

Atraumatic knee pain

REVIEW ARTICLE

ROBIN HOLTEDAHL

E-mail: robi-hol@online.no

Husebybakken

Robin Holtedahl (born 1948), specialist in physical medicine and rehabilitation in private practice, medical advisor to the life insurance company Oslo Pensjonsforsikring (OPF) and the Norwegian Labour and Welfare Administration (NAV), with part-time positions in Unicare Fram, Rykkinn, and the Skogli Rehabilitation Centre, Lillehammer. He is currently participating in a working group under the auspices of South-Eastern Norway Health Authority on outpatient arthroscopic surgery. The author has completed the ICMJE form and declares no conflicts of interest.

BACKGROUND

Knee pain is among the most common reasons for consultations for musculoskeletal pain. The objective of this article is partly to provide an overview of the research undertaken in the area of atraumatic and degenerative knee pain, and partly to assess whether common practices for examination and treatment are consistent with this research.

METHOD

This article is based on a selection of English-language studies of atraumatic and degenerative knee pain found through a search in Medline, manual searches in the lists of references in the articles, and literature suggested by Mendeley. Studies of gonarthrosis, degenerative meniscus injuries and the psychosocial context of knee pain were given preference.

RESULTS

There is frequently little consistency between findings made by diagnostic imaging and the clinical picture, and an extra-articular origin may easily be overlooked. As a rule, symptomatic gonarthrosis may be diagnosed clinically. If there is a need for diagnostic imaging, standing x-ray images are the first

choice. MRI scans are often used uncritically in cases of suspected arthrosis or meniscus injury, but this examination rarely produces clinically useful information. Activity-based forms of treatment yield results that are at least as good as arthroscopic surgery for degenerative knee pain and ought to be the first choice. Atraumatic knee pain is frequently associated with psychosocial problems, in particular in cases for which the clinical and radiological findings are modest.

INTERPRETATION

Many patients with knee pain undergo examinations and therapies that are inconsistent with recent research results. Both the patient and society will be better served by a more evidence-based approach.

Of all musculoskeletal pain, knee pain is among the most frequently occurring. The majority of the patients have an atraumatic knee disorder (1). Pain receptors cannot be detected in joint cartilage, but are abundantly present in other structures, including joint ligaments, the synovium, the subchondral bone and surrounding soft tissue. Experience shows, however, that there will often be doubt regarding the origin of pain in atraumatic knee disorders, even after a thorough examination.

Symptoms that often accompany knee pain include stiffness, knee locking, muscle weakness and instability. In an interview survey of persons who were older than 20 years and resident in Nord-Trøndelag county, one in every seven women and one in every ten men reported to have suffered from knee pain of at least three months' duration in the course of one year (2). There are grounds to assume that knee pain is reported more frequently than before, but this tendency applies to musculoskeletal disorders in general (3, 4).

Radiologically verified gonarthrosis increases in frequency with advancing age (5). Some studies support the notion that the frequency of knee pain also increases somewhat with age, but that the curve levels off around the age of 50 (6–8). Although it is commonly assumed that symptoms of gonarthrosis worsen over time, a prospective study of 1 753 patients with moderate gonarthrosis found that their knee pain remained unchanged over a six-year period, in spite of radiological progression (9).

As for other musculoskeletal pain, patients with knee disorders will as a rule first turn to a general practitioner (6, 7). Commonly, after undertaking a clinical examination, the GP will refer the patient for an MRI scan of the knees and/or to a physiotherapist. If the MRI scan indicates that the condition is operable, the patient will be referred to an orthopaedist and in many cases undergo arthroscopy.

Recent research has cast doubt on the justification for a narrowly biomechanical model of understanding for assessment of patients who suffer from knee disorders that have not been caused by an obvious trauma. It is claimed that current practice partly leads to overdiagnosis as a result of erroneous or over-interpretation of normal variants, and partly to treatment that is of doubtful benefit, which drives up the cost level and involves a risk of

iatrogenic injury (10, 11). Figure 1 provides a schematic overview of precipitating and perpetuating factors in atraumatic knee pain with or without degenerative changes.

