## BMJ Open Which biopsychosocial factors are associated with the development of chronic musculoskeletal pain? Protocol for an umbrella review of systematic reviews

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#### **ABSTRACT**

Introduction Recent systematic reviews have identified many biopsychosocial factors associated with the development of chronic musculoskeletal pain (CMP). Despite often being specific to a particular musculoskeletal condition, findings are similar across systematic reviews. Research is needed to aggregate these findings to identify consistent factors across musculoskeletal disorders that are associated with the development of CMP. The objective of this study is to provide a meta-level synthesis of all biopsychosocial factors associated with the development of CMP.

Methods and analysis An umbrella review and metalevel narrative synthesis±meta-analysis has been designed informed by Joanna Briggs Institute and Cochrane guidance. This protocol is reported in line with the Preferred Reporting Items for Systematic Reviews and Meta-Analysis-P. Sources will include Ovid Medline. Embase, Web of Science Core Collection, Cochrane Database of Systematic Reviews, Database of Abstracts of Reviews of Effects, PsycINFO, CINAHL, PEDro, PROSPERO, Google Scholar and grey literature. Inclusion criteria: any systematic review which investigates biopsychosocial factors which may be associated with the development of CMP through prospective longitudinal methods. The outcome is musculoskeletal pain lasting beyond 3 months. Two independent reviewers will be involved in all stages; screening, selection, data extraction and risk of bias evaluation using the Assessing the Methodological Quality of Systematic Reviews-2 guidelines. A metalevel narrative synthesis will be conducted based on (a) factors associated with development of CMP, (b) the range of musculoskeletal disorders for which the same/similar findings have been established and (c) the quality of studies informing these findings. Where possible, meta-analysis will be performed. The Grading of Recommendations, Assessment, Development and Evaluation guidelines will be followed to determine the level of evidence for each biopsychosocial factor. Ethics and dissemination This umbrella review does not require ethical approval. Findings will be presented at

conferences and published in a peer reviewed journal.

PROSPERO registration number CRD42020193081.

#### Strength and limitations on this study

- Inclusion of a substantial number of original research papers and participants.
- The inclusion of all musculoskeletal conditions will establish findings pertinent to a great number of stakeholders and clinical decision-makers.
- The methods employed for this review are rigorous and in line with the most relevant and up to date Cochrane and Joanna Briggs Institute Guidance.
- Quality of life and disability are common outcomes for systematic reviews of chronic musculoskeletal pain but are not included in this review.
- This umbrella review will be limited to studies whereby the full text is available in the English language.

#### INTRODUCTION

The International Classification of Diseases (ICD) describes chronic musculoskeletal pain (CMP) as pain that arises as part of a disease process directly affecting bones, joints, muscle or related soft tissue lasting longer than 3 months. CMP arises secondary to a complex interaction between the systems of the body including the nervous, endocrine, immunological, autonomic, psychological and musculoskeletal (MSK) systems. This interaction creates a maintained elevated perception of threat within the nociceptive system; thus facilitating a prolonged or repeated experience of pain despite limited evidence of potential or actual tissue damage.<sup>2–5</sup> This transition to a sensitised nociceptive system, termed 'nociplastic pain' by the International Association for the Study of Pain, is thought to explain the mechanisms of CMP.<sup>6</sup>

In the UK, the National Health Service spent £4.7 billion pounds on MSK conditions in 2015<sup>7</sup>—an annual figure which is likely



higher now—causing a loss over 30 million working days each year due to MSK conditions, creating a substantial financial and economic burden. MSK complaints also account for up to 30% of general practitioner consultations. A significant proportion of these likely being for the management of CMP with its estimated prevalence in the UK at 43%. CMP is also a huge burden internationally with low back pain identified as the single greatest cause of years lived with a disability; thus prompting the ICD to recently recognise CMP as a disease in its own right. Given the apparent difficulty in effectively managing CMP, and its increasing prevalence, a better understanding of CMP is needed.

To better understand CMP, there is a need to understand the factors which influence its development. The experience of MSK pain is biopsychosocial and therefore biological factors, such as the type and severity of MSK condition, do not equate to consistent experiences of pain among individuals. 13-15 Rather, many individuals go on to develop CMP where others do not. This discrepancy has frequently been associated with the presence of many biopsychosocial factors such as catastrophised beliefs, lower socioeconomic status and fear of movement. 16-18 However, existing systematic reviews often focus on biological, psychological or social factors individually or in relation to only one form or CMP, for example, low back pain. Given that nociplastic mechanisms, and many biopsychosocial factors, are not specific to a particular MSK condition, synthesis of this evidence base may inform future research and management of CMP conditions as a whole.

