

BMJ Open Interventions for the prediction and management of chronic postsurgical pain after total knee replacement: systematic review of randomised controlled trials

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ABSTRACT

Objectives: Total knee replacement can be a successful operation for pain relief. However, 10–34% of patients experience chronic postsurgical pain. Our aim was to synthesise evidence on the effectiveness of applying predictive models to guide preventive treatment, and for interventions in the management of chronic pain after total knee replacement.

Setting: We conducted a systematic review of randomised controlled trials using appropriate search strategies in the Cochrane Library, MEDLINE and EMBASE from inception to October 2014. No language restrictions were applied.

Participants: Adult patients receiving total knee replacement.

Interventions: Predictive models to guide treatment for prevention of chronic pain. Interventions for management of chronic pain.

Primary and secondary outcome measures: Reporting of specific outcomes was not an eligibility criterion but we sought outcomes relating to pain severity.

Results: No studies evaluated the effectiveness of predictive models in guiding treatment and improving outcomes after total knee replacement. One study evaluated an intervention for the management of chronic pain. The trial evaluated the use of a botulinum toxin A injection with antinociceptive and anticholinergic activity in 49 patients with chronic postsurgical pain after knee replacement. A single injection provided meaningful pain relief for about 40 days and the authors acknowledged the need for a large trial with repeated injections. No trials of multidisciplinary interventions or individualised treatments were identified.

Conclusions: Our systematic review highlights a lack of evidence about the effectiveness of prediction and management strategies for chronic postsurgical pain after total knee replacement. As a large number of people are affected by chronic pain after total knee replacement, development of an evidence base about care for these patients should be a research priority.

Strengths and limitations of this study

- Reproducible systematic review methods.
- Identification of research priorities.
- Interventions for chronic pain after other surgeries may have value in total knee replacement.

INTRODUCTION

Total knee replacement is an increasingly common procedure that aims to reduce pain and functional limitations, particularly for people with osteoarthritis of the knee. In the year to 31 March 2014, nearly 78 000 people received a primary total knee replacement in the UK,¹ and in 2010 approximately 719 000 procedures were performed in the USA.² It is estimated that over half of all people in the USA diagnosed with osteoarthritis will receive a total knee replacement.³

Surgery is a known risk factor for chronic pain⁴ defined as pain ‘present for at least 3 months’.⁵ Chronic postsurgical pain ‘develops after a surgical procedure or increases in intensity after the surgical procedure’.⁶ Although many patients report a good outcome after their total knee replacement, at a time when recovery should have been achieved,⁷ about 10–34% of patients report moderate to severe chronic postsurgical pain.⁸ In the UK, this could mean 7500–25 500 potential new cases of chronic postsurgical pain every year, while in the USA this equates to between 72 000 and a quarter of a million new cases annually. As patients undergo knee replacement in order to relieve knee pain, these estimates are cause for concern.

Given the distress caused by chronic postsurgical pain,⁹ and the predicted increases in prevalence of osteoarthritis,¹⁰ and the need for knee replacement surgery,¹¹ robust

evidence is needed on effective methods for preventing the development of chronic pain, identifying patients at risk of developing chronic pain, and for the management of chronic pain. Inadequately controlled peri-operative pain is a risk factor for long-term pain and, although studied widely, systematic reviews have shown that evidence on long-term benefit is limited.^{12–15}

A large number of preoperative and early post-operative factors are associated with poor pain outcomes, including greater joint pain^{16–18} and pain catastrophisation,^{19–21} poor mental health,^{16 19 21–23} and presence of musculoskeletal comorbidities.^{18 24} As the cause of chronic pain after total knee replacement is likely to be multifactorial, with mechanical, biological and psychological features, simple interventions targeting individual issues will leave a large proportion of patients at risk of developing long-term pain with no appropriate care. The potential value of multivariable risk assessment is clear although the ability of predictive models to identify patients at risk of long-term pain has been highly variable.^{25 26} Furthermore, as with all prognostic models guiding decision-making, evidence on their efficacy and safety in targeting interventions is required before application in clinical practice.²⁷

Owing to the complexity of chronic pain, treatments in appropriate combinations matched to patient characteristics are advocated.^{28 29} As with application of methods for prediction, evidence is required that pain management strategies are effective in patients with chronic postsurgical pain after total knee replacement. This may relate to specific treatments or to multifactorial assessment and management.

Our aim was to conduct a systematic review to identify randomised trials in patients with total knee replacement that have evaluated: (1) the application of predictive models in the targeting of pain management and (2) interventions for the treatment of chronic pain. Relevant outcomes related to pain severity. While our particular interest was total knee replacement, we used a broader search strategy to include any type of knee surgery as appropriate pain prediction, and management methods may have been evaluated in more diverse knee surgeries.

METHODS

We aimed to conduct our literature reviews with transparent and unbiased methods such that they can be considered truly systematic and reproduced on the basis of sources of literature, search processes, study inclusion or exclusion, data analysis if feasible, and study quality assessment. To achieve this, we used methods described in the Cochrane Handbook of Systematic reviews.³⁰ As this review focuses on randomised controlled trials, we conducted the review with reference to Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines which aim to improve the reporting of systematic reviews.³¹

Search strategy

Separate literature search strategies for predictive methods and pain management based on validated searches^{30 32 33} were applied in MEDLINE, EMBASE and the Cochrane Library from inception up to 1 October 2014. We considered that the EMBASE coverage of conference abstracts since 2009 was an appropriate search of ‘grey literature’ in the orthopaedic context. Search strategies as applied in MEDLINE are shown in **box 1** with combinations of terms, such as ‘risk function, risk assessment, randomised trial, knee,’ and ‘pain, post-operative, post-surgical, randomised trial, knee.’ Search terms were in English, but no further language restrictions were applied with funds available to pay translation costs if required. If necessary author contact for additional information was planned.

Eligibility criteria

Eligible studies satisfied PICOS criteria.

- ▶ **Patients:** adults with knee surgery
- ▶ **Intervention:** treatment guided by a predictive model or an intervention for management of chronic pain (pain reported at 3 months or more after surgery)
- ▶ **Control:** a usual care comparison group
- ▶ **Outcome:** an outcome relating to pain severity
- ▶ **Setting:** evaluation in a randomised controlled trial

Data extraction

Articles and inclusion/exclusion decisions were catalogued in Endnote X7. All titles and abstracts were screened independently by two reviewers. Potentially relevant articles were evaluated in detail by two reviewers, independently, with decisions on relevance made after discussion. Data on study and patient characteristics, intervention and control group treatment, follow-up and outcomes and results were extracted onto a summary table.