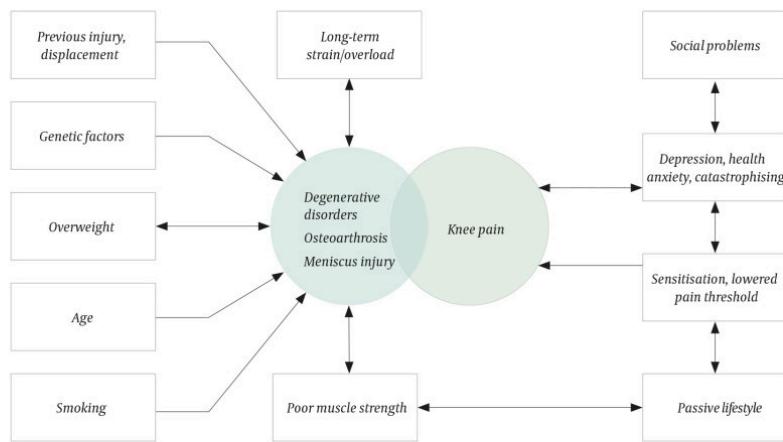


Figure 1 Schematic overview of precipitating and perpetuating factors in atraumatic knee pain with and without degenerative changes. Factors associated with degenerative changes are grouped to the left, factors associated with knee pain without such changes are grouped to the right. The arrows indicate the predominant causal directions. While degenerative knee disorders are mainly associated with biological factors, cognitive, mental and social factors are frequently more relevant in cases of knee pain with no detectable arthropathy.

The objective of this article is partly to provide an overview of recent research on examination and treatment of atraumatic knee pain, and partly to elucidate the degree to which such research results have influenced the health services' approach to this large group of patients.

Method

This review is based on a selection of articles found through literature searches in Medline with the search terms 'knee AND (osteoarthritis* OR meniscus*)' in titles and summaries, in combination with 'degenerative, psychol*', cognitive, radiogr*, MRI, arthros*, diagnos*. Additional articles were identified through suggestions from the Mendeley reference programme and manual searches in lists of references.

The search was restricted to English-language articles on knee pain unrelated to trauma or septic or inflammatory arthritis, published over the last 20 years. It was concluded in September 2017. Altogether 889 articles were identified, of which 141 were reviews. The relevance of each article was determined by discretionary judgment on the basis of the title and summary. Of these 889 articles, 111 were retrieved in full-text versions. A total of 48 of these were included, of which 18 were reviews.

Knee pain and cognitive, mental and social factors

While a biopsychosocial model of understanding has gradually become accepted with regard to musculoskeletal disorders, knee disorders tend to be regarded as an expression of purely biomechanical dysfunction, often as a result of long-term or unaccustomed strain (5).

Patients with knee symptoms are a heterogeneous group, but numerous studies have shown significant mental comorbidity in chronic knee pain. Somewhat paradoxically, the incidence of anxiety, depression, difficult social conditions, poor coping, somatisation and catastrophising is highest in the group with sparse clinical and radiological findings (1), (12–15).

A study of functional impairment in patients with knee pain and radiologically verified osteoarthritis found no association with the x-ray findings (osteophytes and reduced cartilage height), while high body-mass index, perceived helplessness, pain intensity and anxiety explained approximately 60 % of the variance (16).

One study compared a group that had significant gonarthrosis (Kellgren-Lawrence degree 3–4), but little pain, with a group with moderate gonarthrosis (KL degree 1–2), but strong pain. The latter group had increased sensitivity to sensory tests, interpreted as an expression of sensitisation (17).

After knee prosthesis surgery, a considerable minority reports persistent pain one year or more after the operation. This group is characterised by a higher pre-operative incidence of mental problems and catastrophising (18, 19). Such studies underscore that the causal relationships in chronic knee pain can be complex and that the treatment needs to take this into account (Figure 1).