Biomedical healthcare approaches tend to dominate the management of MSK conditions with biopsychosocial approaches often only considered once medical management has failed, as demonstrated by National Institute for Health and Care Excellence guidance for low back pain. <sup>19</sup> This equates to a reactionary approach to CMP whereby biopsychosocial care is usually only implemented once CMP is well established. By this point, CMP is very difficult to effectively treat with 79%–92% of people still experiencing pain when followed-up up to 12 years later. <sup>20–22</sup> A synthesis of the biopsychosocial factors associated with the development of CMP may help identify those at risk

of CMP before it has been established. This provides a window of opportunity for healthcare services to deliver proactive biopsychosocial treatments with an aim of preventing development of CMP. This strategy works well for other chronic diseases such as heart disease <sup>23</sup> and diabetes. <sup>24</sup>

To address the growing prevalence and burden of CMP, it is timely to synthesise the existing literature to identify which biopsychosocial factors are associated with its development. Given the abundance of systematic reviews, an umbrella review will be most appropriate in achieving this.

#### **AIM**

To establish which biopsychosocial factors are associated with the development of CMP.

#### **METHODS**

#### **Design and registration**

This protocol was developed in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses Protocol (PRISMA-P)<sup>25</sup> (see online supplemental file A for completed PRISMA-P checklist). The methodology has been guided by the Joanna Briggs Institute Manual for Evidence Synthesis of Umbrella Reviews<sup>26 27</sup> and the Cochrane handbook for the conduct of systematic reviews.<sup>28</sup>

#### Patient and public involvement

The basis for the need and relevance of this umbrella review is based on the clinical interactions of the authors with previous and current patients with CMP. This includes members of the patient and public involvement and engagement group within the Centre of Precision Rehabilitation for Spinal Pain at the University of Birmingham. The group includes individuals with chronic spinal pain.

#### **Eligibility criteria**

The eligibility criteria were established using a modified PICOS statement (see table 1). 'I' was replaced with 'E'

Table 1 PICOS statement			
Population	Individuals aged 18 and over who have experienced an MSK disorder.		
Exposure	Any biopsychosocial factors which may have influenced outcome, for example, patient reported higher levels of pain at onset.		
Comparator	Those who have not been exposed to the biopsychosocial factor under investigation, for example, the comparator to those who report higher levels of pain at onset would be those who report lower levels of pain at onset.		
Outcome	Persistence in MSK pain beyond 3 months. This may be identified through any MSK pain-related outcome measure or through clinician diagnosis.		
Study designs	Systematic reviews of prospective longitudinal design studies.		

MSK, musculoskeletal.



for exposure due to the epidemiological nature of the review.<sup>29</sup>

#### Inclusion criteria

Systematic reviews of observational studies with/without meta-analysis investigating factors associated with MSK pain lasting longer than 3 months.

#### **Exclusion criteria**

Systematic reviews which include interventional studies, for example, factors associated with successful surgery, populations with a high risk or evidence of poor tissue healing, for example, autoimmune disorders, injuries where tissue healing may be incomplete at 3 months, for example, fractures, draw body region-specific conclusions which are not generalisable to the wider CMP population, for example, a bony heel spur associated with chronic heel pain, pool data with non-MSK chronic pain populations, for example, cancer-related pain, potential systemic drivers of CMP, for example, spondyloarthropathy, do not include an appropriate comparator and systematic reviews where the full text is not available in the English language.

#### **Information sources**

A systematic search of electronic databases and grey literature will be conducted. Ovid MEDLINE, Ovid EMBASE, Web of Science Core Collection and Google Scholar will be searched as suggested by Bramer et al, 30 who found this combination to be most effective for the retrieval of systematic reviews. In addition, the PEDro, CINAHL and Ovid PsycINFO databases will be searched as well as the Cochrane database for systematic reviews, the Database of Abstracts of Reviews of Effects and the PROSPERO register as all are particularly relevant for the subject of MSK pain and/or systematic reviews. There will be no limitation on search dates. Grey literature will be searched using the Canadian Agency for Drugs and Technologies in Health grey literature searching tool. To ensure literature saturation, we will also scan the reference lists of included studies or relevant reviews identified through the search.