Outcomes

We did not exclude studies on the grounds of what outcomes were reported, as the possibility existed that authors might be able to provide unpublished outcome data. However, the outcomes of interest to this review relate to pain severity.

Quality assessment

Study quality was assessed using criteria in the Cochrane risk of bias table,³⁰ and is summarised with other study data in **table 1**.

Analyses

If sufficient studies with similar outcome measures were identified, we intended to conduct an appropriate meta-analysis using Review Manager. If this was not possible, we planned a descriptive overview of studies.

Box 1 Search strategies as applied in MEDLINE

Prediction

- ▶ risk function.mp. or risk assessment
- ▶ risk equation\$.mp.
- ▶ risk chart.mp.
- ▶ (risk adj3 tool\$).mp. [mp=ti, ab, ot, nm, hw, kf, px, rx, ui, sh, tn, dm, mf, dv, kw]
- ▶ risk assessment function.mp.
- ▶ risk assessor.mp.
- ▶ risk appraisal\$.mp.
- ▶ risk calculation\$.mp.
- ▶ risk calculator\$.mp.
- ▶ risk factor\$ calculation\$.mp.
- ▶ risk engine\$.mp.
- ▶ risk equation\$.mp.
- ▶ risk table\$.mp.
- ▶ risk threshold\$.mp.
- ▶ risk scoring method?.mp.
- ▶ scoring scheme?.mp.
- ▶ risk scoring system.mp.
- ▶ risk prediction.mp.
- ▶ predictive instrument.mp.
- ▶ project\$ risk.mp.
- ▶ exp decision support techniques/
Diagnosis, Computer-Assisted/
Decision Support Systems, Clinical/
algorithms/
algorithm?.mp. or Algorithms/
algorith?.mp.
- ▶ decision support?.mp.
- ▶ predictive model.mp.
- ▶ treatment decision.mp.
- ▶ scoring method\$.mp.
- ▶ (prediction\$ adj3 method\$).mp. [mp=ti, ab, ot, nm, hw, kf, px, rx, ui, sh, tn, dm, mf, dv, kw]
- ▶ exp Risk Assessment/
(risk? adj1 assess\$).mp. [mp=ti, ab, ot, nm, hw, kf, px, rx, ui, sh, tn, dm, mf, dv, kw]
- ▶ 1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14 or 15 or 16 or 17 or 18 or 19 or 20 or 21 or 22 or 23 or 24 or 25 or 26 or 27 or 28 or 29 or 30 or 31 or 32 or 33
- ▶ knee.tw.
- ▶ 34 and 35
- ▶ randomized controlled trial.pt.
- ▶ controlled clinical trial.pt.
- ▶ randomized.ab.
- ▶ placebo.ab.
- ▶ randomly.ab.
- ▶ trial.ab.
- ▶ groups.ab.
- ▶ (animals not (humans and animals)).sh.
- ▶ 37 or 38 or 39 or 40 or 41 or 42 or 43
- ▶ 45 not 44
- ▶ 36 and 46

Treatment

- ▶ Pain, Postoperative/
(postoperative adj6 pain*) or (post-operative adj6 pain*) or post-operative-pain*).mp. [mp=title, abstract, original title, name of substance word, subject heading word, keyword heading word, protocol supplementary concept, rare disease supplementary concept, unique identifier]

- ▶ ((post-operative adj6 analgesi*) or (postoperative adj6 analgesi*)).mp. [mp=title, abstract, original title, name of substance word, subject heading word, keyword heading word, protocol supplementary concept, rare disease supplementary concept, unique identifier]
- ▶ ((post-surgical adj6 pain*) or (post surgical adj6 pain*) or (post-surgery adj6 pain*) or (post adj surg* adj pain*)).mp. [mp=title, abstract, original title, name of substance word, subject heading word, keyword heading word, protocol supplementary concept, rare disease supplementary concept, unique identifier]
- ▶ ((post* adj pain*) or pain relief after or pain following surg*).mp. [mp=title, abstract, original title, name of substance word, subject heading word, keyword heading word, protocol supplementary concept, rare disease supplementary concept, unique identifier]
- ▶ ((posttreatment adj6 pain*) or (pain control after adj6 surg*) or ((post-extraction or postextraction or post-surg*) and (pain* or discomfort))).mp. [mp=title, abstract, original title, name of substance word, subject heading word, keyword heading word, protocol supplementary concept, rare disease supplementary concept, unique identifier]
- ▶ randomized controlled trial.pt.
- ▶ controlled clinical trial.pt.
- ▶ randomized.ab.
- ▶ placebo.ab.
- ▶ randomly.ab.
- ▶ trial.ab.
- ▶ 7 or 8 or 9 or 10 or 11 or 12
- ▶ 1 or 2 or 3 or 4 or 5 or 6
- ▶ 13 and 14
- ▶ knee.tw.
- ▶ 15 and 16

RESULTS

Main features of the review process are summarised according to PRISMA guidelines as online supplementary material. As shown in [figure 1](#), searches for evaluations of predictive models and treatments identified 1159 and 1886 articles, respectively.

Application of predictive models

After screening all the titles and abstracts, 16 articles were identified as potentially relevant to the review of predictive models and were evaluated in detail. Reasons for exclusion are summarised with references in the online supplementary material. No studies evaluated the effectiveness of predictive models in guiding treatment and improving long-term outcomes after knee surgery.

Chronic pain management interventions

In the review of treatments for chronic pain after knee surgery, a large majority of studies (66%) reported analgesia or other interventions in the perioperative period. Thirty articles were judged to be potentially relevant. Reasons for exclusion of 29 studies are summarised with references in the online supplementary material.