Knee pain and radiological findings

In cases of typical symptoms of gonarthrosis (weight-bearing pain, transient morning stiffness), clinical findings (reduced mobility, crepitus, joint line tenderness) and risk factors (overweight, female sex and age > 40 years), a diagnosis of osteoarthritis can often be made without any supplementary diagnostic imaging (20).

If such an examination nevertheless is undertaken, standing x-ray images should be made of both knees in three projections: lateral images, frontal images in semiflexion and skyline images of the patellofemoral joint (20). Classical x-ray findings in knees in cases of osteoarthritis include pronounced medial joint space narrowing, osteophytes, subchondral sclerosis and subchondral cysts (21).

Lack of correspondence between radiological findings and knee pain has been frequently described. In a systematic review, the prevalence of radiologically verified gonarthrosis in patients with knee pain varied from 15 % to 76 % in

different studies, while the proportion with knee pain among those who were diagnosed with arthrosis varied from 15 % to 85 % (22).

The association between radiological findings and knee function is also poorly documented (16). An emphasis on radiological findings at the expense of a clinical examination may cause periarticular disorders such as bursitis, tendinosis, regional myofascial pain conditions, pain transferred from proximal structures and neurogenic hyperalgesia to be overlooked (23).

Magnetic resonance imaging in cases of knee pain

In a study of participants in the age group older than 50 years and with no osteoarthritis detected by radiology, an MRI scan showed at least one pathological finding in up to 97 % of those with knee pain and up to 88 % of those with no pain (24). In a study with randomly selected participants in the same age group, a meniscus injury was detected by an MRI scan in 35 % and in more than half of those older than 70 years (25). In the group with moderate to severe osteoarthritis shown radiologically (KL degree 2–4), a meniscus injury was found in 82 %. However, only 39 % of those with a meniscus injury reported any knee symptoms.

The high prevalence and strong association between meniscus injuries and osteoarthritis mean that an MRI examination rarely entails any diagnostic or therapeutic consequences in the age group older than 40 years, unless there are clinical indications of a more severe underlying disorder (26, 27).

In 2002, altogether 95 MRI scans of knees were undertaken per 1 000 inhabitants, increasing to 173 per 1 000 in 2013 (28, 29). The increase was strongest in the age group 50–59 years. Knee pain is currently the third most frequent indication for an MRI scan in Norway, after pain of the head and lumbar region. General practitioners account for 90 % of the referrals for MRI scans to private radiology institutes (29).

A review undertaken by the Office of the Auditor General concluded that the extensive use of MRI scans for knee pain lacked any documented beneficial effects, and there were also inexplicably large variations between the counties (29). The majority of Norwegian radiologists also agree that there is an overconsumption of MRI scans (29, 30). It is reasonable to assume that this also applies to MRI scans for the assessment of knee pain.

Poorly justified MRI scans are costly, may give rise to expectations of a referral to an orthopaedist and contribute to medicalisation (31). As a rule, an adequate clinical examination, supplemented by standing x-ray images will be sufficient to reveal the causal relationships, for the patient as well as the doctor.

Symptoms and findings for meniscus injuries

Mechanical symptoms such as popping, knee locking and catching are traditionally regarded as classical symptoms of meniscus injury with an indication for meniscus surgery. Recent studies cast doubts on this interpretation. In a randomised, placebo-controlled study of patients with knee pain that was degenerative in origin, the prevalence of mechanical symptoms was 49 % after arthroscopic meniscus resection and 43 % in the placebo group (32). In another study, it turned out that pre-operatively self-reported pain and functional impairment were not associated with structural knee pathology, including degenerative meniscus injury, detected by subsequent arthroscopy (33).

