The effectiveness of non-pharmacological sleep interventions for people with chronic pain: a systematic review and meta-analysis

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Abstract

Objective: About two thirds of people with chronic pain report problems sleeping. We aimed to evaluate the effectiveness of non-pharmacological sleep interventions for improving sleep in people with chronic pain.

Design: We conducted a systematic review of non-pharmacological and non-invasive interventions to improve sleep quality or duration for adults with chronic non-cancer pain evaluated in a randomised controlled trial. Our primary outcome of interest was sleep; secondary outcomes included pain, health-related quality of life, and psychological wellbeing. We searched the Cochrane Library, MEDLINE, Embase, PsycINFO and CINAHL from inception to April 2020. After screening, two reviewers evaluated articles and extracted data. Meta-analysis was conducted using a random effects model. Risk of bias was assessed with the Cochrane tool.

Results: We included 42 trials involving 3346 people randomised to 94 groups, of which 56 received an intervention targeting sleep. 10 studies were of fair and 32 of good methodological quality. Overall risk of bias was judged to be low in 11, high in 10 and unclear in 21 studies. In 9 studies with 385 people randomised, cognitive behavioural therapy for insomnia showed benefit post-treatment compared with controls for improved sleep quality, standardised mean difference $\text{−1.23 (95%CI −1.76, −0.70; p < 0.00001)}$. The effect size was only slightly reduced in meta-analysis of 3 studies at low risk of bias. The difference between groups was lower at 3 and 6 months after treatment but still favoured cognitive behavioural therapy for insomnia. Pain, anxiety and depression were reduced post-treatment, but evidence of longer term benefit was lacking. There was no evidence that sleep hygiene interventions were effective in improving sleep and there was some evidence in comparative studies to suggest that cognitive behavioural therapy for insomnia was more effective than sleep hygiene.

Numerous other interventions were evaluated in small numbers of studies, but evidence was insufficient to draw conclusions about effectiveness.

Conclusions: Cognitive behavioural therapy for insomnia is an effective treatment to improve sleep for people with chronic pain, but further high-quality primary research is required to explore combined CBT content that will ensure additional improvements to pain, quality of life and psychological health and longer-term maintenance of benefits. Primary research is also needed to evaluate the effectiveness of interventions for which insufficient evidence exists.

Trial registration: PROSPERO registration number: CRD42019093799.

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Background
While chronic pain may be a primary complaint or secondary to an underlying disease, it is now recognised as a health condition in its own right [1] requiring specific therapy and rehabilitation [2]. In the UK, chronic pain affects between a third and a half of the population and about 10–14% of people report moderate to severely disabling chronic pain [3]. For people affected and their families, chronic pain is associated with a reduced quality of life and impacts on work and social life [4–6]. About two thirds of people with chronic pain report problems sleeping [7], including difficulties falling asleep, staying asleep, or waking early [6, 8–10], and this is evident across a range of conditions associated with chronic pain. In a large US population, 89% of people with chronic pain caused by fibromyalgia reported one, and 63% reported two or more symptoms of sleep disturbance [11]. Sleep disorders are common in people with multiple sclerosis [12, 13], rheumatoid arthritis [14], and osteoarthritis [15–17], with about 60–75% of people affected. Sleep disturbance is greater in people with more severe osteoarthritis symptoms [15, 18]. Other pain conditions with associated sleep disturbance include migraine and frequent headache [19–21], and low back [8, 22] and neck pain [23].

The relationship between sleep and pain is bidirectional [24–27]. Reduced sleep leads to greater pain, and greater pain has a negative impact on sleep. Poor sleep is also associated with the development of chronic pain [24]. In a large Norwegian cohort, women with three symptoms of insomnia (problems falling asleep, waking early and work disruption) were nearly three times more likely to develop fibromyalgia compared with those with no symptoms [28]. In addition, chronic sleeping difficulties are a predictor of acute post-operative pain in patients undergoing total knee replacement [29]. From the other direction, studies have demonstrated that reduced sleep is causally linked to greater pain [26, 30, 31], increasing both the neurotransmitters related to pain sensitivity and the inflammatory markers associated with pain [32, 33]. Restricted total sleep time and frequent waking, similar to the sleep patterns experienced by those with chronic pain, results in high spontaneous pain reports and reduced pain modulation. Improving sleep for people with chronic pain therefore has the potential to reduce pain levels and improve quality of life. The aim of this study was to use systematic review methods and meta-analysis to evaluate the effectiveness of non-pharmacological sleep interventions in improving sleep in people with chronic pain.

Methods
The protocol was registered prospectively with PROSPERO (CRD 42019093799) [34], and the research question formulated according to the PICO principle [35]. Methods were based on those described in the Cochrane Handbook [36], and reporting was in accordance with Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines [37]. (Supplement Table 1).

Patient and public involvement
All of our studies of sleep problems in people with chronic pain are fully supported by patient involvement. This includes regular discussions during development, conduct and reporting of research.

Eligibility criteria
Eligible studies reflected PICOS criteria:

- Population: People aged ≥18 years with chronic non-cancer pain
- Intervention: Non-pharmacological and non-invasive intervention to improve sleep quality or duration
- Comparison: Comparator of standard care, no treatment, attentional, or wait list control
- Outcomes: Primary outcomes of sleep quality and duration, secondary outcomes of other sleep outcomes, pain, health-related quality of life, and psychological wellbeing, and a primary harm outcome of adverse events. Follow up post-treatment and at 3 and 6 months after end of treatment if reported
- Study: Evaluation in a randomised controlled trial

Information sources and searches
We searched MEDLINE, EMBASE, PsycINFO, Cochrane Library, and CINAHL from inception up to 8th April 2020. The search strategy as applied in MEDLINE is included in Supplement Table 2. Citations of key reviews and studies were tracked in Web of Science, reference lists checked and clinical trial records reported in the Cochrane Library followed up. No language restrictions were applied, and relevant non-English articles were translated. Studies reported only as abstracts or that we are unable to acquire using inter-library loans or email contact with authors were excluded.
Study screening and data extraction
Results of searches were imported into Endnote and duplicates removed. After an initial screen by one reviewer to remove clearly off-topic studies, all titles and abstracts were screened independently by two reviewers. Potentially relevant articles were acquired and independently assessed by two reviewers for eligibility with disagreements resolved in discussion with a third reviewer.

One reviewer extracted data from eligible studies into Excel and a second reviewer checked this. Extracted data comprised: country; dates of recruitment; setting; inclusion and exclusion criteria; participant characteristics (chronic pain condition, age, sex); intervention and comparator content, timing, duration and intensity; assessment times; outcome measures; and information on intervention fidelity. We contacted study authors for clarification relating to review eligibility and for missing data.

Risk of bias assessment
Risk of bias was assessed independently by two reviewers using the Cochrane tool [36], specifically relating to: randomisation process; deviations from intended interventions; missing outcome data (>20% considered high risk), measurement of the outcome; and selection of the reported result. Studies with serious concerns relating to risk of bias were considered high risk and those with limited reporting unclear risk. Studies with wait list controls were considered to be at unclear risk of bias due to inherent lack of blinding. Studies with high or unclear risk of bias were excluded from meta-analysis in sensitivity analysis.

Data synthesis
We conducted meta-analyses with Review Manager 5.4 software to compare outcomes across studies with similar interventions and outcome measures. For continuous data, if outcomes were measured identically across studies, an overall mean difference (MD) with 95% confidence intervals (CIs) was calculated. If continuous outcomes were measured differently across studies, overall standardised mean differences (SMDs) and 95% CIs were calculated and presented alongside measures of heterogeneity ($I^2$). Forest plots were generated. Risk of bias as a potential source of heterogeneity was considered in sensitivity analyses. If pooling of outcome data was not appropriate, a narrative synthesis was reported. In interpreting the outcomes from this review we consider effect sizes as described by Cohen: small, $SMD = 0.2$; medium, $SMD = 0.5$; large, $SMD = 0.8$ [38].

Results
Review progress is summarised in Fig. 1. Searches identified 4314 articles of which 305 were considered potentially eligible. After detailed screening we included 42 randomised trials which included 3346 participants. In 33 trials, 2 randomised groups were compared, in 8 there were 3 groups and in 1 there were 4. Overall, 94 groups were compared.

Details of studies and methodological quality scores are summarised in Supplement Table 3 and risk of bias assessments in Supplement Table 4. Methodological quality was assessed as good in 32 studies and fair in ten.

The primary areas addressed by interventions were psychological, physical exercise, physical therapy, and other. Within these groups, comparisons were with untreated controls or alternate active interventions. Sleep outcomes were questionnaires focusing on specific aspects of sleep experience, sleep diaries including aspects of time in bed, nocturnal sleep time, sleep latency, sleep efficiency, wake after sleep onset, total sleep time, and number of awakenings, or measurements with sensors such as actigraphy or polysomnography. Pain outcomes were reported in 38 studies, health-related quality of life outcomes in 24 studies and measures of psychological health in 26 studies. Adverse events were infrequently recorded. All studies reported follow up at the end of intervention or within 2 weeks of completion. Fifteen studies reported follow up at 3 months or longer.

For all studies, effect estimates comparing intervention with control or alternative intervention are summarised in Supplement Table 5 with outcomes reported in multiple studies shown in meta-analysis summaries in Table 2.

Psychological interventions
Cognitive behavioural therapy for insomnia (CBT-I) versus control

In 10 studies including 482 participants, CBT-I was compared with no treatment, attentional control or wait list control [39–48]. Causes of chronic pain were fibromyalgia (2 studies) [41, 43], osteoarthritis (2 studies) [45, 48], spinal pain (1 study) [42], multiple sclerosis (1 study) [39], migraine (1 study) [46], and diverse chronic pain (3 studies) [40, 44, 47]. Three studies were judged to be at low risk of bias [42, 45, 46], but no studies were considered to be at high risk.
At the end of intervention, overall questionnaire assessed sleep quality in 9 studies with 385 participants was improved in people receiving CBT-I compared with untreated controls, SMD -1.23 (95%CI -1.76, –0.70), \(p<0.00001\) (Fig. 2). Heterogeneity was high, (I² 80%). The improvement was sustained but reduced at 6 months in 3 studies with data. In 3 studies at low risk of bias, the benefit for CBT-I over control was slightly reduced, SMD -1.01 (95%CI -1.79, –0.22), \(p=0.01\) and heterogeneity remained high (I² 74%). Exploration of effectiveness in relation to a specific condition was only possible for 2 studies at unclear risk of bias including 79 people with fibromyalgia suggesting no benefit for CBT-I, SMD -0.57 (95%CI -1.44, 0.30), \(p=0.2\), but heterogeneity was high (I² 65%).

Waking after sleep onset, as measured by actigraphy, improved in people receiving CBT-I post-treatment compared with controls and this was also apparent up to at least 6 months in those studies using sleep diaries and polysomnography. Sleep onset latency was improved up to 3 months after CBT-I when assessed by questionnaire, but not with actigraphy or polysomnography. Sleep efficiency was improved up to 6 months after CBT-I compared with control when measured in sleep diaries or by polysomnography, but not by actigraphy. Total sleep time measured by diary, actigraphy or polysomnography was not improved in those receiving CBT-I compared with controls. There were no improvements in diary recorded sleep awakenings or the Epworth Sleepiness Scale in people receiving CBT-I compared with controls. Adverse events were assessed in 5 studies [42, 43, 45–47]. No adverse events were reported in 4 studies. In 1 study 3 adverse event cases were deemed to be study related [45], this included rash from wearing actigraph and tenderness at site of testing.