One intervention fulfilled all inclusion criteria, and study details are summarised in [table 1](#). Singh *et al*³⁴

evaluated the use of a botulinum toxin A injection with antinociceptive and anticholinergic activity in a randomised controlled trial. In the original randomisation, patients with simultaneous bilateral total knee replacement were included, but the published article focused on 49 patients with a unilateral replacement (or first operation in a sequential bilateral replacement). On the basis of criteria specified in the Cochrane risk of bias table, we assessed that this study was of low risk of bias though the small size of the study is a cause for concern. Patients in the trial had received a total knee replacement at least 6 months earlier and had experienced pain in their replaced knee for more than 3 months. Reduced pain intensity was apparent for the intervention compared with placebo after 2 and 3 months, although the authors suggested that meaningful pain relief was evident up to about 40 days with no increase in adverse events. No cost-effectiveness analysis was performed. The authors concluded that the effect of repeated injections should be

assessed in a multicentre trial, but no further study was found on inspection of the Current Controlled Trials database on 5 November 2014.

DISCUSSION

By limiting potential sources of bias, randomised controlled trials provide the best method to assess the effectiveness of healthcare interventions. Systematic reviews aim to appraise evidence from high-quality studies and can have two broad outcomes: a synthesis of knowledge to guide decision-making; or identification of deficits in the evidence base that merit further research.

The main indications for total knee replacement are pain and functional limitations caused by osteoarthritis. The widespread acknowledgement that some people will have chronic postsurgical pain after this potentially curative treatment dates largely from the introduction of patient-reported outcome measures. There is some

Table 1 Characteristics of included study

Author	Singh <i>et al</i> ²⁴
Country	USA
Indication	Total knee replacement >6 months. Chronic pain >3 months (≥ 6 points on 10-point VAS scale). Unsuccessful treatment with oral pain medication, not surgical candidate or infection identified. Mean pain duration 4.5 years
Number of patients	49 patients with 60 total knee replacements (30 intervention: 30 control)
Age	Mean: intervention 67.1 years; control 66.8 years
Sex	Female: intervention 22%; control 12%
Approach	Standardised medial or lateral
Intervention	Intra-articular injection of 100 units botulinum toxin A diluted in 5 mL sterile normal saline
Control	Intra-articular injection of 5 mL sterile normal saline
Follow-up interval	Up to 6 months
Outcome measures	Proportion of responders at 2 months (≥ 2 point VAS reduction) Physicians' global assessment of change Onset and duration of pain (20 point WOMAC pain decrease) WOMAC function Timed-stands test Timed-up-and-go Active knee flexion Medical Outcomes Study Short-Form 36 (SF-36) Short-form McGill Pain Questionnaire Changes in analgesic medications Side effects and adverse outcomes
Economic evaluation	None reported
Risk of bias	Overall: low
Random sequence	Independent
Concealment	Syringes prepared independently
Blinding	Patients, surgeon, investigators, statistician all blind to group allocation
Blind outcome	Assessment blind to group allocation
Complete data	Low losses to follow-up at primary outcome intervals
Selective reporting	Appropriate range of outcomes reported
Other bias	None apparent
Losses to follow-up	2 (1:1) lost to 2 month follow-up. 7 (3:4) lost to 6 month follow-up
Power calculation	Reported to be powered for significant improvement on WOMAC scale
Results summary	Pain severity reduced in 71% of intervention patients compared with 35% in placebo group at 2 months. Benefit also at 3 months but not at 4 months. Duration of meaningful pain relief was 39.6 (SD=50.4) days in intervention group compared with 15.7 (SD=22.6) days in placebo group

VAS, visual analogue scale; WOMAC, Western Ontario and McMaster Universities Osteoarthritis Index.

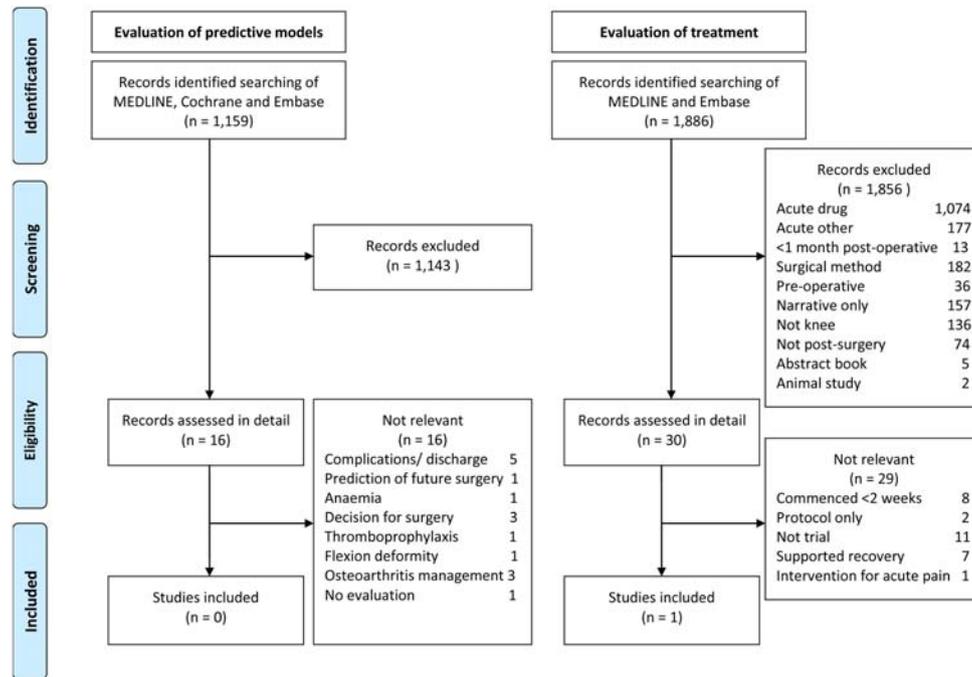


Figure 1 Systematic review flow diagram.

evidence that acute postoperative pain may impact on long-term pain,³⁵ and a considerable number of randomised treatment evaluations have targeted reduction in acute pain with perioperative multimodal anaesthesia.^{12–15} Few studies have followed up patients long term, although the importance of this is now recognised.³⁶ While acknowledging the potential importance of such methods in preventing the development of long-term pain, an appropriate body of research should also explore the issues of prediction and management. The one trial of pain management that we identified showed promise, but further research is needed to confirm the findings. Treatment of chronic pain can be challenging, and there is a need to evaluate multidisciplinary combination treatments and the benefit of matching interventions to patient characteristics.^{28–29}

Our study might be criticised as asking research questions that are too specific and beyond the scope of randomised evaluation. However, evaluation of predictive models in guiding healthcare is recognised in other medical disciplines. For example, risk scoring has been studied in cardiovascular disease in randomised trials both as a guide for appropriate medical treatment of risk factors^{37–38} and lifestyle interventions.³⁹ Without evidence that application of predictive models in total knee replacement is more effective in guiding treatment and improving outcomes than existing care, they have no value in evidence-based clinical practice.