A positive McMurray test, a test in which the tibia is rotated with a simultaneous extension of the knee joint, is often regarded as one of the most reliable clinical signs of a meniscus injury. In controlled studies, on the other hand, inter-rater concordance tends to be low for this test (34). Nor have more recent clinical tests been shown to have a specificity that can reliably distinguish between meniscus injuries and other causes of knee pain (35, 36).

It is thus doubtful whether meniscus injuries can be distinguished from other pathology through detection of specific symptoms or clinical findings in patients with degenerative knee disorders.

Arthroscopy for atraumatic and degenerative knee disorders

In the last couple of decades, knee arthroscopy has become the most common orthopaedic procedure in countries for which there are available data, and this increase has been especially pronounced for middle-aged and elderly patients, among whom degenerative knee pain predominates (37). After two randomised studies showed that arthroscopic lavage and debridement of cartilage and meniscus tissue had no discernible effect on gonarthrosis, some countries have reported a reduction in the use of this procedure.

On the other hand, some increased use of arthroscopic meniscus surgery has been described during the same period (37). Since 2002, a number of randomised, controlled studies of this procedure have been conducted, in Norway as well as in other countries. The majority of the studies have been unable to show any better effect of arthroscopy than of physical exercise under the auspices of a physiotherapist, or placebo surgery (38, 39).

Partial meniscectomy increases the risk of developing radiologic gonarthrosis after one year (40) as well as later in the course of illness (41), although the clinical consequences of this are somewhat uncertain. The risk of pre- or post-operative complications in knee arthroscopy is low, but not negligible (42). The time has come to abandon the myth that arthroscopic surgery is a relevant intervention for the great majority of patients who suffer from atraumatic and degenerative knee disorders (11, 43, 44). We should expect that within a short time, knee arthroscopy will be restricted to indications where it will provide a proven benefit, primarily traumatic meniscus injuries in people younger than 35 years (10).

Conclusion

The 'standard model' applied in examination and treatment of atraumatic knee pain is out of step with new knowledge. Measures to reduce the use of MRI scans in atraumatic knee pain and a further reduction in the use of arthroscopy ought to be given especially high priority in health policy.

Although a conventional pathoanatomical model will not necessarily be irrelevant, the causal relationships in chronic knee pain are often multifactorial. Especially in cases where the symptoms and functional impairment cannot reasonably be related to findings made clinically and by diagnostic imaging, attention should be directed towards psychosocial components.

As a clinician, it is easy to assume a model of 'mechanical failure' and forget the wider context of which the knee symptoms are part (45). By disregarding such factors and continuing with our technically sophisticated, biomedically based diagnostics and treatment, we as healthcare workers risk imposing a further burden on the patient.

Patients may have unrealistic expectations regarding the benefits of diagnostic imaging and orthopaedic procedures (46). Communicating that a thorough clinical examination is often sufficient in cases of atraumatic knee pain is an important, but frequently challenging and time-consuming educational task. The patient should also be informed that 'low-tech' treatment principles – lifestyle change, active coping and individualised exercise therapy – have a documented effect on knee pain and functioning (47, 48). Such measures ought to constitute the mainstay of the approach taken by the health services to patients with atraumatic knee disorders.

I wish to thank May Arna Risberg, Jens Ivar Brox and Knut Arne Holtedahl for their useful comments.

Main message

Atraumatic knee pain is only to a moderate degree associated with findings made by diagnostic imaging, and too many MRI scans are performed

Degenerative meniscus injuries are not normally associated with specific symptoms or findings

Knee pain may have causes other than osteoarthritis or meniscus injury, including soft-tissue pain and pain secondary to underlying psychosocial stress

Self-care, activity-based interventions and weight loss, if appropriate, are the first choice. Arthroscopy is rarely indicated