In 9 studies with 370 people randomised, pain measured by questionnaire was reduced post-treatment in people receiving CBT-I compared with controls, SMD
Table 1  Study and intervention characteristics

| Author | Country, Recruitment dates | Setting | Study design | Inclusion | Intervention/ control | Follow up | Outcomes | Losses to follow up | Risk of bias issues | Methodological quality scorea | Key results |
|--------|---------------------------|---------|-------------|-----------|----------------------|-----------|----------|-------------------|-------------------|--------------------------|------------|
| Abbasi et al. 2016 [39] | Iran, 2014 | Setting not specified | Parallel group RCT | Multiple sclerosis, at least 6 months from time of diagnosis, score of 5 or more for PSQI quality of sleep | CBT targeting sleep quality 8 weekly 90 min sessions Psychiatry nurse Fidelity not reported Group sessions talking about feelings and experiences. Treatment with “common drugs” 3 sessions Therapist not described | Before, immediately and 1 month after treatment PSQI. No pain, HRQoL, psychological health measures. Adverse events not reported. Overall loss to follow up 8% Unclear risk of bias as selected outcomes reported and methodological detail of trial limited. 19/28 | Mean score of sleep quality of patients in the intervention group had a significant difference at 3 stages of before, immediately and 1 month after the intervention. |
| Currie et al. 2000 [40] | Canada, dates not stated | Parallel group RCT | Chronic musculoskeletal pain (excluding fibromyalgia) and sleep difficulties, age < 60 years | CBT. Sleep and insomnia education, behavioural therapy, relaxation training, cognitive restructuring, sleep hygiene education. Coping with chronic pain sleep problems manual. 7 × 2 h group sessions held once a week for 7 weeks Clinical psychology doctoral students or interns with training in CBT. Structured manual. Regular supervision Waiting list controls | Baseline, end of treatment (7 weeks), 3 months Sleep diaries. Actigraphy measured nocturnal activity levels. PSQI. Multidimensional Pain Inventory Pain Severity scale, no HRQoL measure, BDI. Adverse events not reported. Overall loss to follow up 15% at 3 months Unclear risk of bias due to a waiting list control group 18/28 | Authors concluded that short-term CBT improves sleep in people with insomnia secondary to chronic pain. |
| Edinger et al. 2005 [41] | USA, date not stated | Clinic | Parallel group RCT | Fibromyalgia (ACR diagnosis) aged 21–65 with insomnia | CBT for insomnia 6 weekly individual sessions. First 45–60min, then 15–30 min Clinical psychologists Fidelity not reported Usual care Sleep hygiene | Baseline, end of treatment week 3, post-intervention, 6 months Insomnia symptoms questionnaire. McGill Pain questionnaire, no general HRQoL, POMS, SF-36 MH. Adverse events not reported. Overall loss to follow up 45% at 6 months Unclear risk of bias concern due to limited reporting. High risk at 6 months due to high loss to follow up and in comparisons including control due to uneven randomisation. Small study 19/28 | The CBT group achieved nearly a 50% reduction in their nocturnal wake time by study completion compared with 3.5% in usual care and 20% in sleep hygiene groups. 57% of CBT group met strict subjective sleep improvement criteria at end of treatment compared with 0% in usual care and 17% in sleep hygiene groups. |
| Author | Country, Recruitment dates | Setting | Study design | Inclusion | Intervention/ control | Follow up | Outcomes | Losses to follow up | Risk of bias issues | Methodological quality scorea | Key results |
|--------|---------------------------|---------|--------------|-----------|----------------------|-----------|----------|-------------------|-------------------|------------------------|-------------|
| Jungquist et al. 2010 [42] USA, dates not stated Community Parallel group RCT | Chronic spinal pain with insomnia. Age 25+ years 28 (19;9) Mean age 49 years 78% female | CBT for insomnia 8 weekly sessions (30–90 min) Trained masters prepared nurse therapist Fidelity assessed with sleep diaries and actigraphy Appointment with nurse therapist. Reviewed sleep/pain diaries and BDI items above a scale score of 0 for prior week. Discussed rationale that life stress and depression likely to contribute to insomnia and pain. No directed form of therapy provided. 8 weekly sessions, 45–90 min Nurse therapist | Baseline, post-treatment (8 weeks) Sleep diaries, actigraphy, insomnia severity index, Epworth sleepiness scale, Pain disability index, multidimensional pain inventory. No HRQoL or anxiety, BDI Adverse events collected on checklist of symptoms but not reported in article. Overall loss to follow up 25% Low risk of bias. | 23/28 | Significant improvements were found in sleep as well as in the extent to which pain interfered with daily functioning. |
| McCae et al. 2019 [43] USA, 2009–2013 Clinic Parallel group RCT | Fibromyalgia (ACR criteria) and insomnia aged 18 years or older 113 (39; 37; 37) Mean age 53 years 97% female | CBT for insomnia. Sleep education, sleep hygiene and stimulus control; relaxation, sleep restriction, cognitive therapy – monitoring automatic thoughts, cognitive therapy – challenging/replacing dysfunctional thoughts, cognitive therapy – practical recommendations, review of skills and long-term maintenance 8 weekly 50 min sessions Predoctoral students in clinical psychology with training and weekly supervision All treatment sessions audiotaped. Half were randomly selected for scoring by another interventionist, and 25% of the scored tapes were double-scored for reliability by the lead supervising clinical psychologist. Interventionists encouraged adherence and emphasized importance of regular home practice, which was monitored by daily practice logs CBT for pain. Pain education and diaphragmatic breathing, progressive muscle relaxation, activity-rest cycle and autogenic relaxation, visual imagery, cognitive therapy – monitoring automatic thoughts, cognitive therapy – challenging/replacing dysfunctional thoughts, cognitive therapy – balanced thinking, review of skills and long-term maintenance 8 weekly 50 min sessions Predoctoral students in clinical psychology with training and weekly supervision Fidelity as with CBT for insomnia Waiting list control | Baseline, post-treatment, 6 months Self-report sleep diary, dysfunctional beliefs and attitudes about sleep, actigraph, ambulatory polysomnography. McGill Pain questionnaire, VAS pain, Pain Disability Index. No HRQoL, STAI, BDI Adverse events reported. Overall study loss to follow up 33% Unclear risk of bias due to concern for high loss to follow up 24/28 CBT for insomnia and CBT for pain led to improvements in self-reported sleep and this was sustained at 6 months after CBT for insomnia. No differences in pain compared with control group | 23/28 | CBT for insomnia and CBT for pain led to improvements in self-reported sleep and this was sustained at 6 months after CBT for insomnia. No differences in pain compared with control group |
| Author                      | Country, Recruitment dates | Setting       | Study design            | Inclusion                                                                 | Number randomised (intervention; control) | Age | % female | Common treatment                                      | Intervention/ control                                                                                      | Number of sessions/ duration                                                                 | Therapist                                                                                                                                                                                                 | Fidelity                                                                                                                                                                                                 | Follow up                                                                                   | Outcomes                                                                 | Losses to follow up | Risk of bias issues | Methodological quality score | Key results |
|-----------------------------|---------------------------|---------------|-------------------------|---------------------------------------------------------------------------|------------------------------------------|-----|----------|------------------------------------------------------|--------------------------------------------------------------------------------|---------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------|---------------------|----------------------|------------------------|-------------------------|-------------------|
| Pigeon et al. 2012 [44]     | USA, date not reported    | Setting not reported | Parallel group feasibility/ pilot RCT | Chronic non-malignant pain in the spine, shoulders, hips or limbs (unrelated to autoimmune disease or fibromyalgia) and insomnia | 21 (6;6;5;4)                           |     |          |                                                      | CBT for insomnia                                                                                                   | 10 individual weekly sessions                                                                                      | 2 CBT psychologists                                                                                                                                     | Sessions videotaped and rated using treatment fidelity instrument created for study                           | Waiting list control | Baseline, 10 weeks (end of intervention) | Insomnia severity scale, Epworth sleepiness scale, sleep diary, Multi-dimensional Pain Inventory, no HRQoL. Adverse events not reported. No losses to follow up. Unclear risk of bias for wait list controls. Pilot/ feasibility study. | Authors concluded that a combined CBT for insomnia and pain intervention was feasible to deliver. |
| Smith et al. 2015 [45]      | USA, 2008–2013            | Clinic        | Parallel group RCT      | Knee osteoarthritis and insomnia                                          | 100 (50; 50)                           |     |          |                                                      | CBT for insomnia including sleep restriction therapy, stimulus control therapy, cognitive therapy for insomnia, and sleep hygiene education. | 8 × 45 minute weekly sessions                                                                                      | Postdoctoral clinical psychologists, doctoral psychology candidates or faculty with experience in behavioural medicine. | Structured checklist, supervision of intervention providers, sessions taped, and random sample assessed. | Behavioral desensitization – manual adapted to match the CBT insomnia protocol for session number and duration. | Baseline, mid-treatment, post-treatment, 3 months, 6 months | Sleep diaries, polysomnography, actigraphy measures, Insomnia Severity Index, WOMAC pain (also VAS pain), No HRQoL, No physiological health measures. No adverse events reported. Overall loss to follow up 2.7% | Low risk of bias 22/28 | In people with knee osteoarthritis, CBT for insomnia reduced sleep maintenance insomnia and clinical pain compared with active control. |
| Author | Country, Recruitment dates | Setting | Study design | Inclusion | Number randomised (intervention; control) | Age | % female | Common treatment | Intervention/ control | Number of sessions/ duration | Therapist | Fidelity | Follow up | Outcomes | Losses to follow up | Risk of bias issues | Methodological quality score a | Key results |
|--------|---------------------------|---------|--------------|-----------|-----------------------------------------|-----|---------|-----------------|---------------------|--------------------------|-----------|---------|-----------|----------|----------------------|------------------------|-----------------------------|------------|
| Smitherman et al. 2016 [46] USA, 2011–2013 Clinic | Parallel group RCT | Chronic migraine and ICSD-3 criteria for insomnia | 32 (16; 16) | Mean age 31 years | 30.8 (12.9) | 90% female | CBT for insomnia with 4 instructions in stimulus control and 1 in sleep restriction. 3 × 30 min sessions, with 2 weeks between each session. Graduate-level therapists with backgrounds in CBT and behavioural medicine Therapist fidelity to treatment protocols was assessed at each treatment session via therapist self-ratings Lifestyle modification with 5 skills taught and practiced at home including dinner at consistent time each night and consistent liquid intake. 3 × 30 min sessions, with 2 weeks between each session. Graduate-level therapists with backgrounds in CBT and behavioural medicine Therapist fidelity to treatment protocols was assessed at each treatment session via therapist self-ratings | Baseline, 2 and 6 weeks after completing treatment Actiwatch II, PSQI, Epworth Sleepiness Scale. Headache severity (0–10), no HRQoL, GAD-7 anxiety, PHQ-9 depression. Adverse events not reported. Overall loss to follow up 22% Low risk of bias, analysis was ITT |
| Tang et al. 2012 [47] UK, date not stated Clinic | Parallel group pilot RCT | Chronic pain (non-malignant, not neurological) and clinical insomnia. Age 18–65 years | 24 (12;12) | Mean age 48.5 years | 90% female | Hybrid of CBT for insomnia combined with interventions designed to target cognitive-behavioural processes maintaining chronic pain – sleep psychoeducation, stimulus control therapy, sleep restriction therapy, cognitive therapy, individual formulation, goal setting, reducing pain catastrophising, reversing mental defeat 4 weekly 2 h individual sessions Clinician and health psychologist Intervention according to checklist which specified the content of each session. All sessions were video recorded. A sample of the recordings (20%) was independently reviewed for treatment fidelity Wait list control. Monitoring group kept a pain and sleep diary for 4 weeks and then received the hybrid CBT intervention | Baseline, after treatment (4 weeks) Insomnia Severity Index, Anxiety and Preoccupation about sleep questionnaire, dysfunctional beliefs and attitudes about sleep questionnaire, pain-specific sleep beliefs, pre-sleep arousal scale, sleep onset latency, wake after sleep onset, total sleep time, sleep efficiency Brief Pain Inventory. No HRQoL, HADS anxiety and depression. Adverse events not reported. Overall loss to follow up 17% Unclear risk of bias for wait list controls 19/28 Described as pilot RCT but authors reported that compared with symptom monitoring, the hybrid intervention was associated with greater improvement in sleep (as measured with the Insomnia Severity Index and sleep diary) at post-treatment. |
| Vitiello et al. 2009 [48] USA, 2001–2003 Academic medical centre | Parallel group RCT | Osteoarthritis with moderate pain and insomnia. Age 55 or older | 51 (23; 28) | Mean age 69 years | 88% female | CBT for insomnia. Stimulus control, sleep restriction, cognitive restructuring, relaxation training, sleep-hygiene education. 8 weekly 2 h classes Two-clinical psychologists Fidelity not reported Attention-control stress management and wellness. Skill-training on mind-body relationship, reduction of stress and anxiety, effective communication and assertiveness, problem solving and goal setting, nutrition and exercise for individuals with chronic conditions 8 weekly 2 h classes Physician, psychologist, nutritionist, exercise physiologist | Baseline (before treatment), post-treatment. One year not considered as included cross-over patients Total sleep time, naps, sleep latency, wake after sleep onset, sleep efficiency SF-36 bodily pain, no HRQoL, measure GDS. Adverse events not reported. Losses to follow up not reported Unclear risk of bias due to limited reporting 21/28 CBT for insomnia improved immediate self-reported sleep and pain in older patients with osteoarthritis and comorbid insomnia compared with an attention control. |
| Author | Country, Recruitment dates | Setting | Study design | Inclusion | Number randomised (intervention; control | Intervention/ control | Follow up | Outcomes | Losses to follow up | Risk of bias issues | Methodological quality scorea | Key results |
|--------|---------------------------|---------|--------------|-----------|------------------------------------------|-----------------------|-----------|----------|-------------------|------------------|--------------------------|------------|
| Berry et al. 2015 [49] | Canada, dates not stated | Outpatient clinic | Parallel group RCT | Chronic non-cancer pain including neuropathic pain, musculoskeletal pain, complex regional pain syndrome, joint pain, visceral pain and headaches, age 18–80 years | 132 (65;67) | Mean 49 years | 61% female | Education with CBT component. Education session incorporating sleep hygiene and cognitive behavioral strategies. 15 min session once. Subsequently participants contacted weekly for 4 weeks by telephone to address questions or concerns about completing diaries and to ensure they were engaging in treatment strategies. | Trained research assistant | Consistency of intervention delivery monitored | No treatment but offered after study completion | Baseline, week 1, week 2, week 3 during intervention. Week 4 at end of intervention | Daily sleep diary. No pain, HRQoL or psychological measures. | Adverse events not reported | Overall 36% lost to follow up | 20/28 | High risk of bias due to large loss to follow up | No differences between groups except sleep latency improved in Education group |
| Martinez et al. 2014 [50] | Spain, dates not reported | Clinic | Parallel group RCT | Women with fibromyalgia, aged 25–60 years | 64 (32:32) | Mean age 48 years | 100% female | CBT for insomnia 6 group 1.5 h sessions, 1 per week, 5–6 participants per group. Female therapists with experience of pain management and sleep disorders. Fidelity not reported | Sleep hygiene 6 group 1.5 h sessions, 1 per week, 5–6 participants per group. Female therapists with experience of pain management and sleep disorders. Fidelity not reported | Baseline, post-treatment, 3 months, 6 months | PSQI, McGill pain questionnaire (Spanish), FIQ, SCL-90-R. | Adverse events not reported | Overall loss to follow up 27% | Low risk of bias | 19/28 | Authors reported that patients who received CBT for insomnia reported significant, positive and sustained changes in subjective sleep quality, sleep latency, sleep duration, habitual sleep efficiency and sleep disturbances. Patients in the sleep hygiene group only reported a significant improvement in subjective sleep quality and a trend towards improvement in sleep efficiency. |
| Miro et al. 2011 [51] | Spain, dates not reported | Group | Parallel group RCT | Women with fibromyalgia and insomnia | 44 (22:22) | Mean age 46 years | 100% female | CBT for insomnia 6 weekly group sessions lasting 90 mins Female CBT experts with experience in fibromyalgia Fidelity not reported | Sleep hygiene 6 weekly group sessions lasting 90 mins Female CBT experts with experience in fibromyalgia Fidelity not reported | Baseline, 1 week post-intervention | PSQI, McGill pain questionnaire (Spanish). | Adverse events not reported | Overall loss to follow up 9% | Low risk of bias | 23/28 | Authors reported that compared with the sleep hygiene group, patients receiving CBT for insomnia had improved sleep quality. No difference in pain between groups |
| Author                | Country, Recruitment dates | Setting | Study design | Inclusion                                                                 | Intervention/ control                                                                 | Follow up                                                                 | Outcomes                                                                 | Losses to follow up | Risk of bias issues | Methodological quality score | Key results                                                                                   |
|-----------------------|---------------------------|---------|--------------|----------------------------------------------------------------------------|----------------------------------------------------------------------------------------|---------------------------------------------------------------------------|--------------------------------------------------------------------------|-------------------|---------------------|-------------------------------|-----------------------------------------------------------------------------------------------|
| Sanchez et al. 2012   | Spain, dates not reported | Clinic  | Parallel group RCT | Fibromyalgia with chronic insomnia. Age 25–60 years 26 (13;13) Mean age 47 years 100% female | CBT for insomnia 6 weekly group sessions of 90 mins each Female CBT experts with experience in fibromyalgia Fidelity not reported Sleep hygiene 6 weekly group sessions of 90 mins each Female CBT experts with experience in fibromyalgia Fidelity not reported | Baseline, 6 weeks (post-treatment) Polysomnographic parameters. No pain outcome. Adverse events not reported Losses to follow up not reported Low risk of bias 22/28 | Baseline, 6 weeks (post-treatment) Polysomnographic parameters. No pain outcome. Adverse events not reported Losses to follow up not reported Low risk of bias 22/28 | Authors conclude that use of CBT for insomnia in fibromyalgia patients can significantly improve objective sleep parameters | |  |
| Castel et al. 2012    | Spain, dates not stated   | Clinic  | Parallel group RCT | Fibromyalgia, age 18–65 years 93 Mean age 50 years 97% female | CBT for pain and insomnia 14 weekly sessions, 120 mins each, all group except session 2 Not described who delivered No information on fidelity CBT for pain and insomnia plus hypnosis 14 weekly sessions 120 mins each, group except session 2 Session 2 received hypnosis training. Hypnosis exercises performed at end of each session instead of autogenic. Audio compact disc for home practice with analgesic self-hypnosis exercises. Not described who delivered. No information on fidelity Standard care | Baseline, 1 week post-treatment, 3 months, 6 months Medical outcomes: study sleep scale, NRS pain, FIQ, HADS total. No adverse events reported Overall loss to follow up 24% Unclear risk of bias due to limited reporting 17/28 | CBT for pain and insomnia showed greater improvements in sleep than standard care. Adding hypnosis led to increased benefit. | |  |
| Lami et al. 2018      | Spain, dates not reported | Clinic, group | Parallel group RCT | Women with fibromyalgia and insomnia aged 25–65 years 126 (42;42;42) Mean age 50 years 100% female | CBT for insomnia and pain 9 weekly 90 min group (5–7 people) sessions Therapists with high level professional training and experience in chronic pain and sleep disorders Fidelity assessed with video recordings and regular meetings CBT for pain 9 weekly 90 min group (5–7 people) sessions Therapists with high level professional training and experience in chronic pain and sleep disorders Fidelity assessed with video recordings and regular meetings Standard care Control group receiving usual care | Baseline (pre-treatment), 1 week after completion of the intervention, 3 months post-treatment PSQI, McGill Pain Questionnaire Short Form, FIQ, SCL-90-R. No adverse events reported Large losses to follow-up (28%) at post-treatment timepoint High risk of bias due to large loss to follow 20/28 | Authors report significant improvement in sleep variables in CBT for pain and insomnia compared with CBT for pain and CBT groups had improvements in pain outcomes compared with control | |  |
| Pigeon et al. 2012    | Details above             |         |              |                                                                           |                                                                                        |                                                                           |                                                                         |                                 |                                    |                               |                                                                                           |
| Author                  | Country, Recruitment dates | Setting                        | Study design | Inclusion                              | Intervention/ control | Follow up                                                                 | Outcomes                                                                                     | Key results                                                                 |
|------------------------|----------------------------|--------------------------------|--------------|----------------------------------------|-----------------------|---------------------------------------------------------------------------|---------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------|
| Vitiello et al. 2013   | USA, 2009–2011             | Classroom or primary care clinic | Cluster RCT  | Osteoarthritis and insomnia aged 60+   | CBT for pain and insomnia | Baseline, 2 months (post-treatment), 9 months after baseline, 18 months   | Insomnia severity index, sleep efficiency via actigraphy. Graded Chronic pain Scale, no specific HRQoL (AIMS2 symptom subscale reflects pain). Adverse events not reported | Overall loss to follow up 13% Low risk of bias 24/28 At 9 months CBT for pain and insomnia reduced insomnia severity compared with CBT for pain and control. No significant differences between CBT groups and control in sleep or pain outcomes at 18 months |