In total knee replacement, specific biological and mechanical issues, and psychological factors relating to joint replacement should be considered in the treatment of chronic pain. The identification of one randomised trial in our review reflects an understanding that approaches to pain management after total knee replacement have

features that differ from chronic pain attributable to other causes. Furthermore, a range of potential interventions with no robust evaluation were identified in our review, specifically neurostimulation,^{40–43} radiofrequency ablation,⁴⁴ denervation,^{45–46} steroid injection⁴⁷ and secondary resurfacing.⁴⁸ This can only be indicative of an awareness of the issue of treatment of pain after knee surgery as the literature searches were not designed to identify studies that did not report robust evaluations. While these relate specifically to orthopaedic surgery and to underlying musculoskeletal conditions, some strategies will be transferable from more general pain management including analgesic medication, and should be considered as potential interventions in patients with long-term pain after total knee replacement.

In summary, our systematic review highlights the lack of evidence about prediction and management of chronic postsurgical pain after total knee replacement. Given the complexity of chronic postsurgical pain and the range of possible treatment options, screening and adequate referral processes are needed, so that patients can receive appropriate interventions that have the potential to improve outcomes and reduce distress. As a large number of people are affected by chronic pain after total knee replacement, the development of an evidence base about care for these patients should be a research priority.

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Contributors ADB, VW and RG-H conceived and designed the study and produced the search strategy. ADB conducted the searches. ADB and VW screened abstracts and titles, assessed study inclusion and exclusion, extracted data and assessed study quality. ADB drafted the manuscript and VW and RG-H revised it critically for important intellectual content. All authors gave approval to the final manuscript.

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REFERENCES

- National Joint Registry for England and Wales: 11th Annual Report. Hemel Hempstead: NJR Centre, 2014.
- Centers for Disease Control and Prevention. *Number of all-listed procedures for discharges from short-stay hospitals, by procedure category and age: United States, 2010*. Atlanta, GA: Centers for Disease Control and Prevention, 2010. www.cdc.gov/nchs/data/nhds/4procedures/2010pro4numberprocedureage.pdf (accessed 29 Apr 2015)
- Weinstein AM, Rome BN, Reichmann WM, *et al*. Estimating the burden of total knee replacement in the United States. *J Bone Joint Surg Am* 2013;95:385–92.
- Macrae WA. Chronic post-surgical pain: 10 years on. *Br J Anaesth* 2008;101:77–86.
- International Association for the Study of Pain. Classification of chronic pain. Descriptions of chronic pain syndromes and definitions of pain terms. Prepared by the International Association for the Study of Pain, Subcommittee on Taxonomy. *Pain* 1986;3:S1–226.
- Werner MU, Kongsgaard UE. Defining persistent post-surgical pain: is an update required? *Br J Anaesth* 2014;113:1–4.
- Bachmeier CJM, March LM, Cross MJ, *et al*. A comparison of outcomes in osteoarthritis patients undergoing total hip and knee replacement surgery. *Osteoarthritis Cart* 2001;9:137–46.
- Beswick AD, Wylde V, Gooberman-Hill R, *et al*. What proportion of patients report long-term pain after total hip or knee replacement for osteoarthritis? A systematic review of prospective studies in unselected patients. *BMJ Open* 2012;2:e000435.
- Jeffery AE, Wylde V, Blom AW, *et al*. "It's there and I'm stuck with it": Patients' experiences of chronic pain following total knee replacement surgery. *Arthritis Care Res* 2011;63:286–92.
- Turkiewicz A, Petersson IF, Björk J, *et al*. Current and future impact of osteoarthritis on health care: a population-based study with projections to year 2032. *Osteoarthritis Cart* 2014;22:1826–32.
- Kurtz S, Ong K, Lau E, *et al*. Projections of primary and revision hip and knee arthroplasty in the United States from 2005 to 2030. *J Bone Joint Surg Am* 2007;89:780–5.
- Chan EY, Fransen M, Parker David A, *et al*. Femoral nerve blocks for acute postoperative pain after knee replacement surgery. *Cochrane Database Syst Rev* 2014;(5):CD009941.
- Lin J, Zhang L, Yang H. Perioperative administration of selective cyclooxygenase-2 inhibitors for postoperative pain management in patients after total knee arthroplasty. *J Arthroplasty* 2013;28:207–13.
- Marques E, Jones H, Elvers K, *et al*. Local anaesthetic infiltration for peri-operative pain control in total hip and knee replacement: systematic review and meta-analyses of short- and long-term effectiveness. *BMC Musculoskelet Disord* 2014;15:220.
- Keijsers R, van Delft R, van den Bekerom MJ, *et al*. Local infiltration analgesia following total knee arthroplasty: effect on post-operative pain and opioid consumption—a meta-analysis. *Knee Surg Sports Traumatol Arthrosc* 2013. doi:10.1007/s00167-013-2788-1 [Epub ahead of print 30 Nov 2013].
- Lingard EA, Katz JN, Wright EA, *et al*. Predicting the outcome of total knee arthroplasty. *J Bone Joint Surg Am* 2004;86:2179–86.
- Papakostidou I, Dailliana ZH, Papapolychiou T, *et al*. Factors affecting the quality of life after total knee arthroplasties: a prospective study. *BMC Musculoskelet Disord* 2012;13:116.
- Perruccio AV, Power JD, Evans HMK, *et al*. Multiple joint involvement in total knee replacement for osteoarthritis: effects on patient-reported outcomes. *Arthritis Care Res* 2012;64:838–46.
- Sullivan M, Tanzer M, Reardon G, *et al*. The role of presurgical expectancies in predicting pain and function one year following total knee arthroplasty. *Pain* 2011;152:2287–93.
- Burns LC, Ritvo SE, Ferguson MK, *et al*. Pain catastrophizing as a risk factor for chronic pain after total knee arthroplasty: a systematic review. *J Pain Res* 2015;8:21–32.
- Vissers MM, Bussmann JB, Verhaar JAN, *et al*. Psychological factors affecting the outcome of total hip and knee arthroplasty: a systematic review. *Semin Arthritis Rheum* 2012;41:576–88.
- Singh JA, Lewallen DG. Medical and psychological comorbidity predicts poor pain outcomes after total knee arthroplasty. *Rheumatology* 2013;52:916–23.
- Brander VA, Stulberg SD, Adams AD, *et al*. Predicting total knee replacement pain: a prospective, observational study. *Clin Orthop Relat Res* 2003;416:27–36.
- Wylde V, Dixon S, Blom AW. The role of preoperative self-efficacy in predicting outcome after total knee replacement. *Musculoskeletal Care* 2012;10:110–18.
- Judge A, Arden NK, Cooper C, *et al*. Predictors of outcomes of total knee replacement surgery. *Rheumatology* 2012;51:1804–13.
- Althaus A, Hinrichs-Rocker A, Chapman R, *et al*. Development of a risk index for the prediction of chronic post-surgical pain. *Eur J Pain* 2012;16:901–10.
- Wyatt JC, Altman DG. Commentary: prognostic models: clinically useful or quickly forgotten? *BMJ* 1995;311:1539–41.
- Turk DC, Wilson HD, Cahana A. Treatment of chronic non-cancer pain. *Lancet* 2011;377:2226–35.
- Merle C, Brendle S, Wang HL, *et al*. Multidisciplinary treatment in patients with persistent pain following total hip and knee arthroplasty. *J Arthroplasty* 2014;29:28–32.
- Higgins JPT, Green S, eds. *Cochrane handbook for systematic reviews of interventions Version 5.1.0 [Updated March 2011]*. The Cochrane Collaboration, 2011. <http://www.cochrane-handbook.org>
- Moher D, Liberati A, Tetzlaff J, *et al*. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *J Clin Epidemiol* 2009;62:1006–12.
- Chaparro Luis E, Smith Shane A, Moore RA, *et al*. Pharmacotherapy for the prevention of chronic pain after surgery in adults. *Cochrane Database Syst Rev* 2013;(7):CD008307. <http://onlinelibrary.wiley.com/doi/10.1002/14651858.CD008307.pub2/abstract>
- Brindle P, Beswick A, Fahey T, *et al*. Accuracy and impact of risk assessment in the primary prevention of cardiovascular disease: a systematic review. *Heart* 2006;92:1752–9.
- Singh JA, Mahowald ML, Noorbaloochi S. Intraarticular botulinum toxin A for refractory painful total knee arthroplasty: a randomized controlled trial. *J Rheumatol* 2010;37:2377–86.
- Kehlet H, Jensen TS, Woolf CJ. Persistent postsurgical pain: risk factors and prevention. *Lancet* 2006;367:1618–25.
- Wylde V, Lenguerrand E, Gooberman-Hill R, *et al*. The effect of local anaesthetic infiltration on chronic post-surgical pain after total hip and knee replacement: the APEX randomised controlled trials. *Pain* 2015. doi:10.1097/j.pain.000000000000114 [Epub ahead of print 5 Feb 2015].
- Beswick A, Brindle P. Risk scoring in the assessment of cardiovascular risk. *Curr Opin Lipidol* 2006;17:375–86.
- Vagholkar S, Zwar N, Jayasinghe UW, *et al*. Influence of cardiovascular absolute risk assessment on prescribing of antihypertensive and lipid-lowering medications: a cluster randomized controlled trial. *Am Heart J* 2014;167:28–35.
- Willis A, Davies M, Yates T, *et al*. Primary prevention of cardiovascular disease using validated risk scores: a systematic review. *J R Soc Med* 2012;105:348–56.
- Billet B, Hanssens K, Nagels W, *et al*. Treatment of post-surgical knee pain with dorsal root ganglion (DRG) neurostimulation (Abstract). *Pain Practice* 2014;14:112.

