

Subgroups of patients with osteoarthritis and medial meniscus tear or crystal arthropathy benefit from arthroscopic treatment

Katrin Karpinski¹ · Ralf Müller-Rath² · Phillipp Niemeyer³ · Peter Angele⁴ · Wolf Petersen¹

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Abstract

Purpose The purpose of this study was to perform a systematic review of prospective randomized controlled trials comparing arthroscopic treatment for knee osteoarthritis (OA) with either other therapeutic interventions or sham treatment.

Methods A systematic search for randomized controlled trials (RCT) about arthroscopic treatment (AT) for knee OA was performed according to the PRISMA guidelines. Arthroscopic treatment included procedures such as lavage, debridement and partial meniscectomy of the knee. Data source was PubMed central.

Results Fourteen articles could be included. Five studies compared interventive AT with either sham surgery, lavage or diagnostic arthroscopy. Nine trials compared AT with another active intervention (exercise, steroid injection, hyaluronic acid injection). In ten trials, the clinical scores improved after arthroscopic treatment of knee OA in comparison to the baseline. In seven trials, there was a significant difference in the final clinical outcome with higher scores for patients after arthroscopic OA treatment in comparison to a control group. In four trials, the intention to treat analysis revealed no significant difference between arthroscopic OA treatment and the control group. In one of those trials, which compared arthroscopic partial meniscectomy (APM) with exercise, the cross over rate from exercise to AT was 34.9%. The clinical scores of cross-over patients improved after APM. In one study, the subgroup analysis revealed that patients with tears of the anterior two-thirds of the medial meniscus or any lateral meniscus tear had a higher probability of improvement after arthroscopic surgery than did patients with other intraarticular pathology. There was no difference in the side effects between patients with AT and the control group. Despite acceptable scores in the methodological quality assessment, significant flaws could be found in all studies. These flaws include bad description of the exact surgical technique or poor control of postoperative use of non-steroidal anti-inflammatory drugs (NSAID).

Conclusion Results of RCTs comparing AT with other treatment options were heterogeneous. AT in OA patients is not useless because there is evidence that a subgroup of patients with non-traumatic flap tears of the medial meniscus or patients with crystal arthropathy benefit from arthroscopy. This topic has a high relevance because several health insurances do not reimburse arthroscopy for patients with OA anymore. The results of these randomized studies, however, should be interpreted with care because in many studies, the use of other therapeutic variables such as pain killers or NSAIDs was not controlled or reported.

Level of evidence I.

Introduction

Osteoarthritis (OA) is a degenerative joint disease which can affect the whole knee (patellofemoral and tibiofemoral joint) [2]. This degenerative joint disease is a progressive process that can be divided into stages or degrees [2, 26, 34].

With increasing age, OA is the most frequent cause for knee pain [2, 43]. In the fourth and fifth decade of life, light to

⊠ Wolf Petersen

wolf.petersen@pgdiakonie.de

Extended author information available on the last page of the article

moderate stages of OA have the highest prevalence [2]. But the severity of OA increases with aging.

There is no consensus about the criteria for knee OA in the literature. However, in most studies, the radiological classification of Kellgren and Lawrence (KL) is used to stage the progress of OA [26]. According to the American College of Rheumatism, the following clinical criteria should be present: knee pain, osteophytes and one further criterion such as tenderness, age over 50 or crunching of the joint [2].

The main symptoms of OA may have different causes. Pain can be caused mechanically by meniscus lesions, inflammation or subchondral edema. Loss of range of motion can be caused by capsular fibrosis or osteophytes. It has been shown that degenerative meniscal tears are associated with early OA. A non-traumatic meniscus lesion may be the first symptom of knee OA even in the absence of radiological OA signs [10, 11, 25]. Complete loss of the meniscus, however, is an important factor for the progression of OA [12, 44].

For many years, arthroscopic techniques were considered to be the treatment of choice for symptoms of OA because some of the underlying causes can be addressed by AT (partial resection of the meniscus, synovectomy, arthrolysis, removal of free bodies) [8, 25, 37, 38]. Nevertheless, several RCTs have shown that the clinical scores after arthroscopic treatment were not superior in comparison to a control group [29, 37–39]. After publication of these studies, several health care insurances stopped to reimburse AT of knee OA [37]. However, despite these clinical trials, arthroscopy for knee OA has not decreased in every country [8, 50]. One reason for this discrepancy may be that several orthopedic surgeons doubt the results of those trials because of methodological flaws [31].

Aim of this systematic review is to analyze randomized controlled trials of patients with several stages of knee OA and with non-traumatic meniscus lesions to find out if there is any clinically relevant effect of AT in knee OA.

Further objective of this systematic review was to find out if arthroscopy is associated with any side effects in patients with knee OA.

Regarding the outcome, we hypothesize that some subgroups of OA patients (e.g., patients with non-traumatic meniscus lesions) might benefit from arthroscopic surgery.

In contrast to previous systematic reviews, current studies such as the one by Gauffin et al. [15] were included. Furthermore, not only an intention-to-treat analysis of the original study was used to measure the outcome, but also a cross over analysis to identify subgroups of patients who benefit from AT.

Search details

A comprehensive literature search using the PubMed database to identify peer-reviewed articles about AT of knee OA according to the PRISMA statement was conducted. The PRISMA statement consists of a 27-item checklist and a 4-phase flow diagram [19, 36].

Prior to that, the study was registered at PROSPERO, which is an international database of prospectively registered systematic reviews [52]. The corresponding registry number is CRD42016047964.

For this systematic review, different combinations of keywords were utilized: osteoarthritis and arthroscopy, respectively, medial meniscus and arthroscopy. When a study of interest was found, related articles were searched. After identifying those articles, all references were screened for additional relevant publications.

Inclusion and exclusion criteria

The following inclusion criteria were applied:

- prospective randomized trial (level one study),
- trials reporting clinical outcome after AT of patients with any stage of radiological knee OA or of patients with non-traumatic meniscus lesions,
- English language reports,
- publication in a peer-reviewed journal.

All criteria should have been satisfied for inclusion in this systematic review.