1f. Cognitive behavioural therapy focusing on insomnia and pain alone or with additional hypnosis

Lam et al. 2018 [56]
Details above
McCabe et al. 2019 [40]
Details above
Pigeon et al. 2012 [44]
Details above
Vitiello et al. 2013 [55]
Details above

1g. Acceptance and Commitment Therapy based stress management compared with control

Castel et al. 2012 [53]
Details above

1h. Acceptance and Commitment Therapy based stress management compared with exercise
| Author                      | Country, Recruitment dates | Setting                                      | Study design           | Inclusion                                                                 | Intervention/ control                                                                 | Follow up                                                                 | Outcomes                                                                 |
|-----------------------------|---------------------------|----------------------------------------------|------------------------|----------------------------------------------------------------------------|----------------------------------------------------------------------------------------|---------------------------------------------------------------------------|--------------------------------------------------------------------------|
| Wiklund et al. 2018 [56]    | Sweden, dates not reported | Clinic/conference room at hospital/ training facility | Parallel group RCT     | Adults (18–60 years) with chronic (> 3 months) benign neck, low back, and/or generalised pain | Acceptance and commitment therapy (ACT) based stress management 7 weekly 2-h sessions 61% completed 4 or more sessions Group-based exercise: Graded exercises: endurance, coordination, balance, functional strength, and movement training. After 4 weeks: increased aerobic training and decreased range of movement exercises. Individually exercises: graded strength exercises for back, neck, abdomen, shoulders, and arms. Home exercise: One hour, twice a week for 8 weeks. Physiotherapist and physician. | Baseline, immediately post-intervention, 6 and 12 months Insomnia Severity Index. Pain NRS for average pain intensity over past 7 days, no HRQoL measure, HADS. Adverse events not reported. Overall loss to follow up 25% at 6 months 20/28 | Unclear risk of bias due to selective reporting. Unclear risk of bias due to losses to follow up at 6 months No benefit for ACT-based stress management over control or exercise therapy |
| Cash et al. 2015 [57]      | USA, date not specified   | Clinic, group                               | Parallel group RCT     | Women volunteers aged 18+ with physician diagnosed fibromyalgia              | Mindfulness-based stress reduction group. Formal and informal mindfulness practices including attention-focusing, sitting meditation and yoga positions taught to encourage relaxed and focused movements. Home practice guided by a workbook and audiotapes. Half-day meditation retreat Experienced, trained instructor. 1 weekly 2.5-h group session for 8 weeks. Home 45 min per day, 6 days a week No fidelity relating to delivery reported Waiting list controls | Baseline, after 8 week programme, 2 months after completion of programme Stanford Sleep Questionnaire. VAS pain. Adverse events not reported Overall loss to follow up 25% Unclear risk of bias, authors reported ITT with analysis including baseline data for losses to follow up. Wait list controls 20/28 | Authors reported that the mindfulness intervention was associated with significant and maintained reductions of sleep problems and symptom severity compared with controls |
| Author                        | Country, Recruitment dates | Setting            | Study design      | Inclusion Number randomised (intervention; control) | Intervention/ control Number of sessions/ duration | Therapist | Fidelity | Follow up | Outcomes | Losses to follow up | Risk of bias issues | Methodological quality score | Key results                                                                 |
|-------------------------------|----------------------------|--------------------|-------------------|---------------------------------------------------|-------------------------------------------------|-----------|----------|-----------|----------|---------------------|------------------------|---------------------------|---------------------------------------------------------------------------|
| Van Gordon et al. 2017 [58]   | UK, 2012–2014              | Community          | Parallel group RCT| Fibromyalgia. Age 18–65 years                      | Meditation awareness training – Second-generation mindfulness-based intervention. Taught component, facilitated group discussion, guided meditation and/or mindfulness exercises. One-to-one support sessions and compact disc of guided meditations to facilitate daily self-practice. 8 weekly 2 h 25 participant workshops plus CD of guided meditation for self-practice. 2 one-to-one support sessions. Instructors attended a 3-year supervised training programme. |          |          | Baseline, post-treatment, 6 month follow-up FPSQ, Short Form-McGill Pain Questionnaire, DASS. Adverse events not reported | Losses to follow up at final assessment 43% | High risk of bias due to large loss to follow up and retrospective registration | 24/28 | Authors report that Meditation awareness training participants demonstrated significant and sustained improvements over control group participants in sleep quality and pain perception. |
| Soares and Grossi 2002 [59]   | Sweden, dates not reported | Community          | Parallel group RCT| Women with fibromyalgia                           | Behavioural intervention. Individual sessions focused on applied relaxation. Group sessions covering symptoms, stress, behaviour patterns and self-management. 5 individual sessions (1 h each) and 15 group sessions (2 h each/3–5 patients per group) over 10 weeks. Experienced CBT therapist and pain physician. Sessions observed. Educational intervention with a sleep hygiene focus. 2 individual sessions (2 h each) and 15 group sessions (2 h each/3–5 patients per group) over 10 weeks. Experienced physiotherapist and occupational therapist. Sessions observed. Waiting list control. |          |          | Baseline (before treatment), after treatment and 6 months (only post-treatment for wait list controls) | Karolinska Sleep Questionnaire, McGill Pain Questionnaire, The Pain Questionnaire, pain VAS, FQI. No psychological measures. Adverse events not reported. No losses to follow up post-treatment. Overall 23% lost to follow up at 6 months. Unclear risk of bias for relaxation/education comparison due to wait list control | 18/28 | Behavioural intervention associated with long-term improvement in sleep quality. No other differences between groups at long-term follow up. |