41. Bouche B, Eisenberg E, Karmakar MK, *et al.* Facilitation of diagnostic and percutaneous trial lead placement with ultrasound guidance for peripheral subcutaneous field stimulation on infrapatellar branches of saphenous nerve (Abstract). *Reg Anesth Pain Med* 2011;2:E158.
42. Lowry AM, Simopoulos TT, Lowry AM, *et al.* Spinal cord stimulation for the treatment of chronic knee pain following total knee replacement. *Pain Physician* 2010;13:251–6.
43. Koulousakis A. Peripheral subcutaneous nerve stimulation (PNS) in neuropathic pain: seven years of experience. *Neuromodulation* 2013;16:e153.
44. Drakes S, Kim SY, Wahezi SE, *et al.* Ultrasound guided radiofrequency ablation of the saphenous nerve for treatment of saphenous neuropathy: a case report (Abstract). *PM&R* 2013;1:S306.
45. Dellon AL, Mont MA, Krackow KA, *et al.* Partial denervation for persistent neuroma pain after total knee arthroplasty. *Clin Orthop Relat Res* 1995;316:145–50.
46. Nahabedian MY, Mont MA, Hungerford DS. Selective denervation of the knee: experience, case reports, and technical notes. *Am J Knee Surg* 1998;11:175–80.
47. Chaiyakit P, Meknavin S, Pakawattana V, *et al.* Results of peri-articular steroid injection in the treatment of chronic extra-articular pain after total knee arthroplasty. *J Med Assoc Thai* 2012;95(Suppl 10): S48–52.
48. Muoneke HE, Khan AM, Giannikas KA, *et al.* Secondary resurfacing of the patella for persistent anterior knee pain after primary knee arthroplasty. *J Bone Joint Surg Br* 2003;85:675–8.