LITERATURE

1. Urwin M, Symmons D, Allison T et al. Estimating the burden of musculoskeletal disorders in the community: the comparative prevalence of symptoms at different anatomical sites, and the relation to social deprivation. *Ann Rheum Dis* 1998; 57: 649 - 55. [PubMed][CrossRef]
2. Svebak S, Hagen K, Zwart JA. One-Year Prevalence of Chronic Musculoskeletal Pain in a Large Adult Norwegian County Population: Relations with Age and Gender – The HUNT Study. *J Musculoskeletal Pain* 2006; 14: 21 - 8. [CrossRef]
3. Nguyen U-SDT, Zhang Y, Zhu Y et al. Increasing prevalence of knee pain and symptomatic knee osteoarthritis: survey and cohort data. *Ann Intern Med* 2011; 155: 725 - 32. [PubMed][CrossRef]
4. Hagen K, Linde M, Heuch I et al. Increasing prevalence of chronic musculoskeletal complaints. A large 11-year follow-up in the general population (HUNT 2 and 3). *Pain Med* 2011; 12: 1657 - 66. [PubMed] [CrossRef]
5. Arden N, Nevitt MC. Osteoarthritis: epidemiology. *Best Pract Res Clin Rheumatol* 2006; 20: 3 - 25. [PubMed][CrossRef]
6. Jinks C, Jordan K, Ong BN et al. A brief screening tool for knee pain in primary care (KNEST). 2. Results from a survey in the general population aged 50 and over. *Rheumatology (Oxford)* 2004; 43: 55 - 61. [PubMed] [CrossRef]
7. Turkiewicz A, Gerhardsson de Verdier M, Engström G et al. Prevalence of knee pain and knee OA in southern Sweden and the proportion that seeks medical care. *Rheumatology (Oxford)* 2015; 54: 827 - 35. [PubMed] [CrossRef]
8. Hagen K, Svebak S, Zwart J-A. Incidence of musculoskeletal complaints in a large adult Norwegian county population. The HUNT Study. *Spine* 2006; 31: 2146 - 50. [PubMed][CrossRef]
9. Collins JE, Katz JN, Dervan EE et al. Trajectories and risk profiles of pain in persons with radiographic, symptomatic knee osteoarthritis: data from the osteoarthritis initiative. *Osteoarthritis Cartilage* 2014; 22: 622 - 30. [PubMed][CrossRef]
10. Siemieniuk RAC, Harris IA, Agoritsas T et al. Arthroscopic surgery for degenerative knee arthritis and meniscal tears: a clinical practice guideline. *BMJ* 2017; 357: j1982. [PubMed][CrossRef]
11. Randsborg PH, Røtterud JH. Degenerative meniskrupturer bør ikke opereres. *Tidsskr Nor Legeforen* 2017; 137: 258. [PubMed][CrossRef]
12. Creamer P, Lethbridge-Cejku M, Costa P et al. The relationship of anxiety and depression with self-reported knee pain in the community: data from the

Baltimore Longitudinal Study of Aging. *Arthritis Care Res* 1999; 12: 3 - 7. [PubMed][CrossRef]