#### **Search strategy**

The search strategy has been designed by the lead author (MD) with the assistance of coauthors (NRH, ABR and AS) and an experienced health sciences librarian with expertise of systematic review searching. The search strategy includes a validated search strategy for prognostic studies recommended by the Cochrane Prognosis Research Group. The initial search strategy was developed with Ovid MEDLINE using medical subject headings and text words and subsequently adapted to the syntax and subject headings of the other databases to be searched for this review. The search strategy has been peer reviewed by other members of the review team (NRH, ABR, AS and JM) to reduce the risk of search errors and ensure the highest retrieval of eligible studies. Please

see online supplemental file B for the Ovid MEDLINE search strategy.

#### **Screening and selection**

A systematic search of databases using the search strategies will be conducted by two reviewers independently of each other (MD and JM). Titles and abstracts will be retrieved and stored using Microsoft EndNote X9.3.3, will be assessed based on the inclusion/exclusion criteria and allocated into one of two groups: potentially eligible or not eligible. Full-text sources for potentially eligible studies will then be sourced and discussed between both reviewers for confirmation of eligibility. Additional information to determine eligibility will be sought from study authors via email where necessary. Any disagreement of eligibility at this stage will be referred to a third reviewer (NRH). All reasons for excluding reviews will be recorded using a PRISMA flow chart. None of the reviewers will be blinded to the journal titles, study authors or institutions.

#### **Data extraction**

Two reviewers (MD and JM) will independently extract data. All data excluded will be agreed by both reviewers in discussion and any disagreement will be referred to a third reviewer (NRH). Where study data are unclear or missing, the corresponding author will be contacted via email for clarification or further information. A second and final email will be sent if no response has been received after 2 weeks. In the event of a non-response after a further 2 weeks, includable data will be extracted based on agreement between the two reviewers with any disagreement referred to a third reviewer. Data will be extracted using a bespoke and standardised proforma which has been piloted a priori (see online supplemental file C). Data extraction will, where possible, include quantitative data such as demographic information, ORs, relative risk ratios, size of effects and Pearsons  $\chi^2$  scores which will be stored on Microsoft Excel.

#### **Data items**

The data to be extracted are summarised in table 2. As per Joanna Briggs Umbrella Review guidance, data will not be collected from the original research studies but extracted from the included systematic reviews only.

#### Risk of bias assessment

Two reviewers (MD and JM) will independently perform risk of bias assessment of included systematic reviews using the Assessing the Methodological Quality of Systematic Reviews-2 (AMSTAR-2) guidelines and checklist. AMSTAR-2 has been shown to be valid and reliable for the appraisal of systematic reviews and has been used in many high-quality healthcare Umbrella Reviews. AMSTAR-2 includes assessment of study eligibility criteria, identification and selection of studies, data collection methods, study appraisal methods and findings and synthesis methods. Overall confidence in the results of each systematic review will be rated as high, moderate, low or critically low. Any disagreements will be discussed

Table 2 Summary of items to be extracted from included studies			
Study and population characteristics	Reference, type of population (eg, idiopathic, traumatic, mixed), number of studies included, number of participants and demographic information.		
Outcome measures	Name of outcome measures or methods used to determine CMP.		
Methods and findings	Method of data synthesis, biopsychosocial factors associated with development of CMP, methods of data collection for biopsychosocial factors, musculoskeletal conditions for which findings have been established, quality of studies informing findings, appraisal tools used and quantitative results including but not limited to ORs and relative risk ratios.		

CMP, chronic musculoskeletal pain.

and referred to a third reviewer (NRH) for a conclusive decision. Reporting bias will be determined based on whether a protocol exists as a peer review publication or through registration with PROSPERO.

#### **Data synthesis**

Meta-analysis will be performed where two or more included reviews have reported on the same biopsychosocial factor and the data can be suitably pooled for statistical analysis. Statistical analysis will then be performed by computing OR using a random-effects model with a 95% CI. Pearson's  $\chi^2$  test will be used to assess statistical significance of heterogeneity between reviews. Publication bias will be assessed through the use of an inverted funnel plot. All statistical analysis will be conducted using IBM SPSS Statistics V.26 and Microsoft Excel.