All papers qualified for inclusion were read by the reviewers and checked for one of the following exclusion criteria:

- number of patients less than 20,
- Jadad score ≤ 1.

In case of implementation of at least one exclusion criterion the study was excluded.

Two reviewers (WP, KK) performed the initial study identification, secondary study screening, and final determination of eligibility and study inclusion. Both reviewers were also involved in the analysis of the articles.

Analysis

If two separate studies with the same authors and intervention as well as the same patient collective revealed a different follow-up, both publications were counted as one trial. For the analysis, also the appendices of the included study and publications of the study design were deconstructed.

After extraction of all studies' data, a brief tabular narrative of each investigation was presented. Data of this tables included (1) first author and year of publication, (2) number of study centers, (3) country, (4) study groups and number of patients, (5) last follow-up, (6) mean age, (7) OA grade and (8) gender ration, (9) scores (Table 1). Additional tables were added to illustrate the procedures performed in the studies, results of clinical outcome, side effects and study limitations (Tables 2, 3, 4).

Primary and secondary endpoints

Primary endpoint was the group difference in the clinical outcome scores used in the studies.

Secondary endpoints were: (1) subgroup analysis for factors which might have an effect on the outcome after AT of OA, (2) the crossover rate (patients who changed from one treatment group to the other), (3) the rate of side effects and (4) a methodological analysis of the included studies.

Study quality and limitations

Each article was analyzed for limitation and bias by all reviewers. For the quality assessment, information has been extracted from the original article, from published appendices or from published study protocols. Study quality has been analyzed with the Jadad score [17] and with the Coleman methodology score [9].

Jadad score

The Jadad score is a three-point questionnaire that forms the basis of a score [17, 42]. This questionnaire focuses on randomization, blinding and description of dropouts. The questions are as follows: (1) Was the study described as randomized? (2) Was the study described as double blind? (3) Was there any description of withdrawals and dropouts?

For each answer one point is given [17, 42]. Additional points are given if the method of randomization is described in the paper, if that method was appropriate and if the method of blinding was described and appropriate. Points are deducted if the method of randomization or blinding was inappropriate. The highest score a study can receive is, therefore, five points [17].

Coleman methodology scoring system

The Coleman methodology scoring system was developed to analyze the quality of studies reporting surgical treatments of patellar tendinopathy [9]. It's criteria takes into account number of patients, follow-up, number of different treatment procedures, type of study (randomized), diagnostic certainty, description of the surgical procedure, description of postoperative rehabilitation, outcome criteria, procedure for assessing outcome and patient selection [9, 42].

Limitations

These limitations were systematically analyzed: (1) description of the surgical procedure, (2) control of surgical process quality, (3) description of the rate of meniscus extrusion, (4) the rate of varus or valgus malalignment, (5) the outcome score and (6) control of use of pain killers and NSAIDs.

Results

Search results and study design

The search results are shown in Fig. 1 and details of the study design are shown in Table 1. In ten studies, partial meniscectomy was part of the AT. In six of those studies, arthroscopic partial meniscectomy (APM) was the only surgical procedure which was performed. In five studies, multiple procedures were allowed (Table 2). Additional procedures included partial synovectomy, debridement of chondral flaps and resection of osteophytes which blocked joint extension [7, 29, 35, 39]. In three studies, the AT was lavage only [3, 14, 23].

The control groups were also variable (Table 2). In five studies, the control treatment was sham surgery or arthroscopic lavage [7, 22, 23, 39, 48]. In six studies, control treatment was supervised or unsupervised exercise [15, 21, 24, 29, 30, 51].

Clinical outcome scores

Several different outcome scores were used and the results of the different studies were heterogeneous (Tables 1, 3).

WOMAC score

There was no significant difference in the WOMAC total score in both studies with this score as primary endpoint [23, 29]. In one of these studies, however, some secondary endpoints (WOMAC pain and VAS pain) were significantly better in the arthroscopy group (lavage with 3000 ml) in comparison to "placebo" surgery (lavage with 250 ml). In this study, patients with crystals in the synovial fluid had greater improvements in pain [23].

In one study, with the WOMAC pain subscale as primary endpoint, the improvement was significantly greater in the

First author and year	First author and year Number of study centers Country Study groups, no. Last of patients at F/U	Country	Study groups, no. of patients at F/U	F/U	Mean age (years)	Mean age (years) Available information about OA classification and stage	Gender ratio	Scores
Arden et al. (2007)	Dual center	United kingdom	A: 71 C: 79	26 weeks	A: 64.9 C: 67.7	KL score 0–1: A: 16.9%, C: 10.1% KL score II: A: 67.6%, C: 68.4% KL score III-IV: A: 11.3%, C: 20.3%	Male/female A: 19/52 C: 33/46	WOMAC pain (PO) WOMAC physical func- tion (SO) WOMAC stiffness (SO) Patients self-reported improvement (SO) Time to walk 50 m and to climb/descend stairs (SO) Analoetic intake (SO)
Chang et al. (1993)	One center (two sites)	USA	A: 18 C: 14	1 year	A: 61 C: 65	KL score I: A: 22%, C: 14% KL score II: A: 28%, C: 36% KL score III: A: 50%, C. 50%	Male/female A: 5/13 C: 4/10	AIMS 50-foot walk time 50-foot walk time Physician's global assessment of disease activity in the knee Economic measures
Forster et al. (2003)	One center	United kingdom	A: 19 C: 19	1 year	A: 63 C: 60	Inclusion criteria: sympto- matic and radiographic knee OA without mechanical symptoms	Not stated	VAS pain Knee Society function score Lequesne index
Gauffin et al. (2014)	One center	Sweden	A: 75 C: 75	1 year	A: 54 C: 54	KL score 0: A: 49%, C: 43% KL score I: A: 45%, C: 48% KL score II: A: 5%, C: 9%	Male/female A: 53/22 C: 56/19	KOOS pain (PO) KOOS symptoms, QoL, sports, ADL (SO) EQ 5D (SO) VAS (SO) PAS (SO)
Herrlin et al. (2013)	One center	Sweden	A: 47 C: 49	5 years	A: 54 C: 56	Degenerative medial meniscus tear + radio- graphic OA grade 0–I (Ahlbäck)	Male/female A: 28/19 C: 30/19	KOOS (PO) Tegner activity scale (SO) Lysholm (SO) VAS pain (SO)
Hubbard et al. (1996)	One center	United kingdom	A: 40 C: 36	5 years	A: 50 C: <i>5</i> 7	Inclusion of degenera- tive grade 3 or 4 lesions (Outerbridge) of the medial femoral condyle	Male/female A: 28/12 C: 20/16	(Modified) Lysholm score (PO) Pain relief (%) (SO)
Kalunian et al. (2000)	Multi center	USA	A: 41 C: 49	1 year	A: 60.9 C: 58.3	Inclusion of KL score 0-II	Male/female A: 19/22 C: 23/26	Aggregate WOMAC (PO) WOMAC pain (SO) WOMAC physical func- tion (SO) WOMAC stiffness (SO) VAS pain (SO)