Supplementary material. Excluded studies

Study	Intervention	Reason for exclusion Study description	Benefit for intervention
<i>Evaluation of treatment</i>			
Benazzo et al. 2008 ¹	Pulsed electromagnetic fields	Commenced <2weeks Patients with ACL reconstruction irrespective of pain randomised	Shorter recovery time
Billet et al. 2014 ²	Dorsal root ganglion neurostimulation	Not trial 2 patients with <u>post-surgical knee pain</u> case series	Intervention was effective in controlling of intractable, neuropathic pain
Birch et al. 1993 ³	Sodium diclofenac or physiotherapy or control	Commenced <2weeks Patients with arthroscopy irrespective of pain status randomised	Neither intervention showed benefit for Noyes knee score up to 6 weeks
Bouche et al. 2011 ⁴ Bouche et al. 2011 ⁵	Peripheral subcutaneous field stimulation	Not trial 5 patients with <u>intractable pain</u> after knee surgery	Excellent pain relief before and after permanent stimulator implantation
Chaiyakit et al. 2012 ⁶	Peri-articular steroid injection	Not trial 28 patients (29 knees) with <u>chronic pain</u> at mean 5 months after TKR	VAS pain, knee and function scores improved up to 3 months after injection
Cupal and Brewer 2001 ⁷	Relaxation and guided imagery	Supported recovery Patients with ACL reconstruction	Pain and anxiety reduced at 24 weeks

		irrespective of pain status randomised	
Drakes et al. 2013 ⁸	Ultrasound guided radiofrequency ablation of the saphenous nerve	Not trial 1 patient <u>with pain</u> after multiple knee surgeries	Significant pain relief at 9 months
Eastlack et al. 2005 ⁹	Lower body positive-pressure exercise	Not trial Patients with knee surgery irrespective of pain status	Significant pain relief
Giaquinto et al. 2010 ¹⁰	Hydrotherapy compared with gym based rehabilitation	Supported recovery Patients with TKR irrespective of pain status randomised	WOMAC pain, function and stiffness improved
Gooch et al. 2012 ¹¹	Clinical pathway	Supported recovery Patients with TKR irrespective of pain status randomised	Improved WOMAC score
Hall et al. 2012 ¹²	Neuromuscular exercise	Protocol Patients with arthroscopic partial medial meniscectomy irrespective of pain status randomised	Protocol only
Koulousakis 2013 ¹³ Abstract only	Peripheral subcutaneous nerve stimulation	Not trial 405 patients with neuropathic pain including some with <u>post-operative knee pain</u> not randomised	70% showed pain relief up to 3 years
Kramer et al. 2003 ¹⁴	Clinic or home-based rehabilitation	Supported recovery Patients with TKR	Similar outcomes

		irrespective of pain status randomised	
Lee Dellon et al. 1996 ¹⁵	Partial denervation	Not trial 70 patients with <u>persistent pain</u> after TKR, trauma or osteotomy not randomised	Improvement in VAS pain and knee score at around 2 years
Liang et al. 2006 ¹⁶	Traditional Chinese medicine and rehabilitation	<2 weeks Patients with arthroscopic ACL reconstruction irrespective of pain status randomised	Improved pain, swelling and function
Losina et al. 2013 ¹⁷	Motivational- interviewing-based telephone intervention	Protocol Supported recovery Patients with TKR irrespective of pain status randomised	Protocol only
Lowry et al. 2010 ¹⁸	Spinal cord stimulation	Not trial 1 patient with <u>persistent knee pain</u> after TKR	Improvement to pain and function
Madsen et al. 2013 ¹⁹	Group-based rehabilitation or individual supervised home-training	Supported recovery Patients with TKR irrespective of pain status randomised	No difference between groups at 6 months
Mast et al. 1995 ²⁰	Acupuncture	<2 weeks Patients with TKR irrespective of pain status randomised	Pain reduction at 24 hours

Moretti et al. 2012 ²¹	Pulsed electromagnetic fields	<2 weeks Patients with TKR irrespective of pain status randomised	VAS pain, knee score and SF-36 improved
Muoneke et al. 2003 ²²	Secondary patella resurfacing	Not trial 20 patients with <u>chronic anterior knee pain</u> after TKR without patellar resurfacing not randomised	Knee score and function improved but may increase dissatisfaction and need for revision
Nahabedian et al. 1998 ²³	Selective denervation	Not trial 13 patients with <u>chronic knee pain</u> secondary to neuromata from prior surgery or trauma	10/13 rated pain outcome as excellent or good
Naylor et al. 2012 ²⁴	Group-based or monitored home-based rehabilitation	Supported recovery Patients with TKR irrespective of pain status randomised	No difference between groups at 8 weeks
Pellino et al. 2005 ²⁵	Kit of nonpharmacologic strategies for pain and anxiety	<2 weeks Patients with TKR irrespective of pain status randomised	No difference in pain but lower opioid use and anxiety on postoperative day 1
Richards et al. 2013 ²⁶	Morphine plus oxycodone or oxycodone and acetaminophen	Acute pain Patients with <u>moderate to severe pain</u> after TKR randomised	For BPI pain outcomes, flexible dose morphine/oxycodone was superior to low-dose morphine/oxycodone and comparable to oxycodone/

			acetaminophen
Schroer et al. 2011 ²⁷	Prolonged postoperative cyclooxygenase-2 inhibitor	<2 weeks Patients with TKR irrespective of pain status randomised at discharge	Better VAS pain, flexion, knee scores, OKS, SF-12 physical function after 6 weeks treatment
van den Akker-Scheek et al. 2007 ²⁸	Theory-driven support programme	Supported recovery Patients with TKR irrespective of pain status randomised	No difference between groups at 26 weeks
Zorzi et al. 2007 ²⁹	Pulsed electromagnetic fields	<2 weeks Patients receiving arthroscopic treatment of knee cartilage irrespective of pain status randomised	Improved KOOS at 90 days, reduced NSAID use. Better 3 year recovery
<i>Evaluation of predictive model</i>			
Ackland et al. 2010 ³⁰	Cardiac risk index	Prediction of complications Patients with orthopaedic procedure. No evaluation	Risk index predicted postoperative morbidity and hospital stay
Adam et al. 2008 ³¹	Patient decision aids or shared decision making	Decision for surgery Joint replacement. Systematic review	Little evidence on the use of decision aids
Al-Arabi et al. 2009 ³²	Risk classification	Prediction of complications TKR. No evaluation	Risk classification identified patients at risk of complications and longer inpatient stay