Where meta-analysis is not possible, meta-level narrative synthesis will be performed in accordance with guidance by Popay et al (Guidance on the Conduct of Narrative Synthesis in Systematic Reviews. Results of an ESRC funded research project (unpublished report)) and The Cochrane Consumers and Communication Review Group.<sup>39</sup> However, the first step of 'developing a theory of how the intervention works' will be removed due to the non-interventional nature of this umbrella review. Narrative synthesis will be structured around three domains: (#a) the biological, psychological or social factors associated with the development of CMP, (b) the range of MSK disorders for which the same/similar findings have been established and (c) the risk of bias of the studies informing these findings. Using standardised templates, data will be extracted and presented in tabular format. Discussion will be orientated around similarities and differences between the findings of included studies, in line with our three domains. A detailed outline of methodological problems or biases will also be included, alongside an assessment of completeness and applicability.<sup>36</sup>

#### **Confidence in cumulative evidence**

Umbrella reviews have been criticised for lack of consistent or appropriate methods in ascertaining the level of certainty of findings. The Grading of Recommendations Assessment, Development and Evaluation (GRADE) approach is a well-established tool which is commonly used for this purpose with systematic reviews and has been recommended for use with umbrella reviews. The

GRADE approach has been adapted for the assessment of evidence of prognostic factors. 41 GRADE enables evaluation of criteria rating down factors associated with the development of CMP (risk of bias (informed by AMSTAR-2), inconsistency, indirectness, imprecision and publication bias) and criteria of rating up factors associated with the development of CMP (dose–response, large effect or for the nature of plausible biases) to provide a statement of the overall level of evidence as high, moderate, low or very low for each biopsychosocial factor.

#### DISCUSSION

Traditionally, research of CMP has been orientated towards a particular chronic MSK condition rather than considering CMP as an overarching condition in itself. Rather, this umbrella review will consider CMP as a disease in its own right in line with the ICD<sup>1</sup> and seeks to contribute to the body of evidence supporting a shift away from the over-medicalisation of CMP.<sup>42</sup>

Physical assessment and imaging are typically utilised to determine physical 'abnormalities' with these 'abnormalities' then serving as the rationale for the diagnosis and treatment of a specific MSK condition. 43-45 Treatments may include surgery, acupuncture, manual therapy, injections, pharmacology or surgery<sup>19</sup> among others, with the aim of 'fixing' the 'abnormality' and improving the MSK condition. However, research demonstrates that these 'abnormalities' are heavily prevalent in healthy populations with no history of MSK conditions.  $^{43}$   $^{46}$   $^{47}$  This raises concern for the validity of this form of diagnosis. In addition, a large multicentre randomised control trial also demonstrates that sham surgery is equally as effective as surgical intervention for shoulder pain, 48 raising questions for the mechanisms of how effective interventions for MSK pain work. This same finding has also been found with sham surgery for knee, back and elbow pain as evidenced by a high-quality systematic review<sup>49</sup> suggesting that the mechanisms of pain and successful intervention are similar, regardless of the type of MSK condition. Notably, the participants in these studies typically have had symptoms for longer than 3 months and therefore these findings are relevant to the CMP population.

The nociplastic mechanisms by which individuals experience CMP are thought to be contributed to by functional



and anatomical changes in the central nervous system. <sup>50–52</sup> Reorganisation of brain activity takes place with reduced activation of the somatosensory cortex, which is usually highly active during nociception, and an increase in activity of the corticolimbic system, thus creating a shift from nociceptive to emotional circuits. <sup>50</sup> These changes have been shown to be consistent across a range of MSK conditions. <sup>53–56</sup> Furthermore, this persistent activity of the corticolimbic system is responsible for the development of many behavioural and psychological presentations, <sup>57</sup> many of which have separately been identified as risk factors for the development of CMP, such as fear. <sup>16 18</sup> This is further evidenced by the consistent biopsychosocial factors shown to be associated with development of CMP regardless of the type of MSK condition. <sup>16–18</sup>

For these reasons, and that CMP is considered a disease in its own right, this umbrella review recognises the need to aggregate the findings of systemic reviews on the biopsychosocial factors associated with the development of CMP, for all MSK conditions.

#### **AMENDMENTS**

In the event of protocol amendments, the date of each amendment will be accompanied by a description of the change and the rationale for the change.

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Contributors MD conceived, designed and will coordinate the review; will collect, manage, analyse and interpret data; and write the review. JM will collect, manage analyse and interpret data. NRH, ABR and AS contributed to conception and design of the review and will provide general advice on the coordination of the review as well as collection, management, analysis and synthesis of data.