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First author and year	Number of study centers Country	Country	Study groups, no. of patients at F/U	Last F/U	Mean age (years)	Mean age (years) Available information about OA classification and stage	Gender ratio	Scores
Katz et al. (2013)	Multi center	NSA	A: 161 C: 169	1 year	A: 59.0 C: 57.8	KL score 0: A= 34, C= 36 Male/fen KL score I: A= 26, C= 35 A: 71/90 KL score II: A= 37, C= 39 C: 72/97 KL score III: A= 45, C= 39	Male/female A: 71/90 C: 72/97	WOMAC physical func- tion (PO) KOOS pain (SO) SF-36 (SO)
Kise et al. (2016)	Two centers	Norway	A: 70 C: 70	2 years	A: 48.9 C: 50.2	KL score 0: A: 70%, C: 73% KL score I: A: 26%, C: 23% KL score II: A: 3%, C: 4% KL score III: A: 1%, C: 0%	Male sex (%) A: 61 C: 61 C: 61	KOOS (PO) Muscle strength (Biodex 6000 dynamometer) (PO) 5 KOOS subscales (SO) SF 36 (SO) Dynamometer (SO) Lower extremity perfor- mance tests (leg hop test, timed hop test, knee bends test) (SO)
Kirkley et al. (2008)	One center	Canada	A: 92 C: 86	2 years	A: 58.6 C: 60.6	KL score II: A: 46%, C: 42% KL score III: A: 49%, C: 53% KL score IV: A: 5%, C: 5%	Male sex (%) A: 41 C: 33 C: 33	WOMAC total (PO) WOMAC subscales pain, stiffness, and physical function (SO) Physical Component Sum- mary of SF-36 (SO) MACTAR (SO) ASES (SO) ASES (SO) Health-related quality of life/standard gamble util- ity technique (SO)
Moseley et al. (2002)	One center	USA	AD: 59 AL: 61 P: 60	2 years	AL: 51.2 AD: 53.6 C: 52.0	Mild C AL AD (%) 28.3 27.9 30.5 (%) 28.3 27.9 30.5 Mod- 46.7 45.9 45.8 erate 25.0 26.2 23.7 (%) Severe (%) (%)	Male sex (%) AD: 96.6 AL: 88.5 P: 93.3	Knee-Specific Pain Scale (KSPS) (PO) Arthritis Impact Measure- ment Scales AIMS2-P (SO) SF-36-P (SO) PFS (SO)

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Table 1 (continued)

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First author and year	Number of study centers Country	Country	Study groups, no. of patients at F/U	Last F/U	Mean age (years)	Study groups, no. Last F/U Mean age (years) Available information of patients at F/U about OA classification and stage	Gender ratio Scores	Scores
Merchan and Galindo (1993)	One center	Spain	A: 35 C: 38	3 years	A: <i>57</i> C: 56	Patients with limited degenerative OA of the FT joint (radiologically minimal joint space nar- rowing and formation of small osteophytes)	Male/female A: 7/28 C: 13/25	Male/female Hospital for Special Sur- A: 7/28 gery Knee Rating Score C: 13/25
Sihvonen et al. (2013)	Multi center	Finland	A: 70 C: 76	1 year	A: 52 C: 52	KL score 0: A=35, C=36 A: 42/28 KL score 1: A=35, C=40 C: 47/29	A: 42/28 C: 47/29	Lysholm knee score (PO) WOMET (PO) Knee pain score NRS (PO) 15D score (SO)
Yim et al. (2013)	One center	Korea	A: 50 C: 52	2 years	A: 54.9 C: 57.6 KL score 0-1	KL score 0–I	A: 9/41 C: 12/40	Lysholm knee score VAS pain Tegner activity scale Patient subjective knee pain and satisfaction

PO primary outcome measure, *SO* secondary outcome measure, *F/U* follow up, *A* arthroscopy, *C* control, *P* placebo, *D* debridement, *L* lavage, *KL* Kellgren and Lawrance, *KOOS* Knee Osteo-arthritis Outcome Score, *ADL* activities of daily living, *QoL* quality of life, VAS visual analogue scale, *WOMAC* Western Ontario and McMaster Universities Osteoarthritis Index, *SF-36* Short-Form Health Survey, *AIMS* Arthritis Impact Measurement Scales, *WOMET* Western Ontario Meniscal Evaluation Tool, *MACTAR* McMaster-Toronto-Arthritis, *ASES* American Shoulder and Elbow Surgeons