De Achaval et al. 2011 ³³	Patient decision aid and computer-based values clarification exercise	Decision for surgery Elective TKR. No evaluation	Decision aid reduced decisional conflict but no benefit for additional computer-based tool
Enko et al. 2013 ³⁴	Thomas-plot therapeutic algorithm	Prediction of anaemia Elective major orthopaedic surgery. No evaluation	Algorithm predicted need for blood
Fithian et al. 2005 ³⁵	Treatment algorithm	Prediction of future surgery ACL reconstruction. No evaluation	Treatment algorithm effective in predicting risk of further knee surgery
Hunter 2009 ³⁶	Stratification of risk for osteoarthritis progression	Osteoarthritis management Knee osteoarthritis. Narrative review	Risk stratification feasible using a number of different metrics
Jain et al. 2011 ³⁷	Risk profile for post-surgical delirium	Prediction of complications Joint replacement. No evaluation	Risk profile was associated with risk of postoperative delirium
Jain et al. 2013 ³⁸	Flexion deformity algorithm	Prediction of flexion deformity correction TKR. No evaluation	Flexion deformity algorithm predicted postoperative flexion deformity
Lewis et al. 2013 ³⁹	Risk algorithm for prediction of need for knee replacement	Decision for surgery Knee replacement. No evaluation	Algorithm predicted need for knee replacement
Oldmeadow et al. 2004 ⁴⁰	Pre-operative risk assessment	Prediction of complications Elective hip or knee replacement. Before	Use of risk assessment with targeted postoperative care improved time and

		and after study	destination of hospital discharge
Romine et al. 2013 ⁴¹	Peri-operative risk calculator	Prediction of complications TKR. No evaluation	Higher predicted risk associated with more complications
Sanchez-Santos et al. 2014 ⁴²	Clinical risk prediction tool to predict OKS patient reported outcome	Predictive model with no evaluation Primary TKR. No evaluation	Good discrimination and calibration
Saragas et al. 2014 ⁴³	Risk factor score	Decision for thromboprophylaxis Foot and ankle surgery. No evaluation	Specific non-weight-bearing and cast immobilised group require thromboprophylaxis
Simoes et al. 2014 ⁴⁴	Algorithm to select patients for rheumatologist evaluation	Osteoarthritis management Knee osteoarthritis. Not evaluation	Algorithm did not predict under-diagnosis
Van Middelkoop et al. 2014 ⁴⁵	Algorithm to predict success of intra-articular glucocorticoid treatment	Osteoarthritis management. Knee or hip osteoarthritis. No evaluation	Pain and inflammation predicted response to glucocorticoids

ACL anterior cruciate ligament; TKR total knee replacement; VAS visual analogue scale; WOMAC Western Ontario and McMaster Universities Osteoarthritis Index; BPI Brief Pain Inventory; OKS Oxford knee score; KOOS Knee injury and Osteoarthritis Outcome Score; NSAID non-steroidal anti-inflammatory drug

Supplementary material References

1. Benazzo F, Zanon G, Pederzini L et al. Effects of biophysical stimulation in patients undergoing arthroscopic reconstruction of anterior cruciate ligament: prospective, randomized and double blind study. *Knee Surgery, Sports Traumatology, Arthroscopy* 2008;**16**:595-601.
2. Billet B, Hanssens K, Nagels W et al. Treatment of post-surgical knee pain with dorsal root ganglion (DRG) neurostimulation. *Pain Practice* 2014;**14**:112.
3. Birch NC, Sly C, Brooks S et al. Anti-inflammatory drug therapy after arthroscopy of the knee. A prospective, randomised controlled trial of diclofenac or physiotherapy. *Journal of Bone and Joint Surgery - Series B* 1993;**75**:650-652.
4. Bouche B, Eisenberg E, Karmakar MK et al. Facilitation of diagnostic and percutaneous trial lead placement with ultrasound guidance for peripheral subcutaneous field stimulation on infrapatellar branches of saphenous nerve. *Regional Anesthesia and Pain Medicine* 2011;**2**:E158.
5. Bouche B, Eryk E, Michel M et al. Facilitation of diagnostic and percutaneous trial lead placement with ultrasound guidance for peripheral subcutaneous field stimulation on infrapatellar branches of saphenous nerve. *Neuromodulation* 2011;**14 (6)**:564.
6. Chaiyakit P, Meknavin S, Pakawattana V et al. Results of peri-articular steroid injection in the treatment of chronic extra-articular pain after total knee arthroplasty. *Journal of the Medical Association of Thailand* 2012;**95 Suppl 10**:S48-52.
7. Cupal DD, Brewer BW. Effects of relaxation and guided imagery on knee strength, reinjury anxiety, and pain following anterior cruciate ligament reconstruction. *Rehabilitation Psychology* 2001;**46**:28-43.
8. Drakes S, Kim SY, Wahezi SE et al. Ultrasound guided radiofrequency ablation of the saphenous nerve for treatment of saphenous neuropathy: A case report. *PM and R* 2013;**1**:S306.
9. Eastlack RK, Hargens AR, Groppo ER et al. Lower body positive-pressure exercise after knee surgery. *Clinical Orthopaedics & Related Research* 2005;213-219.
10. Giaquinto S, Ciotola E, Dall'Armi V et al. Hydrotherapy after total knee arthroplasty. A follow-up study. *Archives of Gerontology & Geriatrics* 2010;**51**:59-63.
11. Gooch K, Marshall DA, Faris PD et al. Comparative effectiveness of alternative clinical pathways for primary hip and knee joint replacement patients: a pragmatic randomized, controlled trial. *Osteoarthritis & Cartilage* 2012;**20**:1086-1094.

12. Hall M, Hinman RS, Wrigley TV et al. The effects of neuromuscular exercise on medial knee joint load post-arthroscopic partial medial meniscectomy: 'SCOPEX', a randomised control trial protocol. *BMC Musculoskeletal Disorders* 2012;**13**:233.
13. Koulousakis A. Peripheral subcutaneous nerve stimulation (PNS) in neuropathic pain: Seven years of experience. *Neuromodulation* 2013;**16** (5):e153.
14. Kramer JF, Speechley M, Bourne R et al. Comparison of clinic- and home-based rehabilitation programs after total knee arthroplasty. *Clinical Orthopaedics and Related Research* 2003;225-234.
15. Lee Dellon A, Mont MA, Mullick T et al. Partial denervation for persistent neuroma pain around the knee. *Clinical Orthopaedics and Related Research* 1996;216-222.
16. Liang Y, Zhang SM, Hu Y et al. Effects of traditional Chinese medicine and rehabilitation training on knee joint function after anterior cruciate ligament reconstruction in arthroscopy. [Chinese]. *Chinese Journal of Clinical Rehabilitation* 2006;**10**:6-10.
17. Losina E, Collins JE, Daigle ME et al. The AViKA (Adding Value in Knee Arthroplasty) postoperative care navigation trial: rationale and design features. *BMC Musculoskeletal Disorders* 2013;**14**:290.
18. Lowry AM, Simopoulos TT, Lowry AM et al. Spinal cord stimulation for the treatment of chronic knee pain following total knee replacement. *Pain Physician* 2010;**13**:251-256.
19. Madsen M, Larsen K, Madsen IK et al. Late group-based rehabilitation has no advantages compared with supervised home-exercises after total knee arthroplasty. *Danish Medical Journal* 2013;**60**:A4607.
20. Mast R, Schoch T, Scharf HP: [Acupuncture against postoperative pain after total knee replacement - a placebo-controlled trial on immediate effects](in German). In: *Akt Rheumatol.* vol. 20; 1995: 131-134.
21. Moretti B, Notarnicola A, Moretti L et al. I-ONE therapy in patients undergoing total knee arthroplasty: A prospective, randomized and controlled study. *BMC Musculoskeletal Disorders* 2012;**13**:
22. Muoneke HE, Khan AM, Giannikas KA et al. Secondary resurfacing of the patella for persistent anterior knee pain after primary knee arthroplasty. *Journal of Bone & Joint Surgery - British Volume* 2003;**85**:675-678.
23. Nahabedian MY, Mont MA, Hungerford DS. Selective denervation of the knee: experience, case reports, and technical notes. *The American journal of knee surgery* 1998;**11**:175-180.