**Key results**

1. Relaxation compared with control

2. Sleep hygiene compared with control

Edinger et al. 2005 [41]

Details above.
### Table 1 (continued)

| Author                      | Inclusion                                                                 | Intervention/ control                                                                 | Follow up                                                                 |
|-----------------------------|---------------------------------------------------------------------------|----------------------------------------------------------------------------------------|---------------------------------------------------------------------------|
| Soares and Grossi 2002 [59] | Details above                                                             | Aerobic exercise plus relaxation. 30 min graded aerobic exercise in pool followed by 30 mins progressive relaxation | Baseline, at end of 10 week intervention                                   |
| Arcos-Carmona et al. 2011 [60] | Fibromyalgia (ACR diagnosis) in previous 2 years, age 30–60 years, Mean age 44 years | Twice a week for 10 weeks. Not stated who delivered. Fidelity not reported            | PSQI, SF36 bodily pain. Adverse events not reported. Overall 5% lost to follow up |
| Arcos-Carmona et al. 2011 [60] | Fibromyalgia (ACR diagnosis) in previous 2 years, age 30–60 years, Mean age 44 years | Twice a week for 10 weeks. Not stated who delivered. Fidelity not reported            | Unclear risk of bias due to limited methodological reporting 20/28          |
| Arcos-Carmona et al. 2011 [60] | Fibromyalgia (ACR diagnosis) in previous 2 years, age 30–60 years, Mean age 44 years | Twice a week for 10 weeks. Not stated who delivered. Fidelity not reported            | The authors report that the combination of aerobic exercise and progressive relaxation improved night rest |
| Durcan et al. 2014 [61]    | Rheumatoid arthritis, Mean age 60 years, 64% female                        | Home-based cardiovascular exercise, resistance training, flexibility and neuromotor conditioning. 12 weeks. Assessment by doctor and senior physiotherapist every 3 weeks | Baseline and post-treatment (12 weeks)                                     |
| Durcan et al. 2014 [61]    | Rheumatoid arthritis, Mean age 60 years, 64% female                        | Home-based cardiovascular exercise, resistance training, flexibility and neuromotor conditioning. 12 weeks. Assessment by doctor and senior physiotherapist every 3 weeks | PSQI, VAS pain. Adverse events not reported. Overall loss to follow up 3%   |
| Eadie et al. 2013 [62]     | Chronic or recurrent non-specific low back pain, Age 18–70 years, 60 (20 Home-based walking; 20 Control; 20 Supervised exercise) | Home-based walking programme 8 weeks. 30 min moderate intensity physical activity, 5 days/ week by week 5. Physiotherapist 77.5% patient adherence Supervised exercise class Once per week for 8 weeks. Physiotherapist 50% patient adherence Control of usual physiotherapy. 1-to-1 advice, manual therapy and exercise. Number and duration at discretion of treating physiotherapist. Physiotherapist. Fidelity not described (Supervised exercise class) | Baseline, 3 and 6 months (equivalent to post-treatment and 3 months)         |
| Eadie et al. 2013 [62]     | Chronic or recurrent non-specific low back pain, Age 18–70 years, 60 (20 Home-based walking; 20 Control; 20 Supervised exercise) | Home-based walking programme 8 weeks. 30 min moderate intensity physical activity, 5 days/ week by week 5. Physiotherapist 77.5% patient adherence Supervised exercise class Once per week for 8 weeks. Physiotherapist 50% patient adherence Control of usual physiotherapy. 1-to-1 advice, manual therapy and exercise. Number and duration at discretion of treating physiotherapist. Physiotherapist. Fidelity not described (Supervised exercise class) | Baseline, 3 and 6 months (equivalent to post-treatment and 3 months)         |
| Eadie et al. 2013 [62]     | Chronic or recurrent non-specific low back pain, Age 18–70 years, 60 (20 Home-based walking; 20 Control; 20 Supervised exercise) | Home-based walking programme 8 weeks. 30 min moderate intensity physical activity, 5 days/ week by week 5. Physiotherapist 77.5% patient adherence Supervised exercise class Once per week for 8 weeks. Physiotherapist 50% patient adherence Control of usual physiotherapy. 1-to-1 advice, manual therapy and exercise. Number and duration at discretion of treating physiotherapist. Physiotherapist. Fidelity not described (Supervised exercise class) | Baseline, 3 and 6 months (equivalent to post-treatment and 3 months)         |

3. Exercise

3a. Group exercise compared with control

**Follow up**
- Outcomes
- Losses to follow up
- Risk of bias issues
- Methodological quality score
- Key results
| Author | Country, Recruitment dates | Setting | Study design | Inclusion | Number randomised (intervention; control) | Age | % female | Common treatment | Intervention/ control | Number of sessions/ duration | Therapist | Fidelity | Follow up | Outcomes | Losses to follow up | Risk of bias issues | Methodological quality scorea | Key results |
|--------|---------------------------|---------|--------------|-----------|------------------------------------------|-----|---------|------------------|---------------------|---------------------------|-----------|----------|------------|----------|---------------------|-----------------|--------------------------|-------------|
| Freburger et al. 2010 [63] | USA, dates not stated | Community | Parallel group RCT | Self-report arthritis and activity limitations, 18 years or over | 321 (166; 155) | Mean age 70 years | 88% female | Exercise programme. Low-to-moderate-intensity physical activity program: People With Arthritis Can Exercise (PACE)/Arthritis Foundation Exercise Program. Land-based exercise, health education on arthritis self-management and exercise. Activities to promote social interaction, movement, balance, and body awareness. Relaxation techniques. | I hour, twice weekly for 8 weeks | PACE trained instructor | Fidelity not assessed | Waiting list control | Baseline, 8 weeks | Jenkins sleep scale. Pain not reported. Adverse events not reported | Overall 15% lost to follow up | Unclear risk of bias due to wait list control | 21/28 |
| Wiklund et al. 2018 [56] | Details above | 3b. Home-based walking programme versus supervised exercise | Details above |
| Eadie et al. 2013 [62] | Details above | 3c. Moderate aerobic exercise compared with low intensity home-based exercise | Details above |
| Al-Sharman et al. 2019 [64] | Jordan, 2015–2018 | Clinic and control at home | Parallel group pilot RCT | Multiple sclerosis with poor sleep quality, age > 18 years | 40 (20;20) | Mean age 35 years | 77% female | Moderate-intensity aerobic exercise programme. Supervised moderate-intensity aerobic exercise using a recumbent stepper machine. Upper and lower body stretching exercises before and after each exercise session. 40 min per session, 3 times a week for 6 weeks | Weekly exercise logs | Low-intensity home-exercise programme. DVD and exercise manual. DVD showed warm up, cool down activities, flexibility, strength, balance and endurance exercises. Ako, relaxation, stretching and breathing techniques. 50–60 min per session, 3 sessions a week for 6 weeks | Participants asked to demonstrate exercises with examiner. Weekly exercise logs | Baseline, end of intervention (6 weeks) | PSQI, Insomnia Severity Index and Actigraph in subsample of patients. Pain not reported. Adverse events not reported | Overall loss to follow up 25% | High risk of bias due to high and uneven loss to follow up and limited reporting of methods | 19/28 |
| 3d. Aquatic biodance compared with stretching exercises |

*Table 1 (continued)*
### Table 1 (continued)

| Author                          | Country, Recruitment dates | Setting             | Study design          | Inclusion                                                                                      | Intervention/ control                                                                                   | Follow up                                                                                                    |
|---------------------------------|-----------------------------|---------------------|-----------------------|-----------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------|
| Lopez-Rodriguez et al. 2013 [65] | Spain, 2011                | Clinic              | Parallel group RCT    | Fibromyalgia (ACR diagnosis) in previous 2 years, age 18–68 years.                            | Aquatic biodance for 1 h in pool heated to 29°C, 10 min flexibility/ breathing exercises, 40 min creative dance movement to music involving upper/lower limbs, 10 min gentle exercise | Follow up                                                                                                    |
|                                 |                             |                     |                       | Mean age 55 years 100% female                                                                    | Twice weekly 1 h session for 12 weeks Not stated who delivered Fidelity not reported Stretching exercises including neck, trunk, quadriceps and calves. Twice weekly 1 h sessions for 12 weeks | Outcomes                                                                                                    |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           | Losses to follow up                                                                                          |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           | Risk of bias issues                                                                                         |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           | Methodological quality score<sup>a</sup>                                                                       |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           | Key results                                                                                                 |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           |                                                                                                              |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           |                                                                                                              |
| Lu et al. 2017 [66]             | China, 2013                 | Community           | Parallel group RCT    | Knee osteoarthritis aged 60–70 years                                                            | Tai Ji Quan: 8 forms adapted for use in people with osteoarthritis 60 min session 3 times weekly for 24 weeks Two instructors with training and academic specialisation in Tai Ji Quan Specialist monitored fidelity of delivery on a weekly basis Education focusing on wellness and health promotion 60 min class twice per week for 24 weeks Additional 10–15 min weekly check-in phone call from research staff to monitor activity levels, changes in knee pain, and medication usage Fidelity not described | Follow up                                                                                                    |
|                                 |                             |                     |                       | Mean age 65 years 100% female                                                                    |                                                                                                           | Baseline (before intervention), at end of study (after intervention), 24 weeks PSQI, WOMAC pain, SF-36 MH. Adverse events not reported Overall 13% lost to follow up at 24 weeks Low risk of bias 23/28 | Outcomes                                                                                                    |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           | Significant improvement in sleep measures and pain in Tai Ji Quan group compared with controls               | Losses to follow up                                                                                          |
|                                 |                             |                     |                       |                                                                                               |                                                                                                           |                                                                                                              |
| Maddali Bongi et al. 2016 [67]  | Italy, dates not stated     | Community and home  | Parallel group RCT    | Women with fibromyalgia (ACR criteria) 44 (22,22)                                             | Tai Ji Quan: breathing exercises, concentration, postural maintenance, rebalancing, and precise movement. Twice weekly 60 min sessions for 16 weeks Daily DVD led home exercises in two 15 min sessions Not reported who delivered Fidelity not described Education about the disease, symptoms, management and coping Twice weekly 60 min sessions for 16 weeks Fidelity not described | Follow up                                                                                                    |
|                                 |                             |                     |                       | Mean age 52 years 100% female                                                                    |                                                                                                           | Baseline, 16 weeks (end of treatment) PSQI, Widespread pain index, HADS. Adverse events not reported No losses to follow up reported (6 withdrew before study) Unclear risk of bias due to limited reporting 20/28 Improvement in some sleep parameters and widespread pain in Tai Ji Quan group but not in education group | Outcomes                                                                                                    |