Tab	Table 2 Procedures examined in the RCTs	, RCTs	
	First author and year	Arthroscopic procedures	Control treatment
Ξ.	Arden et al. (2007)	Arthroscopic lavage (tidal irrigation)	Corticosteroid injection
5	Chang et al. (1993)	Arthroscopic lavage with partial meniscectomy, partial synovialectomy, removal of loose cartilage fragments	Closed-needle joint lavage
б	Forster et al. (2003)	Arthroscopic lavage	Hyaluronic acid injection $(5x)$
4	Gauffin et al. (2014)	Arthroscopy with partial resection of the medial meniscus + exercise pro- gram	Exercise program
5.	Herrlin et al. (2013)	Arthroscopy with partial meniscectomy and supervised exercise therapy	Supervised exercise therapy
6.	Hubbard et al. (1996)	Arthroscopic debridement (resection of loose cartilage)	Arthroscopic lavage
7.	Kalunian et al. (2000)	Arthroscopic lavage with 3000 ml saline	Diagnostic arthroscopy with 250 ml saline (placebo)
×.	Katz et al. (2013)	Arthroscopy with partial meniscectomy + physical therapy	Physical therapy
9.	Kise et al. (2016)	Arthroscopy with partial meniscectomy	Supervised exercise therapy (12 weeks)
10.	Kirkley et al. (2008)	Arthroscopic treatment (lavage with synovectomy/debridement/partial menisectomy/removal of cartilage fragments/removal of osteophytes) + physical and medical therapy	Physical (supervised and unsupervised exercise at home) and medical therapy (acetaminophen/NSAIDs, i.a. hyaluronic acid, oral glycosamine)
11.	11. Moseley et al. (2002)	$D\acute{e}bridement$ group Arthroscopic lavage (10 l) with partial meniscectomy, cartilage shaving, removal of loose debris Lavage group Arthroscopic lavage (10 l) and if present resection of an unstable bucket handle tear	Placebo: Sham surgery (skin incision)
12.	Merchan and Galindo (1993)	12. Merchan and Galindo (1993) Arthroscopic lavage with partial menisectomy/synovectomy, debridement of Non-steroidal anti-inflammatory drugs and a decrease in the intensity of the loose articular cartilage and removal of loose bodies/osteophytes activities of daily living for a pain free knee	Non-steroidal anti-inflammatory drugs and a decrease in the intensity of the activities of daily living for a pain free knee
13. 14.	 13. Sihvonen et al. (2013) 14. Yim et a. (2013) 	Arthroscopy and partial resection of the medial meniscus Arthroscopy with partial meniscectomy	Sham surgery 2 weeks NSAIDs or muscle relaxants 2 wools envorvious
			8 weeks home exercises

No.	First author and year	Clinical outcome	Subgroup analysis or cross overs
	Arden et al. (2007)	In both treatment groups, over 80% of patients reported improvement at 2 and 4 weeks. After this time, the benefit of steroid injection decreased, whereas that of arthroscopic lavage was maintained	Subgroups: patients with a knee effusion responded better to both treatments, however, this was most apparent for corticosteroid injection
		At 26 weeks, the pain relief afforded by arthroscopic lavage was significantly greater than that of steroid injection. At 26 weeks, 29% of the steroid injection group reported improvement versus 64% of the arthroscopy group ($p < 0.001$)	Patients with less severe radiographic OA also obtained the greatest improve- ment from both treatments
Ċ	Chang et al. (1993)	There were statistically significant differences ($p < 0.05$), favoring the arthroscopy group, in knee tenderness scores and in physician's global assessment scores when a Mann–Whitney U test was used. There were no statistically significant differences in all other clinical, functional, or global outcomes between the subjects receiving arthroscopic surgery and those receiving non-operative management at 3 and 12 months. There was no difference in the mean costs	Subgroups: patients with tears of the anterior two-thirds of the medial menis- cus or any lateral meniscus tear had a higher probability of improvement (by "blinded" physician assessment) after arthroscopic surgery than did patients with other intraarticular pathology
	Forster et al. (2003)	There was no significant difference in VAS, FS or LJ between the two groups at 6 weeks, 3 months, 6 months or 1 year In the Hyalgan group 5 patients underwent TKR or were on a waiting list, in the arthroscopy group 3 underwent TKR or were on a waiting list	None
4.	Gauffin et al. (2014)	KOOS pain: less pain at 3 and 12 months in the surgery group KOOS symptom: fewer symptoms of the surgery group at 12 months Overall KOOs: the changes from baseline to 3 months and from 3 to 12 months were significantly larger for surgery group compared to exercise group EQ5D: no difference PAS: significant improvement from baseline to 12 months for both groups Symptom satisfaction scale: significant improvement from baseline to 12 months for both groups	Cross overs: sixteen patients crossed over from the non-surgery group to receive an operation (21%). Nine patients (12%) that were allocated to the surgery group did not go through with the operation Subgroups: in both groups, older patients exhibited larger improvements in KOOS pain than younger patients did in both groups
S.	Herrlin et al. (2013)	Both groups showed highly significant clinical improvements from baseline to the follow-ups at 24 and 60 months on all subscales of KOOS, Lysholm Knee Scoring Scale and VAS. No group differences were found. There were also no group differences regarding the Tegner score at 24 and 60 months	Cross overs: about one-third of the patients (27.7%) that were treated with exercise therapy alone did not improved in the clinical scores after the treatment, though they crossed over to the arthroscopy group and improved after surgery to the same level
.0	Hubbard et al. (1996)	At 1 year, 32 of the debridement group and five of the washout group were pain free and at five years, 19 of a total of 32 survivors in the debride- ment group and three of the 26 in the washout group were also free from pain. The mean improvement in a modified Lysholm score was 28 for the debridement group at 1 year and 21 at 5 years. In the washout group, it was only 5 at 1 year and 4 at 5 years	None