13. Urquhart DM, Phyomaung PP, Dubowitz J et al. Are cognitive and behavioural factors associated with knee pain? A systematic review. *Semin Arthritis Rheum* 2015; 44: 445 - 55. [PubMed][CrossRef]
14. Kim KW, Han JW, Cho HJ et al. Association between comorbid depression and osteoarthritis symptom severity in patients with knee osteoarthritis. *J Bone Joint Surg Am* 2011; 93: 556 - 63. [PubMed][CrossRef]
15. Williams DA, Farrell MJ, Cunningham J et al. Knee pain and radiographic osteoarthritis interact in the prediction of levels of self-reported disability. *Arthritis Rheum* 2004; 51: 558 - 61. [PubMed][CrossRef]
16. Creamer P, Lethbridge-Cejku M, Hochberg MC. Factors associated with functional impairment in symptomatic knee osteoarthritis. *Rheumatology (Oxford)* 2000; 39: 490 - 6. [PubMed][CrossRef]
17. Finan PH, Buenaver LF, Bounds SC et al. Discordance between pain and radiographic severity in knee osteoarthritis: findings from quantitative sensory testing of central sensitization. *Arthritis Rheum* 2013; 65: 363 - 72. [PubMed][CrossRef]
18. Sharma A, Kudesia P, Shi Q et al. Anxiety and depression in patients with osteoarthritis: impact and management challenges. *Open Access Rheumatol* 2016; 8: 103 - 13. [PubMed][CrossRef]
19. Vissers MM, Bussmann JB, Verhaar JA et al. Psychological factors affecting the outcome of total hip and knee arthroplasty: a systematic review. *Semin Arthritis Rheum* 2012; 41: 576 - 88. [PubMed][CrossRef]
20. Sakellariou G, Conaghan PG, Zhang W et al. Development of the EULAR recommendations for the use of imaging in the clinical management of osteoarthritis. *Ann Rheum Dis* 2017; 76: 1484 - 94. [PubMed][CrossRef]
21. Hayashi D, Roemer FW, Jarraya M et al. Imaging in Osteoarthritis. *Radiol Clin North Am* 2017; 55: 1085 - 102. [PubMed][CrossRef]
22. Bedson J, Croft PR. The discordance between clinical and radiographic knee osteoarthritis: a systematic search and summary of the literature. *BMC Musculoskelet Disord* 2008; 9: 116. [PubMed][CrossRef]
23. Henry R, Cahill CM, Wood G et al. Myofascial pain in patients waitlisted for total knee arthroplasty. *Pain Res Manag* 2012; 17: 321 - 7. [PubMed][CrossRef]
24. Guermazi A, Niu J, Hayashi D et al. Prevalence of abnormalities in knees detected by MRI in adults without knee osteoarthritis: population based observational study (Framingham Osteoarthritis Study). *BMJ* 2012; 345 (aug29 1): e5339. [PubMed][CrossRef]

25. Englund M, Guermazi A, Gale D et al. Incidental meniscal findings on knee MRI in middle-aged and elderly persons. *N Engl J Med* 2008; 359: 1108 - 15. [PubMed][CrossRef]
26. Bhattacharyya T, Gale D, Dewire P et al. The clinical importance of meniscal tears demonstrated by magnetic resonance imaging in osteoarthritis of the knee. *J Bone Joint Surg Am* 2003; 85-A: 4 - 9. [PubMed][CrossRef]
27. Boks SS, Vroegindeweij D, Koes BW et al. Magnetic resonance imaging abnormalities in symptomatic and contralateral knees: prevalence and associations with traumatic history in general practice. *Am J Sports Med* 2006; 34: 1984 - 91. [PubMed][CrossRef]
28. Espeland A, Natvig NL, Løge I et al. Magnetic resonance imaging of the knee in Norway 2002-2004 (national survey): rapid increase, older patients, large geographic differences. *BMC Health Serv Res* 2007; 7: 115. [PubMed] [CrossRef]
29. Riksrevisjonens undersøkelse av bruken av poliklinisk bildediagnostikk. Oslo: Riksrevisjonen, 2016.
<https://www.riksrevisjonen.no/rapporter/Sider/PolikliniskBildediagnostikk.aspx> (5.1.2018).
30. Lysdahl KB, Hofmann BM. What causes increasing and unnecessary use of radiological investigations? A survey of radiologists' perceptions. *BMC Health Serv Res* 2009; 9: 155 - 64. [PubMed][CrossRef]
31. Petron DJ, Greis PE, Aoki SK et al. Use of knee magnetic resonance imaging by primary care physicians in patients aged 40 years and older. *Sports Health* 2010; 2: 385 - 90. [PubMed][CrossRef]
32. Finnish Degenerative Meniscal Lesion Study Group. Mechanical symptoms and arthroscopic partial meniscectomy in patients with degenerative meniscus tear: A secondary analysis of a randomized trial. *Ann Intern Med* 2016; 164: 449 - 55. [PubMed][CrossRef]
33. Tornbjerg SM, Nissen N, Englund M et al. Structural pathology is not related to patient-reported pain and function in patients undergoing meniscal surgery. *Br J Sports Med* 2017; 51: 525 - 30. [PubMed][CrossRef]
34. Décaire S, Ouellet P, Vendittoli PA et al. Reliability of physical examination tests for the diagnosis of knee disorders: Evidence from a systematic review. *Man Ther* 2016; 26: 172 - 82. [PubMed][CrossRef]
35. Solomon DH, Simel DL, Bates DW et al. The rational clinical examination. Does this patient have a torn meniscus or ligament of the knee? Value of the physical examination. *JAMA* 2001; 286: 1610 - 20. [PubMed][CrossRef]
36. Hegedus EJ, Carolina N. Thessaly test is no more accurate than standard clinical tests for meniscal tears. *Evid Based Med* 2016; 21: 39. [PubMed] [CrossRef]

