OA (%)	Feature	Case definition
General population										
Gross ⁸⁷ 2011	Framingham Heart study offsprings + spouses and people from Framingham town	FM	64±9	1094	WORMS			58	Cartilage damage	≥2
				1096				42	Cartilage damage	≥2
Hayes ⁸⁸ 2005	With or without pain or OA	F	46±1	117 (232)	Modified Noyes arthroscopy system		66		Cartilage defect	≥IIA
Kornaat ⁸⁹ 2005	40–70 years with familial generalised OA	FM	60 (43–77)	205	KOSS	66			Cartilage defect	≥1
Sowers ⁹³ 2011	Middle-aged women	F	56±5	360 (724)				89	Cartilage defect	
								45	BML	
Sowers ⁹³ 2011	Middle-aged women	F	56±5	360 (724)						
Teng ⁹⁶ 2015	Without no OA or isolated OA	FM	51±10	61	WORMS			56	Osteophytes	
								46	Cartilage lesion	>1

Where possible, age is presented as mean±SD or mean (range).

*Studies included in subgroup analyses.

†Percentage based on regions.

ACL/R, ACL reconstruction; BOKS, Boston Knee Osteoarthritis Study; BML, bone marrow lesion; F, female; FM, both female and male; FOA, Framingham Osteoarthritis; ICBS, International Cartilage Repair Society; KOSS, Knee Osteoarthritis Scoring System; M, male; MOAKS, MRI Osteoarthritis Knee Score; MOST, Multicenter Knee Osteoarthritis; OA, osteoarthritis; OAI, Osteoarthritis Initiative; TASOAC, Tasmanian Older Adult Cohort. WORMS, Whole-organ MRI Score.

Figure 1 Flow chart of the study selection process. PFOA, patellofemoral osteoarthritis.

performed in women. For any PF OA, the prevalence of any PF OA in high-intensity activity population was 19% (11 to 29) from one study⁵¹ and 19% (3 to 45) in low-intensity activity population based on three studies.^{51 68 76}

Post-traumatic population

The overall prevalence of isolated PF OA from two studies was 17% (5 to 34) from two studies^{49 65} (figure 3D). In the injured knee, the overall prevalence of any PF OA in post-traumatic

population (range: 5 to 22 years) was 27% (19 to 34) from 19 studies^{17 20 23 29 31 34 40 44 47 49 56–58 62 65 66 70 71 73} (figure 3E). For any PF OA, the prevalence of mild OA severity was 26% (18 to 34) from 18 studies.^{17 23 29 31 34 40 44 47 49 56–58 62 65 66 70 71 73} Sensitivity analyses based on age revealed the prevalence of any PF OA was 27% (18 to 36) in individuals under 50 years^{17 20 23 29 31 40 47 49 56–58 65 66 70 71 73} and 26% (17 to 35) in those 50 years or over.^{34 44} In the uninjured knee, overall prevalence of any PF OA was 18% (3 to 42) from three studies,^{20 56 62} with

Figure 2 Prevalence of patellofemoral osteoarthritis (PF OA) in community, knee pain or symptomatic and radiographic and symptomatic OA populations. TF, tibiofemoral.

Figure 3 Prevalence of PF OA in healthy individuals, radiographic OA, occupation-based OA and post-traumatic OA populations. PF OA, patellofemoral osteoarthritis; TF, tibiofemoral.

prevalence of mild OA severity at 25% (2 to 87) from two studies.^{56 62}

Prevalence of PF OA based on MRI

Community-based population

The prevalence of isolated PF structural damage and combined PF and TF structural damage based on cartilage defect were 20% and 44% (respectively) and BML was 18% and 22% (respectively) based on a single study.⁹⁵ The prevalence of any PF structural damage based on cartilage defects was 44% (25 to 65) from three studies^{86 95 99} and BML was 29% (11 to 51) from two studies^{95 99} (figure 4A-B).

Knee pain or symptomatic population

The prevalence of overall isolated PF structural damage and combined PF and TF structural damage could not be calculated for this study population. The prevalence of any PF structural damage was 52% (9 to 93) based on cartilage defect^{81 82 97} and 32% (3 to 72) based on BML^{82 84} (figure 4C, D). Data from one study could not be pooled because of WOMRS definition used for OA diagnosis (cartilage damage defined as ≥ 1 grade in this study compared with ≥ 2 grade used in other studies),⁹⁰ with PF structural damage prevalence of 94%, 16% and 81% based on cartilage defect, BML and osteophytes, respectively. Data stratified based on age revealed that the prevalence of any PF structural damage was 71% (33 to 97) in individuals 50 years or over based on cartilage defect.^{81 82 97}

Radiographic knee osteoarthritis or at risk of developing OA

An overall prevalence of isolated PF structural damage, combined PF and TF structural damage and any PF structural damage based on cartilage defect or BML could not be determined for this

study population. Two studies reported prevalence of isolated PF structural damage in the medial and lateral PF compartments based on cartilage defect^{39 91} and BML.^{39 101} The prevalence of isolated medial and lateral PF structural damage was 56% (29 to 81) and 27% (11 to 46), respectively,^{39 91} based on cartilage defect and 28% (17 to 41) and 15% (11 to 20), respectively,^{39 101} based on BML (figure 4E,F). A single study described PF structural damage prevalence based on PF compartment regions (not based on number of individuals or knees)⁹⁴ and reported prevalence of any PF structural damage based on cartilage defect and BML in women (51% and 29%, respectively) and men (43% and 23%, respectively).⁹⁴ No further analyses could be conducted in this study population.

Healthy individuals

The overall prevalence of any PF structural damage based on cartilage defect was 40% (19 to 63)^{92 100} (figure 4G). Since there were only two studies included in this study population, no further analyses could be conducted.

Radiographic and symptomatic knee OA

The prevalence of combined PF and TF structural damage was 75% based on cartilage defect and osteophytes from a single study,⁸³ and no further analyses could be conducted.

Post-traumatic population

Two studies reported prevalence based on osteophytes in ACL injured or reconstructed,^{85 98} with the prevalence of any PF structural damage at 29%⁹⁸ and compartment-specific prevalence of medial and lateral PF structural damage at 23% and 7%, respectively.⁸⁵ The prevalence of medial and lateral PF structural damage based on BML were 2% and 3%, respectively.⁸⁵ The

Figure 4 Prevalence of MRI-defined PF structural damage in community, knee pain or symptomatic, radiographic OA and healthy individual populations. PF OA, patellofemoral osteoarthritis.

prevalence of any PF structural damage was 36% in an ACL ruptured population based on cartilage defect.⁹⁸ In individuals 2 years post arthroscopic partial medial meniscectomy, the prevalence of isolated PF structural damage was 19% based on cartilage defect.¹⁰⁰

General population

Five studies were included in the general population category.^{87–89 93 96} The cartilage defect based prevalence of any PF structural damage was 49% (36 to 62) from two studies using

the WORMS^{87 96} and was 75% (56 to 91) from three studies using the KOSS.^{88 89 93} The prevalence of any PF structural damage based on BML and osteophytes were 45% and 56%, respectively.⁹³

DISCUSSION

Summary of findings

This systematic review with meta-analysis synthesised prevalence of PF OA and included 85 studies. Meta-analysis revealed

the prevalence of any radiographic PF OA in knee pain or symptomatic, radiographic TF OA or at risk of developing TF OA, and radiographic and symptomatic knee OA cohorts was 43%, 48% and 57%, respectively. The prevalence of any MRI-defined PF structural damage in knee pain or symptomatic population was 32% and 52% based on BML and cartilage defect, respectively. This systematic review and meta-analysis highlights the high prevalence of PF OA/ PF structural damage in a wide range of study populations using different imaging tools.

One half (43%–57%) of people with symptoms and/or established radiographic TF OA had PF OA based on radiography. Similarly, a high prevalence of post-traumatic population exhibited signs of PF OA (~30). With such a high prevalence of PF OA, treatments designed specifically for the PF compartment may be required in the OA management strategy.^{102 103} Clinicians should assess for symptoms of PF pain or PF OA and treat patients accordingly. The few studies that specifically evaluated interventions such as exercise, physical therapy, taping and bracing to address PF OA^{98 99 104} provide some evidence for their use. While some studies hypothesise that there is a potential continuum of PF pain to PF OA¹⁰⁵; no high-quality evidence has supported the association between PF pain in younger individuals to the development of PF OA.¹⁰⁶ Unfortunately, studies included in the knee pain or symptomatic OA population category did not differentiate between PF pain and generalised knee pain. Therefore, in the current systematic review, we were not able to determine the prevalence of PF OA in a PF pain population.

Healthy and community cohorts are also likely to demonstrate some PF OA, with radiographic PF OA evident in 17% and 38%, respectively. Since most studies in the community-based meta-analysis were conducted in individuals over the age of 50 years, it appears that radiographic PF OA may be a natural accompaniment to ageing. The only study with a mean age of ≤50 years (but a large range 20 to 93 years), described a particularly high PF OA prevalence in women (81%) and men (88%). The authors hypothesised that cultural factors in Saudi Arabia, such as sitting cross-legged, squatting and praying with knees fully flexed on the ground, may contribute to the high prevalence. Exclusion of this data from meta-analysis revealed the prevalence of any PF OA was 32% in the community population.

The prevalence of MRI-defined PF structural damage in knee pain or symptomatic population was 52%, which was similar to the healthy (40%), community (44%) and general population (49%) cohorts. The high prevalence of MRI-defined PF structural damage may reflect the ability of MRI to detect early changes in the joint that are not visible on radiographs. However, it is unclear whether these findings represent PF OA, as there is no accepted and validated MRI definition of OA. MRI features such as cartilage damage and BMLs can predict incident radiographic OA,¹⁰⁷ development of knee pain¹⁰⁸ and future total knee replacement.¹⁰⁹ Thus, it is plausible that these MRI findings may represent early stages of the PF OA disease process. Further research is needed to investigate the clinical relevance of MRI-defined PF structural damage.

The current systematic review extends on the results from a prior study.⁷ The previous systematic review reported the radiographic prevalence of PF OA in population-based and symptom-based population; whereas, the current review reported prevalence of PF OA in multiple different populations. Thus, an additional 32 studies were included in the meta-analysis. Further to this, the current review included meta-analysis on prevalence of MRI-defined PF structural damage. Furthermore, the current study extends on the findings from the earlier review by categorising data into multiple study populations and data

pooling with subanalysis based on age, sex, compartment-specific OA pattern and OA severity pattern to obtain more accurate estimations of prevalence.

Limitations

This systematic review is not without limitations. First, a very high level of heterogeneity was noted, particularly in the any PF OA group. The inclusion of isolated PF OA, combined PF OA and TF OA, and unclear PF OA (isolated or combined) data in the any PF OA group may explain the high level of heterogeneity. Other potential sources of heterogeneity include differences in diagnostic criteria, populations and case definitions. Second, all relevant studies were included in this systematic review, regardless of methodological quality. Data from 54 low methodological quality studies were included in this review. While this systematic review is subject to bias through the inclusion of low-quality studies, the levels of evidence applied to the pooled data take into account quality, quantity and homogeneity of studies. Third, we restricted the search to studies published in English. Inclusion of data from non-English language studies may alter the outcomes. Fourth, a number of diagnostic criteria were converted to allow data pooling, which may have influenced the results of this systematic review. Fifth, PF structural damage based on MRI should be interpreted with some caution, as fewer studies contributed to meta-analysis within each study population. Lastly, we recognise that there is no accepted and validated definition of radiographic or MRI defined PF OA. Because of this the prevalence data will largely differ in any given study based on different definitions, which may have influenced the results.

Recommendations

While conducting this systematic review, we identified that prevalence data were not well presented in many studies. We recommend that future studies more clearly describe prevalence data based on OA patterns (eg, isolated PF OA vs combined PF OA and TF OA, medial vs lateral PF OA), OA severity (eg, none, mild and moderate) and subgroups (eg, age, sex). Further to this, discrepancies in diagnostic criteria definitions and reporting were noted; therefore, the PF OA definitions should be clearly stated. Better standardisation of data presentation in future studies will help to better understand PF OA epidemiology.

Implications for research and practice

PF OA is an important source of symptoms in knee OA, and is strongly associated with disability.⁶⁰ Our systematic review and meta-analysis revealed the prevalence of PF OA is highly based on radiography and MRI in community, symptomatic, radiographic knee OA and traumatic knee OA populations. Therefore, well-designed studies are required to evaluate biomechanical, functional and psychological impairments associated with PF OA. Addressing potentially modifiable risk factors for PF OA may reduce the risk of development and progression of PF OA and may have implications for TF disease. This systematic review also revealed a higher prevalence of combined PF OA and TF OA pattern than isolated PF OA; therefore, it is important to explore interventions that target both PF and TF joints.

CONCLUSIONS

Synthesis of prevalence data on PF OA and MRI-defined PF structural damage indicates that signs of PF damage are common and should not be ignored in research or clinical practice. In the future, MRI might become highly relevant to identify patients at

early disease stages where the disease process may still be reversible and amenable to interventions.

What are the new findings?

- Patellofemoral OA is prevalent in individuals in a very wide range of settings—in asymptomatic individuals and in patients with knee pain.
- The prevalence rates are influenced by different diagnostic criteria.
- MRI-defined patellofemoral structural damage criteria may assist in identifying patients at early disease stages.

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