Model Endealment and year Subgroup analysis or tonso overs 7. Kahnian et al. (2000) The study of an ot ennontense an effect of tragition on antitits sverity sis Subgroup analysis patterns with crystals had statistically greater in an ensure by greaterns with crystals had statistically greaterns with crystals had breaterns had breate	Tab	Table 3 (continued)		
 Kalunian et al. (2000) The study did not demonstrate an effect of irrigation on arthritis severity as measured by agregates WOMACC scores, the primary outcome variable; Natz et al. (2013) In the intervition to treat analysis, both groups showed clinical improvements from baseline to the follow-ups at 3, 6 and 12 months on KOOS pain score and WOMAC physical function score. No group differences were found the intervition to treat analysis, both groups showed clinical improvements from baseline to the follow-ups at 3, 6 and 12 months on KOOS pain score and WOMAC physical function score. No group differences were found works at 2, 600 AG (2016) Kise et al. (2016) No clinically relevant difference was found between the two groups in compase in KOOS at 2 years. Kitkley et al. (2003) Months. muscle strength had improved in the exercise group No serious adverse events occurred in either group. After 2 years, the mean (±SD) WOMAC score works for the surgery group was 874±604, as compased with 997±583 for the control group. The SF=36 Physical Component Summary scores were 37.0±11.4 and 37.2±10.6, respectively. Analyses of WOMAC scores that hefore surgery more the strength and addition (115), worse than before improved. For (145), unchanged and 20 (53%) wore timerous. Stote and for the surgery form the strength and addition (116), worse than before treatment. Yim et al. (2013) Both groups showed finical improvements from baseline to all follow-ups at 6 and 12 months on Cy35%) were founder the strength on Cy35% on the treatment from the formet and and strengt in the optical improvements from baseline to all follow-ups at 6 and 20 (53%) were improved. For exploit you cy35% on the physical composed of the exercise using the VAS and 13 months, intelexed of the intervention group in the optical improved from the fine of their last evaluation. 26 (75%) were improved. For (13%) unchanged and 20 (53%) wore than before the groups showed finical improved and for (115%) worse than before	No		Clinical outcome	Subgroup analysis or cross overs
Kaiz et al. (2013) In the interniton to treat analysis, both groups showed clinical improvements from baseline to the follow-ups at 3, 6 and 12 months on KOOS pain score and WOMAC physical function score. No group differences were found the set al. (2016) Kise et al. (2016) No clinically relevant difference was found between the two groups in change in KOOS at 2 years. Kitkley et al. (2008) No clinically relevant difference was found between the two groups vas 874 ± 624, as compared with 897 ± 583 for the control group Noseley et al. (2008) Kitkley et al. (2008) After 2 years, the mean (± S1) WOMAC scores at interim visits and 0712 ± 10.6, respectively. Analyses of WOMAC scores at interim visits and 0712 ± 10.6, respectively. Analyses of WOMAC scores at interim visits and 0712 ± 10.6, respectively. Analyses of WOMAC scores than before the secondary outcomes also failed to show superiority of surgery Moseley et al. (2002) At no point did either of the intervention groups report less pain or better function than the placebo group 20 (53%) worse than before trannowis. (16%) vere improved, 12 (31%) unchanged and 20 (13%) worse than before treatment Silbvonen et al. (2013) Both groups showed marked clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found after exercise using the VAS and 15 D. No group differences were found after exercise using the VAS and 15 D. No group differences were found after exercise using the VAS and 15 D. No group differences were found after exercise using the VAS and 15 D. No group differences were found after exercise using the VAS and 15 D. No group differences for Lysholm score.	7.		The study did not demonstrate an effect of irrigation on arthritis severity as measured by aggregate WOMAC scores, the primary outcome variable. However, full irrigation (arthroscopic lavage) did have a statistically sig- nificant effect on patients' self-reported pain as measured by the WOMAC pain subscale and by a visual analog scale (VAS) (the secondary outcome variables)	Subgroup analysis: patients with crystals had statistically greater improve- ments in pain
Kise et al. (2016)No clinically relevant difference was found between the two groups in change in KOOS at 2 yearsAt 3 months, muscle strength had improved in the exercise group No serious adverse events occurred in either group No serious adverse events occurred in either group No serious adverse events occurred in either group State 524, as compared with 897 ± 583 for the control group The ST-36 Physical Component Summary scores were 37 0.4 ± 11.4 and 37.2 ± 10.6, respectively. Analyses of WOMAC score at interim visits and other secondary outcomes also failed to show superiority of surgery Moseley et al. (2002)Moseley et al. (2002)At no point did either of the intervention groups report less pain or better function than the placebo group mer horoved, five (14%) unchanged and 20 (53%) worse than before surgery. In the nonoperated group, at the time of their last evaluation, six (16%) were improved, five (14%) unchanged and 20 (53%) worse than before treatmentSihvonen et al. (2013)Both groups showed marked clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found after exercise using the YAS and 15 D. No group differences were found after exercise using the YAS and 15 D. No group differences were found after exercise using the groups at 0 and 12 months, the Lysholm score of the treatment group was significantly higher than that of the control group	ś	Katz et al. (2013)	In the intention to treat analysis, both groups showed clinical improvements from baseline to the follow-ups at 3, 6 and 12 months on KOOS pain score and WOMAC physical function score. No group differences were found	Cross overs: after 6 months, 51 patients (30.2%) without improvement after physical therapy had undergone arthroscopic partial meniscectomy. Addi- tional eight patients (4.7%) who were assigned to physical therapy crossed over to arthroscopic partial meniscectomy between 6 and 12 months. After arthroscopic partial meniscectomy, the WOMAC physical-function scores at 12 months were similar to those of patients primarily assigned to the arthroscopic treatment
Kirkley et al. (2008)After 2 years, the mean (±SD) WOMAC score for the surgery group was 874±624, as compared with 897±583 for the control group The SF-36 Physical Component Summary scores were 37.0±11.4 and 37.2±10.6, respectively. Analyses of WOMAC scores at interim visits and other secondary outcomes also failed to show superiority of surgery Moseley et al. (2002)After 2 years, the mean (±SD) WOMAC scores were 37.0±11.4 and 37.2±10.6, respectively. Analyses of WOMAC scores at interim visits and other secondary outcomes also failed to show superiority of surgery Moseley et al. (2002)No At no point did either of the intervention groups report less pain or better function than the placebo group mored, five (14%) unchanged and four (11%) worse than before improved, five (14%) unchanged and for (11%) worse than before treatmentNo Six(16%) Six (16%)Sihvonen et al. (2013)In the nonoperated group, at the time of their last evaluation, is: (16%) were improved, 12 (31%) unchanged and 20 (53%) worse than before treatmentCSihvonen et al. (2013)Both groups showed marked clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, were found after exercise using the VAS and 15 D. No group differences were found so n Lysholm score, Tegner score and pain on VAS, with no difference between the groups. Only at 3 months, the Lysholm score of the treatment group was significantly higher than that of the control group	9.	Kise et al. (2016)	No clinically relevant difference was found between the two groups in change in KOOS at 2 years At 3 months, muscle strength had improved in the exercise group No serious adverse events occurred in either group	Cross over: 19% of the participants allocated to exercise therapy crossed over to surgery during the 2-year follow-up, with no additional benefit
Moseley et al. (2002) At no point did either of the intervention groups report less pain or better function than the placebo group N Merchan and Galindo (1993) In the operative group, at the time of their last evaluation, 26 (75%) were improved, five (14%) unchanged and four (11%) worse than before surgery. In the nonoperated group, at the time of their last evaluation, six (16%) were improved, 12 (31%) unchanged and 20 (53%) worse than before treatment N Sihvonen et al. (2013) Both groups showed marked clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found S C Yim et al. (2013) Both groups showed clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found S N Yim et al. (2013) Both groups showed clinical improvements from baseline to all follow-ups on Lysholm score, Tegner score and pain on VAS, with no difference between the groups. Only at 3 months, the Lysholm score of the treatment group was significantly higher than that of the control group N	10.		After 2 years, the mean (\pm SD) WOMAC score for the surgery group was 874 \pm 624, as compared with 897 \pm 583 for the control group The SF-36 Physical Component Summary scores were 37.0 \pm 11.4 and 37.2 \pm 10.6, respectively. Analyses of WOMAC scores at interim visits and other secondary outcomes also failed to show superiority of surgery	No benefit was conferred by surgery among the subgroup of patients with mechanical symptoms of catching or locking No benefit of arthroscopic treatment in patients with more severe radio- graphic disease (KL III and IV) or less severe radiographic disease (KL II)
Merchan and Galindo (1993) In the operative group, at the time of their last evaluation, 26 (75%) were improved, five (14%) unchanged and four (11%) worse than before surgery. In the nonoperated group, at the time of their last evaluation, six (16%) were improved, 12 (31%) unchanged and 20 (53%) worse than before treatment N Sihvonen et al. (2013) Both groups showed marked clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found S C Yim et al. (2013) Both groups showed clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found S N Yim et al. (2013) Both groups showed clinical improvements from baseline to all follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found S N	11.	. Moseley et al. (2002)	At no point did either of the intervention groups report less pain or better function than the placebo group	No subgroup analysis
Sihvonen et al. (2013)Both groups showed marked clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found SCYim et al. (2013)Both groups showed clinical improvements from baseline to all follow-ups on Lysholm score, Tegner score and pain on VAS, with no difference between the groups. Only at 3 months, the Lysholm score of the treatment group was significantly higher than that of the control groupN	12.		In the operative group, at the time of their last evaluation, 26 (75%) were improved, five (14%) unchanged and four (11%) worse than before surgery. In the nonoperated group, at the time of their last evaluation, six (16%) were improved, 12 (31%) unchanged and 20 (53%) worse than before treatment	No subgroup analysis
Yim et al. (2013) Both groups showed clinical improvements from baseline to all follow-ups on Lysholm score, Tegner score and pain on VAS, with no difference between the groups. Only at 3 months, the Lysholm score of the treatment group was significantly higher than that of the control group	13.		Both groups showed marked clinical improvements from baseline to the follow-ups at 6 and 12 months on Lysholm score, WOMET score, pain after exercise using the VAS and 15 D. No group differences were found	Cross overs: one patient from the treatment group underwent repeat arthros- copy and five patients from control group underwent arthroscopy. One patient of the treatment group received a TKA Subgroups: no difference in outcome of patients with grade 0 or grade I OA (KL). Resection of a torn meniscus has no added benefit over sham surgery to relieve knee catching or occasional locking
	14.		Both groups showed clinical improvements from baseline to all follow-ups on Lysholm score, Tegner score and pain on VAS, with no difference between the groups. Only at 3 months, the Lysholm score of the treatment group was significantly higher than that of the control group	Not reported