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### PRISMA-P 2015 Checklist

This checklist has been adapted for use with protocol submissions to *Systematic Reviews* from Table 3 in Moher D et al: Preferred reporting items for systematic review and meta-analysis protocols (PRISMA-P) 2015 statement. *Systematic Reviews* 2015 **4**:1

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Section/topic	#	Checklist item	Yes	No	number(s)	
ADMINISTRATIVE IN	FORMAT	TION				
Title						
Identification	1a	Identify the report as a protocol of a systematic review			2	
Update	1b	If the protocol is for an update of a previous systematic review, identify as such			N/A	
Registration	2	If registered, provide the name of the registry (e.g., PROSPERO) and registration number in the Abstract			44-45	
Authors						
Contact	3a	Provide name, institutional affiliation, and e-mail address of all protocol authors; provide physical mailing address of corresponding author			5-15	
Contributions	3b	Describe contributions of protocol authors and identify the guarantor of the review			278-282	
Amendments	4	If the protocol represents an amendment of a previously completed or published protocol, identify as such and list changes; otherwise, state plan for documenting important protocol amendments			223-225	
Support						
Sources	5a	Indicate sources of financial or other support for the review			275-277	
Sponsor	5b	Provide name for the review funder and/or sponsor			275-277	
Role of sponsor/funder	5c	Describe roles of funder(s), sponsor(s), and/or institution(s), if any, in developing the protocol			275-277	
INTRODUCTION						
Rationale	6	Describe the rationale for the review in the context of what is already known			60-101	
Objectives	7	Provide an explicit statement of the question(s) the review will address with reference to participants, interventions, comparators, and outcomes (PICO)			102-103 & 116-120	



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Section/topic #		Checklist item		Information reported	
Section/topic	#	Checklist item	Yes	No	number(s)
METHODS					
Eligibility criteria	8	Specify the study characteristics (e.g., PICO, study design, setting, time frame) and report characteristics (e.g., years considered, language, publication status) to be used as criteria for eligibility for the review			116-132
Information sources	9	Describe all intended information sources (e.g., electronic databases, contact with study authors, trial registers, or other grey literature sources) with planned dates of coverage			135-145
Search strategy	10	Present draft of search strategy to be used for at least one electronic database, including planned limits, such that it could be repeated			146-155 & Additional file A
STUDY RECORDS					
Data management	11a	Describe the mechanism(s) that will be used to manage records and data throughout the review			158-159 & 173-176
Selection process	11b	State the process that will be used for selecting studies (e.g., two independent reviewers) through each phase of the review (i.e., screening, eligibility, and inclusion in meta-analysis)			156-165
Data collection process	11c	Describe planned method of extracting data from reports (e.g., piloting forms, done independently, in duplicate), any processes for obtaining and confirming data from investigators			166-176
Data items	12	List and define all variables for which data will be sought (e.g., PICO items, funding sources), any pre-planned data assumptions and simplifications			174-176, 178- 183 & Additional file B
Outcomes and prioritization	13	List and define all outcomes for which data will be sought, including prioritization of main and additional outcomes, with rationale			119-120
Risk of bias in individual studies	14	Describe anticipated methods for assessing risk of bias of individual studies, including whether this will be done at the outcome or study level, or both; state how this information will be used in data synthesis			184-193
DATA					
	15a	Describe criteria under which study data will be quantitatively synthesized			195-200
Synthesis	15b	If data are appropriate for quantitative synthesis, describe planned summary measures, methods of handling data, and methods of combining data from studies, including any planned exploration of consistency (e.g., $I^2$ , Kendall's tau)			196-200



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Section/topic #		Checklist item		Information reported	
Section/topic	#	Checklist item		No	number(s)
	15c	Describe any proposed additional analyses (e.g., sensitivity or subgroup analyses, meta-regression)			N/A
	15d	If quantitative synthesis is not appropriate, describe the type of summary planned			201-211
Meta-bias(es)	16	Specify any planned assessment of meta-bias(es) (e.g., publication bias across studies, selective reporting within studies)			198-199
Confidence in cumulative evidence	17	Describe how the strength of the body of evidence will be assessed (e.g., GRADE)			212-222