### 4. Physical therapy

#### 4.1. Hydrotherapy

[...content of the page continues with more details on hydrotherapy and other physical therapy interventions...]
Table 1 (continued)

| Author                        | Inclusion                                      | Intervention/ control                                                                 | Follow up                                                                                                                                 |
|-------------------------------|------------------------------------------------|----------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------|
| Calandre et al. 2009 [68]     | Fibromyalgia aged 18+ years                    | Hydrotherapy exercise with stretching                                                   | Baseline, end of treatment (6 weeks), 4 weeks after end of treatment, 12 weeks after end of treatment, PSQI, VAS pain, FIQ, STAI, Bdi, Adverse reactions listed, Overall loss to follow up 30%, High risk of bias, large loss to follow up, 19/28 |
| Spain, dates not stated       | 81 (39;42)                                    | 18 sessions of 1 h, 3 times per week for 6 weeks                                        | No differences were found between groups                                                                                                                                                          |
| Clinic                        | Mean age 50 years                              | Physiotherapist                                                                       |                                                                                                                                                                                                     |
| Parallel group RCT            | 90% female                                     | Fidelity not reported                                                                  |                                                                                                                                                                                                     |
|                               |                                                | Hydrotherapy exercise with Tai Chi                                                    |                                                                                                                                                                                                     |
|                               |                                                | 18 sessions of 1 h, 3 times per week for 6 weeks                                        |                                                                                                                                                                                                     |
|                               |                                                | Physiotherapist                                                                       |                                                                                                                                                                                                     |
|                               |                                                | Fidelity not reported                                                                  |                                                                                                                                                                                                     |
|                               |                                                | Hydrotherapy exercise with warm-up, stretching, aerobic exercises and relaxation       | Baseline (pre-treatment), post-treatment. Sleep logs completed for 21 days before and after treatment, Total sleep time, total nap time. SF-36 bodily pain, SF-36 MH. Adverse events not reported, Overall loss to follow up 6%, Low risk of bias, 21/28 |
| Vitorino et al. 2006 [69]     | Women with fibromyalgia                        | 60 mins, 3 times per week for 3 weeks                                                  | Sleep quality showed greater improvement in hydrotherapy group but no difference in improvement in pain outcome \                                                                                     |
| Brazil, dates not reported    | 50 (25;25)                                     | Physiotherapist                                                                       |                                                                                                                                                                                                     |
| Clinic                        | Mean age 48 years                              | Fidelity not described                                                                 |                                                                                                                                                                                                     |
| Parallel group RCT            | 100% female                                    | Conventional physiotherapy with infra-red lamp, stretching, aerobic exercise and relaxation |                                                                                                                                                                                                     |
|                               |                                                | 60 mins, 3 times per week for 3 weeks                                                  |                                                                                                                                                                                                     |
|                               |                                                | Physiotherapist                                                                       |                                                                                                                                                                                                     |
|                               |                                                | Fidelity not described                                                                 |                                                                                                                                                                                                     |
|                               | Intervention/ control                          | Hydrotherapy with warm-up, stretching, aerobic exercises and relaxation                |                                                                                                                                                                                                     |
|                               | Number of sessions/ duration                  | 60 mins, 3 times per week for 3 weeks                                                  |                                                                                                                                                                                                     |
|                               | Therapist                                     | Physiotherapist                                                                       |                                                                                                                                                                                                     |
|                               | Fidelity                                      | Fidelity not described                                                                 |                                                                                                                                                                                                     |
|                               |                                                | Conventional physiotherapy with infra-red lamp, stretching, aerobic exercise and relaxation |                                                                                                                                                                                                     |
|                               |                                                | 60 mins, 3 times per week for 3 weeks                                                  |                                                                                                                                                                                                     |
|                               |                                                | Physiotherapist                                                                       |                                                                                                                                                                                                     |
|                               |                                                | Fidelity not described                                                                 |                                                                                                                                                                                                     |
| 4.2. Massage or Manual therapy compared with relaxation or control | Massage therapy                              | Baseline, 5 weeks (post-treatment)                                                     | Verma and Snyder-Halperin Sleep scale, VAS pain, POMS, STAI, Adverse events not reported, Losses to follow up not reported, Unclear risk of bias for limited reporting, Baseline difference in sleep disturbance, 20/28, Authors report sleep disturbance and pain less in massage therapy group, Baseline, post-intervention (48 h after end of 5 week intervention), PSQI, McGill Pain Questionnaire, Adverse events not reported, No losses to follow up, Unclear risk of bias due to lack of blinding, 23/28, Authors concluded that manual therapy was effective in improving sleep and pain |
| Field et al. 2007 [70]        | Chronic lower back pain                       | Massage therapy                                                                       | Baseline, 5 weeks (post-treatment), Post-treatment, Sleep logs completed for 21 days before and after treatment, Total sleep time, total nap time, SF-36 bodily pain, SF-36 MH. Adverse events not reported, Overall loss to follow up 6%, Low risk of bias, 21/28 |
| USA, dates not stated         | 30 total                                      | 30 mins twice per week for 5 weeks                                                    | Sleep quality showed greater improvement in hydrotherapy group but no difference in improvement in pain outcome \                                                                                     |
| Group clinic and home-based   | Mean age 41 years                              | Trained massage therapists                                                             |                                                                                                                                                                                                     |
| Parallel group RCT            | 47% female                                     | Fidelity not reported                                                                 |                                                                                                                                                                                                     |
| Castro-Sanchez et al. 2014 [71]| Fibromyalgia, age 18–70 years                 | Manual therapy                                                                        | Baseline, 5 weeks (post-treatment), Post-treatment, Sleep logs completed for 21 days before and after treatment, Total sleep time, total nap time, SF-36 bodily pain, SF-36 MH. Adverse events not reported, Overall loss to follow up 6%, Low risk of bias, 21/28 |
| Spain, dates not stated       | 89 (45;44)                                    | 5 weekly sessions of 45 mins each                                                     | Sleep quality showed greater improvement in hydrotherapy group but no difference in improvement in pain outcome \                                                                                     |
| Clinic                        | Mean age 54 years                              | Specialist physiotherapist                                                            |                                                                                                                                                                                                     |
| Parallel group RCT            | 53% female                                     | Fidelity not described                                                                 |                                                                                                                                                                                                     |
|                               |                                                | No treatment                                                                          |                                                                                                                                                                                                     |
| Author                | Country, Recruitment dates | Setting          | Study design    | Inclusion                                                                 | Intervention/ control                                                                 | Follow up                                                                 |
|-----------------------|---------------------------|------------------|-----------------|---------------------------------------------------------------------------|----------------------------------------------------------------------------------------|----------------------------------------------------------------------------|
| Külcü et al. 2009    | Turkey, 2006–2007         | Rehabilitation clinic | Parallel group RCT | Primary fibromyalgia age 18–55 years                                       | Physical therapy programme with hot pack, ultrasound, TENS and low power laser         | Baseline and at end of intervention                                     |
|                       |                           |                  |                 | 60 (40;20)                                                               | 15 sessions                                                                            | Adverse events not reported                                              |
|                       |                           |                  |                 | Mean age 37 year                                                         | Fidelity not described                                                                  | No losses to follow up                                                   |
|                       |                           |                  |                 | 95% female                                                               | No physical therapy                                                                    | High risk of bias due to limited reporting and lack of blinding           |
|                       |                           |                  |                 |                                                                          |                                                                                        |                                                                            |
|                       |                           |                  |                 |                                                                          |                                                                                        | Sleep and pain improved in physical therapy group compared with controls  |
| Correia Moretti et al. 2016 | Brazil, 2011–2013       | Clinic           | Parallel group RCT | Fibromyalgia, aged 18–60 years                                           | Pompage. Global, lymph, trapezius, torso, lumbar, and quad pompage plus stretching and aerobic exercise Twice per week for 12 weeks Not specified who delivered Fidelity not reported Stretching and aerobic exercise Twice per week for 12 weeks Not specified who delivered Fidelity not reported | Baseline, 6 weeks, 12 weeks (post-treatment) Sleep inventory. McGill Pain Questionnaire, no HRQoL or psychological health measures. Adverse events not reported Overall 35% lost to follow up High risk of bias for high and uneven loss to follow up 22/28 Authors report no benefit for pompage for sleep and limited benefit regarding pain |
|                       |                           |                  |                 | 23 (13;10)                                                               | 12 weeks                                                                               | Adverse events not reported                                              |
|                       |                           |                  |                 | Mean age 45 years                                                        | Fidelity not reported                                                                  | High risk of bias for high and uneven loss to follow up                  |
|                       |                           |                  |                 | 100% female                                                              | Participants completed treatment diaries                                               |                                                                            |
|                       |                           |                  |                 |                                                                          |                                                                                        |                                                                            |
|                       |                           |                  |                 |                                                                          |                                                                                        |                                                                            |
| Yeh et al. 2016       | USA, dates not reported   | Clinic/office     | Parallel group RCT | Chronic low back pain                                                     | Auricular point acupuncture. Vaccaria seeds placed on active ear points corresponding to low back pain and alleviation of stress and pain. Participants told to press the seeds on each ear at least 3 times a day for 3 min and whenever they experienced pain. Seed removed after 3 days Four weekly clinic visits to place seeds Not specified who delivered Participants completed treatment diaries | Baseline, during each of the 4 treatments, end of intervention, and 1 month after the last treatment PSQI, daily sleep diary. BPI short form, no HRQoL or psychological health measures. Adverse events not reported Overall 25% lost to follow up High risk of bias for high and uneven loss to follow up 20/28 Authors reported that auricular point acupuncture led to improvement in several sleep parameters and pain |
|                       |                           |                  |                 | 61 (30;31)                                                               | 3 days                                                                                 |                                                                            |
|                       |                           |                  |                 | Mean age 63 years                                                        | Fidelity not reported                                                                  |                                                                            |
|                       |                           |                  |                 | 67% female                                                               | Participants completed treatment diaries                                               |                                                                            |