Table 4Quality assessmentwith the Jadad score andColeman methodology score

No.	First author and year	Blinding	Jadad score (points)	Coleman meth- odology score (points)
1.	Arden et al. (2007)	Single blinded	4	84
2.	Chang et al. (1993)	Single blinded	3	84
3.	Forster et al. (2003)	Not blinded	2	69
4.	Gauffin et al. (2014)	Not blinded	3	85
5.	Herrlin et al. (2013)	Not blinded	3	83
6.	Hubbard et al. (1996)	Not blinded	2	62
7.	Kalunian et al. (2000)	Double blinded	5	77
8.	Katz et al. (2013)	Not blinded	3	91
9.	Kise et al. (2016)	Single blinded	3	98
10.	Kirkley et al. (2008)	Single blinded	3	95
11.	Moseley et al. (2002)	Double blinded	5	100
12.	Merchan and Galindo (1993)	Not blinded	3	78
13.	Sihvonen et al. (2013)	Double blinded	5	95
14.	Yim et al. (2013)	Not blinded	2	88

arthroscopy group (lavage) compared to intraarticular corticoid injections. In this study, patients with a knee effusion or with less severe radiographic OA responded better to both treatments [3].

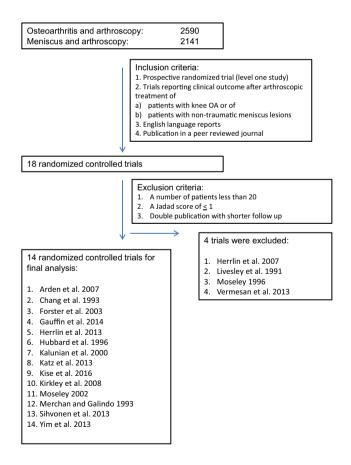


Fig. 1 Flowchart showing the literature review

In one study, the intention-to-treat analysis showed no significant difference in the WOMAC function subscale of knee OA patients after APM or exercise (n.s.). In this study, however, the WOMAC function subscale did not improve in 34.9% of the patients who were assigned to the exercise group. After cross over to APM, the WOMAC function scores at 12 months were similar to those of patients who were primarily assigned to the APM [24].