37. Thorlund JB, Hare KB, Lohmander LS. Large increase in arthroscopic meniscus surgery in the middle-aged and older population in Denmark from 2000 to 2011. *Acta Orthop* 2014; 85: 287 - 92. [PubMed][CrossRef]

38. Thorlund JB, Juhl CB, Roos EM et al. Arthroscopic surgery for degenerative knee: systematic review and meta-analysis of benefits and harms. *Br J Sports Med* 2015; 49: 1229 - 35. [PubMed][CrossRef]

39. Kise NJ, Risberg MA, Stensrud S et al. Exercise therapy versus arthroscopic partial meniscectomy for degenerative meniscal tear in middle aged patients: randomised controlled trial with two year follow-up. *BMJ* 2016; 354: i3740. [PubMed][CrossRef]

40. Roemer FW, Kwoh CK, Hannon MJ et al. Partial meniscectomy is associated with increased risk of incident radiographic osteoarthritis and worsening cartilage damage in the following year. *Eur Radiol* 2017; 27: 404 - 13. [PubMed][CrossRef]

41. Petty CA, Lubowitz JH. Does arthroscopic partial meniscectomy result in knee osteoarthritis? A systematic review with a minimum of 8 years' follow-up. *Arthroscopy* 2011; 27: 419 - 24. [PubMed][CrossRef]

42. Kinsella SD, Carey JL. Complications in brief: Arthroscopic partial meniscectomy. *Clin Orthop Relat Res* 2013; 471: 1427 - 32. [PubMed][CrossRef]

43. Thorlund JB. Deconstructing a popular myth: why knee arthroscopy is no better than placebo surgery for degenerative meniscal tears. *Br J Sports Med* 2017; 51: 1630 - 1. [PubMed][CrossRef]

44. Beaufils P, Becker R, Kopf S et al. Surgical management of degenerative meniscus lesions: the 2016 ESSKA meniscus consensus. *Knee Surg Sports Traumatol Arthrosc* 2017; 25: 335 - 46. [PubMed][CrossRef]

45. Nijs J, Roussel N, Paul van Wilgen C et al. Thinking beyond muscles and joints: therapists' and patients' attitudes and beliefs regarding chronic musculoskeletal pain are key to applying effective treatment. *Man Ther* 2013; 18: 96 - 102. [PubMed][CrossRef]

46. Fagerlin A, Sepucha KR, Couper MP et al. Patients' knowledge about 9 common health conditions: the DECISIONS survey. *Med Decis Making* 2010; 30 (suppl): 35S - 52S.

47. Skou ST, Roos EM. Good Life with osteoArthritis in Denmark (GLA:D™): evidence-based education and supervised neuromuscular exercise delivered by certified physiotherapists nationwide. *BMC Musculoskelet Disord* 2017; 18: 72. [PubMed][CrossRef]

48. Fransen M, McConnell S, Harmer AR et al. Exercise for osteoarthritis of the knee: a Cochrane systematic review. *Br J Sports Med* 2015; 49: 1554 - 7. [PubMed][CrossRef]