24. Naylor JM, Ko V, Naylor JM et al. Heart rate response and factors affecting exercise performance during home- or class-based rehabilitation for knee replacement recipients: lessons for clinical practice. *Journal of Evaluation in Clinical Practice* 2012;**18**:449-458.
25. Pellino TA, Gordon DB, Engelke ZK et al: Use of nonpharmacologic interventions for pain and anxiety after total hip and total knee arthroplasty. In: *Orthopaedic nursing*. vol. 24; 2005: 182-190; quiz 191-182.
26. Richards P, Gimbel JS, Minkowitz HS et al. Comparison of the efficacy and safety of dual-opioid treatment with morphine plus oxycodone versus oxycodone/acetaminophen for moderate to severe acute pain after total knee arthroplasty. *Clinical Therapeutics* 2013;**35**:498-511.
27. Schroer WC, Diesfeld PJ, LeMarr AR et al. Benefits of prolonged postoperative cyclooxygenase-2 inhibitor administration on total knee arthroplasty recovery: A double-blind, placebo-controlled study. *Journal of Arthroplasty* 2011;**26**:2-7.
28. van den Akker-Scheek I, Zijlstra W, Groothoff JW et al. Groningen orthopaedic exit strategy: Validation of a support program after total hip or knee arthroplasty. *Patient Education & Counseling* 2007;**65**:171-179.
29. Zorzi C, Dall'Oca C, Cadossi R et al. Effects of pulsed electromagnetic fields on patients' recovery after arthroscopic surgery: prospective, randomized and double-blind study. *Knee Surgery, Sports Traumatology, Arthroscopy* 2007;**15**:830-834.
30. Ackland GL, Harris S, Ziabari Y et al. Revised cardiac risk index and postoperative morbidity after elective orthopaedic surgery: A prospective cohort study. *British Journal of Anaesthesia* 2010;**105**:744-752.
31. Adam JA, Khaw FM, Thomson RG et al. Patient decision aids in joint replacement surgery: a literature review and an opinion survey of consultant orthopaedic surgeons. *Annals of the Royal College of Surgeons of England* 2008;**90**:198-207.
32. Al-Arabi YB, Al-Arabi YB. Risk classification for primary knee arthroplasty. *Journal of Arthroplasty* 2009;**24**:90-95.
33. De Achaval S, Fraenkel L, Volk R et al. Impact of a patient decision aid with an interactive values component on decisional conflict associated with total knee arthroplasty. *Arthritis and Rheumatism* 2011;**1**):
34. Enko D, Wallner F, Von Goedecke A et al. Algorithm-guided preoperative anemia management using Thomas-plot as diagnostic tool in patients undergoing elective major orthopedic surgery. *Transfusion Medicine and Hemotherapy* 2013;**40**:59.

35. Fithian DC, Paxton EW, Stone ML et al. Prospective trial of a treatment algorithm for the management of the anterior cruciate ligament-injured knee. *American Journal of Sports Medicine* 2005;**33**:335-346.
36. Hunter DJ. Risk stratification for knee osteoarthritis progression: a narrative review. *Osteoarthritis and Cartilage* 2009;**17**:1402-1407.
37. Jain FA, Brooks JO, Larsen KA et al. Individual Risk Profiles for Postoperative Delirium after Joint Replacement Surgery. *Psychosomatics* 2011;**52**:410-416.
38. Jain JK, Sharma RK, Agarwal S. Total knee arthroplasty in patients with fixed flexion deformity: Treatment protocol and outcome. *Current Orthopaedic Practice* 2013;**24**:659-664.
39. Lewis JR, Dhaliwal SS, Zhu K et al. A predictive model for knee joint replacement in older women. *PLoS ONE* 2013;**8**:
40. Oldmeadow LB, McBurney H, Robertson VJ et al. Targeted postoperative care improves discharge outcome after hip or knee arthroplasty. *Archives of Physical Medicine & Rehabilitation* 2004;**85**:1424-1427.
41. Romine LB, May RG, Taylor HD et al. Accuracy and Clinical Utility of a Peri-Operative Risk Calculator for Total Knee Arthroplasty. *Journal of Arthroplasty* 2013;**28**:445-448.
42. Sanchez-Santos MT, Judge A, Batra RN et al. A clinical tool for the prediction of patient-reported outcomes after knee replacement surgery. *Osteoarthritis and Cartilage* 2014;**22**:S412.
43. Saragas NP, Ferrao PNF, Saragas E et al. The impact of risk assessment on the implementation of venous thromboembolism prophylaxis in foot and ankle surgery. *Foot and Ankle Surgery* 2014;**20**:85-89.
44. Simoes D, Andre R, Carmona L et al. Knee osteoarthritis diagnostic gap: Moving beyond classical risk factors. *Osteoporosis International* 2014;**25**:S398.
45. Van Middelkoop M, Arden N, Atchia I et al. The OA trial bank: Meta-analysis of individual patient data show that patients with severe pain or with inflammatory signs detected by ultrasound especially benefit from intra-articular glucocorticoids for knee or hip OA. *Osteoarthritis and Cartilage* 2014;**22**:S474-S475.