#### **ADDITIONAL FILE B**

#### **Ovid Medline Search Strategy**

#### Associated Factors

- 1 Validat\*.mp. or Predict\*.ti. or Rule\*.mp. or (Predict\* and (Outcome\* or Risk\* or Model\*)).mp. or ((History or Variable\* or Criteria or Scor\* or Characteristic\* or Finding\* or Factor\*) and (Predict\* or Model\* or Decision\* or Identif\* or Prognos\*)).mp. or (Decision\*.mp. and ((Model\* or Clinical\*).mp. or Logistic Models/)) or (Prognostic and (History or Variable\* or Criteria or Scor\* or Characteristic\* or Finding\* or Factor\* or Model\*)).mp.
- 2 (Stratification or discrimination or discriminate or c statistic or auc or calibration or indices or algorithm or multivariable).tw.
- 3 (Area under adj1 curve).tw.
- 4 Predict\*.tw. or predictive value of tests/ or Scor\*.tw. or Observ\*.tw. or observer variation/ (4591145)
- 5 Roc curve/ or Forecasting/ or prognosis/ or exp probability/ or exp epidemiologic factors/ or exp regression analysis/
- 6 (Predict\* or forecasting or Prognos\* or probability or determin\* or moderator or effect modifi\* or regression analys?s\* or logistic regression or logistic model\* or odds ratio or risk ratio or relative risk or risk factor\* or protective factor\* or Cox model\* or hazard\* model\* or hazard ratio or reason\* or theme\* or drive\* or indicator\* or sign\* or link\* or correlat\* or cause\* or factor\* or affordance or constraint or obstacle or impediment or enabler or inhibit\* or stimulat\* or facilitate\* or barrier\*).tw,kf.
- 7 (dependenc\* or mobility aid or crutch\* or walking stick\* or walking frame\* or zimmer frame\* or rollator frame\* or wheelchair\* or obese or obesity or overweight or high BMI or underweight or low BMI or age or older or elderly or comorbidit\* or mental health or emotion\* or depression or anxiety or distress or headache\* or poorly controlled pain or pain intensity or education or pre existing pain or catastophi?\* or female or gender or male or psychological or biopsychosocial or psychosocial or smok\*OR sensory disturbance or sensory deficit or altered sensation or paraesthesia or fear\* or avoid\* or occupation or return\* work or genetics or genes or swelling or reduced range motion or reduced ROM or loss range motion or loss range movement or loss ROM or restricted range motion or restricted range movement or restricted ROM or limited weight bearing or reduced weight bearing or ability weight bear or re-injury or stress or passive coping strategies or reduced self efficacy or low self efficacy or somati?ation or resilience or concern\* or physical activity or motivation or persistan\* or guarding or guarded or socioeconomic status or alcohol or high\* income or low\* income or social support or support\* network or propriocept\* or joint position sense).tw,kf.
- 8 (Concomitant adj3 pain).tw,kf.
- 9 (Poor\* adj2 health).tw,kf.
- 10 or/1-9

Chronic pain

- 11 Chronic pain/ OR ((Persistent or chronic or recurrent or continu\* or lasting or long or intractable or endur\*) adj2 pain).tw,kf.
- 12 (exp Outcome Assessment, Health Care/ or patient outcome assessment/ or treatment outcome/ or program evaluation/ or disability evaluation/ or follow-up studies/) and Chronic pain/
- 13 ((Poor\* or reduced or worse or inferior) adj2 outcome\*).tw,kf.
- 14 or/11-13

#### All musculoskeletal conditions

- 15 Knee.tw,kf. or Knee/ or exp knee joint/ or anterior cruciate ligament/
- 16 exp knee injuries/ or Knee dislocation/ or Osteoarthritis, knee/ or Medial Collateral Ligament, Knee/in or Posterior cruciate ligament/in or menisci, tibial/in or Patellofemoral pain syndrome/ or patella/in or patella ligament/in or quadriceps muscle/in
- 17 "sprains and strains"/ and knee.tw,kf.
- ((knee or patellofemoral or tibiofemoral or patella\* or prepatella\* or tibia\* or fibula\* or meniscus or menisci or collateral ligament or medial patellofemoral ligament or cruciate ligament or hamstring\* or quadricep\* or semitendinosus or semimembranosus or biceps femoris or popliteus or popliteal or iliotibial band or chondromalacia or plica) adj3 (sprain\* or strain\* or myalgia or tear\* or bursitis or tend?nopathy or tend?nitis or tend?nosis or tenosynovitis or rupture or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome)).tw,kf.
- 19 (PFP or PFPS or chondromalacia patella).tw,kf.
- 20 (Ankle or foot or toe\*).tw,kf. or Ankle/ or exp foot joints/ or lateral ligament, ankle/
- 21 Ankle injuries/ or lateral ligament, ankle/in or achilles tendon/in or exp foot joints/in or fasciitis, plantar/ or heel spur/ or exp metatarsalgia/ or posterior tibial tendon dysfunction/ or medial tibial stress syndrome/
- 22 "sprains and strains"/ and (ankle or foot).tw,kf.
- 23 ((ankle or foot or talus or talar or calcaneu\* or cuboid or navicular or cuneiform or metatarsal or tarsal or hallux or malleolus or malleolar or talofibular ligament or calcaneofibular ligament or deltoid ligament or tibiotalar ligament or tibiocalcaneal ligament or tibionavicular ligament or achilles or calf or calves or gastroc\* or soleus or tibialis or peroneus or peroneal\* or digitorum or hallucis or digiti minimi or lumbrical\* or interosseus or interossei or retinaculum) adj3 (sprain\* or strain\* or myalgia or tear\* or bursitis or tend?nopathy or tend?nitis or tend?nosis or tenosynovitis or rupture or impingement or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome)).tw,kf. (17499)
- 24 (plantar fasciitis or plantar fascia pain).tw,kf.
- 25 Shoulder.tw,kf. or Shoulder/ or shoulder joint/ or rotator cuff/
- 26 Shoulder pain/ or shoulder impingement syndrome/ or shoulder joint/in or shoulder injuries/ or rotator cuff injuries/ or shoulder dislocation/
- 27 "sprains and strains"/ and shoulder.tw,kf.