KOOS

The KOOS or a KOOS subscale was used in three studies as primary outcome measurement and in two studies as secondary outcome measure [15, 21, 24, 30]. All studies examined the effect of APM or exercise in patients with OA. The results were contradictory. In one study, patients of the surgery group had significantly less pain as measured with the KOOS pain subscale at 3 and 12 months postoperatively [15]. Three studies found no difference in the KOOS pain score [15, 21, 30]. In all three studies, crossover rates from the exercise group to the arthroscopy group have been described (19% [30], 21% [15] and 27.7% [21]). In the Herrlin et al. study, 8 of the 13 cross over patients had flap tears [21].

Lysholm score

Two studies found no statistical difference in the Lysholm score between the APM group and a control treatment (sham surgery or exercise) [48, 51]. In one study, arthroscopy with removal of chondral flaps and trimming of the bed of the flap led to a significantly better Lysholm score than control treatment [22]. In this study, a modified Lysholm score without the instability subscore was used.

Other scores

Four studies used other scores as outcome tools (Table 3). Three of those studies did not differentiate the outcome measures into primary and secondary endpoints [7, 14, 35].

Adverse events

Side effects were analyzed in seven studies. In all studies, the rate of adverse events in both the treatment and control group was low [3, 15, 24, 30, 35, 39, 48]. In four of these studies, AT was compared with a non-operatively treated control group [15, 24, 30, 35]. In three of these studies— with physiotherapy as control group—there was no significant difference in the rate of side effects between the two study groups [15, 24, 30]. In one study, AT was compared to oral NSAIDs [35]. In this study, two deep venous thrombosis, one superficial infection and one hemarthrosis were observed in the arthroscopy group, whereas no adverse effect was observed in the NSAID group [30].

Study quality and limitations

Quality assessment of the studies with the Jadad and the Coleman methodology score is shown in Table 4. The Jadad score ranges from 2 to 5 points. The Coleman methodology score ranges between 59 and 96.

Only three studies addressed varus or valgus malalignment of the participants [3, 29, 35]. No study mentioned the rate of meniscus root tears, but the percentage of participants with meniscus extrusion was described in one study. In this study, the rate of meniscus extrusion was 65% in the arthroscopy group and 50% in the control group [30].

The use of pain killers or NSAIDs was addressed in three studies [3, 29, 39]. In two studies, there was no difference in the consumption of pain killers or NSAIDs during the course of the studies [3, 29]. In one study, the use of pain killers or NSAIDs was described in the baseline characteristics only [39].

Seven of the included studies used a specific OA score as primary outcome measure (KOOS or WOMAC) [3, 15, 21, 23, 24, 29, 30].

Discussion

The most important finding of the present study was that certain subgroups of patients with knee osteoarthritis can benefit from AT.

This systematic review has shown that AT has no major advantage over non-operative treatment for the majority of patients with OA. However, there is evidence in the literature that AT can be a useful option for a subset of OA patients with non-traumatic meniscus lesions or crystal arthropathy.

This statement is in contrast with other previous systematic reviews. In a Cochrane review from 2008, Laupattarakasem et al. have shown that there is 'gold' level evidence that AT has no benefit for the treatment of OA [32]. Two systematic reviews from 2014 could also find no difference in the outcome of OA patients with AT and without AT [6, 37]. An explanation for the contradictory findings is that the study by Gauffin et al. could be not included to these systematic reviews because this study was only published in 2014 [15]. Gauffin et al. could show that patients with mild OA (stage 0-II according to KL [26]) with previous unsuccessful physiotherapy benefit from APM. Gauffin et al. found that the change in KOOS pain was larger in the surgery group compared to the non-surgery group. The difference in improvement between the groups was clinically relevant [15].

A qualitative flaw of these previous systematic reviews was that "the intention to treat analysis" of the original study was used to measure outcome. Katz et al. and Herrlin et al. found in the intention to treat analysis that there was no difference in outcome between patients with APM or physiotherapy [21, 24]. In both studies, however, there was a significant rate of patients in the physiotherapy group (34.9% and 27.7%) who crossed over to the arthroscopy group because they did not improve in clinical scores. After AT, the clinical scores improved in both studies to the same level of patients with initial APM [21, 24]. The studies by Katz et al. and Herrlin et al. have shown that a crossover analysis can be helpful in identifying subgroups of patients who benefit from the procedure [21, 24]. In this context, the "intention-to-treat" analysis popular in clinical research can also be seen critically. This can be illustrated by the following example. Diet A (treatment) is compared to diet B (placebo) in a clinical trial with 40 participants in each group. In group A, 38 participants lost weight, whereas in group B only five participants lost weight. If the weight loss would be analyzed in an "as-treated analysis", the effect of diet A would be underestimated. Therefore, for this trial, an intention-to-treat analysis makes sense. If the same analysis is performed in a RCT about the effect of APM with a crossover rate of approximately one-third of patients with no improvement after physiotherapy, an intention-to-treat analysis is misleading [25].

All studies found that the various clinical scores at follow-up improved significantly after AT of patients with knee OA in comparison to the baseline. Regarding the superiority of AT, the results of the included studies were heterogeneous. Some studies have shown that the outcome after AT is better than control treatment [3, 15, 22, 23, 35]. Other studies have shown that there is no difference in outcome between patients with AT and control treatment [7, 14, 21, 24, 29, 30, 39, 48, 51]. The heterogeneity and discrepancy of the study results can be explained by differences in the stage of OA, type of AT, patient characteristics, study design and study quality.

With regard to the degree of osteoarthritis, very wide inclusion criteria were chosen in the present systematic review to include not only patients with advanced knee osteoarthritis but also patients with early osteoarthritis. Even at stage 0 according to KL [26], a non-traumatic meniscal lesion or a chondral lesion can be seen as an initial process in the development of osteoarthritis [34]. An effect of the AT was found especially in studies with patients in early OA stages (stage KL 0–II) [15, 22, 23, 35]. Two studies showed a benefit of APM even in patients with stage III in OA [7, 24]. Two studies including patients with Grade IV OA after KL failed to demonstrate superiority in AT [29, 39].

The clinical conclusion of these findings is that APM is a useful procedure in knees with stage 0-III OA with initial unsuccessful non-operative treatment. The studies reviewed indicate that the shape of the non-traumatic meniscal lesion may be a prognostic factor for the success of a partial meniscectomy. Yim et al. included only patients with a horizontal tear and found no difference in the outcome of APM in comparison to non-operative treatment [51]. In the Herrlin et al.'s study, the majority of patients who did not benefit from non-operative treatment in the cross over group had flap tears [20, 21]. This statement is in accordance with the 2016 ESSKA meniscus consensus [4]. However, the recommendations of the present paper are broader than the ESSKA meniscus consensus, because the literature did not focus on meniscus studies only. In one study, the removal of chondral flaps had a positive effect on outcome [22] and in one other RCT AT was beneficial for patients with crystal arthropathy [23].

This is a systematic review and flaws of studied RCTs are also flaws of this paper. The quality assessment with the Jadad score the Coleman methodology score shows also heterogeneous results for the 14 trials which were included in this review (Table 4).

The Jadad score was developed for quality assessment of RCTs and this score focuses on aspects as randomization and blinding [17]. Three studies received a maximum score of five points [23, 39, 48]. In all three studies, the control group was sham surgery (placebo). The lack of difference between arthroscopy and placebo suggests that the improvement is not only due to any intrinsic efficacy of the procedures [39]. However, the use of a placebo group has also disadvantages because blinding prevents a change from the control group to the treatment group (cross over). The studies by Katz et al. and Herrlin et al. have shown that a crossover analysis can be helpful in identifying subgroups of patients who benefit from the procedure [20, 21, 24].

The Coleman methodology score was developed for the assessment of orthopedic studies. This score covers additional aspects such as number of patients, follow-up, diagnostic certainty, description of the surgical procedure, description of rehabilitation, outcome criteria, and patient selection. With this score, the studies of Katz et al. [24], Kirckley et al. [29], Moseley et al. [39] and Shivonen et al. [46–49] received the best results.

Other limitations include that most authors give no information about the rate of subchondral edema or varus malalignment. Both factors are predictors for a poorer outcome after arthroscopic surgery.

It is also remarkable that only few studies reported the consumption of pain killers or NSAIDs during the treatment and follow-up period. Good results in the control groups could be the result of a higher NSAID use. The well-known adverse effects of an extensive NSAID use are gastrointestinal bleeding or ulcer [45].

Other flaws that where identified by the reviewers are a selection bias or the use of non-specific scores. Selection bias is a typical limitation of a randomized controlled trial. In the METEOR study, for example, only 26% of eligible patients could be included. That means that no follow-up of those patients preferring not to enter the study was done [24]. Selection bias is assumed when the recruitment rate is below 80% [9]. Therefore, the findings of the studies with a recruitment rate below 80% should only be generalized cautiously [42]. In contrast to the METEOR trial, the Gauffin et al. study has the participation rate of 84% [15].

It is also of concern that four studies used the Lysholm score as outcome measurement. The Lysholm score was originally developed for the assessment of patients with ligamentous instability. To our knowledge, this score is not validated for Finish and Korean. Briggs et al. have shown that there were unacceptable ceiling effects (> 30%) for the Lysholm domains of limp, instability, support and locking [5]. Hence, this score might not be the first choice for the evaluation of outcome after APM. Even the KOOS has floor effects when used for meniscus issues [16]. In this study, the IKDC subjective score showed the best performance on all measurement properties. Unfortunately, the IKDC subjective score was not used in any of the RCTs about APM.

All studies were initially designed to determine the difference between arthroscopy and control treatment for knee OA, but later claimed that the two or three interventions were equivalent. Nevertheless, in many orthopedic studies, improvements have progressed without simultaneously addressing the significant ceiling effect common to many patient-related clinical outcome measures. Alternative statistical strategies such as equivalence or non-inferiority clinical trial designs are needed to circumvent this ceiling-effect problem [33].

Future randomized trials examining surgical procedures should make more effort to describe and standardize the surgical technique. Important surgical details such as the use of tourniquet, the experience of the surgeon, the portals and the use of photos or videos for documentation were only described in few studies. A surgical treatment as variable in a clinical trial is more complex than a pharmacological treatment where all patients of one group receive the same pill [18, 27, 28, 42]. Under this aspect, it is also of concern that the surgical process quality was controlled in none of the studies [42]. If the documentation had been given more attention, meniscal "root tears" should have been discovered and described in any of the studies. This is of concern because several studies have shown that the biomechanical effect of a root tear is comparable to a total meniscectomy [1, 13, 40, 41]. The root injury leads to meniscus extrusion and loss of circular hoop tension [40, 41]. Meniscus extrusion was stated in only one study. In this study, the rate of meniscus extrusion was 50% in the control group and 65% in the arthroscopy group [30]. Kijowski et al. [27] reported poorer clinical outcome when APM was associated with root tears and greater severity of meniscal extrusion.

All these limitations suggest that the results of the RCTs which were included in this systematic review should be interpreted with care and larger randomized trials without the described methodological flaws are needed to make a definite conclusion regarding the value of AT for knee OA in its various stages. Evidence based medicine (EBM) originated primarily in internal medicine. Adapting EBM better to the specifics of clinical orthopedic research should be taken in mind by the orthopedic community.

Conclusion

Despite all limitations, this systematic review shows that the majority of patients with knee OA might not benefit from arthroscopic surgery. Therefore, the indication for this procedure should be given with care. However, this review has also shown that there are subgroups of patients with knee OA who might benefit from AT. Patients who belong to one of these subgroups are people with non-traumatic flap tears of the medial meniscus. Furthermore, there is very low quality evidence that the removal of chondral flaps has a positive effect.

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Compliance with ethical standards

Conflict of interest The authors declare that they have no conflict of interest.

Ethical approval Our article does not contain any studies with human participants performed by any of the authors.

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Affiliations

Katrin Karpinski¹ · Ralf Müller-Rath² · Phillipp Niemeyer³ · Peter Angele⁴ · Wolf Petersen¹

Katrin Karpinski katrin.karpinski@t-online.de

- ³ OCM, Munich, Germany
- ⁴ Sporthopädicum, Straubing, Germany
- ¹ Martin Luther Krankenhaus, Caspar Theyß Str. 27-31, 14193 Berlin, Germany
- ² Orthopädische Gemeinschaftspraxis Neuss, Neuss, Germany