**4.3. Physical therapy programme**

**4.4. Pompage and stretching and aerobic exercise compared with stretching and aerobic exercise**

**5. Miscellaneous interventions**

**5.1. Acupressure**

---

*Table 1 (continued)*
### Table 1 (continued)

| Author                  | Country, Recruitment dates | Setting | Study design   | Inclusion                                                                 | Intervention/ control                                                                 | Follow up       | Outcomes                                      | Losses to follow up | Risk of bias issues                                      | Methodological quality score | Key results                                                                                                                                 |
|-------------------------|----------------------------|---------|----------------|---------------------------------------------------------------------------|----------------------------------------------------------------------------------------|-----------------|-----------------------------------------------|----------------------|------------------------------------------------------------|----------------------------|------------------------------------------------------------------------------------------------------------------------------------------------|
| Murphy et al. 2019 [75] | USA, 2013–2016             | Clinic  | Parallel group RCT | Chronic nonspecific low back pain and fatigue. Aged 18+ years              | Self-administered relaxing acupressure at 19 points. Pressure applied to each point in circular motion 27–30 min per day for 6 weeks, each point for 3 min Trained acupressure educators Standardised training of educators, proper enactment of intervention and methods to track adherence | Baseline, post-treatment (6 weeks) PSQI, Brief Pain Inventory: Adverse events reported Overall loss to follow up 18% Unclear risk of bias due to limited reporting 23/28 Authors report no improvement in sleep quality between acupressure groups and compared with control. Pain reduced in acupressure groups but no change in control. |
| Pearl et al. 1996 [76]  | Canada, 1992–1993           | Home    | Crossover RCT    | Fibromyalgia. Aged 21–65 years                                             | Bright light therapy delivered by a visor with Krypton incandescent bulbs with a mean of 4750 (SD 2337) lux 4 weeks of 1 condition, break week, and 4 weeks alternative treatment Administered at home Fidelity not reported No light condition. Visor system fitted with opaque filter (exposed Kodak film) 4 weeks of 1 condition, break week, and 4 weeks alternative treatment Administered at home Fidelity not reported | Pre-light, light week 4, pre-no light, no light week 4 Daily post sleep questionnaire, VAS sleep quality, daily sleep diary Daily NRS for pain in 10 body regions, FIQ anxiety and depression. Adverse events not reported Overall 26% did not complete crossover protocol High risk of bias due to large loss to follow up and limited reporting of methods 21/28 Authors reported no significant differences between light and no light conditions on sleep and pain. |
| Bakir et al. 2018 [77]  | Turkey, 2015               | Clinic  | Parallel group RCT | Rheumatoid arthritis aged 18+ years                                          | Foot reflexology 60 min repeated once a week for 6 weeks Certified researcher Treatment fidelity not reported Routine polyclinic monitoring and information | Baseline, 1 week, 6 weeks (pain recorded weekly) PSQI, VAS pain. No HRQoL or psychological health. Adverse events not reported 12% lost to follow up Unclear risk of binding due to lack of blinding 20/28 Authors report that sleep and pain improved in the foot reflexology group |
### Table 1 (continued)

| Author | Country, Recruitment dates | Setting | Study design | Intervention/ control | Follow up | Outcomes | Losses to follow up | Risk of bias issues | Methodological quality score\(^a\) | Key results |
|--------|-----------------------------|---------|--------------|-----------------------|-----------|----------|---------------------|---------------------|----------------------|-------------|
| Harvey et al. 2017 [78] | Canada, dates not stated | Laboratory | Parallel group feasibility RCT | Transcranial direct current stimulation (tDCS) applied over the primary motor cortex (2 mA, 20 min) 5 daily sessions of 20 min given in afternoon or evening | Baseline, day 12 (post-treatment), and day 19 (7 days post-treatment). PSQI, Pain and sleep log books completed each day at home, actigraph days 1–19 (only available for 4 participants). VAS pain. No HRQoL, or psychological health. Adverse events not reported 13% lost to follow up | Feasibility study, unclear risk of bias due to limited reporting 21/28 | Study provides guidelines for future studies. Authors report no difference in sleep parameters between groups |
| Colbert et al. 1999 [79] | USA, 1997 | Home | Parallel group RCT | Magnetic mattress pad strength 1100 G which delivered 200–600 G to the skin surface Delivered to patients with instructions on placement Each night for 16 weeks No information on treatment fidelity Sham mattress pad, Delivered to patients with instructions on placement Each night for 16 weeks No information on treatment fidelity | Baseline, 16 weeks | VAS (sleep, fatigue, tiredness on waking, total sleep time). Pain VAS, FIQ-ADL, no psychological measure. Diary of adverse reactions. Low risk of bias Overall 17% lost to follow up. 23/28 | Patients sleeping on the intervention magnetic mattress pad had improvement in reported sleep (p<0.01) and decrease in pain (p<0.001) with no adverse events related to the magnetic mattress pad. |
| Minetto et al. 2018 [80] | Italy, dates not reported | Home | Parallel group pilot RCT | Mattress overlay. Aiartex overlay with suspensory mono-filaments to support a person lying in bed No supervision. Each night for 2 months | Baseline, end of intervention at 2 months | PSQI, VAS pain. No HRQoL, or psychological health. Adverse events not reported. Losses to follow up not reported High risk of bias due to randomization procedure, non-blinding and reporting. 18/28 | Authors report that the mattress overlay was associated with better and clinically meaningful sleep and pain compared with controls. |

\(^a\) Downs and Black Score modification as described in Hooper P, Jutai JW, Strong G, Russell-Minda E. Age-related macular degeneration and low-vision rehabilitation: a systematic review. Can J Ophthalmol. 2008 Apr;43(2):180–7. doi: https://doi.org/10.3129/08-001. Quality levels: excellent (26–28); good (20–25); fair (15–19); poor (<14)
| Follow up     | Effect measure | Number of studies (participants) | Effect estimate (95%CI), p-value | \( I^2 \) |
|--------------|----------------|----------------------------------|------------------------------------|--------|
| Cognitive behavioural therapy for insomnia versus control | Sleep quality from questionnaire | Post-treatment SMD 9 (203:183) | -1.23 (−1.76, −0.70), p < 0.00001 80% |
|              | 3 months SMD 2 (65:68) | −0.80 (−2.17, 0.57), p = 0.25 93% |
|              | 6 months SMD 3 (64:70) | −0.51 (−1.02, −0.01), p = 0.04 41% |
|              | Sleep - onset latency (diary) | Post-treatment MD 7 (168:160) | −17.42 (−28.41, −6.43), p = 0.002 73% |
|              | 3 months MD 2 (64:70) | −14.58 (−21.67, −7.50), < 0.0001 0% |
|              | 6 months MD 3 (64:70) | −6.48 (−14.36, 1.40), p = 0.11 0% |
|              | Sleep - efficiency (diary) | Post-treatment MD 8 (174:164) | −9.86 (−14.06, −5.66), p < 0.00001 67% |
|              | 3 months MD 2 (64:70) | −9.36 (−15.10, −3.61), p = 0.001 53% |
|              | 6 months MD 3 (62:68) | −6.16 (−10.31, −2.01), p = 0.04 46% |
|              | Sleep – wake after sleep onset (diary) | Post-treatment MD 7 (168:160) | −31.12 (−43.55, −18.69), p < 0.00001 71% |
|              | 3 months MD 2 (64:70) | −29.15 (−55.10, −3.19), p = 0.03 67% |
|              | 6 months MD 4 (85:96) | −18.58 (−30.57, −6.58), p = 0.002 51% |
|              | Sleep awakenings (diary) | Post-treatment MD 2 (51:37) | −0.23 (−0.88, 0.42), p = 0.48 0% |
|              | Sleep - total sleep time (diary) | Post-treatment MD 8 (174:164) | −6.70 (−31.98, 18.58), p = 0.60 58% |
|              | 3 months MD 2 (64:70) | −22.10 (−69.11, 24.91), p = 0.36 73% |
|              | 6 months MD 4 (85:96) | −2.44 (−22.15, 17.26), p = 0.81 0% |
|              | Sleep - onset latency (actigraph) | Post-treatment MD 3 (80:81) | −3.89 (−14.55, 6.77), p = 0.47 63% |
|              | 6 months MD 3 (56:64) | −4.15 (−9.97, 1.68), p = 0.16 0% |
|              | Sleep - efficiency (actigraph) | Post-treatment MD 4 (96:96) | −1.58 (−5.59, 2.42), p = 0.44 50% |
|              | 6 months MD 3 (55:65) | −2.17 (−5.42, 1.09), p = 0.19 0% |
|              | Sleep – wake after sleep onset (actigraph) | Post-treatment MD 3 (80:81) | −13.12 (−24.39, −1.85), p = 0.02 0% |
|              | 6 months MD 3 (56:64) | −8.31 (−20.16, 3.53), p = 0.17 0% |
|              | Sleep - Total sleep time (actigraph) | Post-treatment MD 4 (96:96) | 13.66 (−18.58, 45.91), p = 0.41 51% |
|              | 6 months MD 3 (56:64) | 17.50 (−9.93, 44.93), p = 0.21 11% |
|              | Sleep - onset latency (polysomnograph) | Post-treatment MD 2 (65:74) | 5.89 (−9.16, 20.93), p = 0.44 0% |
|              | 6 months MD 2 (56:59) | −5.24 (−23.33, 12.84), p = 0.57 0% |
|              | Sleep - efficiency (polysomnograph) | Post-treatment MD 2 (65:74) | −5.06 (−9.55, −0.57), p = 0.03 0% |
|              | 6 months MD 2 (56:59) | −4.39 (−9.11, 0.33), p = 0.07 0% |
|              | Sleep – wake after sleep onset (polysomnograph) | Post-treatment MD 2 (65:74) | −33.91 (−51.82, −16.00), p = 0.0002 0% |
|              | 6 months MD 2 (56:59) | −22.45 (−41.93, −2.97), p = 0.02 0% |
|              | Sleep - Total sleep time (polysomnograph) | Post-treatment MD 2 (65:74) | 16.41 (−23.60, 56.42), p = 0.42 29% |
|              | 6 months MD 2 (56:59) | 18.63 (−12.36, 49.62), p = 0.24 0% |
|              | Sleep - Epworth Sleepiness Scale | Post-treatment MD 2 (22:19) | −0.58 (−3.30, 2.14), p = 0.68 0% |
Table 2 (continued)

| Follow up            | Effect measure | Number of studies (participants) | Effect estimate (95%CI), p-value | I² |
|----------------------|----------------|-----------------------------------|----------------------------------|----|
| **Pain**             |                |                                   |                                  |    |
| Post-treatment       | SMD            | 9 (192:178)                      | −0.24 (−0.45, −0.03), p = 0.02  | 0% |
| 3 months             | SMD            | 3 (83:83)                        | −0.31 (−0.69, 0.07), p = 0.11   | 31%|
| 6 months             | SMD            | 3 (66:70)                        | 0.07 (−0.27, 0.41), p = 0.68    | 0% |
| **Anxiety**          |                |                                   |                                  |    |
| Post-treatment       | SMD            | 3 (53:53)                        | −0.54 (−1.01, −0.06), p = 0.03  | 28%|
| **Depression**       |                |                                   |                                  |    |
| Post-treatment       | SMD            | 6 (123:113)                      | −0.57 (−1.05, −0.08), p = 0.008 | 65%|
| 3 months             | SMD            | 2 (48:43)                        | −0.37 (−0.79, 0.05), p = 0.08   | 0% |
| **Cognitive behavioural therapy for insomnia versus sleep hygiene** | | | | |
| Sleep quality from questionnaire | | | | |
| Post-treatment       | SMD            | 3 (65:64)                        | −0.25 (−0.82, 0.33), p = 0.40   | 61%|
| 6 months             | SMD            | 2 (33:27)                        | −0.12 (−0.84, 0.59), p = 0.73   | 34%|
| **Pain**             |                |                                   |                                  |    |
| Post-treatment       | SMD            | 3 (65:64)                        | −0.51 (−1.17, 0.15), p = 0.13   | 70%|
| 6 months             | SMD            | 2 (33:27)                        | 0.54 (−1.30, 2.38), p = 0.57    | 85%|
| **Health related quality of life** | | | | |
| Post-treatment       | SMD            | 2 (50:47)                        | −0.79 (−1.20, −0.37), p = 0.0002| 0% |
| **Anxiety**          |                |                                   |                                  |    |
| Post-treatment       | SMD            | 2 (50:47)                        | −0.32 (−0.72, 0.08), p = 0.12   | 0% |
| **Depression**       |                |                                   |                                  |    |
| Post-treatment       | SMD            | 2 (50:47)                        | −0.61 (−1.05, −0.18), p = 0.006 | 11%|
| **Cognitive behavioural therapy for insomnia and pain versus no treatment, wait list or attentional control** | | | | |
| Sleep quality from questionnaire | | | | |
| Post-treatment       | SMD            | 4 (180:192)                      | −0.79 (−1.58, 0.00), p = 0.05   | 88%|
| 3 months             | SMD            | 2 (56:56)                        | −0.41 (−1.96, 1.15), p = 0.61   | 94%|
| 6 months             | SMD            | 2 (142:150)                      | −0.76 (−1.85, 0.33), p = 0.17   | 92%|
| **Sleep - Total sleep time (diary)** | | | | |
| Post-treatment       | MD             | 2 (40:34)                        | −61.58 (−105.25, −17.91), p = 0.006| 0% |
| **Pain**             |                |                                   |                                  |    |
| Post-treatment       | SMD            | 4 (180:182)                      | −0.13 (−0.36, 0.10), p = 0.28   | 10%|
| 3 months             | SMD            | 2 (56:56)                        | −0.48 (−0.86, −0.10), p = 0.01  | 0% |
| 6 months             | SMD            | 3 (161:395)                      | −0.25 (−0.62, 0.13), p = 0.20   | 58%|
| **Health related quality of life** | | | | |
| Post-treatment       | MD             | 2 (58:65)                        | −1.77 (−5.33, 1.78), p = 0.33   | 98%|
| 3 months             | MD             | 2 (54:49)                        | −1.84 (−5.90, 2.22), p = 0.37   | 98%|
| **Depression**       |                |                                   |                                  |    |
| Post-treatment       | SMD            | 2 (33:42)                        | 0.14 (−0.54, 0.83), p = 0.68    | 33%|
| **Cognitive behavioural therapy for insomnia and pain versus cognitive behavioural therapy for pain** | | | | |
| Sleep quality from questionnaire | | | | |
| Post-treatment       | SMD            | 3 (146:150)                      | −0.47 (−1.19, 0.25), p = 0.20   | 75%|
| **Pain**             |                |                                   |                                  |    |
| Post-treatment only  | SMD            | 3 (146:150)                      | −0.03 (−0.26, 0.20), p = 0.78   | 0% |
| **Health related quality of life** | | | | |
| Post-treatment       | SMD            | 2 (139:145)                      | 0.03 (−0.20, 0.26), p = 0.81    | 0% |
| **Depression**       |                |                                   |                                  |    |
| Post-treatment       | SMD            | 2 (33:33)                        | −0.34 (−1.05, 0.37), p = 0.35   | 28%|