- ((Shoulder or glenohumeral or GHJ or humerus or humeral or clavicle or clavicular or scapula\* or acromioclavicular or ACJ or scapulothoracic or subacrom\* or rotator cuff or supraspinatus or infraspinatus or subscapularis or teres minor or teres major or trapezius or deltoid\* or bicep\* or bicipital or latissimus dorsi or rhomboid\* or labrum or labral or glenoid or supraglenoid or infraglenoid) adj3 (sprain\* or strain\* or myalgia or tear\* or bursitis or tend?nopathy or tend?nitis or tend?nosis or tenosynovitis or impingement or rupture or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome)).tw,kf.
- 29 (Elbow or forearm or wrist or hand or thumb or finger).tw,kf. or Elbow joint/ or exp hand joints/ or exp upper extremity/
- 30 exp hand joints/in or upper extremity/in or exp elbow tendinopathy/ or exp hand injuries/ or De Quervain disease/
- 31 ((Elbow\* or forearm\* or arm\* or wrist\* or hand\* or finger\* or thumb\* or radioulnar or metacarpal or interphalangeal or phalanx or thenar or hypothenar or humerus or olecranon or radius or radial or ulna or scaphoid or lunate or triquetrum or pisiform or trapezium or trapezoid or capitate or hamate or scapholunate ligament or collateral ligament or Bicep\* or Bicipital or coracobrachialis or brachioradialis or brachialis or tricep\* or carpi radialis or carpi ulnaris or pronator teres or digitorum or Pronator quadratus or palmaris longus or supinator or pollicis or lumbrical\* or digiti minimi or interosseus or interossei or triangular cartilage or triangular fibrocartilage complex or TFCC) adj3 (sprain\* or strain\* or myalgia or tear\* or bursitis or tend?nopathy or tend?nitis or tend?nosis or tenosynovitis or rupture or impingement or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome)).tw,kf.
- 32 (de quervain\* or golfers elbow or tennis elbow).tw,kf.
- 33 ((medial or lateral) adj1 (epicondylitis or epicondylalgia or epicondylopathy or epicondylosis or tendinopathy or elbow pain)).tw,kf.
- 34 (Thoracic or mid back or middle back or upper back).tw,kf. or Thoracic vertebrae/ or thorax/ or Intercostal muscles/ or intermediate back muscles/ or paraspinal muscles/
- 35 (osteoarthritis/ or osteoarthritis, spine/ or myofascial pain syndromes/ or exp spinal osteophytosis/ or spondylolisthesis/ or intervertebral disc displacement/) and (thoracic or mid back or middle back or upper back).tw,kf.
- 36 Tietze's Syndrome/
- 37 ((thoracic or thoracic spin\* or T spin\* or thoracic wall or thorax or thoracolumbar or thoracic intervertebral disc or thoracic disc or thoracic muscle\* or erector spin\* or intercostal or costovertebral or costotransverse or costochondral or rib or pectoral or T1 or T2 or T3 or T4 or T5 or T6 or T7 or T8 or T9 or T10 or T11 or T12) adj3 (strain\* or impingement or compression or stenosis or narrowing or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome or spondylitis or spondylolysis or spondylolisthesis or spondylosis or bulge or herniation or prolapse or slipped)).tw,kf.
- 38 (Lumb?r or low\* back).tw,kf. or Lumbar vertebrae/ or exp back/ or exp back muscles/ or sacroiliac joint/
- 39 (Osteoarthritis/ or osteoarthritis, spine/ or myofascial pain syndromes/ or exp osteophytosis/ or spondylolisthesis/ or intervertebral disc displacement/) and (lumb?r or back).tw,kf.

