RESEARCH



Effects of manual lymphatic drainage on total knee replacement: a systematic review and meta-analysis of randomized controlled trials

Hongyuan Lu^{1,4†}, Quanwei Shao^{2†}, Wenyao Li^{3†}, Fei Li^{4,5}, Weiyi Xiong⁵, Kunpeng Li^{4*} and Wei Feng^{4,6*}

Abstract

Background Total knee joint replacement (TKR) is an effective method for the treatment of severe knee osteoarthritis. With an increasing number of surgeries, complications such as lower limb edema, pain, and limited mobility have caused a heavy burden. Manual lymphatic drainage (MLD) may be a solution to solve the problem. The study aims to evaluate the efficacy of MLD in reducing knee edema, pain, and improving range of motion (ROM) in patients after TKR.

Methods A search was conducted in PubMed, Embase, Cochrane Library, Web of Science, CNKI, VIPs, WanFang database, and Google Scholar from inception to June 2023. Only randomized controlled trials (RCTs) that compared the effects of MLD and non-MLD (or another physiotherapy) on improving knee edema, pain, and ROM after TKR were included. Stata 16.0 was used for meta-analysis. GRADE was used to assess the quality of evidence.

Results In total, 7 RCTs with 285 patients were identified. There were no significant differences found in the ROM of knee flexion (standardized mean difference (SMD) = 0.03, 95% confidence interval (Cl): -0.22, 0.28, P = 0.812) and the ROM of knee extension (SMD= -0.30, 95%CI: -0.64, 0.04, P = 0.084). No differences were observed in the lower extremity circumference after TKR (SMD= -0.09, 95%CI: -0.27, 0.09, P = 0.324). For postoperative pain, there was no significant advantage between the MLD and non-MLD groups (SMD= -0.33, 95%CI: -0.71, 0.04, P = 0.083).

Conclusions Based on the current evidence from RCTs, manual lymphatic drainage is not recommended for the rehabilitation of patients following total knee replacement.

Keywords Manual lymphatic drainage, Total knee replacement, Edema, Range of motion, Pain

[†]Hongyuan Lu, Quanwei Shao, and Wenyao Li contributed equally to this work.

*Correspondence: Kunpeng Li xyfyli@163.com Wei Feng fwginger@126.com ¹Guanghua Integrated Traditional Chinese and Western Medicine Hospital, Shanghai 200052, China
²Yangpu District Central Hospital, Shanghai 200090, China
³Shanghai No.4 Rehabilitation Hospital, Shanghai 200040, China
⁴School of Rehabilitation Science, Shanghai University of Traditional Chinese Medicine, Shanghai 201203, China
⁵Shanghai Fourth People's Hospital, Shanghai 200040, China
⁶The Second Rehabilitation Hospital of Shanghai, Shanghai 202441, China



© The Author(s) 2023. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/. The Creative Commons Dedication waiver (http://creativecommons.org/publicdomain/zero/1.0/) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

Introduction

Osteoarthritis of the knee is a prominent factor contributing to disability among adults aged 65 years old and over [1]. Progressive joint destruction is caused by osteoarthritis, which impacts all anatomical components of the joint [2]. End-stage knee osteoarthritis can be effectively treated with total knee replacement (TKR) [3]. An estimated 700,000 TKR are performed annually in the United States, with projections indicating that the number will increase to 3.5 million per year by 2030 [4]. The efficacy and affordability of TKR have been demonstrated in clinical studies [5]. However, a considerable proportion of individuals express dissatisfaction with the outcomes of TKR [6]. Firstly, untreated chronic swelling could exacerbate discomfort and lead to difficulties in mobility and flexibility [7]. Moreover, edema potentially increases the risk of infection in the impacted region, reduces the flow of blood, and impacts the elasticity of the arteries [8]. Individuals are greatly affected by these complaints, causing both physical and psychological harm.

Typically, it is advised to undergo some type of early rehabilitation (within 0–6 weeks) prior to being discharged from the hospital. Physiotherapy following TKR involves the utilization of techniques to alleviate pain and edema, targeted physical activities to enhance the flexibility of the joint, and exercises to improve muscle strength and endurance [9, 10]. Nevertheless, the current evidence for the use of early rehabilitation after TKR is limited, and some studies indicated that certain techniques like cryotherapy, compression, and pulsed electromagnetic fields have shown no impact on knee swelling [11, 12].

Manual lymphatic drainage (MLD) involves specialized rhythmic pumping techniques employed to massage the affected region and improve the flow of lymph [13]. MLD has the potential to decrease swelling by encouraging the pumping action of lymphangion and redirecting lymph away from stagnant areas to functional lymphatic vessels by reducing hydrostatic pressure [14]. Some studies reported the effectiveness of MLD in decreasing lymphatic swelling in females related to breast cancer [15, 16]. The application of MLD in musculoskeletal diseases has also garnered significant interest. Nevertheless, there are still some disputes regarding the effectiveness of MLD in patients receiving TKR. It has been shown in several randomized controlled trials (RCTs) that MLD, in combination with standard rehabilitation protocols, has positive effects on early edema and pain levels [17–19]. However, it has also been reported in several articles that the application of MLD in the early postoperative period after TKR does not reduce swelling [12, 20]. Can MLD be a routine therapy added following TKR or just be applied when the appearance of lower limb edema or pain? Despite the reporting of several RCTs that applied additional MLD after the surgery, there has been a lack of relevant meta-analysis and systematic review. An immediate study is necessary to assess whether MLD is more effective than standard routine rehabilitation or another physiotherapy in improving the following aspects: (1) lower limb circumferences or volume; (2) range of motion (ROM) of the knee joint; (3) knee joint pain.

Methods

The study was performed following relevant requirements suggested by the Cochrane Handbook [21] and was reported following the guidelines of the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA) statement [22]. The protocol has been registered on PROSPERO: CRD42023441423.

Search strategy

PubMed, Embase, Cochrane Library, Web of Science, CNKI, VIPs, WanFang database, and Google Scholar were utilized for the search from inception to June 2023. Specific databases were customized with tailored keywords and corresponding Medical Subject Headings (MeSH) terms for "Manual Lymphatic Drainage" AND "Arthroplasty, Replacement, Knee". All search terms were utilized due to the interchangeable use of different terms in the literature including TKR, TKA, knee arthroplasty, manual lymph drainage, decongestive lymphatic therapy, and Foldi or Vodder method. The detailed search strategy can be found in supplementary file 1. Additional eligible studies were manually examined by referring to reference lists obtained from retrieved articles. After removing duplicates, the titles and abstracts were reviewed independently by two authors (HYL and QWS) based on the inclusion criteria. In July 2023, the search was repeated in the same databases to identify any new trials or previously overlooked studies. Since MLD is mostly used for the treatment of postoperative lymphedema of breast cancer, such trials related to TKR were fewer, other resources were searched but were not limited to (1)monthly search of the Cochrane Controlled Trial Center Register (CENTRAL); (2) manual search of MLD related journals and major TKR conference documents; (3) contact researchers who are known to have participated in previous experiments to seek information about unpublished or incomplete experiments; (4) reference lists for review articles and related experiments in case of any possible omission.

Inclusion and exclusion criteria

Inclusion criteria (1) Population: patients who underwent TKA/TKR; (2) Intervention: application of MLD; (3) Comparison: no application of MLD or other physical therapy; (4) At least one of the following outcomes: knee edema (knee circumferences or volume); ROM of the

knee joint; pain and knee function. Only studies defined MLD as the application of a gentle massage technique by trained therapists, following the lymphatic pathways in a certain direction (from proximal to distal and then following the opposite way [23] or claimed to have followed the 'Foldi or Vodder method' were included. To meet the inclusion criteria, MLD needed to be employed as the primary intervention, not as a co-intervention with other physical therapy treatments in studies. In addition, if a study designed a comprehensive rehabilitation program, MLD had to possess over 50% of the program as the main treatment to be included.

The outcome measures included knee edema (knee circumferences or volume), ROM of the knee joint, and pain [using the Visual Analogue Scale (VAS), McGill Pain Questionnaire, and Numerical Rating Scale (NRS)]. For knee function, Knee Injury and Osteoarthritis Outcome Score (KOOS), EuroQol Five Dimensions Questionnaire (EQ-5D), Functional independence measurement (FIM), and Knee Society Score (KSS) were extracted.

Exclusion criteria (1) Reviews, meetings, and case reports; (2) Duplicate publications or studies with similar data; (3) Data that is incomplete, ambiguous, or unable to be extracted or compared.

Study selection

Two researchers (HYL and QWS) conducted separate searches of the databases and examined the titles and abstracts of the studies that were retrieved. Any ambiguities between the reviewers (HYL and QWS) were bought and discussed with the third reviewer (KPL). The eligible studies under the inclusion and exclusion criteria were carried out after reading the complete text.

Data extraction

HYL and QWS performed data extraction, and any disagreements were settled by KPL. Data extraction included: first author, publication year, country, sample size, the average age of control and experimental patients, diagnosis, study type, frequency and intervention time of control and experimental group, outcome assessments, and follow-up.

Risk of bias assessment

The risk of bias assessment was evaluated by two independent reviewers (HYL and QWS) respectively. If there were any disagreements to reach the final assessment, a third author (KPL) was consulted. Regarding RCTs, the Cochrane Risk of Bias tool ver.2 for RCTs (Cochrane RoB 2 tool) was employed [24]. Five domains of bias were assessed: (1) bias in selection; (2) bias in performance; (3) bias in attrition; (4) bias in detection; and (5) bias in selective outcome reporting [24, 25]. The Grading of Recommendations, Assessment, Development, and Evaluations approach (GRADE) was employed to assess the certainty of the evidence [26].

Statistical analysis

Stata 16.0 (STATA Corporation, Lakeway, Texas, USA) was utilized for all statistical analyses. Standardized mean differences (SMD) with 95% CIs for continuous outcomes were pooled. The I² statistic and Q test (χ^2) were used to quantify the heterogeneity of included studies, with a significance level of $P \le 0.050$ [27]. Based on the analysis of I^2 values, heterogeneity may not be important: 0–40%; moderate heterogeneity: 30-60%; substantial heterogeneity: 50-90%, and considerable heterogeneity: 75-100% [21]. To account for clinical heterogeneity, a randomeffects model was employed to combine outcome data. If possible to check various factors affecting MLD, a subgroup analysis may be performed according to (1) the time point and dosage for starting MLD after surgery; (2) different control interventions (other forms of exercises, kinesio tape, mixed physical therapy techniques or no intervention). Separate meta-analyses should be performed for each subgroup.

Given the inadequate quantity of research conducted, publication bias was evaluated through the utilization of a funnel plot and further assessed using the linear regression tests of Egger and Begg [28]. A sensitivity analysis is presented. Besides, there were reports of potential conflicts of interest and disclosure of the funding source.

Results

Literature search

The detailed literature screening process is shown in Fig. 1. A total of 150 articles were retrieved, after removing duplicates from 39 citations, there were 111 articles remaining. Then, 61 studies including 18 case reports, 21 irrelevant articles, 17 systematic reviews, and 5 pilot studies were excluded. After that, 50 studies were thought to be potentially eligible and 43 articles were excluded for reasons. Finally, 7 RCTs from 5 countries were confirmed after full-text review and included in the systematic review and meta-analysis.

Study characteristics

The basic characteristics of the included studies are summarized in Table 1. Among the 7 included trials, 3 studies were conducted in Turkey [12, 17, 18] and one each in Switzerland [20], Australia [29], Japan [30], and Italy [31]. The included trials were published in English between 2013 and 2022, with a total of 285 patients enrolled. There were 6 of 7 trials that compared MLD with non-MLD and one trial compared MLD with kinesiotape. The age of the patients ranged from 48 to 89, with an average age (standard deviation) of 54.38 (9.81). 5 of 7 studies

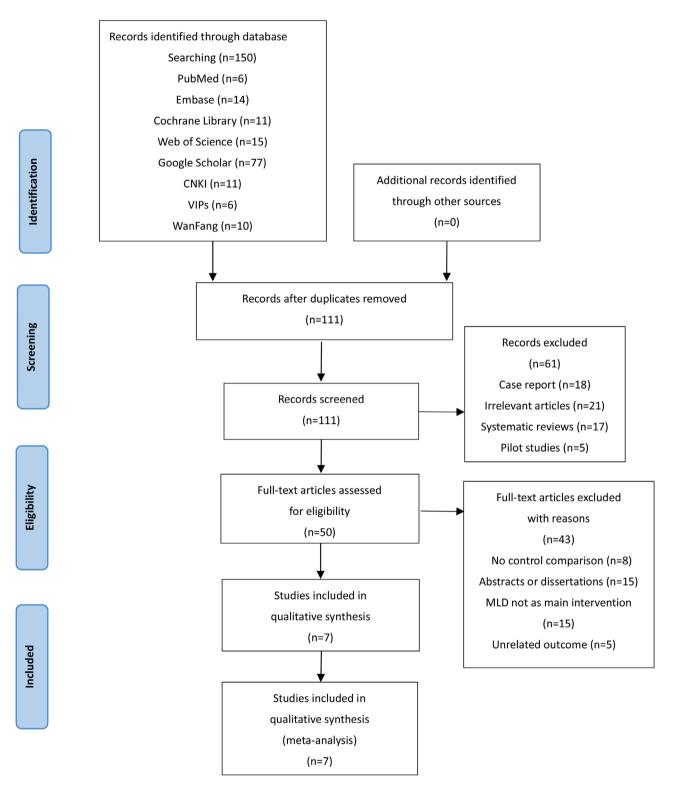
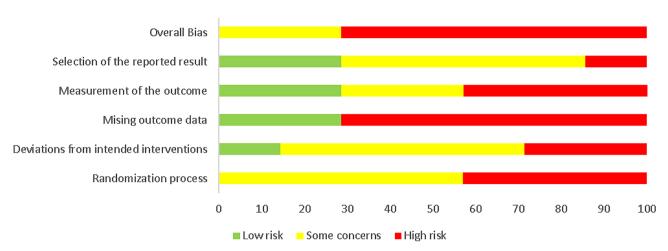


Fig. 1 The flow diagram of study selection

included 228 patients (80%) who had a primary diagnosis of knee osteoarthritis before the surgery while 57 patients of 2 studies had unclear diagnoses before surgery. All patients in the included studies underwent TKR/TKA by an experienced orthopedic surgeon. In all 7 RCTs, MLD treatment was initiated 2 days after surgery. The duration of the MLD treatment course was reported as 2–4 days postoperative in 2 studies (28.6%), over 6 days of

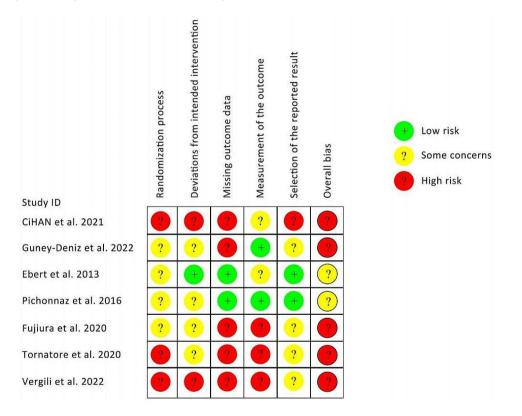
First Author, Year	Country of	Intervention	Control	Intervention	Control group	Intervention time	Follow up time	Outcomes measures
	study	group(Sample Size)	group(Sample Size)	group				
Vergili et al., 2022 [12]	Turkey	8	8	MLD+ physical therapy	NO MLD+ physi- cal therapy	30 min/day on the second and fourth day after surgery	D2, D4 and D6 after surgery	ROM; pain:VAS; edema: knee volume; function: FIM
CiHAN et al., 2021 [17]	Turkey	11	11	MLD + physical therapy	NO MLD+ physi- cal therapy	NO MLD + physi- No technical details cal therapy	D3 and 6 weeks after surgery	pain:VAS; function:NHP; fear:TKS
Guney-Deniz et al. 2022 [18]	Turkey	13	15	MLD+ exercises	NO MLD + exercises	30 min/day, 2–4 days after surgery	D2, D3, D4, 2 weeks and 6 weeks after surgery	ROM; pain:VAS; function:KOOS; edema:knee circumference
Pichonnaz et al., 2016 [20]	Switzerland	29	27	MLD+ physical therapy	NO MLD+ physi- cal therapy	NO MLD+ physi- 30 min/day, 2–7 days after cal therapy surgery	D2, D7, and 3 months after surgery	ROM; pain:VAS; function:KSS,WOMAC; edema:BIS limb volume
Ebert et al., 2013 [29]	Australia	24	26	MLD+physical therapy	NO MLD+ physi- cal therapy	NO MLD+ physi- 30 min/day, 2–4 days after cal therapy surgery	D2, D3, D4, and 6 weeks after surgery	ROM; pain:NRS; function:KOOS; edema:knee girths
Fujiura et al., 2020 [30]	Japan	20	20	MLD+ physical therapy	NO MLD + physi- cal therapy	20 min/day, 2–6 days after surgery	Immediately and the fifth MLD application after surgery	ROM; pain:VAS; edema:knee circumference
Tornatore et al., 2020 [31]	Italy	33	33	MLD+education Kinesio tape+e	Kinesio tape + education	Kinesio 30 min/day on the second tape + education and fourth day after surgery	D4, D6 after surgery	ROM; pain:NRS; edema:knee circumference
Note: MLD: Manual lymphatic drainage; VAS: Visual Analogue Scale; NRS: Score; NHP: Nottingham Health Profile; BIS: Bioimpedance Spectroscop)	natic drainage; VAS: Health Profile; BIS: B	Visual Analogue Sca 3ioimpedance Spect	ile; NRS: Numeric Ra roscopy; WOMAC: M	ting Scale; FIM: Funct /estern Ontario and <i>I</i>	tional independence McMaster Universitie	Note: MLD: Manual lymphatic drainage; VAS: Visual Analogue Scale; NRS: Numeric Rating Scale; FIM: Functional independence measurement; TKS: Tampa Kinesiophobia Scale; KOOS: Score; NHP: Nottingham Health Profile; BIS: Bioimpedance Spectroscopy, WOMAC: Western Ontario and McMaster Universities Osteoarthritis Index; KSS: Knee Society Score; D: Day	iophobia Scale; KOOS: Kne : Society Score; D: Day	Numeric Rating Scale; FIM: Functional independence measurement; TKS: Tampa Kinesiophobia Scale; KOOS: Knee Injury and Osteoarthritis Outcome ; WOMAC: Western Ontario and McMaster Universities Osteoarthritis Index; KSS: Knee Society Score; D: Day

Table 1General characteristics of the included studiesFirst Author, YearCountry ofInterventionCo



As percentage (intention-to-treat)

Fig. 2 Risk of bias graph showing each risk of bias item as percentages across all included studies





treatment in 2 studies (28.6%), the second day postoperative in one study (14.3%), and the second and fourth day postoperative in 2 studies (28.6%).

Risk of bias assessment

A summary of the risk of bias assessment is presented in Fig. 2. Figure 3 provides the bias risk for each study included, as well as for each domain. In general, most of the studies (71%; n=5) were judged as 'high risk of bias' whereas two studies (29%) were considered to have 'some concern'. An inadequate randomized sequence and allocation concealment of three studies were considered to be 'high risk'. The majority of studies (71%) failed to sufficiently balance baseline covariates among the groups. Five studies did not report blinding of outcome assessments or provide information about its absence. In addition, only three studies (42.8%) registered their protocol following a prespecified analysis plan. Refer to supplementary file 2 for detailed information regarding the quality of evidence for the measured outcomes.

Funding and conflicts of interest

Two studies were funded by university or hospital research foundations [20, 29]. Two studies [12, 18] did not receive any forms of foundations while the other three studies were unclear [17, 30, 31]. The authors of six studies [12, 18, 20, 29–31] declared no conflicts of interest, and one study [17] had no official disclosure of conflicts of interest report.

Effects of interventions

ROM of the knee joint

Six RCTs [12, 18, 20, 29–31] involving 256 patients provided data on the ROM of the knee joint. The follow-up time is mostly concentrated on the fourth day after the operation [12, 18, 29, 31], and the seventh day in one RCT [20]. Six RCTs [12, 18, 20, 29–31] compared the ROM of knee joint flexion, and three RCTs [18, 20, 29] compared the ROM of knee joint extension. The heterogeneity was low ($I^2=0\%$, P=0.636), which was analyzed using a fixed-effects model. The MLD therapy demonstrated no significant advantage over non-MLD groups both in knee flexion (SMD=0.03, 95%CI: -0.22, 0.28, P=0.812, Fig. 4) and knee extension (SMD=-0.30, 95%CI: -0.64, 0.04, P=0.084, Fig. 5).

Pain after Surgery

Seven RCTs [12, 17, 18, 20, 29–31] with 278 patients reported postoperative pain. The follow-up time of four

RCTs was the fourth day after TKR [12, 18, 29, 31], the third day after the operation [17], the seventh day after the operation in one study [20], and after the fifth application of MLD in one study [30]. Five RCTs [12, 17, 18, 20, 30] used the Visual Analogue Scale (VAS) and two RCTs [29, 31] used the Numeric Rating Scale (NRS). Subgroup analysis of the knee joint pain was conducted based on different measurements. The heterogeneity was moderate (I^2 =55.8%, *P*=0.035), which was analyzed using a random-effects model. Patients treated with MLD had no benefit in relieving pain after TKR than non-MLD groups (SMD=-0.33, 95%CI: -0.71, 0.04, *P*=0.083, Fig. 6).

Knee edema

The knee edema measurements included thigh, calf, and ankle circumferences. Four RCTs [18, 29-31] with 184 patients reported lower limb lymphedema. Four RCTs [18, 29-31] provided data on thigh circumferences, and three RCTs [18, 29, 31] provided calf and ankle circumferences. The heterogeneity was low ($I^2=0\%$, P=0.889), which was analyzed using a fixed-effects model. No significant differences were found between the MLD and non-MLD groups in terms of thigh circumferences (SMD= -0.15, 95%CI: -0.44, 0.14, P=0.302), calf circumferences (SMD= -0.04, 95%CI: -0.37, 0.29, P=0.813) and ankle circumferences (SMD= -0.06, 95%CI: -0.39, 0.26, P=0.701). The final results showed that compared with non-MLD groups, MLD did not improve knee edema after TKR (SMD= -0.09, 95%CI: -0.27, 0.09, P=0.324, $I^2=0\%$, Fixed Effect Model, Fig. 7).

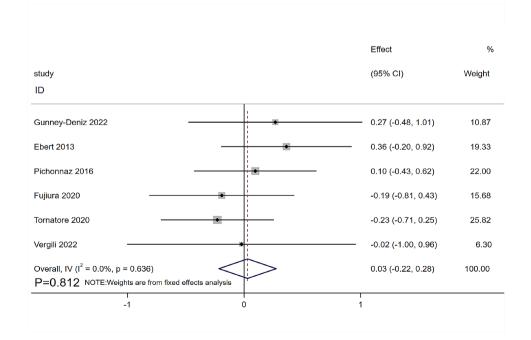


Fig. 4 Forest plot of knee flexion range of motion

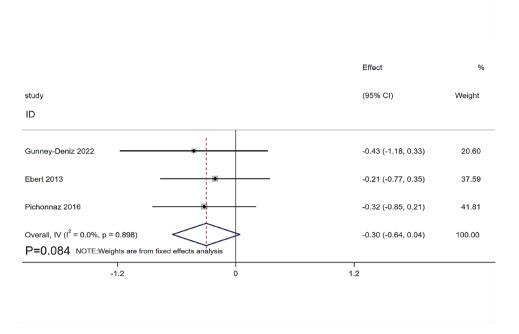


Fig. 5 Forest plot of knee extension range of motion

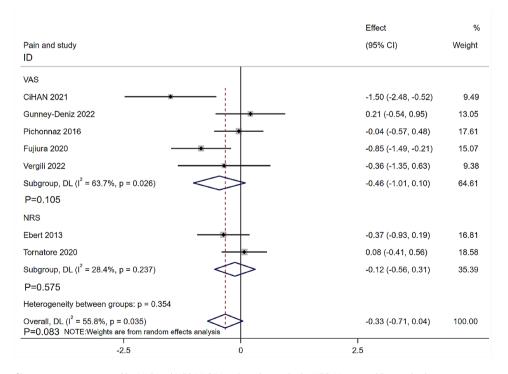


Fig. 6 Forest plot of knee joint pain assessed by VAS and NRS. VAS: Visual Analogue Scale; NRS: Numerical Rating Scale

Evaluation of publication bias

There was no clear evidence of publication bias when examining the funnel plot (Fig. 8) for the impact of MLD compared to non-MLD groups on postoperative pain. The results of Egger's test showed no publication bias on knee joint pain (P=0.200). Additionally, a sensitivity analysis was conducted to evaluate the stability of the

combined findings. The sensitivity analysis indicated that the results were statistically reliable (Fig. 9) after sequentially omitting each study and the outcomes were not altered.

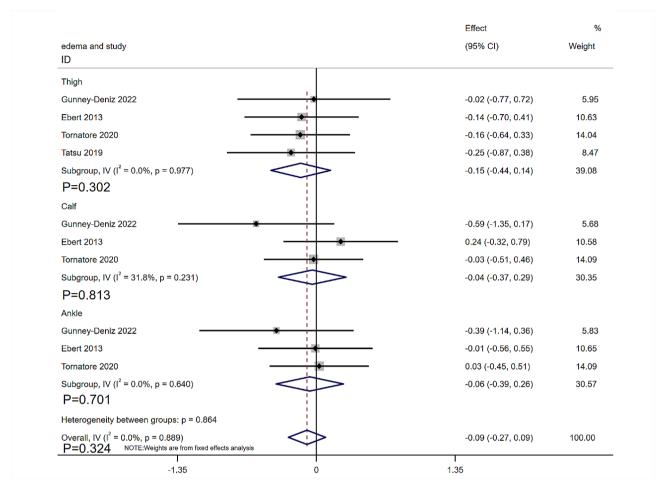


Fig. 7 Forest plot of knee edema assessed by thigh, calf, and ankle circumferences

Discussion

Seven RCTs assessing the efficacy of MLD in patients with pain, limited ROM, and lower limb edema after TKR were included. The findings of the current study indicated that MLD did not demonstrate significant reductions in lower limb edema, pain, or improve joint mobility when compared to the absence of MLD or other physiotherapy interventions after TKR.

Furthermore, previous studies also indicated that early MLD for patients undergoing breast cancer surgery cannot significantly reduce or prevent lymphedema [13, 32, 33]. Additionally, a separate systematic review expressed uncertainty regarding the effectiveness of MLD [34]. However, Provencher AM et al. [35] have provided support for the use of MLD in reducing swelling and pain, improving mobility, and enhancing patient satisfaction and quality of life following musculoskeletal injuries. Gutiérrez-Espinoza H's systematic review and meta-analysis [36] concluded that the inclusion of mobilization with movement and MLD resulted in significant improvements in the wrist, upper limb function, and hand edema among individuals diagnosed with distal radius fracture. Klein et al. [7] revealed that only the application of multiple layers and prolonged compression effectively decreases swelling following orthopedic injury or surgical procedures. Schingale et al. [37] conducted a narrative review and found that MLD has the potential to be utilized for the symptomatic management of different diseases, including multiple sclerosis and Parkinson's disease. The potential effectiveness of MLD in animal models was shown in three animal experiments [38–40], but its applicability in humans is currently uncertain.

There is a lack of consensus among published studies regarding the efficacy of MLD therapy. Most studies focused on the application of MLD for treating lymphedema in females following breast cancer surgery. Apart from the negative outcomes indicated in the aforementioned studies [13, 32, 33], Ezzo J et al. [41] included six trials and the findings were contradictory for function (ROM). Regardless of the treatment they received, 60–80% of participants reported improved sensations, including pain relief and reduced heaviness. One RCT revealed that standard therapy and MLD in combination with multilayer compressive bandage treatment

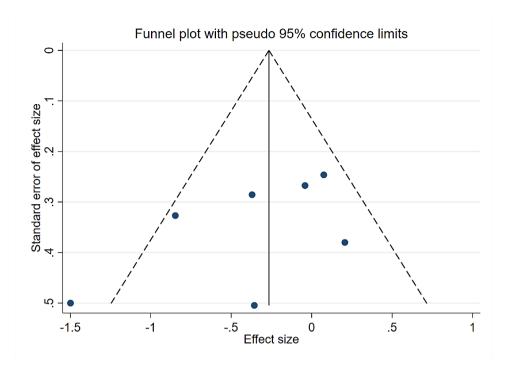


Fig. 8 Funnel plot of knee joint pain after TKR surgery. TKR: Total knee replacement

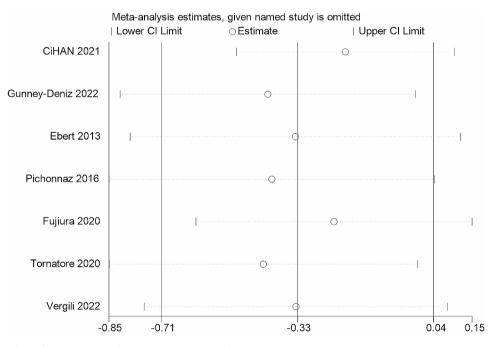


Fig. 9 Sensitivity analysis of knee joint pain after TKR surgery. TKR: Total knee replacement

were both effective [42]. Nevertheless, there were no further effects observed in terms of a decrease in arm volume percentage during the period of intensive treatment after breast cancer surgery. On the contrary, a systematic review [43] indicated that MLD may yield positive outcomes in terms of reducing volume, enhancing quality of life, and alleviating symptoms following breast cancer surgery. However, the author also indicated that MLD may not provide additional benefits for patients with moderate to severe lymphedema. Doubblestein and colleagues' review [44] suggested the utilization of MLD to reduce the incidence of lymphedema in early rehabilitation post-surgery. The author included 5 studies (192 patients) focusing on acute musculoskeletal disorders

and subacute edema. In contrast with the present systematic review and meta-analysis, the author found that MLD could potentially be beneficial in reducing edema and enhancing ROM when combined with auxiliary therapies. Nevertheless, an absence of ample evidence and high-quality RCTs to endorse the utilization of MLD in enhancing the overall clinical manifestation.

The present review conducted the comparison between MLD and non-MLD (or another physiotherapy), with only one RCT utilizing MLD and kinesiotape for comparison. The effectiveness between MLD and other compression therapy remained unclear. One systematic review and meta-analysis evaluating the effects of MLD and compression therapy found that compression bandaging was effective in managing breast cancer-related lymphedema [13]. One RCT conducted by Rigoni et al. [45] compared MLD and connective tissue techniques for the treatment of post-surgical inflammation in patients with TKA. The findings indicated that both approaches had a beneficial impact on the outcomes and demonstrated comparable effectiveness in regulating the erythrosedimentation rate (ESR) for both methods. Bertinchamp et al. [19] found that both MLD and kinesiotape improved ROM and decreased pain, but MLD demonstrated greater quality of life scores in patients after TKA. Preneuf-Pauthiera et al. [46] concluded that no significant statistical difference was found between MLD and usual physiotherapy regarding ROM, but potential effects on decreasing pain were found in the MLD group. Given the trend of pain reduction, the author considered a large sample and the type of edema (superficial or deep) should be taken into consideration to verify this finding.

The meta-analysis of the results related to knee joint pain showed moderate heterogeneity, while the heterogeneity in the remaining outcomes was relatively low. The heterogeneity observed among the knee joint pain examined may be attributed to differences in the clinical setting and variations in measurement techniques. Firstly, the included studies differed in terms of the duration of MLD usage. Five studies [12, 18, 20, 29, 31] applied MLD lasting 30 min, one study [30] applied MLD lasting 20 min and one study [17] did not specify the technical details of the MLD method. Secondly, the use of MLD was mainly applied between 2 and 6 days while two studies [12, 31] applied MLD on the second and fourth day after the surgery. Finally, the outcomes could be affected by the experience of the physiotherapist or the specific operation of the MLD method. Four studies [12, 18, 29, 30] listed the specific process of the MLD method and tended to be similar. One study [20] claimed that their research was based on the recommendations of Földi and Kubik [47]. While the other one [17] claimed that their MLD was performed by a physiotherapist with a medical certificate. However, another study [31] did not specify the method used for MLD.

Clinical implications

Swelling following TKR leads to pain, inflammation, limited mobility, hindered quadriceps function, and a negative perception of recovery [48]. The prevention and treatment of initial edema after TKR is very important. The strength of this research lies in the potential benefit of MLD as a singular method to relieve patients' burdens, since MLD is convenient and economical. The study fills the blank of the meta-analysis and systematic review of the efficacy of MLD for patients receiving TKR. However, the effect sizes did not reach a statistical difference when using MLD compared to not using MLD, which could be attributed to the small sample size of the included studies, with only one RCT involving over 60 participants [31]. Large-scale, high-sample clinical trials in the future may provide valuable evidence to support the widespread application of MLD in clinical practice. Furthermore, studies investigating MLD use alone (without concurrent treatments) were limited [49], making it challenging to obtain conclusive evidence supporting the standalone efficacy of MLD as a swelling reduction technique. In summary, based on the available RCT evidence, there is insufficient support for MLD as a stand-alone edema reduction technique after TKR in terms of reducing lower limb edema, pain and improving ROM of the knee joint.

Limitations and future scope

Certain limitations should be noted in the current study. Firstly, the total samples of the included studies were small, of which one study only recruited 8 patients in each group [12]. Secondly, only one of the included studies involved kinesiotape as a control group [31], making it difficult to be generalized to other forms of compression therapy. Finally, the follow-up periods in the majority of the studies were relatively short, mainly concentrated within the first week after surgery [12, 30, 31]. To address these limitations, future research is needed to focus on larger sample sizes, more rigorous, and higher-quality RCTs of a single application of MLD. Furthermore, future studies could be conducted to explore the inclusion of other compression techniques in the control group and compare these with MLD. Lastly, further studies are necessary to investigate the long-term efficacy of MLD on postoperative edema and pain after TKR.

Conclusion

Manual Lymphatic Drainage (MLD) does not significantly decrease lymphedema, enhance range of motion, or alleviate pain following Total Knee Replacement (TKR). Consequently, further high-quality research is imperative to accurately evaluate the potential advantages of integrating MLD with standard rehabilitation protocols or other compression techniques in TKR patients.

Abbreviations

TKR	Total knee replacement
MLD	Manual lymphatic drainage
ROM	Range of motion
VAS	Visual analog scale
NRS	Numerical Rating Scale
FIM	Functional independence measurement
TKS	Tampa Kinesiophobia Scale
KOOS	Knee Injury and Osteoarthritis Outcome Score
NHP	Nottingham Health Profile
BIS	Bioimpedance Spectroscopy
WOMAC	Western Ontario and McMaster Universities Osteoarthritis Index
KSS	Knee Society Score

Supplementary Information

The online version contains supplementary material available at https://doi. org/10.1186/s12891-023-07153-8.

Supplementary File 1: Searching strategy for Each Database

Supplementary File 2: Grade evidence for the outcomes

Acknowledgements

Not applicable.

Author contributions

HYL, QWS, and WYL conceived the initial concept for the study and suggested the study's design. The literature search was conducted by HYL, QWS, and KPL, who also screened and selected the studies that were initially identified. HYL, QWS, and KPL assessed and appraised the caliber of the studies incorporated. FL and WYX conducted the meta-analysis. KPL and WF edited and revised the manuscript. All the authors agreed to be accountable for all aspects of this work.

Funding

Not applicable.

Data availability

All data and materials are contained within the manuscript.

Declarations

Ethics approval and consent to participate Not applicable.

Consent for publication

Not applicable.

Competing interests

The authors declare no competing interests.

Received: 1 August 2023 / Accepted: 26 December 2023 Published online: 02 January 2024

References

- 1. Neogi T. The epidemiology and impact of pain in osteoarthritis. Osteoarthritis Cartilage. 2013;21(9):1145–53.
- Nguyen C, Boutron I, Roren A, Anract P, Beaudreuil J, Biau D, et al. Effect of Prehabilitation before total knee replacement for knee osteoarthritis on functional outcomes. JAMA Netw Open. 2022;5(3):e221462.

- Carr AJ, Robertsson O, Graves S, Price AJ, Arden NK, Judge A, et al. Knee replacement. Lancet. 2012;379(9823):1331–40.
- Kurtz S, Ong K, Lau E, Mowat F, Halpern M. Projections of primary and revision hip and knee arthroplasty in the United States from 2005 to 2030. J Bone Jt Surg Am. 2007;89(4):780–5.
- Nwachukwu BU, Bozic KJ, Schairer WW, Bernstein JL, Jevsevar DS, Marx RG, et al. Current status of cost utility analyses in total joint arthroplasty: a systematic review. Clin Orthop Relat Res. 2015;473(5):1815–27.
- Bourne RB, Chesworth BM, Davis AM, Mahomed NN, Charron KDJ. Patient satisfaction after total knee arthroplasty: who is satisfied and who is not? Clin Orthop Relat Res. 2010;468(1):57–63.
- Klein I, Tidhar D, Kalichman L. Lymphatic treatments after orthopedic Surgery or injury: a systematic review. J Bodyw Mov Ther. 2020;24(4):109–17.
- Földi E, Földi M, Weissleder H. Conservative treatment of lymphoedema of the limbs. Angiology. 1985;36(3):171–80.
- Naylor J, Harmer A, Fransen M, Crosbie J, Innes L. The status of physiotherapy rehabilitation following total knee replacement in Australia. Physiother Res Int. 2006;11(1):35–47.
- Brander VA, Stulberg SD, Chang RW. Rehabilitation following hip and knee arthroplasty. Phys Med Rehabil Clinics North Am. 1994;5(4):815–36.
- Han ASY, Nairn L, Harmer AR, Crosbie J, March L, Parker D, et al. Early rehabilitation after total knee replacement Surgery: a multicenter, noninferiority, randomized clinical trial comparing a home exercise program with usual outpatient care. Arthritis Care Res (Hoboken). 2015;67(2):196–202.
- Vergili Ö, Canbeyli İD, Özsar BK, Oktaş B, Keskin S. The effect of manual lymphatic drainage on the postoperative recovery process following total knee arthroplasty. J Med Palliat Care. 2022;3(1):66–70.
- Huang TW, Tseng SH, Lin CC, Bai CH, Chen CS, Hung CS, et al. Effects of manual lymphatic drainage on breast cancer-related lymphedema: a systematic review and meta-analysis of randomized controlled trials. World J Surg Oncol. 2013;11:15.
- 14. Leduc O, Leduc A, Bourgeois P, Belgrado JP. The physical treatment of upper limb edema. Cancer. 1998;83:2835–9.
- Williams AF, Vadgama A, Franks PJ, Mortimer PS. A randomized controlled crossover study of manual lymphatic drainage therapy in women with breast cancer-related lymphedema. Eur Eur J Cancer Care. 2002;11(4):254–61.
- Williams A. Manual lymphatic drainage: exploring the history and evidence base. Br J Community Nurs. 2010;15(4):18–24.
- CİHAN E, YILDIRIM NÜN, BİLGE O, BAKAR Y, DORAL MN. Outcomes with additional manual lymphatic drainage to rehabilitation protocol in primary total knee arthroplasty patients: preliminary clinical results. J Health Sci. 2021;12(3):319–29.
- Guney-Deniz H, Kinikli GI, Aykar S, Sevinc C, Caglar O, Atilla B, et al. Manual lymphatic drainage and Kinesio taping applications reduce early-stage lower extremity edema and pain following total knee arthroplasty. Physiother Theory Pract. 2023;39(8):1582–90.
- Bertinchampa U, Brossardb V, Faouzi M. Comparison between the effectiveness of manual lymph drainage versus Kinesio Taping® on postoperative edema after total knee replacement Surgery: a randomized controlled trial. Kinesither Rev. 2021;21(234):22–35.
- Pichonnaz C, Bassin JP, Lécureux E, Christe G, Currat D, Aminian K, et al. The effect of manual lymphatic drainage following total knee arthroplasty: a randomized controlled trial. Arch Phys Med Rehabil. 2016;97(5):674–82.
- 21. Higgins JPT, Altman DG. Assessing risk of bias in included studies. Cochrane Handb Syst Reviews Interventions. 2008:187–241.
- Moher D, Liberati A, Tetzlaf J, Altman DG. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. J Clin Epidemiol. 2009;62(10):1006–12.
- Howard SB, Krishnagiri S. The use of manual edema mobilization for the reduction of persistent edema in the upper limb. J Hand Ther. 2001;14(4):291–301.
- Sterne JAC, Savović J, Page MJ, Elbers RG, Blencowe NS, Boutron I, et al. Rob 2: a revised tool for assessing risk of bias in randomised trials. BMJ. 2019;366:14898.
- Mansournia MA, Higgins JPT, Sterne JAC, Hernán MA. Biases in randomized trials: a conversation between trialists and epidemiologists. Epidemiology. 2017;28(1):54–9.
- Guyatt GH, Oxman AD, Vist GE, Kunz R, Falck-Ytter Y, Alonso-Coello P, et al. Grade: an emerging consensus on rating quality of evidence and strength of recommendations. BMJ. 2008;336(7650):924–6.
- 27. Grindem H, Mansournia MA, Øiestad BE, Ardern CL. Was it a good idea to combine the studies? Why clinicians should care about heterogeneity

when making decisions based on systematic reviews. Br J Sports Med. 2019;53(7):399–401.

- Harbord RM, Harris RJ, Sterne JA. Updated tests for small-study effects in meta-analyses. Stata J. 2009;9(2):197–210.
- 29. Ebert JR, Joss B, Jardine B, Wood DJ. Randomized trial investigating the efficacy of manual lymphatic drainage to improve early outcome after total knee arthroplasty. Arch Phys Med Rehabil. 2013;94(11):2103–11.
- Fujiura T, Nagasawa H, Wakabayashi H. Effect of manual lymph drainage for up to 10 days after total knee arthroplasty: a randomized controlled trial. Phys Ther Res. 2020;23(1):39–46.
- Tornatore L, De Luca ML, Ciccarello M, Benedetti MG. Effects of combining manual lymphatic drainage and kinesiotaping on pain, edema, and range of motion in patients with total knee replacement: a randomized clinical trial. Int J Rehabil Res. 2020;43(3):240–6.
- Lin Y, Yang Y, Zhang X, Li W, Li H, Mu D. Manual lymphatic drainage for breast Cancer-related Lymphedema: a systematic review and Meta-analysis of Randomized controlled trials. Clin Breast Cancer. 2022;22(5):e664–73.
- Liang M, Chen Q, Peng K, Deng L, He L, Hou Y, et al. Manual lymphatic drainage for lymphedema in patients after Breast cancer Surgery: a systematic review and meta-analysis of randomized controlled trials. Med (Baltim). 2020;99(49):e23192.
- Wanchai A, Armer JM. Manual Lymphedema drainage for reducing risk for and managing breast Cancer-related Lymphedema after breast Surgery: a systematic review. Nurs Women Health. 2021;25(5):377–83.
- Provencher AM, Giguère-Lemieux É, Croteau É, Ruchat SM, Corbin-Berrigan LA. The use of manual lymphatic drainage on clinical presentation of musculoskeletal injuries: a systematic review. Complement Ther Clin Pract. 2021;45:101469.
- Gutiérrez-Espinoza H, Araya-Quintanilla F, Olguín-Huerta C, Valenzuela-Fuenzalida J, Gutiérrez-Monclus R, Moncada-Ramírez V. Effectiveness of manual therapy in patients with distal radius fracture: a systematic review and meta-analysis. J Man Manip Ther. 2022;30(1):33–45.
- Schingale FJ, Esmer M, Küpeli B, Ünal D. Investigation of the less known effects of Manual Lymphatic drainage: a narrative review. Lymphat Res Biol. 2022;20(1):7–10.
- Déry MA, Yonuschot G, Winterson BJ. The effects of manually applied intermittent pulsation pressure to a rat ventral thorax on lymph transport. Lymphology. 2000;33(2):58–61.
- Knott EM, Tune JD, Stoll ST, Downey HF. Increased lymphatic flow in the thoracic duct during manipulative intervention. J Am Osteopath Assoc. 2005;105(10):447–56.

- Hodge LM, King HH, Williams AG Jr, Reder SJ, Belavadi T, Simecka JW, et al. Abdominal lymphatic pump treatment increases leukocyte count and flux in thoracic duct lymph. Lymphat Res Biol. 2007;5(2):127–33.
- Ezzo J, Manheimer E, McNeely ML, Howell DM, Weiss R, Johansson KI, et al. Manual lymphatic drainage for lymphedema following breast cancer treatment. Cochrane Database Syst Rev. 2015;(5):CD003475.
- Sen El, Arman S, Zure M, Yavuz H, Sindel D, Oral A. Manual lymphatic drainage may not have an additional effect on the intensive phase of breast Cancerrelated Lymphedema: a Randomized Controlled Trial. Lymphat Res Biol. 2021;19(2):141–50.
- Thompson B, Gaitatzis K, Janse de Jonge X, Blackwell R, Koelmeyer L. Manual lymphatic drainage treatment for lymphedema: a systematic review of the literature. J Cancer Surviv. 2021;15(2):244–58.
- Doubblestein B, Sublett S, Huang M. Effects of manual lymphatic drainage techniques on conditions affecting the musculoskeletal system: a systematic review. Orthop Pract. 2020;32(2):92–101.
- 45. Rigoni S, Tagliaro L, Baù D, Scapin M. Effectiveness of two rehabilitation treatments in the modulation of inflammation during the acute phase in patients with knee prostheses and assessment of the role of the diet in determining post-surgical inflammation. J Orthop. 2021;25:237–43.
- 46. Preneuf-Pauthiera B, Redona V, Marchata N, Toniolo J, Rousseau V, Delaide V, et al. Impact of a manual veno-lymphatic stimulation according to the Schiltz method on the recovery of knee mobility following total knee arthroplasty: a randomized controlled trial. Kinesither Rev. 2022;22(250):14–20.
- 47. Földi M, Kubik S. Treatment of the leg. Textbook of lymphology, for Physicians and Lymphedema therapists. 3rd ed. München: Urban & Fischer; 2005.
- Yau LK, Henry FU, Man Hong C, Amy C, Wai Kwan Vincent C, Ping Keung C, et al. Swelling assessment after total knee arthroplasty. J Orthop Surg (Hong Kong). 2022;30(3):10225536221127668.
- Gradalski T, Ochalek K, Kurpiewska J. Complex decongestive lymphatic therapy with or without Vodder II Manual Lymph Drainage in more severe chronic Postmastectomy Upper Limb Lymphedema: a randomized noninferiority prospective study. J Pain Symptom Manage. 2015;50(6):750–7.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.