SMD: Standardised Mean Difference, CI: Confidence Interval, MD: Mean Difference, p: p-value
-0.24 (95% CI -0.45, -0.03; p = 0.02), but this was not apparent in 3 studies at low risk of bias, SMD -0.19 (95% CI -0.59, 0.220), p = 0.36, or sustained at 3 months or longer (Fig. 3). There was no evidence of heterogeneity. In 2 studies at unclear risk of bias including 79 people specifically with fibromyalgia, there was no benefit for CBT-I, SMD -0.31 (95% CI -0.75, 0.140), p = 0.18. There was no suggestion of heterogeneity.

No studies reported health-related quality of life outcomes. For psychological health, information from studies was mainly limited to post-treatment with benefit suggested for anxiety SMD -0.54 (95% CI -1.01, -0.06), p = 0.03 with slight heterogeneity (I² 28%), and depression SMD -0.57 (95% CI -1.05, -0.08), p = 0.08 with high heterogeneity (I² 65%). In 1 study exclusively including people with fibromyalgia, anxiety and depression were reduced in the Group receiving CBT-I compared with controls.

In 2 studies, a further comparison was made between CBT-I and CBT solely for pain (CBT-P). One was a small pilot study with 11 people with chronic pain randomised to the 2 interventions [44]. In a larger study with unclear risk of bias due to high losses to follow up, there was no suggestion of benefit for any outcome for CBT-I compared with CBT-P in people with fibromyalgia [43].

**Brief education with CBT component versus no treatment**
In 1 study with 132 people with chronic non-cancer pain randomised, a brief educational intervention incorporating sleep hygiene and cognitive behavioural strategies was compared with wait list controls [49]. The study was at high risk of bias due to large losses to follow up. Only sleep outcomes were reported and there was no difference between groups in sleep quality or sleep diary measures excepting diary recorded sleep latency which favoured the intervention.

**Cognitive behavioural therapy for insomnia (CBT-I) versus sleep hygiene**
CBT-I was compared with a sleep hygiene intervention in 4 randomised trials [41, 50–52]. Studies included 270 participants, all with chronic pain from fibromyalgia. Risk of bias was low in 3 studies [50–52] and unclear in 1 due to limited reporting of methods [41].
Data on overall sleep quality was available for 3 studies [41, 50, 51]. There was no difference between randomised groups post-treatment, SMD -0.25 (95%CI -0.82, 0.33), \( p = 0.40 \) but heterogeneity was high (I\(^2\) 61%). Excluding the study at unclear risk of bias removed heterogeneity and there was an improvement in overall sleep quality after CBT-I compared with sleep hygiene, SMD -0.53 (95%CI -0.94, −0.12), \( p = 0.01 \) [50, 51]. Evidence relating to longer term outcomes was limited but with no clear suggestion of benefit for CBT-I over sleep hygiene.

In 3 studies, results for pain outcome were similar in direction to sleep quality [41, 50, 51] but a difference favouring CBT-I post-treatment was only apparent in the 2 studies at low risk of bias, SMD -0.85 (95%CI -1.26, −0.43), \( p < 0.0001 \) [50, 51] with no evidence of heterogeneity. Health-related quality of life was improved in people receiving CBT-I compared with sleep hygiene in 2 studies with 97 people randomised, both at low risk of bias and with no heterogeneity, SMD -0.79 (95%CI -1.20, −0.37), \( p = 0.0002 \). Improvements in pain and health-related quality of life were not evident at longer follow up. Evidence relating to psychological health was limited to 2 studies at low risk of bias with 97 patients randomised. There was no benefit for CBT-I compared with sleep hygiene for anxiety, SMD -0.32 (95%CI -0.72, 0.08), \( p = 0.12 \) with no heterogeneity, but depression was reduced, SMD -0.61 (95%CI-1.05, −0.18), \( p = 0.006 \) with slight heterogeneity (I\(^2\) 11%).

**Cognitive behavioural therapy for insomnia and pain (CBT-IP) versus control**

In 4 studies with 432 participants randomised, cognitive behavioural therapy focusing on insomnia and pain (CBT-IP) was compared with no treatment, wait list or attentional control [44, 53–55]. In 2 studies, the cause of pain was fibromyalgia [53, 54], and in 1 each, osteoarthritis [55], or diverse causes [44]. Risk of bias was high in 1 study due to large losses to follow up at the end of treatment [54], and unclear in 1 due to lack of methodological detail [53]. A third was a small pilot study [44]. For
sleep quality, data for meta-analysis was available from all studies post-treatment (Fig. 4). Compared with controls, people receiving CBT-IP had marginally improved sleep quality and improved diary recorded total sleep time, SMD -0.79 (95%CI -1.58, 0.00), \( p = 0.05 \), and MD -61.58 min (95%CI -105.25, -17.91), \( p = 0.006 \), respectively. In the 1 study at low risk of bias, the difference in sleep quality was smaller and in 2 studies with data, the benefit relating to sleep quality was not sustained at 3 and 6 months. In 2 studies with 299 people exclusively with fibromyalgia, there was no evidence for a difference in sleep quality, SMD -0.88 (95%CI -2.16, 0.41), \( p = 0.18 \) but heterogeneity was high (I² 94%).

All 4 studies reported a pain outcome post-treatment (Fig. 5) [44, 53–55]. There was no benefit for CBT-IP compared with control except in 2 studies at 3 months [53, 54]. No benefit was seen in the 2 studies of people with fibromyalgia or the study at low risk of bias [55]. Information was limited relating to health-related quality of life and psychological health but in 2 studies there was no difference post-treatment in quality of life or depression between groups.

In 4 studies with 415 people randomised, CBT-IP was compared with a control condition that included cognitive behavioural therapy for pain (CBT-P) with no specific focus on sleep [43, 44, 54, 55]. In studies with data suitable for meta-analysis there was no difference in sleep quality, pain or health-related quality of life post-treatment, or anxiety or depression post-treatment or at 3 months, and this was not changed if restricted to studies at low or unclear risk of bias or in a study exclusively including people with fibromyalgia. Heterogeneity was high (I² 75%).

In 1 study at unclear risk of bias due to limited reporting of methods with 95 people with fibromyalgia randomised, hypnosis additional to CBT-IP was evaluated [53]. Compared with CBT-IP alone, there were no differences in sleep quality, diary assessed total sleep time, or pain post-treatment and at 3 and 6 months. Health-related quality of life was improved after CBT-IP with
hypnosis compared with CBT-IP but only post-treatment and at 3 months. At 6 months, CBT-IP without hypnosis showed a favourable outcome. General psychologi cal health was improved in people receiving CBT-IP and hypnosis compared with CBT-IP post-treatment and at 3 and 6 months.

Acceptance and commitment therapy versus attentional control or exercise
In 1 study, 299 people with chronic pain were randomised to a 7-week course of acceptance and commitment therapy based stress management, or to a control discussion group of similar intensity and duration, or to group-based exercise [56]. The study was at high risk of bias due to high losses to follow up. There were no differences between randomised groups in insomnia severity, pain, anxiety or depression at the end of treatment or at 6 month follow up.

Mindfulness versus control
In 2 studies including 239 people with fibromyalgia pain, a mindfulness-based intervention was compared with waiting list controls [57], or CBT with no specific focus on sleep [58]. Risk of bias was unclear in the former mainly because of wait list controls [57], while in the latter, risk of bias was high mainly due to large losses to follow up [58]. In meta-analysis, there was benefit for improved sleep quality post-treatment after mindfulness intervention compared with wait list controls or CBT, SMD -0.41 (95%CI -0.72, -0.11), \( p = 0.008 \), and this was consistent at 3 [57], and 6 months [58]. For pain, and health-related quality of life, there was no consistent evidence for benefit of mindfulness intervention over CBT or wait list control.

Relaxation versus control
In 1 study with 53 people with fibromyalgia pain, group and individualised applied relaxation was compared with a sleep hygiene-based educational intervention, and a wait list control [59]. Risk of bias was low for the comparison of interventions, but unclear in relation to the wait list control. There were no differences post-treatment between sleep quality, pain, or health-related quality of life in relaxation and wait list control groups. This was also the case for relaxation compared with education, with the exception of improved pain in the relaxation group at 6 month follow up. Risk of bias was high at this
follow up time due to high and uneven losses to follow up.

Sleep hygiene versus control
Further to the studies comparing sleep hygiene intervention with CBT-I [41, 50–52], in 2 studies with 54 people with fibromyalgia pain, sleep hygiene was compared with untreated controls [41, 59]. Risk of bias in 1 study with a wait list control was unclear [59], and high in the other due to uneven randomisation and large loss to follow up [41]. In meta-analysis, there was no benefit post-treatment for sleep hygiene compared with controls for sleep quality or pain (Table 2). A difference in general psychological health favouring sleep hygiene over control was limited to a single study at high risk of bias.

Physical exercise
Group-based exercise versus control
In 5 studies with 697 people randomised, exercise programmes were compared with usual care or an attentional control [56, 60–63]. Pain conditions were, arthritis, rheumatoid arthritis, fibromyalgia, low back pain and general chronic pain. In 4, the programme was delivered at a clinic or in a group [56, 60, 62, 63], and in 1 at home [61]. Studies were at low [61], unclear [60, 62, 63], or high risk [56] of bias. One was a feasibility study [62]. For the 3 studies with data [56, 61, 63], questionnaire assessed sleep quality was not improved in the exercise groups compared with controls, SMD -0.10 (95%CI -0.31, 0.12), p = 0.39 and this was consistent in the study at low risk of bias. There was no difference in 1 study with sleep measures at 6 months [56]. One study including 321 people with arthritis comparing a low to moderate intensity physical activity programme with wait list controls and at unclear risk of bias, presented sleep data dichotomised into groups of people with no problems and those with moderate to severe problems [63]. In intention to treat analyses, the authors reported post-treatment benefit for the intervention compared with controls for the outcome waking up tired, p < 0.001, but not trouble falling asleep or staying asleep, waking up at night, or trouble staying asleep. Benefit was not maintained at 3 or 6 months. In one study including 53 people with fibromyalgia, the authors reported that sleep quality was improved in the group receiving group-based exercise compared with controls, p = 0.051 [56, 60].

Pain in 3 studies was reduced in the exercise group compared with controls, SMD -0.52 (95%CI -0.76,
at low risk of bias. No heterogeneity was evident. Also in the study at low risk of bias, sleep efficiency measured by questionnaire and diary was improved in people receiving Tai Ji Quan [66], but there were no differences in other sleep, pain or psychological measures.

**Physical therapy**
Physical therapy modalities were evaluated in 6 studies.

**Hydrotherapy**
Two studies described treatment comparisons including hydrotherapy [68, 69]. In 1 study including 81 people with fibromyalgia pain, a course of hydrotherapy with stretching was compared against hydrotherapy with Tai Chi [68]. Risk of bias was high due to large losses to follow up. Results were presented as graphs and interpreted by the authors as showing no differences between randomised groups.

A course of hydrotherapy was compared with conventional physiotherapy in 1 study at low risk of bias with 50 people with fibromyalgia randomised [69]. Post-treatment, total sleep time was marginally higher and total nap time lower in people who received hydrotherapy compared with conventional physiotherapy. Pain and general psychological health did not differ between groups post-treatment. Adverse events were assessed in 1 study [68], 3 patients dropped out of the intervention group due to pain exacerbation (n = 2) and chlorine hypersensitivity (n = 1).

**Massage or manual therapy**
A course of massage therapy was compared with relaxation therapy in 1 study with 30 people with low back pain [70]. Reporting was limited and risk of bias unclear. A large difference in sleep disturbance between groups was apparent at baseline. Post-treatment, there were no differences between groups. A possible favourable sleep disturbance outcome for massage therapy at follow up may have been masked by the difference at baseline.

In 1 study with 89 people with fibromyalgia pain randomised, a course of manual therapy was compared with no treatment [71]. The study was at unclear risk of bias due to lack of blinding of the intervention. Results reported separately for men and women suggested improved sleep quality, pain, health-related quality of life and depression in people who received manual therapy compared with controls.

**Physical therapy programme**
In 1 study including 60 people with fibromyalgia, a physical therapy programme with hot pack, ultrasound, transcutaneous electrical nerve stimulation and low power laser was compared with an untreated control group [72].
Risk of bias was high mainly through lack of blinding. The authors reported improvements in sleep quality, pain and health-related quality of life in people receiving the physical therapy programme compared with controls.

**Pompage**
A course of pompage was compared with controls in 1 study with 23 people with fibromyalgia pain randomised [81]. All participants received stretching and aerobic exercises. Risk of bias was high to large losses to follow up. There was no improvement in sleep quality or pain post-treatment in people receiving pompage compared with controls.

**Other interventions**

**Acupressure**
Auricular point acupressure was compared with sham auricular point acupressure in 1 study including 61 people with chronic low back pain [74]. Risk of bias was high due to large and uneven losses to follow up. People receiving active intervention reported improved sleep quality post-treatment compared with the sham group. Differences in other sleep measures were marginal. In another study, self administered relaxing or stimulating acupressure was compared with usual care in 67 people with low back pain [75]. Risk of bias was unclear due to limited reporting of methods. With results shown as graphs, the authors reported no improvement in sleep quality between acupressure groups, and compared with controls. Pain was reduced in acupressure groups but not in controls. Adverse events were assessed in 1 study [75] with four mild events related to acupressure treatments involving too much pressure to acupoints.

**Bright light treatment**
In 1 study, visor delivered bright light treatment was compared with sham therapy in 19 people with fibromyalgia pain randomised in a crossover trial [76]. The study was at high risk of bias due to concerns about the randomisation procedure and blinding. Post-treatment, sleep quality was similar between groups. Presented as graphs, the authors reported improved pain in the mattress overlay group compared with controls but no difference in sleep quality.

**Foot reflexology versus control**
In 1 study, 68 people with pain from rheumatoid arthritis were randomised to a course of foot reflexology or control [77]. Risk of bias was unclear due to lack of blinding. Actigraphy was only completed for 25% of participants. Sleep quality and pain were improved in people receiving reflexology compared with controls.

**Transcranial stimulation versus sham**
One study compared transcranial stimulation with sham in 16 patients with musculoskeletal pain [78]. This was a small feasibility study with unclear risk of bias due to limited reporting. People receiving transcranial stimulation had reduced pain after the intervention compared with controls, but there was no difference in sleep quality. The authors described aspects of study conduct to advise future evaluations of transcranial stimulation.

**Mattress interventions versus control**
In 1 study at low risk of bias with 30 people with fibromyalgia pain randomised, a period of sleep on a magnetic mattress pad was compared with sham [79]. Sleep, pain and ADL focused quality of life were improved in the intervention group compared with sham control. Adverse events were assessed with none reported. A period of sleep on a supportive mattress overlay was compared with untreated controls in 1 study with 38 people with low back pain randomised [80]. The study was at high risk of bias due to concerns about the randomisation procedure and blinding. Post-treatment, sleep quality was similar between groups. Presented as graphs, the authors reported improved pain in the mattress overlay group compared with controls but no difference in sleep quality.

**Discussion**
We identified 42 randomised trials evaluating sleep interventions for people with chronic pain. CBT interventions provided the largest evidence base with CBT-I interventions demonstrating benefit post-treatment compared with controls for improved sleep quality, however evidence for a longer-term sustained benefit was lacking. Evidence in people with specific conditions was limited due to primary study limitations and statistical heterogeneity. Numerous interventions were evaluated in small numbers of studies, but evidence was insufficient to draw conclusions on effectiveness.

Findings from meta-analysis found that CBT-I (9 studies) and CBT-IP (4 studies) demonstrated a medium to large (−0.79) effect compared with control for sleep quality at post-treatment. Differences between groups at 6 months was slightly reduced with a medium effect size for CBT-I and was not sustained for CBT-IP. CBT-I showed a small improvement in pain outcome post-treatment but this was not sustained. However, there was high heterogeneity in studies which should be considered when interpreting results. In 4 studies comparing CBT-IP with CBT-P only, no differences in pain outcome were found. Due to the active control it is therefore only possible to infer that adding insomnia specific content to CBT-P does not have an additional impact on pain.
outcomes. Our findings regarding the impact of CBT-I interventions on sleep quality reflect the existing literature. A recent systematic review of CBT-I therapies in patients with chronic non-malignant pain showed significant treatment effects immediately post-treatment for global measures of sleep [73]. Condition specific reviews show similar results with CBT therapies improving sleep outcomes in the short-term for patients with lower back pain, fibromyalgia, and osteoarthritis [82–84].

Our results demonstrate that improving sleep for people with chronic pain is possible, and that CBT approaches have the strongest evidence base. Poor sleep has a negative impact on optimism, sociability, and psychological functioning [85]. Poor sleep also has clear links with depression and pain catastrophising, both of which can affect pain management and coping. Pain catastrophising is linked with maladaptive coping techniques and depression is linked to lack of engagement in treatment [86, 87]. CBT approaches are already widely used in pain management with a focus on coping strategies and behavioural rehearsal [88]. This systematic review demonstrates that additional focus on sleep improvement could be of benefit. As CBT is an established treatment approach, future work should focus on how best to implement CBT sleep interventions for people with chronic pain and foster more equitable access to support, particularly for underserved populations. Results from the National Pain audit highlighted that service provision for the management of chronic pain in the UK is inadequate [89]. Only 40% of pain clinics in the UK are multidisciplinary which presents a challenge for implementation of psychological interventions such as CBT.

Evaluation of the longer-term effectiveness of sleep interventions for people with chronic pain is lacking. Although evidence suggests that CBT interventions improve sleep immediately post treatment, effects reduce over time. In addition, most studies had follow-up data collected at 3 months or less post-intervention, with 9 studies reporting 6-month outcomes, and 2 studies reporting 12-month outcomes. Understanding longer-term effectiveness of these interventions is crucial for people with chronic pain. Due to the nature of the condition, individuals with chronic pain may experience disturbed sleep for many months and years, therefore effective interventions need to have sustained effects.

Assessment of outcomes within the trials included in this review varied considerably, and this was particularly notable for the secondary outcomes. This heterogeneity limits comparison between studies, particularly for health-related quality of life and psychological wellbeing, because outcome measures assess different aspects of these constructs, for example general mood assessment versus specific anxiety or depression measures. The issue of heterogeneity across randomised trials is well established and initiatives such as COMET have been addressing this through the development of core outcome sets [90, 91]. Core outcome sets provide a minimum set out of outcomes to be used in all trials of a certain focus, ensuring comparability across multiple studies. The challenge of a review of this scope is that the interventions included are varied and include both generic and condition specific measures. As the evidence base builds, focused reviews on areas of promise, such as CBT and third wave therapies, could offer benefit. The feasibility of developing a sleep core outcome set could also be explored as this would provide opportunity for greater comparison across interventions.

Nine of the 42 studies in this review provided data on adverse events with no serious adverse events reported [42, 43, 45–47, 62, 68, 75, 79]. Assessment of adverse events is vital for patient safety, however unlike in trials of pharmacological treatments where monitoring and reporting of harm outcomes is mandatory, behavioural and psychological interventions are not held to the same account [92]. In 2004 the CONSORT group provided ten recommendations for reporting harm outcomes in trials [93]. All except 5 of the studies included in our review were published after the recommendations. This demonstrates a need for evaluations of psychological and behavioural interventions to improve reporting of harm outcomes.

**Strengths and limitations**

This review has strengths and limitations that should be considered when interpreting the findings. The method used in this review was robust and systematic, following Cochrane guidance [36]. The review provides a comprehensive overview of the existing literature on sleep interventions for patients with chronic pain, but as included studies addressed a wide variety of interventions with small numbers of studies for each intervention, this limited opportunity for meta-analysis. The population of people with chronic pain that we considered is heterogeneous with a range of underlying medical conditions. However, chronicity reflects pain that persists for 3 months or longer, and chronic pain is usually the affected person’s main clinical problem. Secondary outcome measure tools used were highly heterogeneous, which limited comparison between studies.

**Conclusion**

CBT approaches have the potential to be an effective treatment to improve sleep for people with chronic pain, but further high-quality primary research is required to explore refinements that will ensure parallel improvements to pain, quality of life, psychological health and
maintain all benefits in the long term. Individuals who experience depression and pain catastrophising may particularly benefit from sleep interventions. As CBT is an established treatment approach, future work should focus on how best to deliver these interventions, for instance by exploring any difference between online or face-to-face delivery or differences between delivery professionals. Importantly, future research could focus on how best to facilitate equal access and outcomes in underserved populations.

Primary research is also needed to evaluate the effectiveness of interventions including mindfulness, aquatic exercise and hydrotherapy, Tai Ji Quan, manual therapy, physical therapy programmes, acupressure, foot reflexology and magnetic mattress pads. Individuals who experience chronic pain could benefit from interventions that address sleep, and research is needed to assess any impact of sleep interventions on pain.

Supplementary Information

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Additional file 1.
Additional file 2.
Additional file 3.
Additional file 4.
Additional file 5.

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Authors' contributions

KW: lead and corresponding author, study conception and design, data screening, data extraction, manuscript writing and preparation. JD: study design, data searches, data screening, data extraction, meta-analysis, review of the manuscript. VW: study conception and design, data screening, data extraction, review of the manuscript. AB: study conception and design, data searches, data screening, data extraction, meta-analysis, narrative synthesis, manuscript writing and review. RGH: study conception and design, review of the manuscript. All authors: approval of the submitted versions and accountability for their contributions.

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Availability of data and materials

No additional data are available. Extracted data is included within the manuscript and supplementary materials.

Declarations

Ethics approval and consent to participate

This study did not recruit any participants or collect any primary data, therefore ethical approval was not required.
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