- 40 back pain/ or low back pain/ or exp back injuries/
- 41 ((low\* back or lumb?r or sacroiliac or lumbar facet or SIJ or L1 or L2 or L3 or L4 or L5 or S1) adj3 (strain\* or impingement or compression or stenosis or narrowing or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome or spondylitis or spondylolysis or spondylolisthesis or spondylosis or bulge or herniation or prolapse or slipped)).tw,kf.
- 42 non specific low\* back pain.tw,kf.
- 43 (Hip or pelvi?).tw,kf. or Hip/ or hip joint/ or exp pelvis/
- 44 Osteoarthritis, hip/ or Femoroacetabular impingement/ or hip dislocation/ or hip injuries/ or pelvic girdle pain/ or piriformis syndrome/
- 45 "sprains and strains"/ and hip.tw,kf.
- ((hip or labrum or labral or greater trochanter\* or GT or lesser trochanter\* or acetabulum or femur or femoral or groin or thigh or hip flexor\* or pelvi? or iliac or ilium or ischium or ischial or pubi? or ASIS or anterior superior iiliac spine or rectus femoris or vastus medialis or vastus lateralis or vastus intermedius or iliacus or psoas major or iliopsoas or gluteus or gluteal or adductor or piriformis) adj3 (sprain\* or strain\* or myalgia or tear\* or bursitis or tend?nopathy or tend?nitis or tend?nosis or tenosynovitis or rupture or impingement or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome)).tw,kf.
- 47 (FAI or PGP).tw,kf.
- 48 (neck or cervical or temporomandibular).tw,kf. or Neck/ or neck muscles/ or cervical plexus/ or exp cervical vertebrae/ or atlanto-axial joint/ or atlanto-occipital joint/ or temporomandibular joint/
- 49 (osteoarthritis/ or osteoarthritis, spine/ or myofascial pain syndromes/ or exp spinal osteophytosis/ or spondylolisthesis/ or intervertebral disc displacement/) and (cervical or neck).tw,kf.
- 50 exp neck pain/ or temporomandibular joint dysfunction syndrome/ or exp neck injuries/ or cervical rib syndrome/ or torticollis/
- ((neck or odontoid or cervical or occip\* or atlant\* or cervical disc or cervical interverteb\* or cervical facet or C1 or C2 or C3 or C4 or C5 or C6 or C7 or C8) adj3 (strain\* or impingement or compression or stenosis or narrowing or injur\* or impair\* or disorder or dysfunction or trauma or arthritis or osteoarthritis or syndrome or spondylitis or spondylolysis or spondylolisthesis or spondylosis or bulge or herniation or prolapse or slipped)).tw,kf.
- 52 Musculoskeletal pain/ OR myofascial pain syndromes/
- 53 or/15-52

Systematic review/meta-analysis

54 (systematic review or meta analysis).tw,kf.

All Strings Combined

55 10 and 14 and 53 and 54

# Which biopsychosocial factors are associated with the development of chronic musculoskeletal pain? An umbrella review of systematic reviews

#### **Data Extraction Template**

<ol> <li>General information and citation d</li> </ol>	letail	S
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Reviewer:	
Date form completed:	
(DD/MM/YYYY)	
Study ID:	
(Surname of first author and year	
of publication e.g. Smith 2001)	
Title of study:	
Notes:	

2. Population, databases, studies and participants

MSK conditions:	
(e.g. neck pain, WAD, shoulder pain etc)	
Population:	
(idiopathic, traumatic or mixed)	
Notes:	
Number of studies included in review:	
Number of participants included in review:	
Age range of participants:	
Notes:	

3. Methods, appraisal and findings

Methods of data			
analysis/synthesis:			
Appraisal tool used:			
Notes:			
Outcome measures			
to determine CMP:			
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Factor associated with	aevelopment of	Quality of studies	MSK condition
chronic musculoskelet		which inform findings	IVISK condition
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		· · · · · · · · · · · · · · · · · · ·	NISK condition

Factor NOT associated with development of CMP	Quality of studies which inform findings	MSK condition

4. Notes or comments: