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## Psychological therapies for the management of chronic and recurrent pain in children and adolescents (Review)

Eccleston C, Palermo TM, Williams ACDC, Lewandowski Holley A, Morley S, Fisher E, Law E

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[Intervention Review]

# Psychological therapies for the management of chronic and recurrent pain in children and adolescents

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## ABSTRACT

### Background

This is an update of the original Cochrane review first published in Issue 1, 2003, and previously updated in 2009 and 2012. Chronic pain affects many children, who report severe pain, disability, and distressed mood. Psychological therapies are emerging as effective interventions to treat children with chronic or recurrent pain. This update focuses specifically on psychological therapies delivered face-to-face, adds new randomised controlled trials (RCTs), and additional data from previously included trials.

### Objectives

There were three objectives to this review. First, to determine the effectiveness on clinical outcomes of pain severity, disability, depression, and anxiety of psychological therapy delivered face-to-face for chronic and recurrent pain in children and adolescents compared with active treatment, waiting-list, or standard medical care. Second, to evaluate the impact of psychological therapies on depression and anxiety, which were previously combined as 'mood'. Third, we assessed the risk of bias of the included studies and the quality of outcomes using the GRADE criteria.

### Search methods

Searches were undertaken of CENTRAL, MEDLINE, EMBASE, and PsycINFO. We searched for further RCTs in the references of all identified studies, meta-analyses, and reviews. Trial registry databases were also searched. The date of most recent search was January 2014.

### Selection criteria

RCTs with at least 10 participants in each arm post-treatment comparing psychological therapies with active treatment, standard medical care, or waiting-list control for children or adolescents with episodic, recurrent or persistent pain were eligible for inclusion. Only trials conducted in person (face-to-face) were considered. Studies that delivered treatment remotely were excluded from this update.

## Data collection and analysis

All included studies were analysed and the quality of outcomes were assessed. All treatments were combined into one class, psychological treatments. Pain conditions were split into headache and non-headache. Both conditions were assessed on four outcomes: pain, disability, depression, and anxiety. Data were extracted at two time points; post-treatment (immediately or the earliest data available following end of treatment) and at follow-up (between three and 12 months post-treatment).

## Main results

Seven papers were identified in the updated search. Of these papers, five presented new trials and two presented follow-up data for previously included trials. Five studies that were previously included in this review were excluded as therapy was delivered remotely. The review thus included a total of 37 studies. The total number of participants completing treatments was 2111. Twenty studies addressed treatments for headache (including migraine); nine for abdominal pain; two for mixed pain conditions including headache pain, two for fibromyalgia, two for recurrent abdominal pain or irritable bowel syndrome, and two for pain associated with sickle cell disease.

Analyses revealed psychological therapies to be beneficial for children with chronic pain on seven outcomes. For headache pain, psychological therapies reduced pain post-treatment and at follow-up respectively (risk ratio (RR) 2.47, 95% confidence interval (CI) 1.97 to 3.09,  $z = 7.87$ ,  $p < 0.01$ , number needed to treat to benefit (NNTB) = 2.94; RR 2.89, 95% CI 1.03 to 8.07,  $z = 2.02$ ,  $p < 0.05$ , NNTB = 3.67). Psychological therapies also had a small beneficial effect at reducing disability in headache conditions post-treatment and at follow-up respectively (standardised mean difference (SMD) -0.49, 95% CI -0.74 to -0.24,  $z = 3.90$ ,  $p < 0.01$ ; SMD -0.46, 95% CI -0.78 to -0.13,  $z = 2.72$ ,  $p < 0.01$ ). No beneficial effect was found on depression post-treatment (SMD -0.18, 95% CI -0.49 to 0.14,  $z = 1.11$ ,  $p > 0.05$ ). At follow-up, only one study was eligible, therefore no analysis was possible and no conclusions can be drawn. Analyses revealed a small beneficial effect for anxiety post-treatment (SMD -0.33, 95% CI -0.61 to -0.04,  $z = 2.25$ ,  $p < 0.05$ ). However, this was not maintained at follow-up (SMD -0.28, 95% CI -1.00 to 0.45;  $z = 0.75$ ,  $p > 0.05$ ).

Analyses revealed two beneficial effects of psychological treatment for children with non-headache pain. Pain was found to improve post-treatment (SMD -0.57, 95% CI -0.86 to -0.27,  $z = 3.74$ ,  $p < 0.01$ ), but not at follow-up (SMD -0.11, 95% CI -0.41 to 0.19,  $z = 0.73$ ,  $p > 0.05$ ). Psychological therapies also had a beneficial effect for disability post-treatment (SMD -0.45, 95% CI -0.71 to -0.19,  $z = 3.40$ ,  $p < 0.01$ ), but this was not maintained at follow-up (SMD -0.35, 95% CI -0.71 to 0.02,  $z = 1.87$ ,  $p > 0.05$ ). No effect was found for depression or anxiety post-treatment (SMD -0.07, 95% CI -0.30 to 0.17,  $z = 0.54$ ,  $p > 0.05$ ; SMD -0.15, 95% CI -0.36 to 0.07,  $z = 1.33$ ,  $p > 0.05$ ) or at follow-up (SMD 0.06, 95% CI -0.16 to 0.28,  $z = 0.53$ ,  $p > 0.05$ ; SMD 0.05, 95% CI -0.24 to 0.33,  $z = 0.32$ ,  $p > 0.05$ ).

## Authors' conclusions

Psychological treatments delivered face-to-face are effective in reducing pain intensity and disability for children and adolescents (<18 years) with headache, and therapeutic gains appear to be maintained, although this should be treated with caution for the disability outcome as only two studies could be included in the follow-up analysis. Psychological therapies are also beneficial at reducing anxiety post-treatment for headache. For non-headache conditions, psychological treatments were found to be beneficial for pain and disability post-treatment but these effects were not maintained at follow-up. There is limited evidence available to estimate the effects of psychological therapies on depression and anxiety for children and adolescents with headache and non-headache pain. The conclusions of this update replicate and add to those of the previous review which found that psychological therapies were effective in reducing pain intensity for children with headache and non-headache pain conditions, and these effects were maintained at follow-up for children with headache conditions.

## PLAIN LANGUAGE SUMMARY

### Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Psychological therapies (e.g. relaxation, hypnosis, coping skills training, biofeedback, and cognitive behavioural therapy) may help people manage pain and its disabling consequences. Therapies can be delivered face-to-face by a therapist, via the Internet, by telephone call, or by computer programme. This review focuses on treatments that are delivered face-to-face by a therapist. For children and adolescents there is evidence that both relaxation and cognitive behavioural therapy (treatment that helps people test and revise their thoughts and actions) are effective in reducing the intensity of pain in chronic headache, recurrent abdominal pain, fibromyalgia, and sickle cell disease immediately after treatment.

Psychological therapies also have a lasting effect in reducing pain and disability for chronic headache. Fifty-six per cent of children who were treated with psychological therapies reported less pain compared with 22% of children who did not receive a psychological therapy. Anxiety was also reduced for children with headaches immediately following treatment. Psychological therapies also reduce pain and disability for children with mixed pain conditions (excluding headache) immediately following treatment. However, we did not find that any treatment effects were maintained at follow-up (between 3-12 months after the end of treatment) for children with mixed pain conditions. Psychological therapies did not produce changes in depression in children with either headache or non-headache conditions, and anxiety did not change in children with non-headache conditions receiving psychological therapies.

More studies are needed to understand whether psychological therapies can improve depression and anxiety and have more lasting effects on pain and disability in other groups of young people who have chronic pain.

## SUMMARY OF FINDINGS FOR THE MAIN COMPARISON *[Explanation]*

Psychological therapies compared with any control for children with frequent headache						
<b>Patient or population:</b> children and adolescents with frequent headache <b>Settings:</b> Community <b>Intervention:</b> Psychological therapies <b>Comparison:</b> Any control						
Outcome	Probable outcome with control	Probable outcome with intervention	NNT and/or relative effect (95% CI)	No of participants	Quality of the evidence (GRADE)	Comments
<b>Pain</b> (low scores mean lower pain ratings)	<b>220 in 1000</b>	<b>560 in 1000</b>	<b>NNT = 2.94</b> <b>RR 2.47</b> (1.97 to 3.09)	714 participants, 302 events (15 studies)	⊕⊕○○ <b>low</b>	Majority of studies included in analysis had high risk of bias, and mostly wait-list controls
<b>Pain (at follow-up)</b> (low scores mean lower pain ratings)	<b>478 in 1000</b>	<b>750 in 1000</b>	<b>NNT = 3.67</b> <b>RR 2.89</b> (1.03 to 8.07)	251 participants, 158 events (5 studies)	⊕○○○ <b>very low</b>	Majority of studies included in analysis had high risk of bias, wide confidence intervals, heterogeneity >45%, low number of participants, and some studies did not report full outcomes in published paper
<b>Disability</b> (low scores mean lower disability ratings)		The mean disability in the intervention groups was <b>0.49 standard deviations lower</b> (0.74 to 0.24 lower)		263 participants (3 studies)	⊕⊕○○ <b>low</b>	A low number of participants could be included in the analysis and some studies did not report full outcomes in published paper SMD -0.49 (-0.74 to -0.24)

<p><b>Disability (at follow-up)</b> (low scores mean lower disability ratings)</p>	<p>The mean disability (at follow-up) in the intervention groups was <b>0.46 standard deviations lower</b> (0.78 to 0.13 lower)</p>	<p>148 participants (2 studies)</p>	<p>⊕⊕○○ <b>low</b></p>	<p>A low number of participants could be included in the analysis SMD -0.46 (-0.78 to -0.13)</p>
<p><b>Depression</b> (low scores mean lower depression ratings)</p>	<p>The mean depression in the intervention groups was <b>0.18 standard deviations lower</b> (0.49 lower to 0.14 higher)</p>	<p>164 participants (3 studies)</p>	<p>⊕⊕⊕○ <b>moderate</b></p>	<p>A low number of participants could be included in the analysis SMD -0.18 (-0.49 to 0.14)</p>
<p><b>Anxiety</b> (low scores mean lower anxiety ratings)</p>	<p>The mean anxiety in the intervention groups was <b>0.33 standard deviations lower</b> (0.61 to 0.04 lower)</p>	<p>203 participants (4 studies)</p>	<p>⊕⊕○○ <b>low</b></p>	<p>A low number of participants could be included in the analysis and some studies did not report full outcomes in published paper SMD -0.33 (-0.61 to -0.04)</p>
<p><b>Anxiety (at follow-up)</b> (low scores mean lower anxiety ratings)</p>	<p>The mean anxiety (at follow-up) in the intervention groups was <b>0.28 standard deviations lower</b> (1 lower to 0.45 higher)</p>	<p>67 participants (2 studies)</p>	<p>⊕○○○ <b>very low</b></p>	<p>The analysis included wide confidence intervals, heterogeneity &gt;45%, low number of participants, and some studies did not report full outcomes in published paper SMD -0.28 (-1.00 to 0.45)</p>

GRADE Working Group grades of evidence

**High quality:** Further research is very unlikely to change our confidence in the estimate of effect.

**Moderate quality:** Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.

**Low quality:** Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

**Very low quality:** We are very uncertain about the estimate.

NNT: number needed to treat for an additional beneficial effect; RR: risk ratio; CI: Confidence interval; SMD: Standardised Mean Difference.



## BACKGROUND

### Description of the condition

This review is an update of a previously published review in the *The Cochrane Library* on 'Psychological therapies for the management of chronic and recurrent pain in children and adolescents' (Eccleston 2012). Chronic and recurrent pain (pain lasting more than three months) is a common problem in young people. Recent epidemiology gives a prevalence of 15% to 30%, with 8% of children described as having severe and frequent pain (Perquin 2000; Perquin 2001; Stanford 2008). The most common location for pain is in the head, abdomen, and limbs (Perquin 2000). All types of chronic and recurrent pain are more commonly reported by girls, and there is a peak in incidence at ages 14 to 15 years (Stanford 2008). Young people report pain to be distressing and interfering, and in some cases this can be severely debilitating, affecting all aspects of a child's life (Bursch 1998; Palermo 2000), and the lives of their parents and family members (Palermo 2005; Walker 1989). The deleterious effects of untreated pain in childhood can also extend to adulthood (Fearon 2001).

### Description of the intervention

There is a broad family of treatments included in the general term 'psychological'. In essence, treatments are specifically designed to alter psychological processes thought to underlie or significantly contribute to pain, distress, and/or disability. The design of psychological treatments is normally informed by specific theories of the aetiology of human behaviour, or treatments have developed pragmatically through observation and study of response to intervention. Behavioural and cognitive treatments designed to ameliorate pain, distress, and disability were first introduced in adults over 40 years ago and have become well established (Fordyce 1968; Keefe 2004). A companion review of psychological treatments for the management of chronic pain in adults is also published (Williams 2012). Treatments were originally developed to be delivered in a face-to-face delivery format in which the patients and therapists work together in person to implement therapeutic strategies. Methods of remote delivery of psychological treatments have been developed. These are the subject of a separate Cochrane review ('Psychological therapies (remotely delivered) for the management of chronic and recurrent pain in children and adolescents' (protocol in press)).

### How the intervention might work

In paediatric practice, the treatments have a shorter history and different therapeutic aims and components than those for adults. In general, psychological treatments aim to control pain and modify situational, emotional, familial, and behavioural factors that play a

role in pain or related consequences (e.g. McGrath 1990). A variety of intervention strategies have been designed to reduce pain experience, increase comfort, and/or reduce associated disability and dysfunction in children with pain conditions. Behavioural strategies include relaxation training, biofeedback, and behavioural management programmes (e.g. teaching parents operant strategies to reinforce adaptive behaviours such as school attendance). Cognitive strategies include hypnosis, stress management, guided imagery, and cognitive coping skills (Palermo 2012).

Cognitive behavioural programmes incorporate elements of both behavioural and cognitive strategies. Given that headache and abdominal pain are the most common types of recurrent pain in children, most of the treatment literature has focused on these two populations. By far the most commonly described treatment is relaxation training and/or biofeedback for headache, and recommendations have been made to offer psychological treatment as a matter of routine care for children with headaches (Masek 1999). In an effort to enhance the efficiency of psychological treatments for children with headache, more recent treatment developments have compared different elements of relaxation training and biofeedback with a variation in treatment formats (individual and group), treatment dose, and treatment setting (clinic, school, and home).

Psychological therapies have also been developed to treat children with non-headache chronic and recurrent pain including children with abdominal, musculoskeletal, and disease-related pain. Multidisciplinary pain treatment programmes for children have recently become a standard of care (McGrath 1999a), and now many specialised pain clinics are available for children with chronic or recurrent pain, which may involve outpatient care or intensive inpatient rehabilitation. Such programmes offer physical rehabilitation, psychological treatment, and medical strategies, and aim to restore function rather than provide pain relief. Case series and uncontrolled studies provide evidence for the effectiveness of multidisciplinary treatment with psychological therapy for paediatric chronic and recurrent pain (Eccleston 2003b).

### Why it is important to do this review

Several reviews have documented the effectiveness of psychological therapies for children with headache, abdominal, and disease-related pain (Holden 1999; Huertas-Ceballos 2008; Janicke 1999; Kibby 1998; Walco 1999; Weydert 2003). Four reviews have used data pooling techniques for studies of children with headache (Eccleston 2012; Fisher 2014; Hermann 1995; Trautmann 2006). In their review of paediatric migraine, Hermann 1995 found that biofeedback and muscle relaxation are more effective than placebo treatments and prophylactic drug treatments in controlling headache. In the previously published Cochrane review (Eccleston 2012), we found that psychological treatments were effective in reducing pain intensity in youths with headache and non-headache pain. Fisher 2014 reported similar findings for chil-

dren and adolescents with headache. Trautmann 2006 conducted a meta-analysis of psychological treatment for recurrent headache in children, finding small effect sizes across three headache variables: frequency, duration, and intensity, although reduction in pain intensity at post-treatment was a statistically significant effect. A large binomial effect size of 50% or greater reduction in headache symptoms was reported.

Developments in paediatric psychology have led to new populations of children being treated. The aim of this review is to update the published evidence on the efficacy of psychological treatments for chronic and recurrent pain in children and adolescents. In this review, we aim to focus specifically on therapy delivered in person (face-to-face) rather than remotely to the child in order to estimate treatment effects among studies using a relatively homogenous delivery method. A separate review for *The Cochrane Library* focused on remotely delivered treatments for youth with chronic pain is currently in progress ('Psychological therapies (remotely delivered) for the management of chronic and recurrent pain in children and adolescents' (protocol in press)). In this review, we also aim to examine the impact of psychological therapies on 'mood' in more detail than previous reviews by separating depression and anxiety into discrete outcome domains.

## OBJECTIVES

- The primary objective of this updated review was to determine the effectiveness on clinical outcomes of pain severity, disability, depression, and anxiety of psychological therapy delivered face-to-face for chronic and recurrent pain in children and adolescents compared with active treatment, waiting-list, or standard medical care.
- The secondary objective was to examine the impact of psychological therapies on children's mood symptoms with more specificity by evaluating depression and anxiety as discrete outcomes.
- The third objective was to describe the risk of bias of included studies and the quality of outcomes using the GRADE criteria.

## METHODS

### Criteria for considering studies for this review

#### Types of studies

Randomised controlled trials (RCTs) comparing a credible psychological treatment, or a compound treatment with credible primary psychological content, to an active treatment, treatment as usual, or waiting-list control. Content was judged credible if it was based on an extant psychological theory or framework. Studies were excluded if the pain was associated with cancer or other medical conditions (e.g. diabetes) or the therapy was delivered remotely using methods such as telephone or Internet.

Studies were included if they:

- were available as a full report of a RCT;
- had a design that placed a psychological treatment as an active treatment of primary interest;
- had a psychological treatment with definable psychotherapeutic content (although not necessarily delivered by someone with psychological qualifications);
  - were published (or electronically pre-published) in a peer-reviewed scientific journal;
  - participants reported chronic (i.e. at least three months duration) or recurrent (episodic) pain;
  - had 10 or more participants in each treatment arm at the end-of-treatment assessment; and
  - included a psychological intervention that was delivered in person (face-to-face treatment).

#### Types of participants

Children and adolescents (<18 years) reporting persistent, recurrent, or episodic pain in any body site, not associated with cancer or other medical conditions (e.g. diabetes).

#### Types of interventions

Studies were included if at least one trial arm consisted of a psychological intervention delivered face-to-face, and a comparator arm consisted of active treatment, treatment as usual, or waiting-list control. Primary interventions that were delivered remotely via other methods (e.g., Internet, telephone) were excluded.

#### Types of outcome measures

- Data were collected on descriptive characteristics of patients and characteristics of the treatments, including treatment setting and treatment dose (duration).
  - All measurement instruments reported in each study were assessed and recorded. The most appropriate measurement instruments for the four domains of pain, disability, depression, and anxiety were selected.
  - Any mention of adverse events was also recorded.

### Search methods for identification of studies

#### Electronic searches

RCTs of any psychological therapy for paediatric chronic or recurrent pain were identified by searching CENTRAL, MEDLINE, EMBASE, and PsycINFO from their inception to January 2014. Four separate searches have been undertaken. The first search was undertaken from inception of the abstracting services to the end of 1999 (Eccleston 2003a), the second searched databases from 1999 to 2008 (Eccleston 2009), the third searched databases from 2008 to March 2012 (Eccleston 2012), and the fourth from 2012 to 21st January 2014.

Further, trial registries were searched for possible ongoing or complete trials in this area. Reference lists of included studies and relevant systematic reviews were examined for other potential RCTs.

## Data collection and analysis

### Selection of studies

The selection of included studies was made using the following criteria; the study had to be RCT in design and published in a peer-reviewed journal, include children (<18 years of age) who have chronic pain (non-cancer pain), include a psychological intervention as an active treatment, and have  $\geq 10$  participants in each arm at each extraction time-point. Studies that have not been peer reviewed were excluded in order to keep the quality of included studies high. For this update, psychological therapies delivered remotely (e.g., Internet, telephone) were excluded. Psychological interventions were considered for inclusion if they had credible, recognisable psychological/psychotherapeutic content and were specifically designed to change the child's behaviour, cognition, and/or mood. The trials used in the previous systematic review and meta-analysis were considered automatically eligible for inclusion (Eccleston 2012).

### Data extraction and management

Data extracted included: details relating to the design of the study, the participants, primary diagnosis, method of treatment, adverse events, outcome measurement tools used, and outcome data for computation of effect sizes. When data were missing for primary outcomes of interest, we contacted trial authors via email to obtain data necessary for effect size calculations. Data suitable for pooling were entered into RevMan 5.2 (RevMan 2012).

### Assessment of risk of bias in included studies

The risk of bias was measured using the recommended Cochrane 'Risk of bias' tool (Higgins 2011). We assessed five categories from this tool; random sequence generation (selection bias), allocation concealment (selection bias), blinding of outcome assessment (detection bias), incomplete outcome data (attrition bias), and selective reporting (reporting bias). 'Blinding of participant or personnel' (performance bias) was excluded for the purposes of this re-

view as we deemed it redundant because of the nature of delivering or receiving a psychological intervention.

Judgements were made on the categories using the following rules. Random sequence generation judgements were based on whether authors gave a convincing method of randomisation. Allocation concealment bias judgements were based on whether there were convincing methods used for random allocation to take place. Participants being stratified by age or gender were not deemed as biased. Blinding of outcome assessment was judged on whether the measures were taken by a third party who was blind to the treatment condition. Incomplete outcome data bias judgements were based on whether attrition was fully reported. Authors had to report attrition at each measurement time point (post-treatment and follow-up), and state whether there were any significant differences between completers and non-completers. Finally, selective reporting bias was judged on whether data could be fully extracted for analyses in this review. If authors provided data when requested, we would have marked this category as 'unclear bias'. Summary of findings tables using the GRADE criteria are presented separately for outcomes for children with headache and non-headache pain conditions (Summary of findings for the main comparison, Summary of findings 2). The GRADE table presents 'probable outcomes' for the control and intervention group, rather than 'assumed risk' and 'corresponding risk' as presented in traditional GRADE tables. The probable outcome of events was calculated per 1000 for both the control group and those receiving psychological therapies, similar to other reviews including patients with pain conditions (e.g. Moore 2014). The studies included for each outcome were judged using five criteria: risk of bias, indirectness, inconsistency, imprecision, and publication bias. Limitations in the design and implementation were used to assess the overall risk of bias of included studies for each outcome. An outcome was downgraded if the majority of studies had unclear or high risk of bias. Indirectness was assessed if a population, intervention, or outcome was not of direct interest to the review (e.g. using mostly wait-list controls). Inconsistency was determined by the heterogeneity of results. If an outcome had a heterogeneity outcome of >45%, the outcome quality was downgraded. Imprecision was assessed by the number of participants included in an outcome and confidence intervals. Outcomes were downgraded when only a small number of participants could be included in the analysis, or the analysis had wide confidence intervals. Finally, publication bias was downgraded if studies failed to report outcomes in the published manuscript or if there was a suspicion that null findings had not been published or reported (Higgins 2011).

Each outcome was given a quality marking ranging from 'very low' to 'high'. High quality ratings are given when "further research is unlikely to change our estimate of effect". Moderate ratings are given when "further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate". Low quality is given when "further research is very likely to have an important impact on our confidence in the estimate

of effect and is likely to change the estimate". Finally, very low quality is given when "we are very uncertain about the estimate" (p. 404, [Balslem 2011](#)). The seven 'most important outcomes' were reported in each table ([Guyatt 2013](#)). Therefore, the seven outcomes that reported the largest amount of participants were included in each summary of findings table.

### Measures of treatment effect

All treatments labelled as psychological were combined in the following meta-analyses, and designated "Treatment". Similarly, all control conditions were combined and designated "Control". Where more than one intervention or control group was reported the intervention or control arms were combined to create a single pairwise comparison in accordance with the *Cochrane Handbook for Systematic Reviews of Interventions* ([Higgins 2011](#)). The studies were divided into two groups based on pain condition. The first group was labelled "headache" and the second group "non-headache". Two assessment points were also selected: post-treatment and follow-up. Post-treatment is the assessment point occurring soonest following treatment (often after a delay of several weeks to allow for recording of episodic pain), and follow-up is the assessment point at least three months after the post-treatment assessment point, but not more than 12 months, and the longer time point was selected if there were two follow-up assessments within this time frame. Therefore, four separate comparisons were designed comprising two forms of comparator (Treatment, Control) and two assessment time points (post-treatment and follow-up). They were labelled as follows.

1. Treatment versus control (headache) post-treatment.
2. Treatment versus control (headache) follow-up.
3. Treatment versus control (non-headache) post-treatment.
4. Treatment versus control (non-headache) follow-up.

Multiple measurement tools were typically used in each study. For each comparison, four outcomes were identified and labelled 'Pain', 'Disability', 'Depression', and 'Anxiety'. From each trial we selected the measure considered most appropriate for each outcome. To guide the choice of outcome measure, we applied two rules. First, if an outcome measure was established and occurred frequently among studies it was selected over more novel instruments. Second, given a choice between single item and multi-item self-report tools, multi-item tools were chosen on the basis of inferred increased reliability. Studies did not necessarily report data in all four outcomes. For headache treatments, the data for pain outcomes were dichotomous so relative ratios or risk ratios (RR) were used, and we calculated numbers needed to treat to benefit (NNTBs). For disability, depression, and anxiety outcomes, continuous data were used. Continuous data were used for pain, dis-

ability, depression, and anxiety for non-headache studies. Effect sizes can be interpreted as follows; small = 0.2, medium = 0.5, large = 0.8 ([Cohen 1992](#)).

### Data synthesis

For dichotomous outcomes, such as achieved (or failed to achieve) 50% reduction in pain, we calculated the RR using 95% confidence intervals (CI) and a random-effects model. For ease of interpretation, the risk ratio (RR) and NNTB are reported. For continuous outcomes (such as rating scales) we calculated the standardised mean differences using a 95% CI and a random-effects model. The heterogeneity of the findings are also reported.

## RESULTS

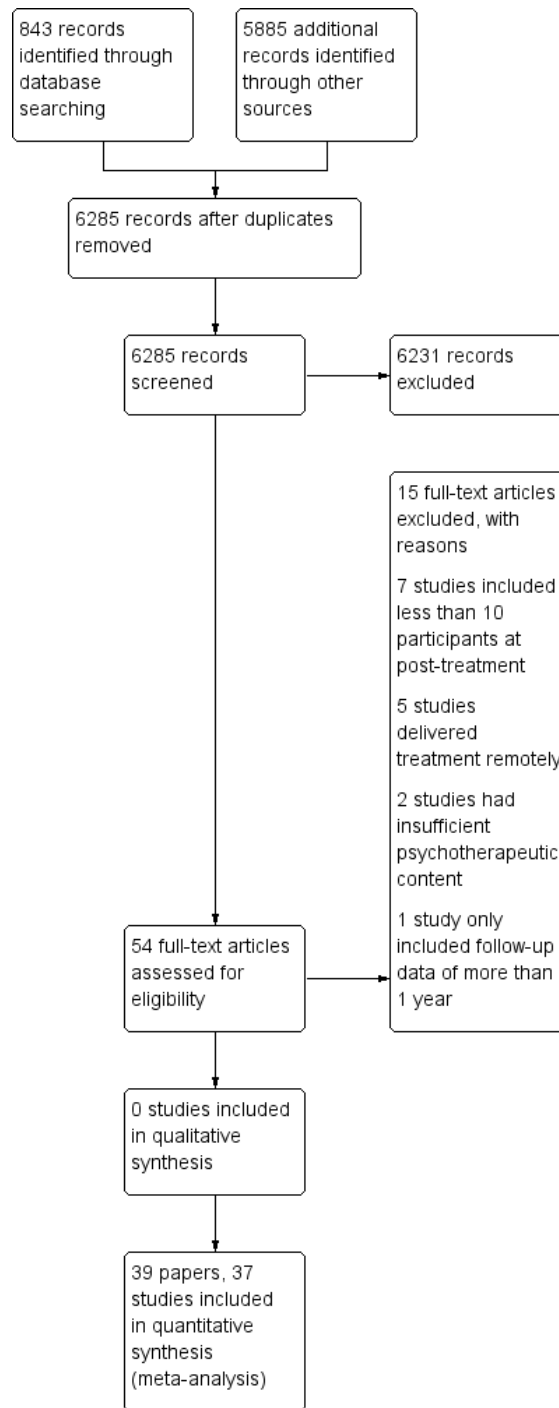
### Description of studies

See: [Characteristics of included studies](#); [Characteristics of excluded studies](#).

### Results of the search

Four separate searches have been undertaken using databases from inception to January 2014 (see [Figure 1](#)). Details of the previous three searches can be found in [Appendix 2](#). In the most recent search, databases were searched from March 2012 to January 2014. In total from the four searches, 6285 abstracts were screened. The current search yielded 443 abstracts and seven papers were included ([Grob 2013](#); [Gulewitsch 2013](#); [Hechler 2014](#); [Kashikar-Zuck 2012](#); [Levy 2010](#); [Powers 2013](#); [van der Veek 2013](#)). [Kashikar-Zuck 2012](#) and [Levy 2010](#) provided additional data for studies previously included in this review. Five studies that were previously included, were excluded from this review since treatment was delivered remotely ([Connelly 2006](#); [Hicks 2006](#); [Palermo 2009](#); [Stinson 2010](#); [Trautmann 2010](#)). Therefore, a total of 37 RCTs are included (39 papers) ([Abram 2007](#); [Alfven 2007](#); [Barakat 2010](#); [Barry 1997](#); [Bussone 1998](#); [Duarte 2006](#); [Fichtel 2001](#); [Gil 1997](#); [Griffiths 1996](#); [Grob 2013](#); [Gulewitsch 2013](#); [Hechler 2014](#); [Humphreys 2000](#); [Kashikar-Zuck 2005](#); [Kashikar-Zuck 2012](#); [Kroener-Herwig 2002](#); [Labbe 1984](#); [Labbe 1995](#); [Larsson 1987a](#); [Larsson 1987b](#); [Larsson 1990](#); [Larsson 1996](#); [Levy 2010](#); [McGrath 1988](#); [McGrath 1992](#); [Osterhaus 1997](#); [Passchier 1990](#); [Powers 2013](#); [Richter 1986](#); [Robins 2005](#); [Sanders 1994](#); [Sartory 1998](#); [Scharff 2002](#); [van der Veek 2013](#); [van Tilburg 2009](#); [Vlioger 2007](#); [Wicksell 2009](#)).

**Figure 1. Study flow diagram.**



## Included studies

The total number of participants completing treatments from the 37 studies was 2111. Of the 37 studies, one had four treatment arms, 10 had three arms, and 26 had two arms. The mean number of participants per study at the end of treatment was 57 (standard deviation (SD) 37). Girls outnumbered boys in 29 studies, and boys outnumbered girls in eight (Mean = 68% girls, range 22% to 100%). Child age was reported in 34 studies (Mean 12.45 years, SD 2.2 years). Only 16 studies reported the duration of pain, with a mean of 3.2 years.

Participants were recruited from a range of healthcare settings and other sources. Twenty-one studies recruited from hospital or clinic settings, four from schools, six recruited volunteers from school or hospital, referrals, or recruited through advertisements, one from the community, and five did not report the source. There were 20 studies of treatments for children with headache (including migraine). Of the remainder, nine were for abdominal pain (Alfven 2007; Duarte 2006; Grob 2013; Humphreys 2000; Levy 2010; Robins 2005; Sanders 1994; van der Veek 2013; van Tilburg 2009), and two studies treated participants with either a primary diagnosis of abdominal pain or a primary diagnosis of irritable bowel syndrome (Gulewitsch 2013; Vlieger 2007). Two studies treated children with fibromyalgia (Kashikar-Zuck 2005; Kashikar-Zuck 2012), two were for the treatment of pain associated with sickle cell disease (Barakat 2010; Gil 1997), and a further two studies included mixed pain conditions including headache and non-headache pain (Hechler 2014; Wicksell 2009), and so data were included in both analyses as appropriate.

Treatment arms were classified on the basis of their content and of the label given by the study authors. The interventions were categorised into three broad groups. The first is best described as behavioural, typically relaxation-based, with or without biofeedback, and including autogenic or hypnotherapeutic content (Bussone 1998; Fichtel 2001; Labbe 1984; Labbe 1995; Larsson 1987a; Larsson 1987b; Larsson 1990; Larsson 1996; McGrath 1988; McGrath 1992; Passchier 1990; Vlieger 2007). The second is best described as cognitive behavioural therapy, including cognitive coping, coping skills training, and parent behavioural strategies (Abram 2007; Alfven 2007; Barakat 2010; Barry 1997; Duarte 2006; Gil 1997; Griffiths 1996; Grob 2013; Gulewitsch 2013; Humphreys 2000; Kashikar-Zuck 2005; Kashikar-Zuck 2012; Kroener-Herwig 2002; Levy 2010; McGrath 1992; Osterhaus 1997; Powers 2013; Richter 1986; Robins 2005; Sanders 1994; Sartory 1998; Scharff 2002; van der Veek 2013; van Tilburg 2009; Wicksell 2009). The third, used a three week interdisciplinary pain programme consisting of paediatricians, psychologists, psy-

chiatrists, nurses, physiotherapists, occupational therapists, and social workers with treatment delivered in an inpatient setting. The number of psychological content hours within this programme was 24-31 hours (Hechler 2014). Psychological therapy delivered in this group was based on cognitive-behavioural principles.

Different control conditions were employed and were categorised into either active control (e.g. treatment as usual, education, n = 25) or wait-list (n = 12). Twenty-nine studies reported extractable post-treatment data, and 13 studies reported extractable follow-up data of between three months and a year. Thirty-three studies reported the treatment length; this was typically short (Mean = 6 hours 37 minutes for headache studies, Mean = 6 hours 41 minutes for non-headache studies, Table 1). Three studies did not report the duration of psychological treatment (Alfven 2007; Humphreys 2000; Sartory 1998).

The setting of treatment delivery varied between studies (Table 1). Twenty-three studies delivered treatment in a clinic, three studies delivered treatment at home (e.g. with a therapist, following a manual), and three were based either in a clinic or at home, so exposure to treatment was uncontrolled. A further three were based in schools and five were unknown. Home maintenance or practice of treatment was a common and important feature of many studies, but overall treatment exposure including home practice was not reported.

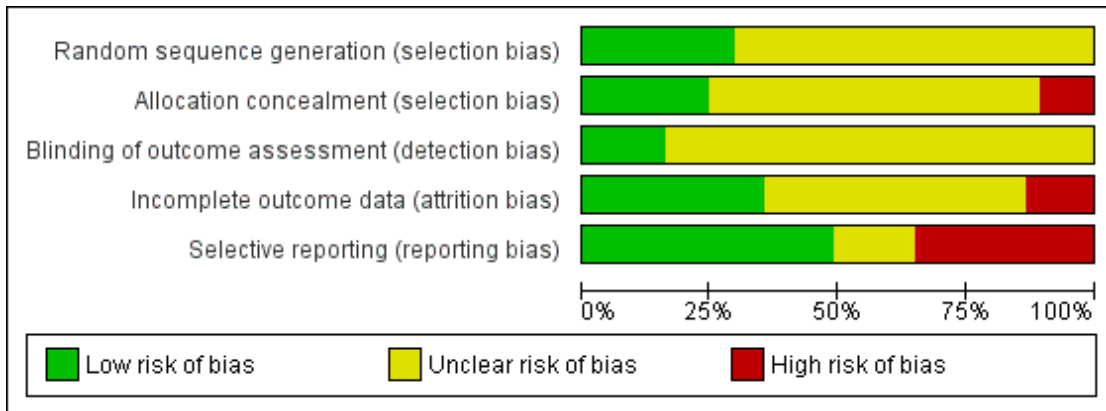
## Excluded studies

Fifteen studies were excluded, of which six are new to this update (Connelly 2006; Hicks 2006; Koenig 2013; Palermo 2009; Stinson 2010; Trautmann 2010). Connelly 2006, Hicks 2006, Palermo 2009, Stinson 2010, and Trautmann 2010 were excluded as they were delivered remotely, so did not meet the new inclusion criteria. Seven studies were excluded as they had fewer than 10 participants in a treatment arm at the end of treatment (Fentress 1986; Kroener-Herwig 1998; Larsson 1986; Sanders 1989; Trautmann 2008; Weydert 2006; Youssef 2009), two studies were judged to have insufficient psychological content in the treatment (Koenig 2013; Olness 1987), and one study reported only follow-up data of more than one year (Vlieger 2012).

## Risk of bias in included studies

All included studies were rated for risk of bias on five categories; random sequence generation (selection bias), allocation concealment (selection bias), blinding of outcome assessment (detection bias), incomplete outcome data (attrition bias), and selective reporting (reporting bias) (Figure 2; Figure 3).

**Figure 2. Methodological quality graph: review authors' judgements about each methodological quality item presented as percentages across all included studies.**





**Figure 3. Methodological quality summary: review authors' judgements about each methodological quality item for each included study.**

	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding of outcome assessment (detection bias)	Incomplete outcome data (attrition bias)	Selective reporting (reporting bias)
Abram 2007	+	?	?	?	+
Aifven 2007	?	?	?	+	+
Barakat 2010	?	?	?	?	+
Barny 1997	?	+	?	?	?
Bussone 1998	?	+	?	+	+
Duarte 2006	?	?	?	+	+
Fichtel 2001	?	?	?	+	+
Gil 1997	?	?	?	+	+
Griffiths 1996	?	?	?	+	+
Grob 2013	?	+	?	?	+
Gulewitsch 2013	+	?	?	?	+
Hechler 2014	+	+	?	?	?
Humphreys 2000	?	?	?	+	+
Kashikar-Zuck 2005	+	+	+	?	+
Kashikar-Zuck 2012	+	+	+	+	+
Kroener-Herwig 2002	?	?	?	?	+
Labbe 1984	?	?	?	+	+
Labbe 1995	?	?	?	+	+
Larsson 1987a	?	+	?	?	+
Larsson 1987b	?	+	?	+	+
Larsson 1990	?	?	?	?	+
Larsson 1996	?	?	?	+	+
Levy 2010	+	+	?	?	?
McGrath 1988	?	?	?	+	+
McGrath 1992	?	?	?	?	+
Osterhaus 1997	?	?	?	+	+
Passchier 1990	?	?	?	+	+
Powers 2013	+	+	?	?	?
Richter 1986	?	?	?	+	+
Robins 2005	+	?	?	?	+
Sanders 1994	?	?	?	+	+
Sartory 1998	?	?	?	?	+
Scharff 2002	+	?	?	+	+
van der Veek 2013	?	?	?	?	?
van Tilburg 2009	?	+	?	?	?
Vlieger 2007	+	+	?	?	+
Wicksell 2009	+	+	?	?	+



Eleven studies were scored as low risk of bias and gave a convincing method of randomisation, a further 26 studies were judged unclear on random sequence generation as they did not provide an adequate method of randomisation. None was scored as having high risk of bias. For allocation, nine studies were judged to have a low risk of bias and gave a convincing method, 24 studies were unclear and four studies had a high risk of bias. For outcome assessment, six studies used a third person blinded to the group allocation when taking measurements, 31 studies did not report this and so were unclear. Thirteen studies reported attrition fully, reporting that there was no significant difference between completers and non-completers. Nineteen studies only partially reported attrition and so we judged them to be unclear and five studies did not report attrition so we judged them to have a high risk of bias. Seventeen studies reported data fully, which could be extracted and used in analyses; six studies did not fully report data in the published trial, but provided data when contacted via email; 14 studies did not provide full extractable data and we judged them to have high risk of bias for selective reporting.

We attempted 16 analyses for this update (pain, disability, depression, and anxiety outcomes for headache and non-headache conditions post-treatment and at follow-up). One comparison had only one eligible study and so we did not perform analysis. Of the remaining 15 comparisons, four showed low heterogeneity ( $I^2$  value below 25%), four showed modest heterogeneity ( $I^2$  value over 25% to below 50%), and seven showed large heterogeneity ( $I^2$  value 50% or more).

The quality of evidence was assessed separately for headache and non-headache outcomes using the GRADE criteria. For headache conditions, two outcomes scored very low quality meaning we were very uncertain of the estimates of pain at follow-up, and anxiety at follow-up. Four outcomes (pain post-treatment, disability

post-treatment and at follow-up, and anxiety post-treatment) scored low quality meaning further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate. Depression post-treatment scored moderate quality, meaning further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate ([Summary of findings table 1](#)). For non-headache outcomes, the quality was higher. Two outcomes (pain and disability post-treatment) scored very low quality. Pain and disability at follow-up were deemed to be of low quality. All other outcomes scored moderate quality ([Summary of findings table 2](#)).

## Effects of interventions

See: [Summary of findings for the main comparison](#); [Summary of findings 2](#)

### Treatment versus control (headache) post-treatment

Fifteen studies with 714 participants into an analysis of the effects of treatment on pain post-treatment ([Barry 1997](#); [Fichtel 2001](#); [Griffiths 1996](#); [Kroener-Herwig 2002](#); [Labbe 1984](#); [Labbe 1995](#); [Larsson 1987a](#); [Larsson 1987b](#); [Larsson 1990](#); [Larsson 1996](#); [McGrath 1992](#); [Osterhaus 1997](#); [Powers 2013](#); [Sartory 1998](#); [Scharff 2002](#)). This analysis gave a risk ratio (RR) of 2.47 (95% confidence interval (CI) 1.97 to 3.09;  $z = 7.87$ ,  $p < 0.01$ ) for a beneficial reduction in headache pain (number needed to treat to benefit (NNTB) = 2.94) ([Analysis 1.1](#); [Figure 4](#); [Figure 5](#)). However, the GRADE quality rating for this outcome was low, meaning further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

**Figure 4. Forest plot of comparison: I Treatment versus control (headache) post-treatment, outcome: I.I Pain.**

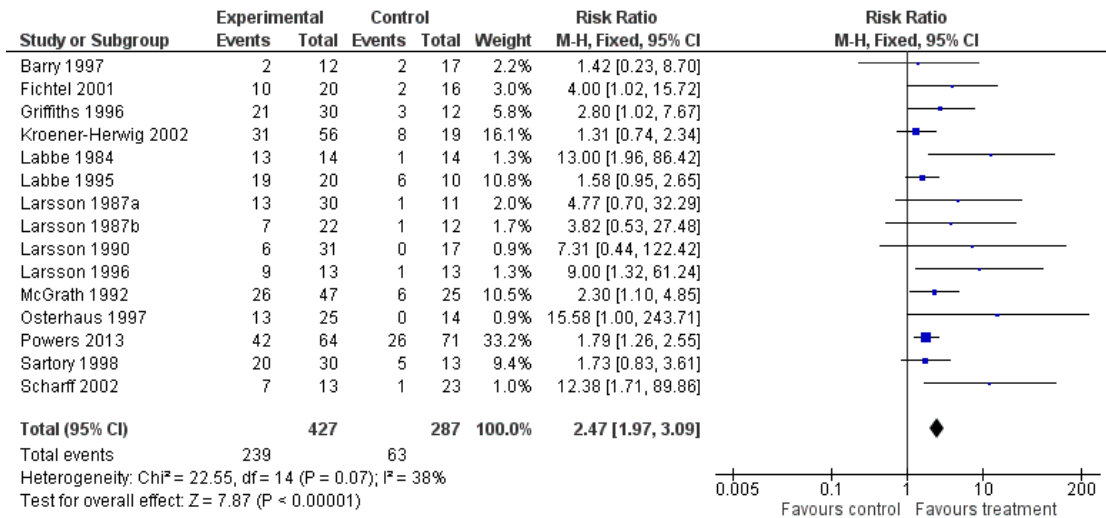
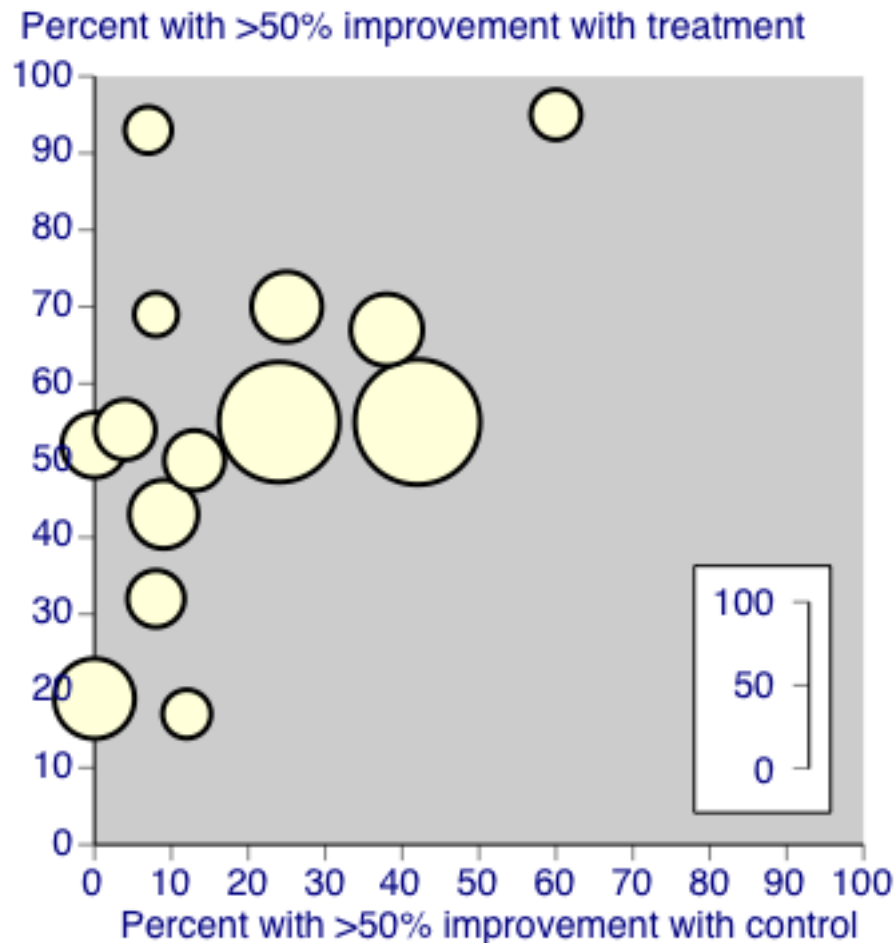


Figure 5.



Three studies with 263 participants were included in the analysis of the effects of treatment on disability (Hechler 2014; Powers 2013; Wicksell 2009). The analysis revealed that psychological therapies were beneficial at reducing disability in children with headache, with a small effect size (Standardised mean difference (SMD) -0.49, 95% CI -0.74 to -0.24,  $z = 3.90$ ,  $p < 0.01$ ; Analysis 1.2). The quality of this outcome was scored low, meaning further research is very likely to have an important impact on the effect. Three studies with 164 participants were entered into an analysis of the effects of treatment on depression (Griffiths 1996; Hechler 2014; Wicksell 2009). The analysis revealed that psychological therapies did not show a beneficial effect for reducing depression for children with headache (SMD -0.18, 95% CI -0.49 to 0.14,  $z = 1.11$ ,  $p > 0.05$ ; Analysis 1.3). A moderate quality rating was judged for this outcome, meaning further research is likely to have an important impact on our estimate of effect. Four studies with 203 participants were entered into an anal-

ysis of the effects of treatment on anxiety at post-treatment (Bussone 1998; Griffiths 1996; Hechler 2014; Wicksell 2009) which showed a small beneficial effect for psychological therapies (SMD -0.33, 95% CI -0.61 to -0.04,  $z = 2.25$ ,  $p < 0.05$ ; Analysis 1.4). We have low confidence in this estimate of effect. Out of the 20 headache studies, only Powers 2013 reported adverse events. The study authors categorised adverse events into different grades dependent on severity. There were 199 adverse events in total, although the authors do not state how many were due to the intervention. There was no difference in the severity of events between the CBT and headache education group.

#### Treatment versus control (headache) follow-up

Five studies of 251 participants were entered into analysis of the effects of treatment on pain at follow-up (Labbe 1984; Larsson

1987a; Larsson 1987b; Larsson 1996; Powers 2013). This analysis produced a RR of 2.89 (95% CI 1.03 to 8.07;  $z = 2.02$ ,  $p < 0.05$ ; Analysis 2.1), for a clinically beneficial change in pain (NNTB = 3.67). Using the GRADE criteria, pain at follow-up scored very low, meaning we were very uncertain of the estimate of effect.

Two studies with 148 participants were included in the analysis to determine the effects of treatment on disability at follow-up (Powers 2013; Wicksell 2009). Psychological therapies showed a small beneficial effect for reducing disability at follow-up (SMD -0.46, 95% CI -0.78 to -0.13,  $z = 2.72$ ,  $p < 0.01$ ; Analysis 2.2). Similar to disability post-treatment, we have low confidence in this estimate of effect.

Only one study could be included in the analysis on depression at follow-up Wicksell 2009, therefore no conclusion could be drawn. We were very uncertain of this estimate of effect.

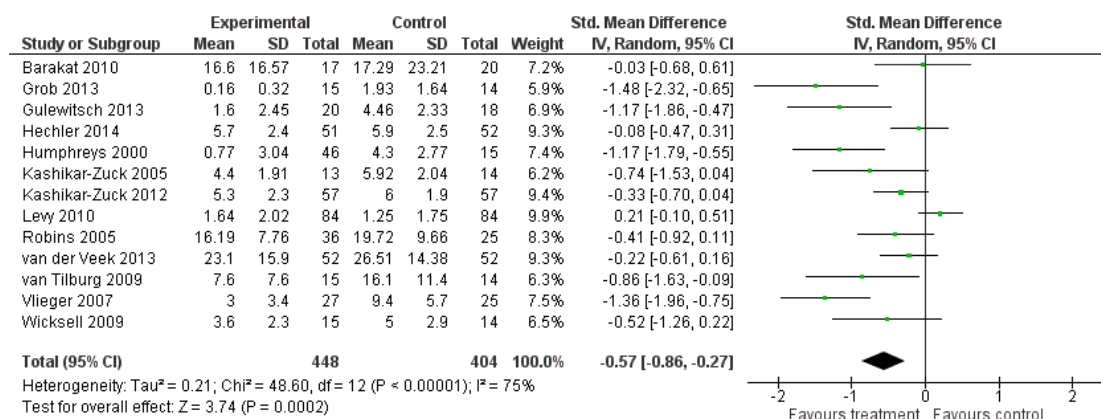
Two studies with 67 participants were entered into an analysis of the effects of treatment on anxiety at follow-up (Bussone 1998;

Wicksell 2009) finding no beneficial effect of psychological therapies (SMD -0.28, 95% CI -1.00 to 0.45;  $z = 0.75$ ,  $p > 0.05$ ; Analysis 2.4).

### Treatment versus control (non-headache) post-treatment

Thirteen studies of 852 participants were entered into an analysis of the effects of psychological treatment on continuous pain outcomes immediately post-treatment (Barakat 2010; Grob 2013; Gulewitsch 2013; Hechler 2014; Humphreys 2000; Kashikar-Zuck 2005; Kashikar-Zuck 2012; Levy 2010; Robins 2005; van der Veek 2013; van Tilburg 2009; Vlieger 2007; Wicksell 2009). Psychological therapies had a medium size beneficial effect on pain (SMD -0.57, 95% CI -0.86 to -0.27,  $z = 3.74$ ,  $p < 0.01$ ; Analysis 3.1; Figure 6). According to the GRADE criteria for assessing quality of outcomes, pain post-treatment scored very low quality, meaning we were very uncertain of the estimate of effect.

**Figure 6. Forest plot of comparison: 3 Treatment versus control (non-headache) post-treatment, outcome: 3.1 Pain.**



Eleven studies with 764 participants were entered into analysis of the effects of treatment on disability (Grob 2013; Gulewitsch 2013; Hechler 2014; Humphreys 2000; Kashikar-Zuck 2005; Kashikar-Zuck 2012; Levy 2010; Robins 2005; van der Veek 2013; van Tilburg 2009; Wicksell 2009). Psychological therapies had a small beneficial effect on reducing disability for children with chronic pain (SMD -0.45, 95% CI -0.71 to -0.19,  $z = 3.40$ ,  $p < 0.01$ ; Analysis 3.2). However, we were very uncertain of this estimate of effect.

Six studies with 538 participants were entered into analysis of the effects of treatment on depression (Hechler 2014; Kashikar-Zuck

2005; Kashikar-Zuck 2012; Levy 2010; van der Veek 2013; Wicksell 2009). The analysis revealed no beneficial effect of psychological therapies on depression (SMD -0.07, 95% CI -0.30 to 0.17,  $z = 0.54$ ,  $p > 0.05$ ; Analysis 3.3). We were moderately confident in the estimate of effect, meaning further research is likely to have an important impact on our confidence in the estimate of effect.

Five studies including 498 participants were entered into an analysis to determine the effects of treatment on anxiety immediately post-treatment (Hechler 2014; Kashikar-Zuck 2012; Levy 2010; van der Veek 2013; Wicksell 2009). The results revealed no bene-

ficial effect of psychological therapies on anxiety in children with chronic pain (SMD -0.15, 95% CI -0.36 to 0.07,  $z = 1.33$ ,  $p > 0.05$ ; [Analysis 3.4](#)). Similar to depression, we were moderately confident in the estimate of effect.

Of the 17 non-headache studies, four reported adverse events. [Gulewitsch 2013](#), [Kashikar-Zuck 2012](#), and [van der Veek 2013](#) reported no adverse events that were study-related. [Wicksell 2009](#) reported that two participants withdrew due to adverse effects of amitriptyline, which was part of the study condition.

#### **Treatment versus control (non-headache) follow-up**

Seven studies of 543 participants had data available for analysis of the effects of treatment on pain at follow-up ([Barakat 2010](#); [Grob 2013](#); [Hechler 2014](#); [Kashikar-Zuck 2012](#); [Levy 2010](#); [van der Veek 2013](#); [Wicksell 2009](#)). Analysis revealed no beneficial effect for psychological therapies on pain at follow-up (SMD -0.11, 95% CI -0.41 to 0.19,  $z = 0.73$ ,  $p > 0.05$ ; [Analysis 4.1](#)). The quality of outcome was low for this outcome, meaning further research is very likely to have an important impact on our confidence in the estimate of effect.

Six studies of 508 participants were entered into an analysis of the effects of treatment on disability ([Grob 2013](#); [Hechler 2014](#); [Kashikar-Zuck 2012](#); [Levy 2010](#); [van der Veek 2013](#); [Wicksell 2009](#)). No beneficial effect was found for psychological therapies on disability at follow-up (SMD -0.35, 95% CI -0.71 to 0.02,  $z = 1.87$ ,  $p > 0.05$ ; [Analysis 4.2](#)). We have low confidence in the estimate of effect.

Five studies with 473 participants were entered into an analysis of the effects of treatment on depression ([Hechler 2014](#); [Kashikar-Zuck 2012](#); [Levy 2010](#); [van der Veek 2013](#); [Wicksell 2009](#)). No beneficial effect was found for psychological therapies on depression at follow-up (SMD 0.06, 95% CI -0.16 to 0.28,  $z = 0.53$ ,  $p > 0.05$ ; [Analysis 4.3](#)). Similar to depression post-treatment, we were moderately confident in the effect.

Five studies with 452 participants were entered into an analysis of anxiety at follow-up ([Hechler 2014](#); [Kashikar-Zuck 2012](#); [Levy 2010](#); [van der Veek 2013](#); [Wicksell 2009](#)). Similar to depression, no beneficial effect was found for psychological therapies on anxiety at follow-up (SMD 0.05, 95% CI -0.24 to 0.33,  $z = 0.32$ ,  $p > 0.05$ ; [Analysis 4.4](#)).

## ADDITIONAL SUMMARY OF FINDINGS *[Explanation]*

Psychological therapies compared with any control for children with non-headache pain						
<b>Patient or population:</b> children and adolescents with non-headache pain <b>Settings:</b> Community <b>Intervention:</b> Psychological therapies <b>Comparison:</b> Any control						
Outcome	Probable outcome with control	Probable outcome with intervention	NNT and/or relative effect (95% CI)	No of participants	Quality of the evidence (GRADE)	Comments
<b>Pain</b> (low scores mean lower pain ratings)		The mean pain in the intervention groups was <b>0.57 standard deviations lower</b> (0.86 to 0.27 lower)		852 participants (13 studies)	⊕○○○ <b>very low</b>	Majority of studies had high risk of bias, heterogeneity >45%, some studies did not fully report outcomes in published paper SMD -0.57 (-0.86 to -0.27)
<b>Pain (at follow-up)</b> (low scores mean lower pain ratings)		The mean pain (at follow-up) in the intervention groups was <b>0.11 standard deviations lower</b> (0.41 lower to 0.19 higher)		543 participants (7 studies)	⊕⊕○○ <b>low</b>	Heterogeneity >45% and some studies did not fully report outcomes in published paper SMD -0.11 (-0.41 to 0.19)
<b>Disability</b> (low scores mean lower disability ratings)		The mean disability in the intervention groups was <b>0.45 standard deviations lower</b> (-0.71 to -0.19)		764 participants (11 studies)	⊕○○○ <b>very low</b>	Majority of studies had high risk of bias, heterogeneity >45%, some studies did not fully report outcomes in published paper SMD -0.45 (-0.71 to -0.19)

<b>Disability (at follow-up)</b> (low scores mean lower disability ratings)	The mean disability in the intervention groups was <b>0.35 standard deviations lower</b> (0.71 lower to 0.02 higher)	508 participants (6 studies)	⊕⊕○○ <b>low</b>	Heterogeneity >45% and some studies did not fully report outcomes in published paper SMD -0.17 (-0.71 to 0.02)
<b>Depression</b> (low scores mean lower depression ratings)	The mean depression in the intervention groups was <b>0.07 standard deviations lower</b> (0.3 lower to 0.17 higher)	538 participants (6 studies)	⊕⊕⊕○ <b>moderate</b>	Some studies did not fully report outcomes in published paper SMD -0.07 (-0.3 to 0.17)
<b>Depression (at follow-up)</b> (low scores mean lower depression ratings)	The mean anxiety in the intervention groups was <b>0.06 standard deviations higher</b> (0.16 lower to 0.28 higher)	473 participants (5 studies)	⊕⊕⊕○ <b>moderate</b>	Some studies did not fully report outcomes in published paper SMD 0.06 (-0.16 to 0.28)
<b>Anxiety</b> (low scores mean lower anxiety ratings)	The mean anxiety in the intervention groups was <b>0.15 standard deviations lower</b> (0.36 lower to 0.07 higher)	498 participants (5 studies)	⊕⊕⊕○ <b>moderate</b>	Some studies did not fully report outcomes in published paper SMD 0.15 (-0.36 to 0.07)

GRADE Working Group grades of evidence

**High quality:** Further research is very unlikely to change our confidence in the estimate of effect.

**Moderate quality:** Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.

**Low quality:** Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

**Very low quality:** We are very uncertain about the estimate.

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CI: Confidence interval; SMD: Standardised Mean Difference.



## DISCUSSION

### Evidence base

Thirty-seven studies (end of treatment N = 2111) were included in this updated review. In multi-arm trials involving more than one treatment or control group, we combined similar treatments or control groups for the purposes of the analyses. The majority of studies used one or two treatment conditions in comparison to a waiting-list or to a treatment as usual control group. As in the previous review, we categorised treatments as behavioural or cognitive behavioural, although these were combined for all analyses. The average length of treatment in studies of headache conditions and non-headache studies was very similar, between six and seven hours. Follow-up data are increasingly being reported in more recent studies and were included when relevant.

The inclusion of further studies has extended the evidence base. Of the 16 possible analyses, psychological therapies were beneficial for seven outcomes. Psychological therapies were beneficial at reducing pain intensity for headache and non-headache groups post-treatment, and for the headache group at follow-up. Fifty-six per cent of children with headaches reduced their pain scores post-treatment compared with only 22% in the control groups. Similar findings were demonstrated for disability, for which the findings on disability for the headache group are new to this update. Psychological therapies were beneficial at reducing disability in children with headache pain and non-headache pain post-treatment, and for headache groups at follow-up, although all effect sizes were small. Although we previously found a beneficial effect for treatment effect on mood findings at follow-up in the headache group (Eccleston 2012), several changes in our protocol have modified this effect. We have now separated mood into depression and anxiety, and have included only trials that delivered treatment face-to-face (rather than remotely). Psychological therapies were only found to have a small beneficial effect for anxiety post-treatment for the headache group. No other beneficial effects were found for depression and anxiety in children with chronic pain.

Pain intensity was the most common treatment outcome assessed, with 15 studies of children with headache and 13 studies of children with non-headache pain providing data. An NNTB of 2.94 for psychological therapies to produce more than 50% relief in pain in children with headaches was found. An NNTB of 3.67 was found for the smaller number of trials reporting on headache pain at follow-up. Medium effect sizes were also found for reduction in pain intensity in non-headache chronic and recurrent pain at post-treatment. However, the confidence intervals around the effects are large.

### Issues for consideration

More recent trials typically use cognitive behavioural therapy rather than behavioural therapy, likely reflecting changes in prac-

tice by psychologists entering the field of paediatric pain management.

In regard to pain condition, this review included 20 trials of children and adolescents with headache pain, nine abdominal pain studies, two abdominal pain and irritable bowel syndrome studies, two fibromyalgia studies, two sickle cell disease studies, and two mixed pain studies (including headache and non-headache pain conditions). There is limited evidence to draw conclusions about the effects of psychological treatment on disability in headache conditions. Although psychological therapies were shown to be beneficial, only three studies could be included in this analysis post-treatment, and two at follow-up. There is also limited evidence for treatment affecting depression and anxiety as outcomes. Previously, we reported that mood and disability outcomes in trials of children with chronic pain were an increasing focus for trials (McGrath 2008). This seems still to be the case, and with more studies reporting these outcomes it would be helpful for consensus on the best measurement instruments to be used consistently across the field of paediatric chronic pain, particularly treatment trials.

One limitation of this review is that we are unable to discuss fully the effectiveness of psychological interventions as they were compared with a control group that combined active (e.g. education) and waiting-list controls. Most studies used active controls, yet we did not feel that it was an appropriate sample to separate for analysis as has been done in a companion review of treatments for adults with pain (Williams 2012). This limitation may contribute to an overestimation of the treatment effects since it is not possible to separate differences specific to treatment versus active treatment or waiting-list control.

## AUTHORS' CONCLUSIONS

### Implications for practice

Psychological treatments, principally relaxation and cognitive behavioural therapies delivered face-to-face, are effective treatments producing change in pain, disability, and anxiety for children with headache conditions post-treatment. There is also evidence that the positive changes in pain and disability continue at follow-up. However, the overall quality of evidence for headache conditions was low/very low, meaning we are not confident in the estimate of effect. Further research is necessary to increase this confidence. Behavioural and cognitive behavioural treatments are also effective in reducing non-headache pain and disability post-treatment, but these beneficial effects were not maintained at follow-up. The quality of outcomes was higher for non-headache conditions, but further research is likely to have an important impact on our confidence of the estimate of effect. There is some evidence to support reductions in anxiety in response to behavioural pain treatment, particularly for children and adolescents with headache conditions

at post-treatment. There is insufficient evidence to comment on the effectiveness of psychological interventions for specific non-headache pain conditions due to the limited number of studies for each condition, although this has been attempted in a recent review by Fisher 2014.

Taken together, these findings suggest that behavioural treatment should be considered as part of standard care for all children and adolescents with chronic pain. Although there was a small effect for anxiety reduction in children and adolescents with headache conditions at post-treatment, this was not maintained at follow-up and there were no effects on depression at either time-point. This lack of effect may be due to the fact that anxiety and depression are typically not a specific intervention target of cognitive and behavioural pain management interventions.

### Implications for research

Since the original version of this review there has been an improvement in the evidence base by the addition of new studies, and the extension into non-headache pain conditions and treatments that rely on more complex methods. However, this structure limits our understanding of whether psychological therapies are unique in their improvement of symptoms in comparison to active or waiting-list control groups, yet we judged it important to present combined groups before introducing further analyses. The author team is considering the following changes for the next version of the review.

1. Increasing the current criterion from 10 to 20 participants in either arm at the point of analysis.
2. Splitting the title into two: one for headache only and one for non-headache (e.g. mixed pain conditions).
3. Exploring the possibility of subgroup analyses to try to identify variance attributable to non-specific factors which can

nevertheless affect treatment outcome, such as type of therapy, dose of therapy, setting of therapy, and therapeutic change agents (e.g. interventions delivered to parents).

Primary research is needed in the following areas.

1. To establish the efficacy of CBT in outcomes other than pain. In particular, it is important to establish whether CBT can improve mood outcomes and important functional outcomes (such as return to normal schooling), and can reduce the demand for healthcare resources. CBT often has a broad focus beyond pain. Additional pain and non-pain endpoints are desirable, in particular those relating to mood, disability, and social role functioning (see McGrath 2008).
2. To establish the efficacy of CBT in non-headache conditions, in particular idiopathic musculoskeletal pain such as fibromyalgia, and complex regional pain disorders. Randomised controlled trials are possible and desirable.
3. To establish the efficacy of CBT delivered to and/or via other significant therapeutic change agents such as parents, teachers, or peers. Randomised controlled trials are possible and desirable.

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\* Indicates the major publication for the study

## CHARACTERISTICS OF STUDIES

### Characteristics of included studies [ordered by study ID]

#### Abram 2007

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment (3-month follow-up), 6 months	
Participants	End of treatment n = 50 Start of treatment n = 81 Sex: 45 F, 36 M Mean age = 12.7 (range 10 to 18) Source = hospital and clinic Diagnosis = headache Mean years of pain = not given	
Interventions	“Headache Clinical Model: behavioural intervention” “Headache Traditional Model: consultation with neurologist”	
Outcomes	<b>Primary pain outcome: none</b> <b>Primary disability outcome: Ped-MIDAS</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Pediatric Migraine Disability Assessment (Ped-MIDAS) 2. FDI-C 3. Headache Knowledge test 4. Use of Healthcare measure	
Notes	Updated study 2009 Total quality = 22/35 Treatment quality = 7/9 Design quality = 15/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors’ judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Low risk	“If the family was interested in the study, they were randomised (using a random number table) to either a TCM appointment or a HCM appointment.” Comment: probably done
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done



**Abram 2007** (Continued)

Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however no significant differences between completers and non-completers were reported
Selective reporting (reporting bias)	High risk	Data were incompletely reported

**Alfven 2007**

Methods	RCT. 2 arms. Assessed at pre-treatment and 1-year follow-up	
Participants	End of treatment n = 48 Start of treatment n = 48 Sex: 61 F, 22 M (of entire sample in 3 treatment conditions, 1 post-randomisation) Mean age = 9.9 (range 6 to 18) Source = hospital Diagnosis = recurrent abdominal pain Mean years of pain = 2.5	
Interventions	“psychological treatment and physiotherapy” “Physiotherapy alone”	
Outcomes	<b>Primary pain outcome: pain score</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Pain intensity (VAS) 2. Pain score a) frequency b) intensity c) duration 3. Tender points (algometer)	
Notes	Updated study 2009 Total quality = 13/35 Treatment quality = 2/9 Design quality = 11/26	
<b><i>Risk of bias</i></b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“The children recruited during 1996-1999 were randomised” Comment: probably done, method not described

**Alfven 2007** (Continued)

Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts reported
Selective reporting (reporting bias)	High risk	Data were incompletely reported

**Barakat 2010**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment and 12 months	
Participants	End of treatment n = 42; follow-up 1 year n = 34 Start of treatment n = 42 Sex: 12 F, 15 M Mean age = 14.17 (1.75) Source = sickle cell centre Diagnosis = sickle cell disease Mean years of pain = lifetime	
Interventions	“Pain Management Intervention” “Disease Education Intervention”	
Outcomes	<b>Primary pain outcome: pain diary</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Pain diary 2. Health-related Hindrance Inventory 3. Child Health Questionnaire 4. Family Cohesion Scale 5. Disease Self-efficacy Scale 6. Coping Strategies Inventory 7. SCD Transition Knowledge Questionnaire 8. Medical chart review 9. School attendance	
Notes	Updated study 2012 Total quality = 27/35 Treatment quality = 9/9 Design quality = 18/26	

***Risk of bias***

Barakat 2010 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"A 2-group, randomised treatment design was used." Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described; no significant differences between completers and non-completers were reported
Selective reporting (reporting bias)	Low risk	Data were fully reported

Barry 1997

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 3 months
Participants	End of treatment n = 29 Start of treatment n = 36 Sex: 19 F, 10 M Mean age = 9.4 Source = volunteers via school and primary healthcare settings; referrals invited from primary and secondary care Diagnosis = headache Mean years of pain not given
Interventions	"Cognitive behaviour therapy" "waiting-list control"
Outcomes	<b>Primary pain outcome: headache intensity</b> <b>Primary disability outcome: school absence</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Headache intensity 2. Headache duration 3. Mood 4. School absence due to headache 5. Activities missed due to headache 6. Medication intake 7. Pain management strategies used

Notes	Original study Total quality = 14/35 Treatment quality = 3/9 Design quality = 11/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“Each parent-child pair was initially matched with another pair based on the child’s age, sex and headache pain as indicated by the parents’ ratings of average duration, frequency, and intensity of headaches. Subsequently, one of each of the matched parent-child pairs was randomly assigned to either the treatment condition or the waiting-list control condition.” Comment: probably done, method not described
Allocation concealment (selection bias)	High risk	“Each parent-child pair was initially matched with another pair based on the child’s age, sex and headache pain as indicated by the parents’ ratings of average duration, frequency, and intensity of headaches. Subsequently, one of each of the matched parent-child pairs was randomly assigned to either the treatment condition or the waiting-list control condition.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described; no significant differences between completers and non-completers were reported
Selective reporting (reporting bias)	Unclear risk	Data were completely reported on request

**Bussone 1998**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 6 months, 12 months
Participants	End of treatment n = 35 Start of treatment n = 35 Sex: 17 F, 18 M Mean age = 11.4 (range 11 to 15) Source = specialised headache clinic Diagnosis = headache Mean years of pain (mean) = 2.6
Interventions	“Biofeedback (assisted relaxation)” “Relaxation”
Outcomes	<b>Primary pain outcome: pain index</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: State Trait Anxiety Index</b> 1. Pain Total Index (headache diary) 2. State Trait Anxiety Index (STAI) 3. Analgesic use
Notes	Updated study 2009 Total quality = 18/35 Treatment quality = 5/9 Design quality = 13/26

***Risk of bias***

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	“Patients were randomly assigned to one of two experimental conditions” Comment: probably done, method not described
Allocation concealment (selection bias)	High risk	“... with the constraint that subjects be over-sampled in BFB-REL treatment (2:1 ratio) in order to make actual treatment available to as many children as possible.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts reported in study
Selective reporting (reporting bias)	High risk	Data incompletely reported

**Duarte 2006**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment
Participants	End of treatment n = 32 Start of treatment n = 32 Sex: 22 F, 10 M Mean age = 9.1 (SD 2.1) Source = paediatric gastroenterology service Diagnosis = recurrent abdominal pain Mean years of pain = 2.1
Interventions	“Cognitive behavioural family intervention” “Standard paediatric care, 4 sessions”
Outcomes	<b>Primary pain outcome: pain intensity VAS</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Pain VAS (reduced to 4 categories), completed daily 2. Parent estimate of frequency over last month 3. Pressure point threshold using algometer
Notes	Updated study 2009 Total quality = 15/35 Treatment quality = 5/9 Design quality = 10/26

***Risk of bias***

<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“Randomly allocated to 2 groups.” Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts were reported in the study
Selective reporting (reporting bias)	High risk	Data were incompletely reported

**Fichtel 2001**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 8 to 12 months
Participants	End of treatment n = 36 Start of treatment n = 36 Sex: 25 F, 11 M Mean age = 15.4 (range 13 to 18) Source = school Diagnosis = headache Mean years of pain = not given
Interventions	“Relaxation” “waiting-list control”
Outcomes	<b>Primary pain outcome: total headache score</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Total headache score (headache diary) 2. Medication consumption
Notes	Updated study 2009 Total quality = 15/35 Treatment quality = 4/9 Design quality = 11/26

***Risk of bias***

<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“The subjects were randomly assigned to the relaxation treatment or waiting-list groups” Comment: probably done, no method is described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts were reported in the study
Selective reporting (reporting bias)	Low risk	Data were fully reported

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment	
Participants	<p>End of treatment n = 49          Start of treatment n = 49          Sex: 23 F, 26 M          Mean age = 11.9          Source = university medical centre, sickle cell centre          Diagnosis = sickle cell anaemia (SS), sickle cell disease (SC), sickle beta thalassaemia          Mean years of pain = not given</p>	
Interventions	<p>“Cognitive coping skills”          “Standard care control”</p>	
Outcomes	<p><b>Primary pain outcome: none</b>  <b>Primary disability outcome: none</b>  <b>Primary depression outcome: none</b>  <b>Primary anxiety outcome: none</b>          1. Pain sensitivity (pressure stimulator)          2. Coping strategy questionnaire          3. Disease severity: acute and chronic complications in past 12 months</p>	
Notes	<p>Original study          Total quality = 16/35          Treatment quality = 8/9          Design quality = 8/26</p>	
<b><i>Risk of bias</i></b>		
<b>Bias</b>	<b>Authors’ judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“Participants were then randomly assigned to one of two conditions.” Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts reported in study
Selective reporting (reporting bias)	High risk	Data not fully reported



**Griffiths 1996**

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment and 9 weeks post-treatment
Participants	End of treatment n = 42; follow-up n = 42 Start of treatment n = 51 Sex: 21 F, 21 M Mean age = 11.3 Source = not known Diagnosis = migraine Mean years of pain = not given: minimum 6 months
Interventions	“Cognitive behavioural therapy (clinic based)” “Cognitive behavioural therapy (home based)” “Self monitoring”
Outcomes	<b>Primary pain outcome: headache index</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: Child Depression Scale</b> <b>Primary anxiety outcome: Child Manifest Anxiety Scale (CMAS)</b> 1. Headache index (averaged intensity) 2. Medication used 3. Child Manifest Anxiety Scale (CMAS) 4. Children’s Depression Scale (CDS) 5. Self efficacy 6. Coping responses from Children’s Headache Assessment Scale (CHAS)
Notes	Original study Total quality = 18/35 Treatment quality = 5/9 Design quality = 13/26

***Risk of bias***

Bias	Authors’ judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	“It was decided to assign children to groups by true randomisation rather than on the basis of headache diagnosis” Comment: probably done, no method is described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	High risk	Attrition was not described

Griffiths 1996 (Continued)

Selective reporting (reporting bias)	Low risk	Data were fully reported
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Grob 2013

Methods	RCT. 2 arms. Assessed pre-treatment, post-treatment and at 3 months
Participants	End of treatment n = 28; follow-up n = 28 Start of treatment n = 29 Sex: 25 F, 4 M Mean age = 9.6 (SD = 1.47) Source = schools Diagnosis = chronic abdominal pain Mean years of pain = 2.8 years (SD = 1.71)
Interventions	“Stop the pain with Happy Pingu” CBT “Wait-list control”
Outcomes	<b>Primary pain outcome: pain intensity</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Pain diary (intensity, frequency, duration) 2. KINDL-R disease-specific module 3. PedsQL 4. Self administered questionnaire based on Itch-questionnaire for pain-related cognitions
Notes	-

*Risk of bias*

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	“Computer-aided randomization was performed by a person who was not involved in the study” Comment: probably done, no method is described
Allocation concealment (selection bias)	Low risk	“Computer-aided randomization was performed by a person who was not involved in the study” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done

Grob 2013 (Continued)

Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition was described; differences between completers and non-completers were not described
Selective reporting (reporting bias)	Low risk	Data were fully reported

Gulewitsch 2013

Methods	RCT. 2 arms. Assessed pre-treatment, post-treatment (3 months)
Participants	End of treatment n = 37 Start of treatment n = 38 Sex: 24 F, 14 M Mean age = 9.4 (SD = 1.72) Source = adverts in local newspapers and paediatricians' offices Diagnosis = functional abdominal pain or irritable bowel syndrome Mean years of pain = 34.84 months (SD = 40.7)
Interventions	"Hypnotherapeutic therapy" (hypnotherapeutic and behavioural methods) "Wait-list control group"
Outcomes	<b>Primary pain outcome: mean pain intensity</b> <b>Primary disability outcome: Paediatric Pain Disability Index</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Mean pain intensity 2. Number of days with AP 3. Mean duration of pain episodes 4. School absence 5. Paediatric Pain Disability Index 6. Parent report of Abdominal Pain Index 7. Parent report of Paediatric Pain Disability Index 8. KINDL child report (health-related quality of life) 9. KINDL parent report (health-related quality of life)
Notes	-

*Risk of bias*

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Families were randomly assigned following simple randomization procedures (computerized random number generator)" Comment: probably done

**Gulewitsch 2013** (Continued)

Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition was described; differences between completers and non-completers were not described
Selective reporting (reporting bias)	Low risk	Data fully reported

**Hechler 2014**

Methods	RCT. 2 arms. Assessed pre-treatment, post-treatment, 6 months, and 12 months
Participants	End of treatment n = 108 Start of treatment n = 120 Sex: 87 F, 27 M Mean age = 14 (SD 2.85) Source = clinic Diagnosis = chronic pain (mixed conditions) Mean years of pain = median of 18 months (intervention group) and 13.5 months (control group)
Interventions	“Intensive interdisciplinary pain treatment” “Wait-list control”
Outcomes	<b>Primary pain outcome: mean pain intensity</b> <b>Primary disability outcome: Paediatric Pain Disability Index</b> <b>Primary depression outcome: Depression Inventory for Children and Adolescents (DIKJ)</b> <b>Primary anxiety outcome: Pain-Related Cognitions Questionnaire for Children (catastrophising sub-scale)</b> 1. Mean pain intensity 2. Paediatric Pain Disability Index 3. School absence 4. Anxiety Questionnaire for Pupils 5. Pain-Related Cognitions Questionnaire for Children (Catastrophising sub-scale) 6. Depression Inventory for Children and Adolescents (DIKJ) 7. Questionnaire to assess the economic effects of chronic pain 8. Utilisation of healthcare services 9. Parental work absenteeism 10. Work days lost 11. Subjective financial burden
Notes	-

**Hechler 2014** (Continued)

<i>Risk of bias</i>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Low risk	“Randomisation was conducted with a 1:1 approach and in blocks of 4 and blocks of 6 for both groups and was stratified for gender” Comment: probably done
Allocation concealment (selection bias)	Low risk	“The individual who carried out the randomization procedure was blinded to the treatment condition” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition was described; differences between completers and non-completers were not described
Selective reporting (reporting bias)	Unclear risk	Data fully reported on request

**Humphreys 2000**

Methods	RCT. 4 arms. Assessed at pre-treatment, post-treatment
Participants	End of treatment n = 61 Start of treatment n = 64 Sex: 38 F, 26 M Mean age = 9.8 (SD 2.5) Source = advertisement and physician referral Diagnosis = recurrent abdominal pain Mean years of pain = none given
Interventions	“CBT + biofeedback + parental support + fibre” “CBT + biofeedback + fibre” “Biofeedback + fibre” “fibre”
Outcomes	<b>Primary pain outcome: pain diary</b> <b>Primary disability outcome: school attendance</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Child pain diary 2. Parental observation record

**Humphreys 2000** (Continued)

	3. Health care utilisation record 4. Medical record 5. School attendance	
Notes	Updated study 2009 Total quality = 14/35 Treatment quality = 5/9 Design quality = 9/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"Patients were randomly assigned to one of the four groups" Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	High risk	Attrition not described; significant differences between completers and non-completers not reported
Selective reporting (reporting bias)	Low risk	Data fully reported

**Kashikar-Zuck 2005**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment (week 8), 6 weeks
Participants	End of treatment n = 27 Start of treatment n = 30 Sex: 30 F, 0 M Median age = 15.8 (SD 1.3) Source = paediatric rheumatology clinic of a children's hospital Diagnosis = juvenile primary fibromyalgia (JPFM criteria; Yunus) Mean years of pain = 19 for > 2 years, 11 for 6 months to 2 years
Interventions	"Coping skills training" "Self-monitoring"

Outcomes	<b>Primary pain outcome: average pain VAS</b> <b>Primary disability outcome: Functional Disability Inventory</b> <b>Primary depression outcome: Children's Depression Inventory</b> <b>Primary anxiety outcome: none</b> 1. Average pain VAS 0 to 100 2. Highest pain VAS 0 to 100 3. Functional Disability Inventory (FDI) 4. Children's Depression Inventory (CDI) 5. Pain Coping Questionnaire (PCQ) 6. Pain Coping Efficacy (items from PCQ) 7. Tender points	
Notes	Updated study 2009 Total quality = 25/35 Treatment quality = 7/9 Design quality = 18/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Low risk	“A computer generated pseudo-random number list was used. A simple randomisation technique was used with a 1:1 allocation ratio for 30 subjects as a single block.” Comment: probably done
Allocation concealment (selection bias)	Low risk	“A computer generated pseudo-random number list was used. A simple randomisation technique was used with a 1:1 allocation ratio for 30 subjects as a single block.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Low risk	“A research assistant who was blind to the study objectives and to the subjects' treatment assignment administered the self-report measures. The rheumatologist or occupational therapist who conducted the tender point assessments was blind to the subjects' treatment assignment.” Comment: probably done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not reported

**Kashikar-Zuck 2005** (Continued)

Selective reporting (reporting bias)	Low risk	Data were fully reported on request for additional data
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**Kashikar-Zuck 2012**

Methods	RCT. 2 arms. Assessed pre-treatment, post-treatment, 6-month follow-up
Participants	End of treatment n = 106; follow-up 6 months n = 100 Start of treatment n = 114 Sex: 105 F, 9 M Mean age = 15.0 (1.8) Source = paediatric rheumatology centres in Midwestern USA Diagnosis = fibromyalgia syndrome Mean years of pain = 2 years, 10 months (2 years, 6 months)
Interventions	“Cognitive behavioural therapy” “Fibromyalgia education”
Outcomes	<b>Primary pain outcome: pain severity VAS (averaged over 7 days)</b> <b>Primary disability outcome: Functional Disability Scale</b> <b>Primary depression outcome: Children’s Depression Inventory</b> <b>Primary anxiety outcome: Pain Coping Questionnaire</b> 1. Pain severity VAS (averaged over 7 days) 2. Functional Disability Scale 3. Children’s Depression Inventory 4. Tender point sensitivity 5. Pediatric Quality of Life Inventory 6. Sleep quality VAS (averaged over 7 days) 7. Physician’s global assessment VAS
Notes	Updated study 2012 Total quality = 32/35 Treatment quality = 9/9 Design quality = 23/26

***Risk of bias***

Bias	Authors’ judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	“Eligible patients were randomly assigned to 1 of the 2 treatment arms based upon a computer-generated randomisation list. Randomisation was stratified by site.” Comment: probably done
Allocation concealment (selection bias)	Low risk	“When a patient was enrolled, the study therapist contacted the biostatistician to obtain the subject identification number



**Kashikar-Zuck 2012** (Continued)

		and treatment allocation.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Low risk	“The principal investigator, study physicians, study coordinator, and assessment staff were all blinded to the patients’ treatment condition throughout the trial. Patients were asked not to divulge what treatment they were receiving to the study physician.” Comment: probably done
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition is described; no significant differences between completers and non-completers were reported
Selective reporting (reporting bias)	Low risk	Data were fully reported

**Kroener-Herwig 2002**

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment, 6 months
Participants	End of treatment n = 75 Start of treatment n = 78 Sex: 35 F, 40 M Mean age = 12.1 (SD 1.3) Source = newspaper advertisement - 2 or more headaches per month reported by parents Diagnosis = paediatric headache: migraine (30%), tension-type (40%), combined (30%) Mean years of pain = 4.0 (SD 2.6)
Interventions	“Cognitive behavioural training group” “Self-help” “waiting-list control”
Outcomes	<b>Primary pain outcome: pain intensity</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Headache frequency (mean no. per day) 2. Pain intensity (mean daily) 3. Headache duration (mean no hours per day)
Notes	Updated study 2012 Total quality = 19/35 Treatment quality = 7/9 Design quality = 12/26
<b>Risk of bias</b>	

**Kroener-Herwig 2002** (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Assignment to the treatment groups was random." Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not reported
Selective reporting (reporting bias)	Low risk	Data were fully reported

**Labbe 1984**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment (1 month after end of treatment), 6 months
Participants	End of treatment n = 28 Start of treatment n = 28 Sex: 14 F, 14 M Mean age = 10.8 Source = community paediatrician referral, newspaper advertisement Diagnosis = migraine headache Mean years of pain = 4.3
Interventions	"Autogenic feedback training" "waiting-list control"
Outcomes	<b>Primary pain outcome: headache diary</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Headache index 2. Headache frequency 3. Headache duration 4. Headache peak intensity 5. Medication use

Labbe 1984 (Continued)

Notes	Original study Total quality = 16/35 Treatment quality = 4/9 Design quality = 12/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"The children who attended the first session were matched on age, sex, and baseline headache index and then randomly assigned to either a treatment group or waiting-list control group." Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts reported in study
Selective reporting (reporting bias)	Low risk	Data were reported fully

Labbe 1995

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment, 6 months
Participants	End of treatment n = 30 Start of treatment n = 46 Sex: 17 F, 13 M Mean age = 12.0 Source = not given Diagnosis = vascular or migraine headache Mean years of pain = not given
Interventions	"Skin temperature biofeedback and autogenic relaxation" "Autogenic relaxation" "waiting-list control"
Outcomes	<b>Primary pain outcome: headache diary</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: Childhood Depression Inventory</b>

Labbe 1995 (Continued)

	<b>Primary anxiety outcome: How-I-Feel questionnaire</b>	
	1. Headache index 2. Headache frequency 3. Headache duration 4. Child aggression parent-rated (Myth Type A) 5. Childhood Depression Inventory 6. How-I-Feel questionnaire: anxiety	
Notes	Original study Total quality = 11/35 Treatment quality = 2/9 Design quality = 9/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"Children were matched by age, sex, and baseline headache activity and then randomly assigned to one of three groups." Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	"Data on the dropouts were compared to those children participating in the treatment sessions. No differences were found in sex, age or headache history." Comment: probably done
Selective reporting (reporting bias)	Low risk	Data were fully reported

Larsson 1987a

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment, 5 months
Participants	End of treatment n = 46 Start of treatment n = 46 Sex: 40 F, 6 M Mean age = not given: range 16 to 18 Source = not given Diagnosis = headache (migraine, tension, or both)

	Mean years of pain = most 1 to 5 years
Interventions	“Therapist assisted relaxation” “Self-help relaxation” “Self monitoring group”
Outcomes	<b>Primary pain outcome: headache sum</b> <b>Primary disability outcome: school absence</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Headache sum 2. Headache frequency 3. Headache-free days 4. Headache duration 5. Peak headache intensity 6. Medication 7. School absence 8. Significant other rating of headache improvement 9. Cost-effectiveness
Notes	Original study Total quality = 21/35 Treatment quality = 6/9 Design quality = 15/26

**Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	“In the randomisation procedure” Comment: probably done, no method described
Allocation concealment (selection bias)	High risk	“In the randomisation procedure the following restrictions were applied: (a) class mates were assigned to the same treatment group in order to lessen the risk of treatment contamination, (b) subjects were evenly distributed across groups within separate schools.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not reported

Larsson 1987a (Continued)

Selective reporting (reporting bias)	High risk	Data were not fully reported
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Larsson 1987b

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment, 5 months
Participants	End of treatment n = 36; follow-up n = 34 Start of treatment n = 36 Sex: 32 F, 2 M Mean age = 17 Source = not given Diagnosis = headache Mean years of pain = most 1 to 5 years
Interventions	“Self-help relaxation” “Problem discussion group” “Self monitoring (control)”
Outcomes	<b>Primary pain outcome: headache sum</b> <b>Primary disability outcome: school absence</b> <b>Primary depression outcome: Depression Scale for Female Adolescents</b> <b>Primary anxiety outcome: Swedish translation of Children’s Manifest Anxiety Scale</b> 1. Headache sum 2. Headache frequency 3. Headache-free days 4. Headache duration 5. Peak headache intensity 6. Medicine consumption 7. School absence 8. Headache annoyance 9. Depression/anxiety 10. Social relationship-competence questionnaire 11. Significant other rating of headache improvement
Notes	Original study Total quality = 16/35 Treatment quality = 5/9 Design quality = 11/26

*Risk of bias*

Bias	Authors’ judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	“Finally, 36 students were randomly assigned to the three experimental conditions.” Comment: probably done, no method de-

Larsson 1987b (Continued)

		scribed
Allocation concealment (selection bias)	High risk	“The allocation of subjects was conducted with two restrictions on the procedure: (a) Classmates were assigned to the same treatment condition (to lessen the risk of treatment contamination), and (b) students with a high frequency of headaches were identified and evenly distributed across groups.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	High risk	Attrition is not described
Selective reporting (reporting bias)	High risk	Data were not fully reported

Larsson 1990

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment
Participants	End of treatment n = 43 Start of treatment n = 49 Sex: 44 F, 5 M Mean age = 17 Source = school Diagnosis = headache Mean years of pain = median 2 to 5 years
Interventions	“Self help relaxation” “waiting-list control”
Outcomes	<b>Primary pain outcome: headache activity</b> <b>Primary disability outcome: none given</b> <b>Primary depression outcome: Beck Depression Inventory</b> <b>Primary anxiety outcome: Modified Child Manifest Anxiety Scale</b> 1. Headache index 2. Medication use 3. Headache annoyance 4. Modified Child Manifest Anxiety Scale (CMAS) 5. Depression - Beck Depression Inventory 6. Somatic complaints (composite of multiple complaints) 7. Stress (4-point scale)

Larsson 1990 (Continued)

Notes	Original study Total quality = 12/35 Treatment quality = 4/9 Design quality = 8/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"...the outlines of the study including the use of randomisation and a placebo treatment period." Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"A graduate student in psychology administered the assessment instruments and the treatment material used in the study." Comment: unsure
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not reported
Selective reporting (reporting bias)	Low risk	Data were fully reported

Larsson 1996

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 6 months
Participants	End of treatment n = 26 Start of treatment n = 26 Sex: 25 F, 1 M Mean age = not given: range 10 to 15 years Source = school Diagnosis = headache Mean years of pain = 2.1
Interventions	"Relaxation treatment" "No treatment"
Outcomes	<b>Primary pain outcome: headache intensity</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b>



Larsson 1996 (Continued)

	1. Headache intensity ('sum')	
	2. Headache-free days	
	3. Headache frequency	
Notes	Original study Total quality = 20/35 Treatment quality = 6/9 Design quality = 14/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"Thus, 26 pupils were randomly allocated into a relaxation training group or to a no-treatment control group". Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	There were no dropouts reported in the study
Selective reporting (reporting bias)	Low risk	Data were fully reported

Levy 2010

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 3-month follow-up, 6-month follow-up
Participants	End of treatment n = 168; follow-up 3 months n = 143; follow-up 6 months = 154 Start of treatment n = 200 Sex: 145 F, 55 M Mean age = 11.21 (2.55) Source = paediatric gastroenterology clinics at Seattle Children's Hospital and the Atlantic Health System in Morristown, New Jersey. Seattle participants were also recruited through local area clinics and community-posted flyers Diagnosis = functional abdominal pain Mean years of pain = 3+ episodes of abdominal pain during a 3-month period
Interventions	"Cognitive-behavioural treatment" "Educational intervention"

Outcomes	<p><b>Primary pain outcome: Faces Pain Scale-Revised</b>  <b>Primary disability outcome: Functional Disability Inventory</b>  <b>Primary depression outcome: Children's Depression Inventory</b>  <b>Primary anxiety outcome: Multidimensional Anxiety Scale for Children</b></p> <ol style="list-style-type: none"> <li>1. Faces Pain Scale-Revised</li> <li>2. Functional Disability Inventory</li> <li>3. Children's Depression Inventory</li> <li>4. Children's Somatization Inventory</li> <li>5. Multidimensional Anxiety Scale for Children</li> </ol>	
Notes	<p>Updated study 2012  Total quality = 27/35  Treatment quality = 7/9  Design quality = 20/26</p>	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Low risk	<p>"Randomisation was then performed by a different researcher using a computerised random-number generator, stratifying by age."</p> <p>Comment: probably done</p>
Allocation concealment (selection bias)	Low risk	<p>"Randomisation was then performed by a different researcher using a computerised random-number generator, stratifying by age."</p> <p>Comment: probably done</p>
Blinding of outcome assessment (detection bias) All outcomes	Low risk	<p>"Nurse assessors were blind to the treatment assignment of the children."</p> <p>Comment: probably done</p>
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described; significant differences between completers and non-completers are not reported
Selective reporting (reporting bias)	Unclear risk	Data were fully reported when requested

**McGrath 1988**

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment, 3 months, 12 months
Participants	End of treatment n = 99 Start of treatment n = 136 Sex: 69 F, 30 M Mean age = 13.1 (range 11 to 18) Source = hospital Diagnosis = headache Mean years of pain = not given: minimum 3 months
Interventions	“Relaxation training” “Attention control” “Own best efforts”
Outcomes	<b>Primary pain outcome: headache index</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Headache index 2. Headache-free days 3. Highest pain intensity
Notes	Original study Total quality = 23/35 Treatment quality = 7/9 Design quality = 16/26

***Risk of bias***

<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“Randomly assigned to one of three groups” Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition is described, however significant differences between completers and non-completers are not reported
Selective reporting (reporting bias)	Low risk	Data were completely reported

**McGrath 1992**

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment, 3 months and 1-year follow-up
Participants	End of treatment n = 74 Start of treatment n = 87 Sex: 63 F, 24 M Mean age = not given: range 11 to 18 years Source = paediatricians and family physicians Diagnosis = migraine Mean years of pain not given: minimum 3 months
Interventions	“Therapist administered cognitive behavioural/stress coping/relaxation training” “Self-administered cognitive behavioural/ stress coping/relaxation training” “Information and support”
Outcomes	<b>Primary pain outcome: headache diary</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: Poznanski Depression Scale</b> <b>Primary anxiety outcome: none</b> 1. Headache index 2. Efficiency of treatment 3. Poznanski Depression Scale
Notes	Original study Total quality = 15/35 Treatment quality = 2/9 Design quality = 13/26

***Risk of bias***

<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“Randomised to 1 of the 8-week treatments” Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers are not reported
Selective reporting (reporting bias)	High risk	Data were incompletely reported

**Osterhaus 1997**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment and 1-year follow-up
Participants	End of treatment n = 39, 1-year follow-up n = 21 Start of treatment n = 39 Sex: 29 F, 10 M Mean age = 15.2 (SD 3.3) Source = newspaper article Diagnosis = headache (migraine, tension-type, mixed) Mean years of pain = 5.6
Interventions	“Behavioural treatment package” “waiting-list control”
Outcomes	<b>Primary pain outcome: headache index</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Headache index 2. Headache frequency 3. Headache duration 4. Headache intensity
Notes	Original study Total quality = 18/35 Treatment quality = 6/9 Design quality = 12/26

***Risk of bias***

<b>Bias</b>	<b>Authors’ judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“The participants were randomly assigned to one of two groups” Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	High risk	Attrition is not described
Selective reporting (reporting bias)	Low risk	Data were fully reported

**Passchier 1990**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment
Participants	End of treatment n = 119 Start of treatment n = 119 Sex: 65 F, 54 M Mean age = 13.7 (SD 1.4) Source = school Diagnosis = headache (at least weekly) Mean years of pain = none given
Interventions	“Progressive relaxation training” “Placebo physical concentration training”
Outcomes	<b>Primary pain outcome: headache intensity</b> <b>Primary disability outcome: school problems</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: Fear of Failure</b> 1. Headache intensity 2. Headache frequency 3. Headache duration 4. School problems (composite) 5. Fear of failure (from Hermans’ Debilitating Anxiety of Achievement Motivation Test)
Notes	Original study Total quality = 15/35 Treatment quality = 5/9 Design quality = 10/26

***Risk of bias***

<b>Bias</b>	<b>Authors’ judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“The 19 classes of the participating teachers were allocated at random to a Progressive Relaxation Training or a Placebo Training group.” Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts were reported

Passchier 1990 (Continued)

Selective reporting (reporting bias)	High risk	Data were incompletely reported
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Powers 2013

Methods	RCT. 2 arms. Assessed pre-treatment, post-treatment, and 12 months
Participants	End of treatment n = 124 Start of treatment n = 135 Sex: 107 F, 28 M Mean age = 14.4 (SD 2.0) Source = clinic Diagnosis = migraine Mean years of pain = none given
Interventions	“Cognitive behavioral therapy plus amitriptyline” “Headache education plus amitriptyline”
Outcomes	<b>Primary pain outcome: headache frequency</b> <b>Primary disability outcome: PedMIDAS</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Headache diary (use of abortive medication, headache occurrence, intensity, duration, associated symptoms for migraine) 2. PedMIDAS 3. Treatment integrity 4. Treatment credibility
Notes	-

*Risk of bias*

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	“Block randomization (with varying block sizes of 4-10) was used, and participants were stratified by age. Randomization was computer generated and supplied via secure e-mail to the study therapist” Comment: probably done
Allocation concealment (selection bias)	Low risk	“Randomization was computer generated and supplied via secure e-mail to the study therapist.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Low risk	“Outcome assessments were conducted by blinded study personnel.”

**Powers 2013** (Continued)

		Comment: probably done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not described
Selective reporting (reporting bias)	Unclear risk	Data fully reported on request

**Richter 1986**

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment
Participants	End of treatment n = 43 Start of treatment n = 51 Sex: 34 F, 17 M Mean age = 12.9 Source = referred by physicians to children's hospital Diagnosis = migraine Mean years of pain = not given: most over 2 years
Interventions	"Relaxation training" "Cognitive coping" "Attention control"
Outcomes	<b>Primary pain outcome: headache diary</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: Child Depression Rating Scale</b> <b>Primary anxiety outcome: State Trait Anxiety Inventory</b> 1. Headache index (intensity, frequency, duration, medication taken: diary) 2. State Trait Anxiety Inventory (STAI) or State-Trait Anxiety Inventory for Children (STAI-C) 3. Children's Depression Rating Scale
Notes	Original study Total quality = 20/35 Treatment quality = 6/9 Design quality = 14/26

***Risk of bias***

<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"... and randomly assigned to treatment" Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done



**Richter 1986** (Continued)

Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	“Over the course of treatment there were 8 drop-outs. A chi-square analysis comparing attrition rates across interventions was not significant.” Comment: attrition adequately reported and no significant differences between completers and non-completers reported
Selective reporting (reporting bias)	High risk	Data were incompletely reported

**Robins 2005**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment (3 months after start), 6 to 12 months	
Participants	End of treatment n = 69 Start of treatment n = 86 Sex: 39 F, 30 M Mean age = 11.4 (SD 2.4) Source = paediatric gastroenterology outpatient clinic of children's hospital Diagnosis = recurrent abdominal pain Mean years of pain = not stated	
Interventions	“Short term cognitive behavioural family treatment plus standard medical care” “Standard medical care”	
Outcomes	<p><b>Primary pain outcome: Abdominal Pain Index</b></p> <p><b>Primary disability outcome: Functional Disability Inventory</b></p> <p><b>Primary depression outcome: none</b></p> <p><b>Primary anxiety outcome: none</b></p> <ol style="list-style-type: none"> <li>1. Abdominal Pain Index</li> <li>2. Child Somatization Inventory</li> <li>3. Functional Disability Inventory</li> <li>4. Abdominal Pain Index (parent)</li> <li>5. Child Somatization Inventory (parent)</li> </ol>	
Notes	Updated study 2009 Total quality = 27/35 Treatment quality = 7/9 Design quality = 20/26	
<b><i>Risk of bias</i></b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>

**Robins 2005** (Continued)

Random sequence generation (selection bias)	Low risk	“The remaining sample of 86 were randomly assigned using a coin-flip method.” Comment: probably done
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not described
Selective reporting (reporting bias)	High risk	Data were incompletely reported

**Sanders 1994**

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 6 months, 1 year
Participants	End of treatment n = 44 Start of treatment n = 44 Sex: 28 F, 16 M Mean age = 9.2 (SD 1.9) Source = not given Diagnosis = recurrent abdominal pain Mean years of pain = 3.7
Interventions	“Cognitive behaviour therapy” “Standard paediatric care”
Outcomes	<b>Primary pain outcome: pain diary</b> <b>Primary disability outcome: interference with child activity</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Pain intensity diary 2. Parent observation of child pain behaviour (POR) 3. Child behaviour checklist (CBCL '83) 4. Relapse versus pain-free 5. Interference with child activity (child report) 6. Interference with child activity (parent report)
Notes	Original study Total quality = 19/35 Treatment quality = 4/9 Design quality = 15/26

Sanders 1994 (Continued)

<i>Risk of bias</i>		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"The study used a randomised group comparison design with two treatment conditions." Comment: method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	High risk	Attrition was not described and significant differences between completers and non-completers were not reported
Selective reporting (reporting bias)	High risk	Data were incompletely reported

Sartory 1998

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment (4 weeks after end of intervention), 8 months follow-up
Participants	End of treatment n = 43 Start of treatment n = 43 Sex: 17 F, 26 M Mean age = 11.3 (SD 2.1) Source = outpatient clinic of paediatric hospital and advertising in press Diagnosis = migraine Mean years of pain = 4.6
Interventions	"Cephalic vasomotor training + stress management" "Relaxation training + stress management" "Beta-blocker (metoprolol)"
Outcomes	<b>Primary pain outcome: headache index</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: mood faces scale</b> <b>Primary anxiety outcome: none</b> 1. Headache index 2. Episodes/week when analgesics taken 3. Mood faces scale, 5-point smiling - upset

**Sartory 1998** (Continued)

Notes	Updated study 2009 Total quality = 19/35 Treatment quality = 6/9 Design quality = 13/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"Children were allocated randomly to one of three treatment groups" Comment: probably done, no method described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not described
Selective reporting (reporting bias)	Low risk	Data were fully reported

**Scharff 2002**

Methods	RCT. 3 arms. Assessed at pre-treatment, post-treatment, 3 months, 6 months, 12 months
Participants	End of treatment n = 34 Start of treatment n = 36 Sex: 24 F, 12 M Mean age 12.8 (SD 2.4) Source = children's hospital Diagnosis = migraine (all), tension-type headache (minority) Mean years of pain = 2.4 (SD 2.1)
Interventions	"Handwarming biofeedback and stress management" "Handcooling attention control" "Waitlist control"
Outcomes	<b>Primary pain outcome: headache index</b> <b>Primary disability outcome: none</b> <b>Primary depression outcome: Child Depression Inventory</b> <b>Primary anxiety outcome: State Trait Anxiety Inventory for Children</b> 1. Headache index

Scharff 2002 (Continued)

	2. Days with headache 3. Highest headache rating 4. Child Depression Inventory (CDI) 5. State-Trait Anxiety Inventory for Children (STAIC)	
Notes	Updated study 2009 Total quality = 19/35 Treatment quality = 4/9 Design quality = 15/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Low risk	"At the assessment visit children were randomised into three groups using a randomisation table" Comment: probably done
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Low risk	Attrition is described; there were no significant differences between completers and non-completers
Selective reporting (reporting bias)	High risk	Data were incompletely reported

van der Veek 2013

Methods	RCT. 2 arms. Assessed pre-treatment, post-treatment, 6 months, and 12 months follow-up
Participants	End of treatment n = 92; n = 88 at 12 months follow-up Start of treatment n = 104 Sex: 24 F, 12 M Mean age 11.9 (SD 2.77) Source = children's hospital Diagnosis = abdominal pain Mean months of pain = 34.01 (SD 37.54)
Interventions	"Cognitive behavior therapy" "Intensive medical care"

Outcomes	<p><b>Primary pain outcome: Abdominal Pain Index (child report)</b>  <b>Primary disability outcome: Functional Disability Inventory (child report)</b>  <b>Primary depression outcome: Revised Child Anxiety and Depression Scale - Short Version (child report)</b>  <b>Primary anxiety outcome: Revised Child Anxiety and Depression Scale - Short Version (child report)</b></p> <ol style="list-style-type: none"> <li>1. Abdominal pain index (completed by child and parent)</li> <li>2. Functional disability inventory (completed by child and parent)</li> <li>3. Revised Child Anxiety and Depression Scale - Short Version (completed by child and parent)</li> <li>4. KIDSCREEN (quality of life) (completed by child and parent)</li> <li>5. Satisfaction with treatment and therapist/doctor (completed by child and parent)</li> <li>6. Pain diary (child report)</li> <li>7. Health care use (follow-up only)</li> </ol>	
Notes	-	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	"The first author randomized the children using a computerized randomization program" Comment: probably done, method not described
Allocation concealment (selection bias)	Unclear risk	No description found in text Comment: probably not done
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	"Diary data were entered in SPSS by students who were blinded to treatment." Comment: probably not done but no description given for other measures
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers were not described
Selective reporting (reporting bias)	Unclear risk	Data fully reported when requested

van Tilburg 2009

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 6 months	
Participants	End of treatment n = 29; follow-up n = 24 Start of treatment n = 34 Sex: 25 F, 9 M Mean age = 10.25 (SD 2.6) Source = University of North Carolina and Duke University Medical Centres Diagnosis = functional abdominal pain Mean years of pain = unknown	
Interventions	“Guided imagery treatment” “Standard medical care”	
Outcomes	<b>Primary pain outcome: Abdominal Pain Index</b> <b>Primary disability outcome: Functional Disability Inventory</b> <b>Primary depression outcome: none</b> <b>Primary anxiety outcome: none</b> 1. Abdominal pain index 2. Functional disability inventory 3. School attendance 4. Pediatric quality of life inventory 5. Global rating of change in abdominal pain 6. Treatment compliance 7. Questionnaire of paediatric gastrointestinal symptoms 8. Health care utilisation	
Notes	Updated study 2012 Total quality = 21/35 Treatment quality = 8/9 Design quality = 13/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors’ judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Unclear risk	“Thirty-four children were assigned randomly to receive 2 months of standard medical care with or without home-based, guided imagery treatment.” Comment: probably done, method not described
Allocation concealment (selection bias)	Low risk	“Children picked a closed envelope that determined whether they would receive standard medical care with or without guided imagery treatment.” Comment: probably done

van Tilburg 2009 (Continued)

Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	No description found in text Comment: probably not done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers are not reported
Selective reporting (reporting bias)	Unclear risk	Data were fully reported

Vlieger 2007

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 6 months, 1 year	
Participants	End of treatment n = 51 Start of treatment n = 52 Sex: 39 F, 13 M Mean age = 13.3 (SD 2.7) Source = paediatric gastroenterology department in hospital Diagnosis = functional abdominal pain (n = 31) and irritable bowel syndrome (IBS) (n = 22) Mean years of pain = 3.4	
Interventions	“Gut-directed hypnotherapy” “Standard medical care plus supportive therapy”	
Outcomes	<p><b>Primary pain outcome: weekly pain intensity</b>  <b>Primary disability outcome: none</b>  <b>Primary depression outcome: none</b>  <b>Primary anxiety outcome: none</b></p> <p>1. Total pain intensity over 1 week (9-point faces affective pain intensity scale, reduced to 0 to 3 points hence 0 to 21)  2. Total pain frequency over 1 week (frequency reduced to 0 to 3 scale per day)  3. Associated symptoms (nausea, vomiting, loss of appetite, flatus, nocturnal pain, pain on wakening, pain related to meals)</p>	
Notes	Updated study 2009 Total quality = 24/35 Treatment quality = 6/9 Design quality = 18/26	

**Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	“Patients were randomly allocated using a computerised random-number generator



Vlieger 2007 (Continued)

		for concealment to either HT or standard medical care.” Comment: probably done
Allocation concealment (selection bias)	Low risk	“Patients were randomly allocated using a computerised random-number generator for concealment to either HT or standard medical care.” Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Low risk	“Pain diaries were analysed by S. W. (medical student), who was blinded to the treatment arm.” Comment: probably done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers are not reported
Selective reporting (reporting bias)	Low risk	Data were fully reported

Wicksell 2009

Methods	RCT. 2 arms. Assessed at pre-treatment, post-treatment, 3.5 months, 6.8 months
Participants	End of treatment n = 29; follow-up 3.5 months n = 24; follow-up 6.8 months = 24 Start of treatment n = 32 Sex: 25 F, 7 M Mean age = 14.8 (SD 2.4) Source = Astrid Lindgren Children’s Hospital, Karolinska University Hospital Diagnosis = mixed pain (headache, back/neck, widespread musculoskeletal, complex regional pain syndrome, visceral, lower extremities, postherpetic type cheek pain) Mean years of pain = 2.7
Interventions	“Exposure and acceptance” “Multidisciplinary treatment and amitriptyline”
Outcomes	<b>Primary pain outcome: pain intensity</b> <b>Primary disability outcome: Functional Disability Inventory</b> <b>Primary depression outcome: Center for Epidemiological Studies Depression Scale for Children</b> <b>Primary anxiety outcome: Pain Coping Scale (catastrophising sub-scale)</b> 1. Pain intensity 2. Functional disability inventory 3. Center for Epidemiological Studies Depression Scale for Children 4. Multidimensional Pain Inventory (interference scale) 5. Brief pain inventory (pain interference items) 6. Pain and impairment relationship scale

	7. Short form-36 health survey 8. Tampa scale of Kinesiophobia 9. Pain coping questionnaire (internalising and catastrophising) 10. 5 author-generated questions on pain-related discomfort	
Notes	Updated study 2012 Total quality = 20/35 Treatment quality = 6/9 Design quality = 14/26	
<b>Risk of bias</b>		
<b>Bias</b>	<b>Authors' judgement</b>	<b>Support for judgement</b>
Random sequence generation (selection bias)	Low risk	"A total of 32 participants were included in the study and randomised to one of the two treatment conditions. A simple randomisation technique was used." Comment: probably done
Allocation concealment (selection bias)	Low risk	"A sealed envelope (prepared by a secretary blind to the objective of the study) containing a code for 'exposure and acceptance' or 'MDT' was opened, assigning the participant to one of the treatment conditions." Comment: probably done
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"All assessments were conducted by a nurse who was not involved in delivering the treatment protocol." Comment: probably done
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Attrition is described, however significant differences between completers and non-completers are not reported
Selective reporting (reporting bias)	Low risk	Data are fully reported

AP: abdominal pain

CBT: cognitive behavioural therapy

F: female

FDI-C: Functional Disability Inventory - Children

HT: hypnotherapy

JPFM: juvenile primary fibromyalgia

M: male

NRS: numeric rating scale

Ped-MIDAS: Pediatric Migraine Disability Assessment

PEDSQL: Paediatric Scale Quality of Life Inventory

RCT: randomised controlled trial

SCD: sickle cell disease

SD: standard deviation

VAS: visual analogue scale

### Characteristics of excluded studies *[ordered by study ID]*

Study	Reason for exclusion
<a href="#">Connelly 2006</a>	Intervention delivered remotely
<a href="#">Fentress 1986</a>	Inadequate sample size (n < 10 in 1 arm of study design)
<a href="#">Hicks 2006</a>	Intervention delivered remotely
<a href="#">Koenig 2013</a>	Insufficient psychological treatment
<a href="#">Kroener-Herwig 1998</a>	Inadequate sample size (n < 10 in 1 arm of study design)
<a href="#">Larsson 1986</a>	Inadequate sample size (n < 10 in 1 arm of study design)
<a href="#">Olness 1987</a>	Insufficient psychological treatment
<a href="#">Palermo 2009</a>	Intervention delivered remotely
<a href="#">Sanders 1989</a>	Inadequate sample size (n < 10 in 1 arm of study design)
<a href="#">Stinson 2010</a>	Intervention delivered remotely
<a href="#">Trautmann 2008</a>	Inadequate sample size (n < 10 in 1 arm of study design)
<a href="#">Trautmann 2010</a>	Intervention delivered remotely
<a href="#">Vliieger 2012</a>	Follow-up period more than 1 year
<a href="#">Weydert 2006</a>	Inadequate sample size (n < 10 in 1 arm of study design)
<a href="#">Youssef 2009</a>	Inadequate sample size (n < 10 in 1 arm of study design)

## DATA AND ANALYSES

### Comparison 1. Treatment versus control (headache) post-treatment

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Pain	15	714	Risk Ratio (M-H, Fixed, 95% CI)	2.47 [1.97, 3.09]
2 Disability	3	263	Std. Mean Difference (IV, Random, 95% CI)	-0.49 [-0.74, -0.24]
3 Depression	3	164	Std. Mean Difference (IV, Random, 95% CI)	-0.18 [-0.49, 0.14]
4 Anxiety	4	203	Std. Mean Difference (IV, Random, 95% CI)	-0.33 [-0.61, -0.04]

### Comparison 2. Treatment versus control (headache) follow-up

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Pain	5	251	Risk Ratio (M-H, Random, 95% CI)	2.89 [1.03, 8.07]
2 Disability	2	148	Std. Mean Difference (IV, Random, 95% CI)	-0.46 [-0.78, -0.13]
3 Depression	1	24	Std. Mean Difference (IV, Random, 95% CI)	-0.54 [-1.36, 0.28]
4 Anxiety	2	67	Std. Mean Difference (IV, Random, 95% CI)	-0.28 [-1.00, 0.45]

### Comparison 3. Treatment versus control (non-headache) post-treatment

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Pain	13	852	Std. Mean Difference (IV, Random, 95% CI)	-0.57 [-0.86, -0.27]
2 Disability	11	764	Std. Mean Difference (IV, Random, 95% CI)	-0.45 [-0.71, -0.19]
3 Depression	6	538	Std. Mean Difference (IV, Random, 95% CI)	-0.07 [-0.30, 0.17]
4 Anxiety	5	498	Std. Mean Difference (IV, Random, 95% CI)	-0.15 [-0.36, 0.07]

### Comparison 4. Treatment versus control (non-headache) follow-up

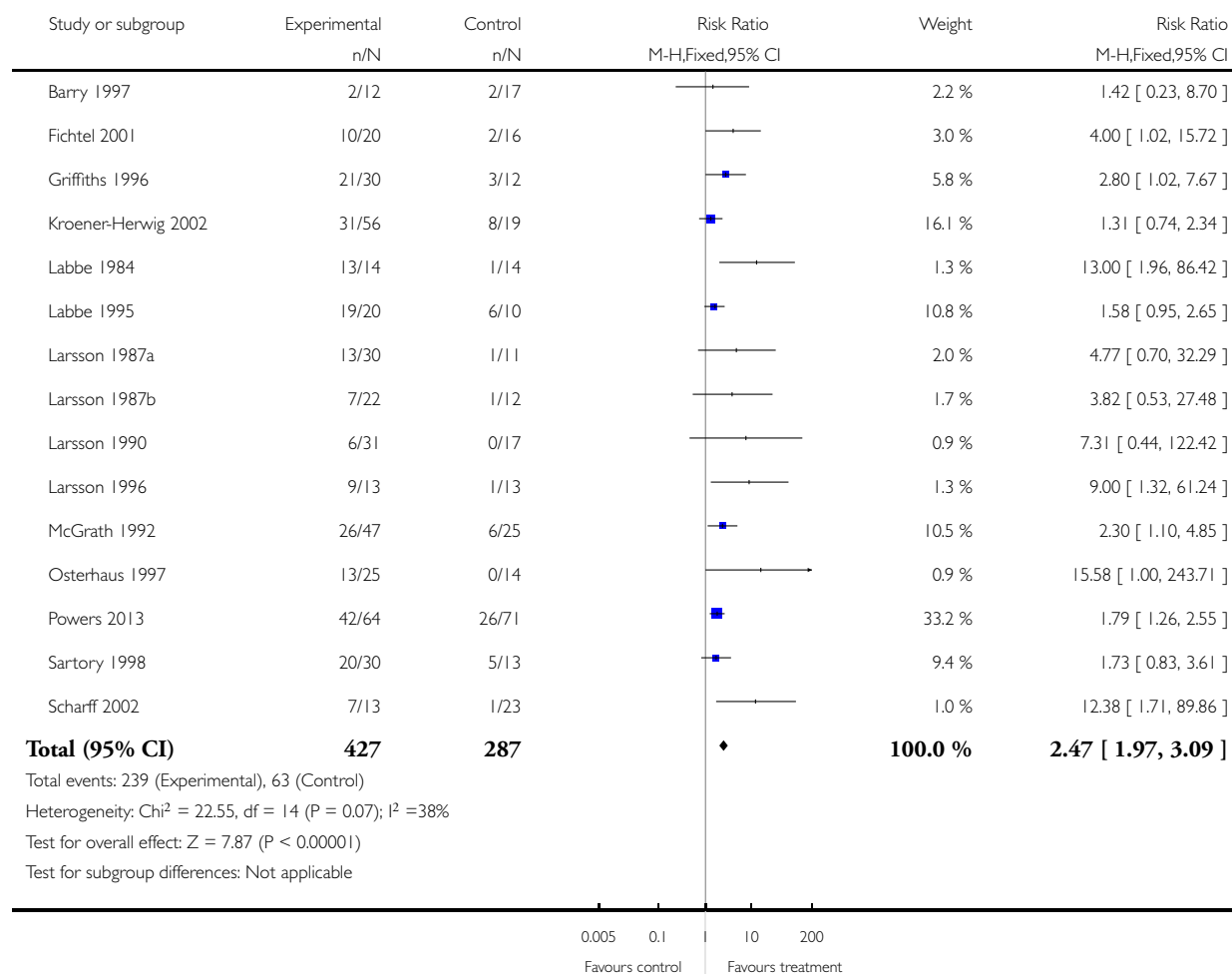
Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Pain	7	543	Std. Mean Difference (IV, Random, 95% CI)	-0.11 [-0.41, 0.19]
2 Disability	6	508	Std. Mean Difference (IV, Random, 95% CI)	-0.35 [-0.71, 0.02]
3 Depression	5	473	Std. Mean Difference (IV, Random, 95% CI)	0.06 [-0.16, 0.28]
4 Anxiety	5	452	Std. Mean Difference (IV, Random, 95% CI)	0.05 [-0.24, 0.33]

### Analysis 1.1. Comparison 1 Treatment versus control (headache) post-treatment, Outcome 1 Pain.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 1 Treatment versus control (headache) post-treatment

Outcome: 1 Pain

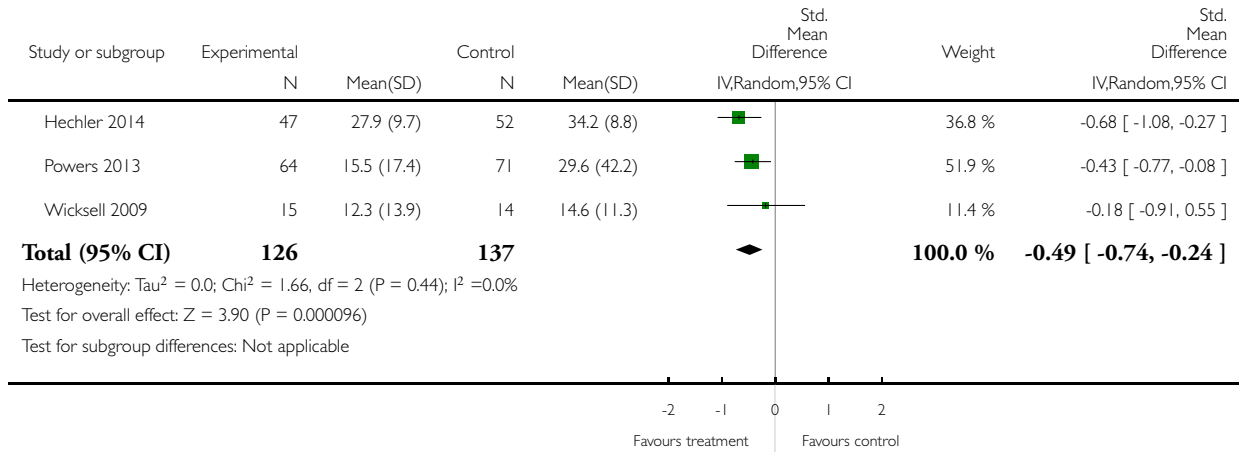


## Analysis 1.2. Comparison 1 Treatment versus control (headache) post-treatment, Outcome 2 Disability.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 1 Treatment versus control (headache) post-treatment

Outcome: 2 Disability

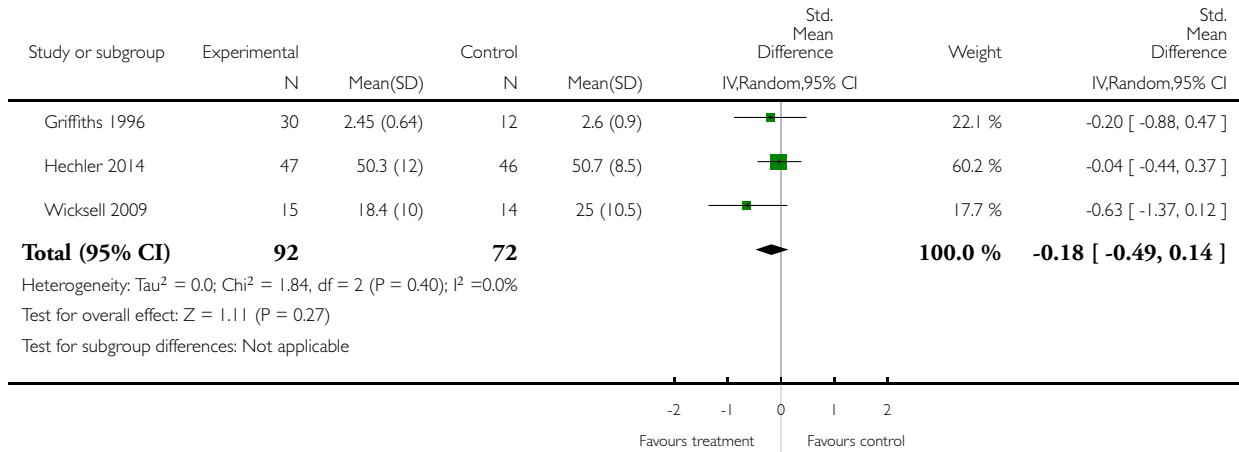


### Analysis 1.3. Comparison 1 Treatment versus control (headache) post-treatment, Outcome 3 Depression.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 1 Treatment versus control (headache) post-treatment

Outcome: 3 Depression

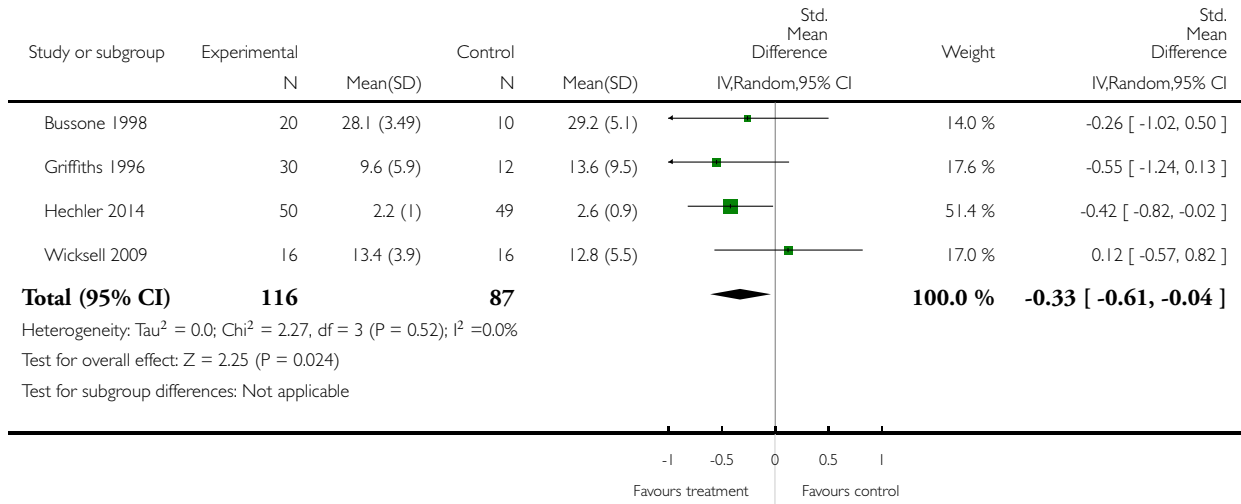


### Analysis 1.4. Comparison 1 Treatment versus control (headache) post-treatment, Outcome 4 Anxiety.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 1 Treatment versus control (headache) post-treatment

Outcome: 4 Anxiety



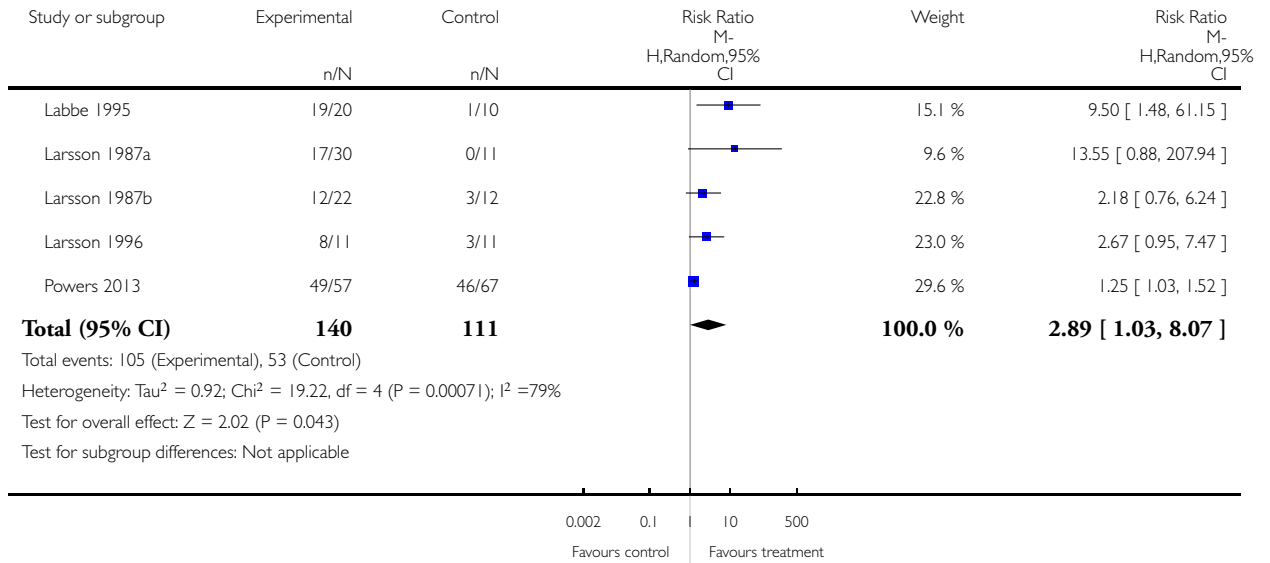


## Analysis 2.1. Comparison 2 Treatment versus control (headache) follow-up, Outcome 1 Pain.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 2 Treatment versus control (headache) follow-up

Outcome: 1 Pain

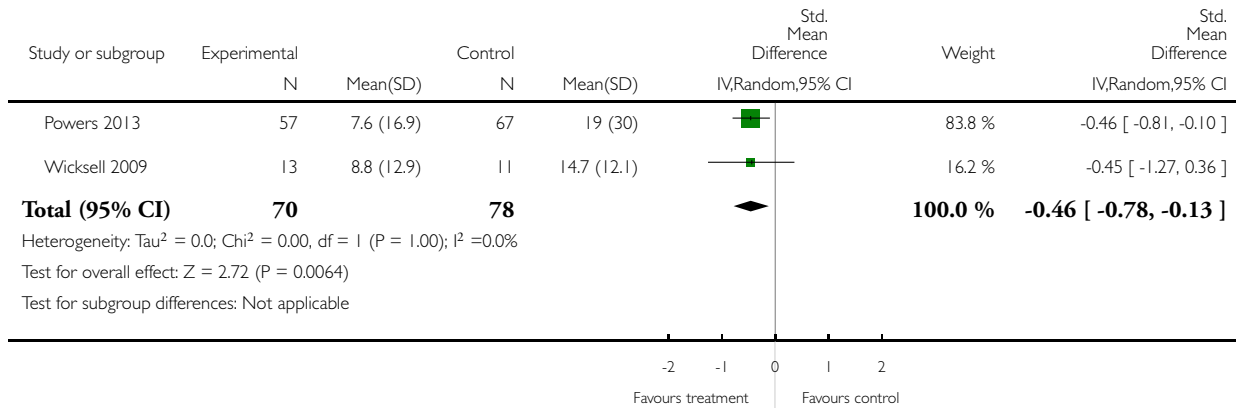


### Analysis 2.2. Comparison 2 Treatment versus control (headache) follow-up, Outcome 2 Disability.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 2 Treatment versus control (headache) follow-up

Outcome: 2 Disability

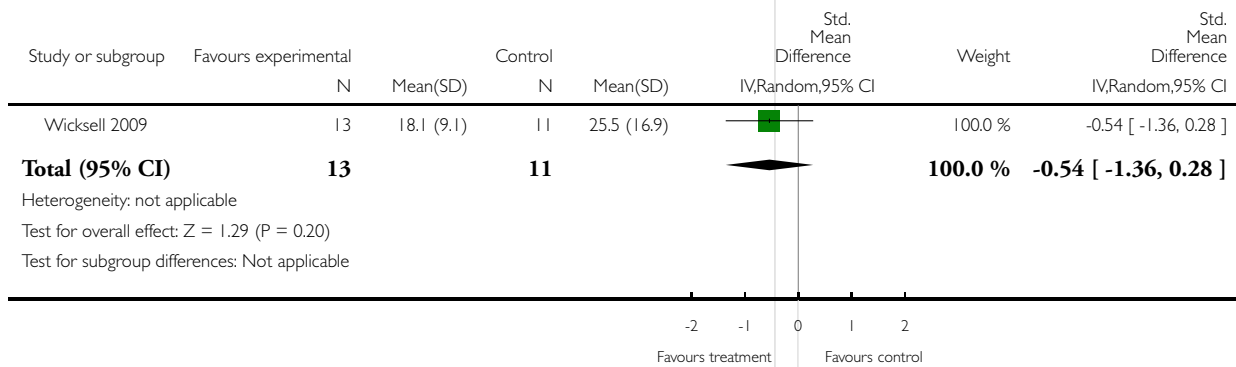


### Analysis 2.3. Comparison 2 Treatment versus control (headache) follow-up, Outcome 3 Depression.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 2 Treatment versus control (headache) follow-up

Outcome: 3 Depression

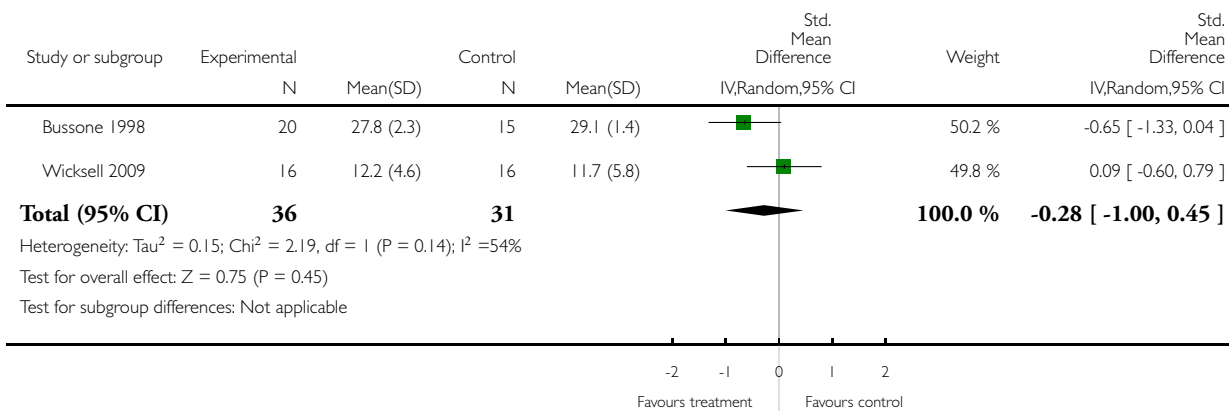


## Analysis 2.4. Comparison 2 Treatment versus control (headache) follow-up, Outcome 4 Anxiety.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 2 Treatment versus control (headache) follow-up

Outcome: 4 Anxiety

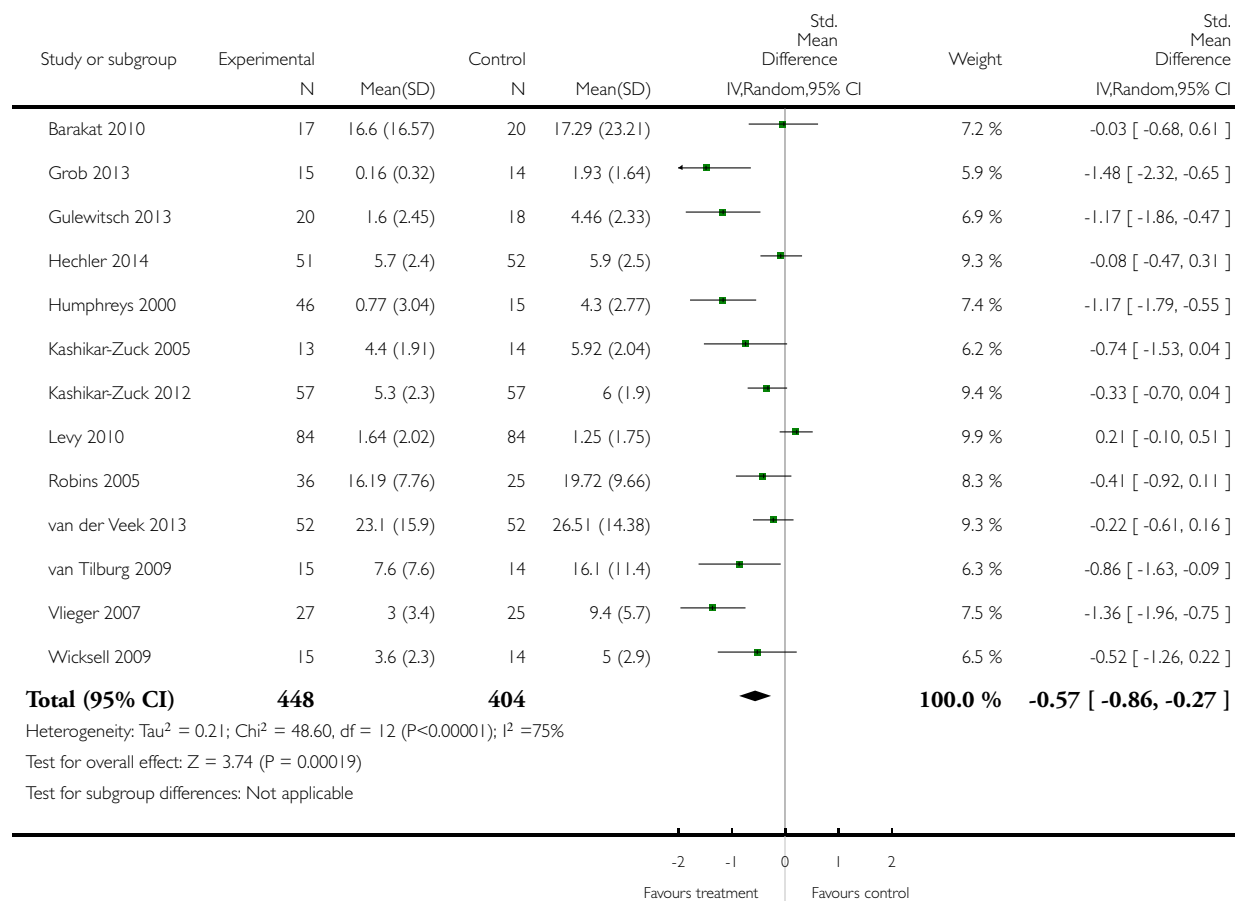


### Analysis 3.1. Comparison 3 Treatment versus control (non-headache) post-treatment, Outcome 1 Pain.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 3 Treatment versus control (non-headache) post-treatment

Outcome: 1 Pain

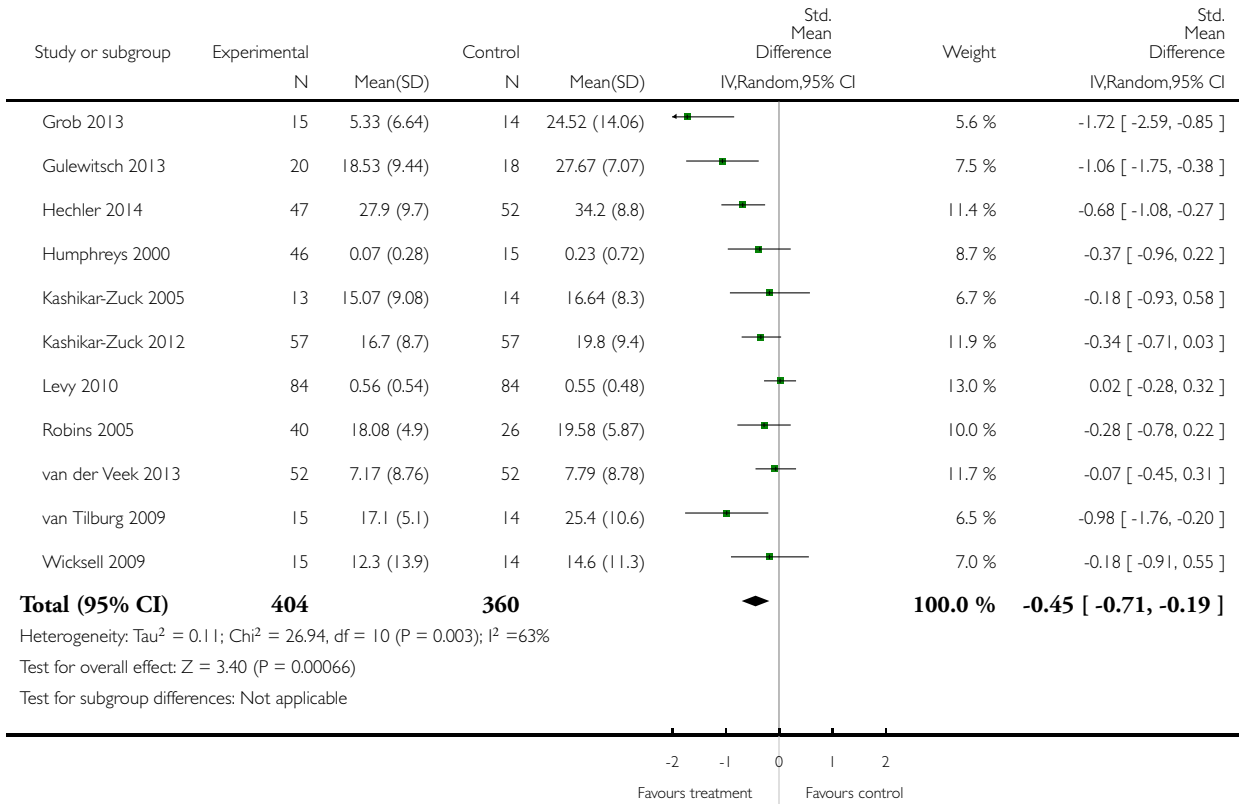


### Analysis 3.2. Comparison 3 Treatment versus control (non-headache) post-treatment, Outcome 2 Disability.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 3 Treatment versus control (non-headache) post-treatment

Outcome: 2 Disability

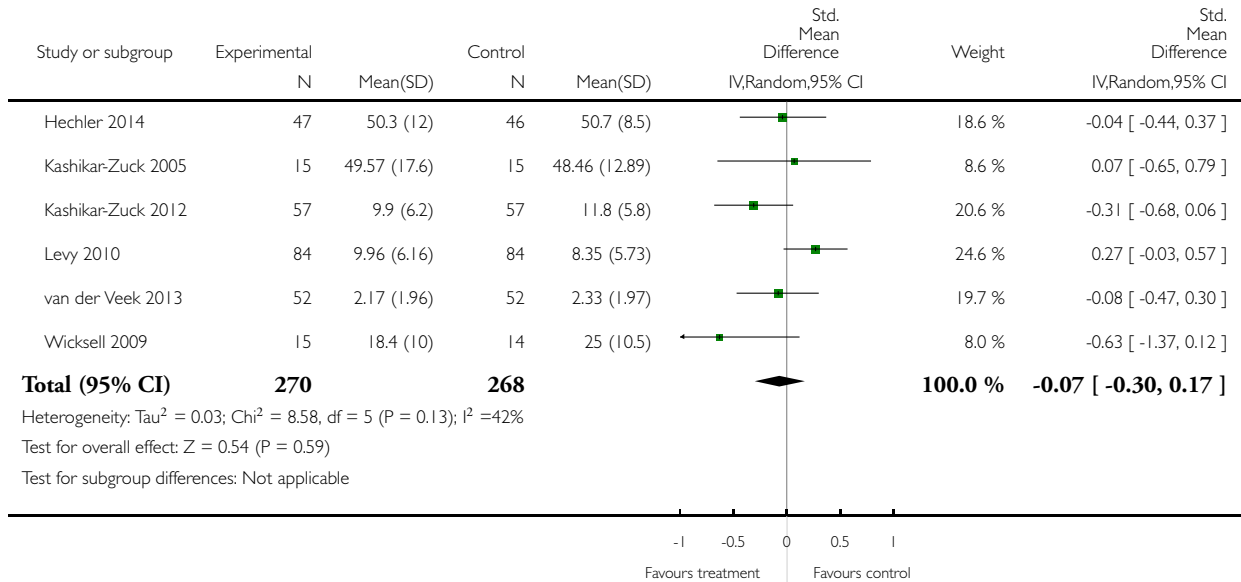


### Analysis 3.3. Comparison 3 Treatment versus control (non-headache) post-treatment, Outcome 3 Depression.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 3 Treatment versus control (non-headache) post-treatment

Outcome: 3 Depression

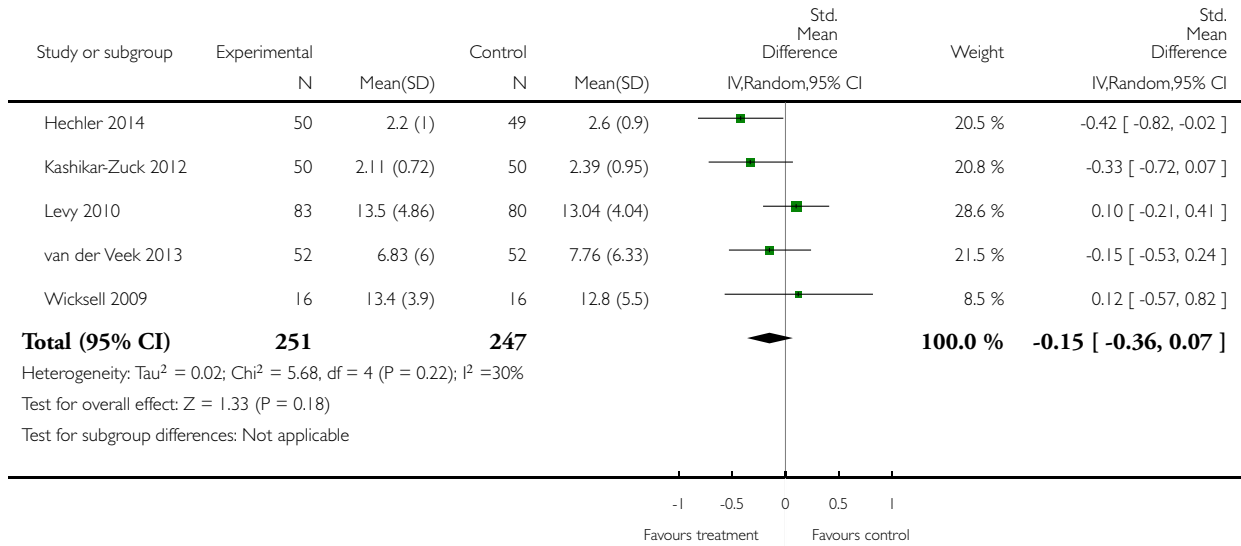


### Analysis 3.4. Comparison 3 Treatment versus control (non-headache) post-treatment, Outcome 4 Anxiety.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 3 Treatment versus control (non-headache) post-treatment

Outcome: 4 Anxiety

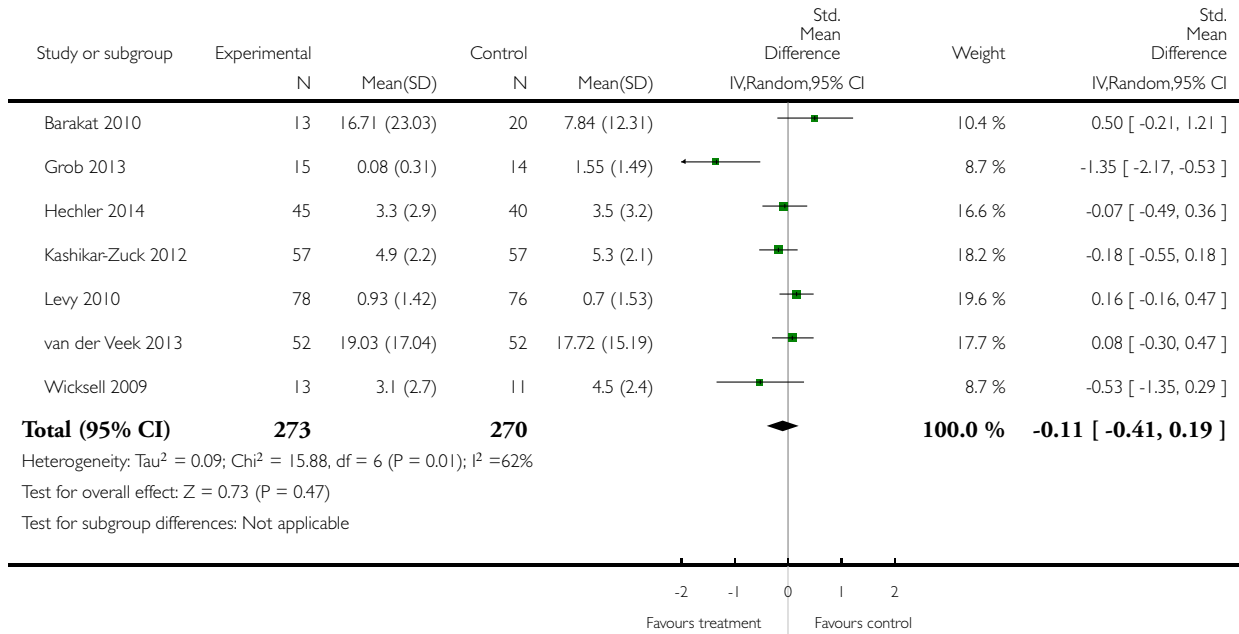


### Analysis 4.1. Comparison 4 Treatment versus control (non-headache) follow-up, Outcome 1 Pain.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 4 Treatment versus control (non-headache) follow-up

Outcome: 1 Pain



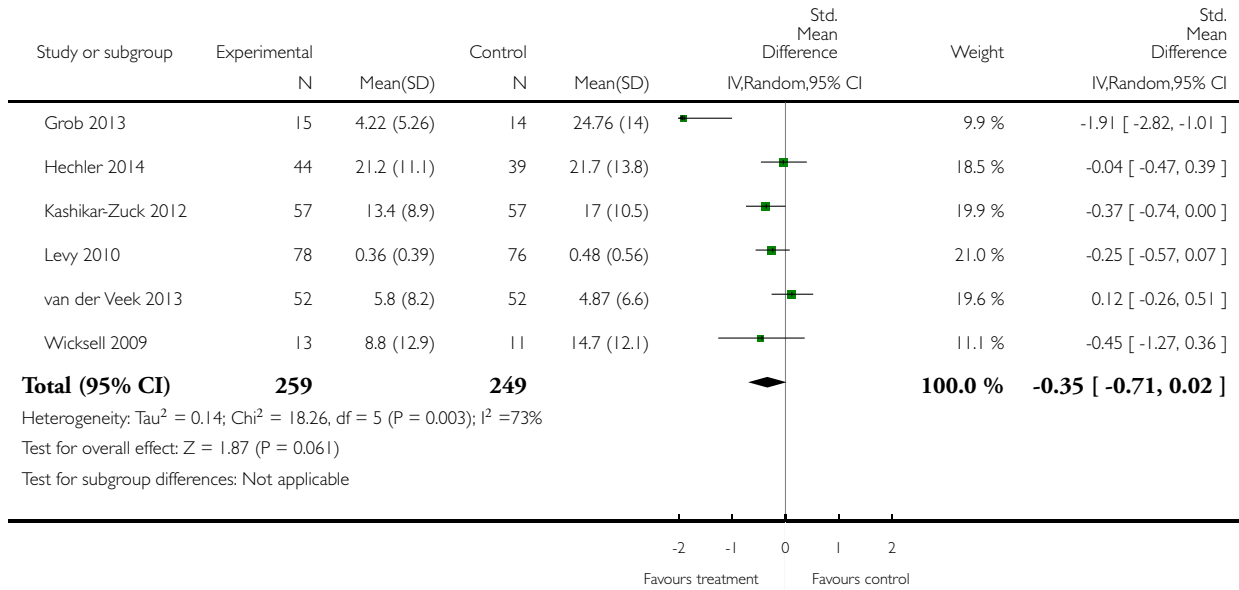


### Analysis 4.2. Comparison 4 Treatment versus control (non-headache) follow-up, Outcome 2 Disability.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 4 Treatment versus control (non-headache) follow-up

Outcome: 2 Disability

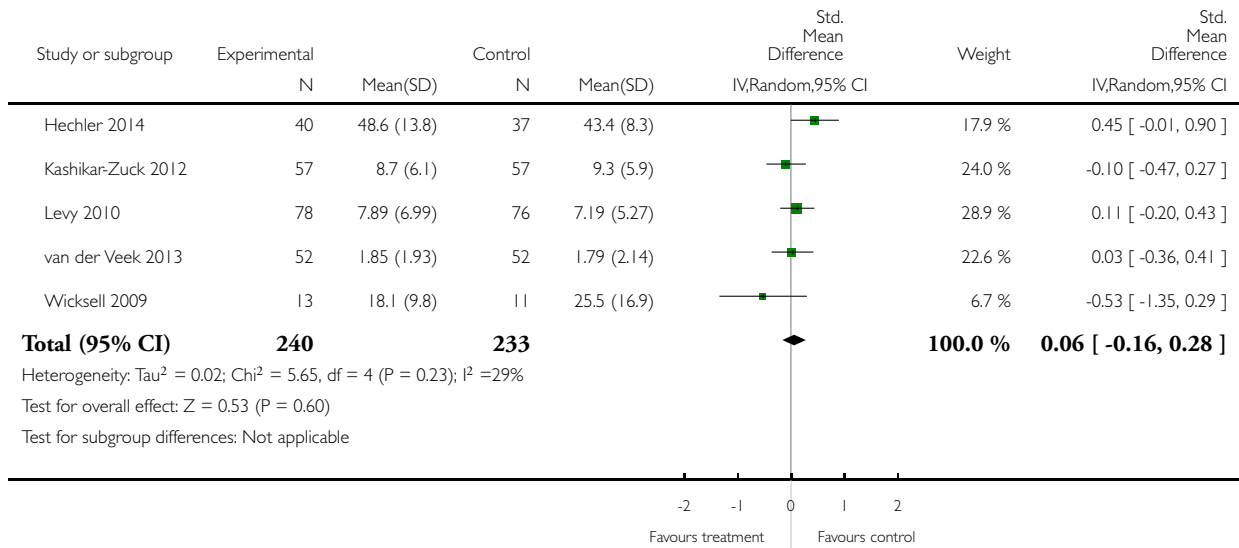


### Analysis 4.3. Comparison 4 Treatment versus control (non-headache) follow-up, Outcome 3 Depression.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 4 Treatment versus control (non-headache) follow-up

Outcome: 3 Depression

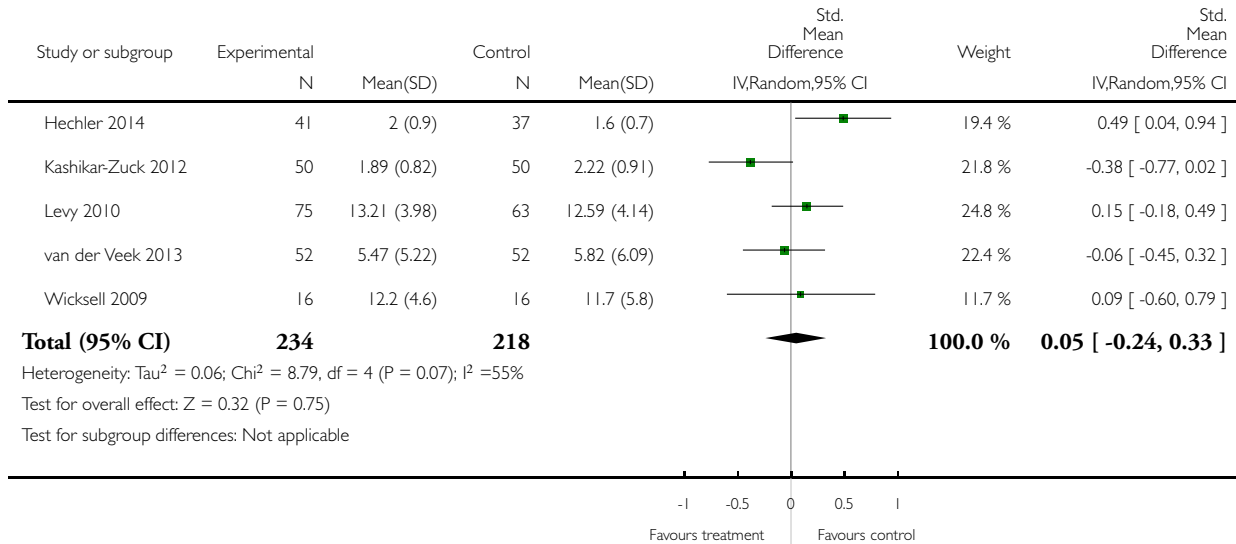


#### Analysis 4.4. Comparison 4 Treatment versus control (non-headache) follow-up, Outcome 4 Anxiety.

Review: Psychological therapies for the management of chronic and recurrent pain in children and adolescents

Comparison: 4 Treatment versus control (non-headache) follow-up

Outcome: 4 Anxiety



## ADDITIONAL TABLES

Table 1. Duration of treatment and setting by condition

Headache studies			
Author	Illness	Treatment duration (hours)	Setting
<a href="#">Abram 2007</a>	Headache	1.5	Clinic
<a href="#">Barry 1997</a>	Headache	3	Unknown
<a href="#">Bussone 1998</a>	Headache	7	Clinic
<a href="#">Fichtel 2001</a>	Headache	6.75	Clinic
<a href="#">Griffiths 1996</a>	Headache	12	Clinic/home
<a href="#">Hechler 2014</a>	Mixed	136.5 (3-week intensive therapy)	Clinic
<a href="#">Kroener-Herwig 2002</a>	Headache	12	Clinic

**Table 1. Duration of treatment and setting by condition** (Continued)

Labbe 1984	Headache	6.7	Clinic
Labbe 1995	Headache	7.5	Clinic
Larsson 1987a	Headache	6.75	School
Larsson 1987b	Headache	5	School
Larsson 1990	Headache	1.7	Home
Larsson 1996	Headache	3.3	Clinic
McGrath 1988	Headache	6	Unknown
McGrath 1992	Headache	8	Home/clinic
Osterhaus 1997	Headache	9.3	Clinic
Passchier 1990	Headache	2.5	School
Powers 2013	Headache	13	Clinic
Richter 1986	Headache	9	Unknown
Sartory 1998	Headache	Unknown	Clinic
Scharff 2002	Headache	4	Clinic
Wicksell 2009*	Mixed	10	Clinic
<b>Non-headache studies</b>			
<b>Author</b>	<b>Illness</b>	<b>Treatment duration (hours)</b>	<b>Setting</b>
Alfven 2007	RAP	Unknown	Clinic
Barakat 2010	SCD	6	Home
Duarte 2006	RAP	3.3	Unknown
Gil 1997	SCD	0.75	Clinic
Grob 2013	RAP	9	Clinic
Gulewitsch 2013	RAP/IBS	2	Clinic
Hechler 2014	Mixed	136.5 (3-week intensive therapy, psychological content unknown)	Clinic
Humphreys 2000	RAP	Unknown	Clinic

**Table 1. Duration of treatment and setting by condition** (Continued)

Kashikar-Zuck 2005	Fibromyalgia	6	Clinic
Kashikar-Zuck 2012	Fibromyalgia	7.5	Unknown
Levy 2010	RAP	4	Home/clinic
Robins 2005	RAP	3.5	Clinic
Sanders 1994	RAP	6	Clinic
van der Veek 2013	RAP	4.5	Clinic
van Tilburg 2009	RAP	1.8	Home
Vlioger 2007	RAP/IBS	5	Clinic
Wicksell 2009*	Mixed	10	Clinic

\*Mixed headache and non-headache studies are entered twice.

Recurrent abdominal pain (RAP), sickle cell disease (SCD), juvenile idiopathic arthritis (JIA), irritable bowel syndrome (IBS).

## APPENDICES

### Appendix I. Search strategies

#### MEDLINE via Ovid search strategy

1. exp child/
2. Infant/
3. Adolescent/
4. (child\$ or adolescent\$ or infant\$ or juvenil\$ or pediatric\$ or paediatric\$ or “young person\$” or “young people” or youth\$ or “young adult\$”).ab,it,kf.
5. 1 or 2 or 3 or 4
6. exp Psychology/
7. exp Psychotherapy/
8. exp Behavior Therapy/
9. (psycholog\$ or (behavio?r and therapy) or hypnos\$ or relaxation\$ or ((family or color or colour or music or play) adj therap\$) or imagery or cogniti\$ or psychotherap\$).ab,it,kf.
10. 6 or 7 or 8 or 9
11. (pain\$ or headache\$ or “head ache\$” or head-ache\$ or migraine\$ or cephalalg\$ or “stomach ache\$” or “tummy ache\$” or “abdominal ache\$” or “belly ache\$” or earache\$ or ear-ache\$ or toothache\$ or tooth-ache\$ or odontalg\$ or dysmenorrh\$ or neuralgi\$).ab,it,kf.
12. exp Pain/
13. exp Headache Disorders/

14. 11 or 12 or 13
15. 5 and 10 and 14
- 16 randomized controlled trial.pt.
- 17 controlled clinical trial.pt.
- 18 randomized.ab.
- 19 placebo.ab.
- 20 drug therapy.fs.
- 21 randomly.ab.
- 22 trial.ab.
- 23 or/16-22
- 24 exp animals/ not humans.sh.
- 25 23 not 24
- 26 25 and 15

### **EMBASE via Ovid search strategy**

1. Child/
2. Infant/
3. Adolescent/
4. (child\$ or adolescent\$ or infant\$ or juvenil\$ or pediatric\$ or paediatric\$ or “young person\$” or “young people” or youth\$ or “young adult\$”).ab,it.
5. 1 or 2 or 3 or 4
6. exp PSYCHOLOGY/
7. exp PSYCHOTHERAPY/
8. behavior therapy/
9. (psycholog\$ or (behavio?r and therapy) or hypnos\$ or relaxation\$ or ((family or color or colour or music or play) adj therap\$) or imagery or cogniti\$ or psychotherap\$).ab,it.
10. 6 or 7 or 8 or 9
11. (pain\$ or headache\$ or “head ache\$” or head-ache\$ or migraine\$ or cephalalgi\$ or “stomach ache\$” or “tummy ache\$” or “abdominal ache\$” or “belly ache\$” or earache\$ or ear-ache\$ or toothache\$ or tooth-ache\$ or odontalgi\$ or dysmenorrh\$ or neuralgi\$).ab,it.
12. exp Pain/
13. exp “Headache and Facial Pain”/
14. 11 or 12 or 13
15. 5 and 10 and 14
- 16 random\$.tw.
- 17 factorial\$.tw.
- 18 crossover\$.tw.
- 19 cross over\$.tw.
- 20 cross-over\$.tw.
- 21 placebo\$.tw.
- 22 (doubl\$ adj blind\$).tw.
- 23 (singl\$ adj blind\$).tw.
- 24 assign\$.tw.
- 25 allocat\$.tw.
- 26 volunteer\$.tw.
- 27 Crossover Procedure/
- 28 double-blind procedure.tw.
- 29 Randomized Controlled Trial/
- 30 Single Blind Procedure/
- 31 or/16-30
- 32 (animal/ or nonhuman/) not human/
- 33 31 not 32
- 34 15 and 33

## PsycINFO via OVID

1. (child\$ or adolescent\$ or infant\$ or juvenil\$ or pediatric\$ or paediatric\$ or "young person\$" or "young people" or youth\$ or "young adult\$").ab,it.
2. exp PSYCHOLOGY/
3. exp PSYCHOTHERAPY/
4. behavior therapy/
5. (psycholog\$ or (behavio?r and therapy) or hypnos\$ or relaxation\$ or ((family or color or colour or music or play) adj therap\$) or imagery or cogniti\$ or psychotherap\$).ab,it.
6. 2 or 3 or 4 or 5
7. (pain\$ or headache\$ or "head ache\$" or head-ache\$ or migraine\$ or cephalalg\$ or "stomach ache\$" or "tummy ache\$" or "abdominal ache\$" or "belly ache\$" or earache\$ or ear-ache\$ or toothache\$ or tooth-ache\$ or odontalg\$ or dysmenorrh\$ or neuralgi\$).ab,it.
8. exp Pain/
9. Headache/
10. Migraine Headache/
11. Muscle Contraction Headache/
12. 7 or 8 or 9 or 10 or 11
13. 1 and 6 and 12
- 14 clinical trials/
- 15 (randomis\* or randomiz\*).tw.
- 16 (random\$ adj3 (allocat\$ or assign\$)).tw.
- 17 ((clinic\$ or control\$) adj trial\$).tw.
- 18 ((singl\$ or doubl\$ or trebl\$ or tripl\$) adj3 (blind\$ or mask\$)).tw.
- 19 (crossover\$ or "cross over\$").tw.
- 20 random sampling/
- 21 Experiment Controls/
- 22 Placebo/
- 23 placebo\$.tw.
- 24 exp program evaluation/
- 25 treatment effectiveness evaluation/
- 26 ((effectiveness or evaluat\$) adj3 (stud\$ or research\$)).tw.
- 27 or/14-26
- 28 13 and 27

## CENTRAL (The Cochrane Library)

- #1 MeSH descriptor: [Child] explode all trees
- #2 MeSH descriptor: [Infant] explode all trees
- #3 MeSH descriptor: [Adolescent] explode all trees
- #4 (child\* or adolescent\* or infant\* or juvenil\* or pediatric\* or paediatric\* or "young person\*" or "young people" or youth\* or "young adult\*"):it,ab,kw (Word variations have been searched)
- #5 #1 or #2 or #3 or #4
- #6 MeSH descriptor: [Psychology] explode all trees
- #7 MeSH descriptor: [Psychotherapy] explode all trees
- #8 MeSH descriptor: [Behavior Therapy] explode all trees
- #9 (psycholog\* or (behavio?r and therapy) or hypnos\* or relaxation\* or ((family or color or colour or music or play) next therap\*) or imagery or cogniti\* or psychotherap\*):it,ab,kw (Word variations have been searched)
- #10 #6 or #7 or #8 or #9
- #11 (pain\* or headache\* or "head ache\*" or head-ache\* or migraine\* or cephalalg\* or "stomach ache\*" or "tummy ache\*" or "abdominal ache\*" or "belly ache\*" or earache\* or ear-ache\* or toothache\* or tooth-ache\* or odontalg\* or dysmenorrh\* or neuralgi\*):it,ab,kw (Word variations have been searched)
- #12 MeSH descriptor: [Pain] explode all trees
- #13 MeSH descriptor: [Headache Disorders] explode all trees
- #14 #11 or #12 or #13

#15 #5 and #10 and #14

## Appendix 2. Previous search results

Four separate searches have been undertaken. The first search was undertaken from inception of the abstracting services to the end of 1999 (Eccleston 2003a). This yielded 3715 abstracts, of which 123 were read in full, identifying 18 RCTs. The second search, which updated the original review, was undertaken focusing on the 10 years since the previous search, overlapping by one year (from 1999 to 2008) and was later published (Eccleston 2009). This yielded 1319 abstracts, of which 45 papers were read in full, identifying a further 16 RCTs, giving a total set of 34. However, five studies were later excluded because they did not meet the minimum criteria of 10 participants in each arm, therefore, leaving 29 studies. The third, which searched databases from 2008 to March 2012 yielded 851 abstracts, of which 25 papers were read in full, and eight further RCTs were included in the review (Eccleston 2012). The fourth searched databases from March 2012 to January 2014 yielding 443 abstracts, of which 19 were read in full, and seven papers were included (Groß 2013; Gulewitsch 2013; Hechler 2014; Kashikar-Zuck 2012; Levy 2010; Powers 2013; van der Veen 2013). Kashikar-Zuck 2012 and Levy 2010 provided additional data to previously included studies. Five studies, which were previously included, were excluded from this review since treatment was delivered remotely (Connelly 2006; Hicks 2006; Palermo 2009; Stinson 2010; Trautmann 2010). Therefore, a total of 37 RCTs are included (39 papers).

## WHAT'S NEW

Date	Event	Description
9 February 2016	Review declared as stable	See <a href="#">Published notes</a> .

## HISTORY

Date	Event	Description
14 May 2014	Amended	Minor change to the GRADE assessment wording.
30 April 2014	New citation required but conclusions have not changed	A new search was run in January 2014.
14 March 2014	New search has been performed	Five new studies were added. Two trials containing additional information for previously included studies were included. Five studies that were previously included were excluded as they delivered treatment remotely. These will be included in the new Cochrane review ('Psychological therapies (remotely delivered) for the management of chronic and recurrent pain in children and adolescents'). 'Mood' outcome was split into two discrete domains; anxiety and depression
21 August 2013	Amended	'Summary of findings' tables have been updated.



(Continued)

24 October 2012	New citation required and conclusions have changed	The previous review reported that psychological treatments were effective for headache and non-headache groups at post-treatment and effects were maintained at follow-up. Updated studies have altered the previous results. The current update found that pain improved at post-treatment for headache and non-headache groups, and for headache groups at follow-up. An additional significant finding for disability at post-treatment for the non-headache group was found. Conclusions have been updated accordingly
24 October 2012	New search has been performed	New authors have been added to this review. A new search was run in March 2012. Eight new studies were added ( <a href="#">Barakat 2010</a> ; <a href="#">Kashikar-Zuck 2012</a> ; <a href="#">Levy 2010</a> ; <a href="#">Palermo 2009</a> ; <a href="#">Stinson 2010</a> ; <a href="#">Trautmann 2010</a> ; <a href="#">van Tilburg 2009</a> ; <a href="#">Wicksell 2009</a> ), and four new studies were excluded ( <a href="#">Trautmann 2008</a> ; <a href="#">Vlieger 2012</a> ; <a href="#">Weydert 2006</a> ; <a href="#">Youssef 2009</a> ).
16 May 2008	Amended	Converted to new review format.

## CONTRIBUTIONS OF AUTHORS

Christopher Eccleston oversaw the project, contributed to the design, analysis and authoring of the text, and is responsible for any future update of this review.

Amy Lewandowski Holley, Emma Fisher, Emily Law, Stephen Morley, Tonya Palermo, and Amanda Williams all contributed to the design, analysis, and authoring of the text.

## DECLARATIONS OF INTEREST

None known.

## DIFFERENCES BETWEEN PROTOCOL AND REVIEW

1. In [Eccleston 2009](#), odds ratios and risk ratios were reported for dichotomous outcomes. In this review we only report risk ratio.
2. In this review, therapy that was delivered remotely (e.g. via Internet, telephone) has been removed and the 'mood' outcome has been separated into two discrete outcomes: depression and anxiety.

## NOTES

At February 2016, there are no new potentially relevant studies likely to change the conclusions. Therefore, this review has now been stabilised following discussion with the authors and editors. The review will be re-assessed for updating in 2017.

## INDEX TERMS

### Medical Subject Headings (MeSH)

Abdominal Pain [therapy]; Chronic Pain [etiology; psychology; \*therapy]; Cognitive Therapy; Fibromyalgia [therapy]; Headache [therapy]; Hemoglobin SC Disease [complications]; Mood Disorders [therapy]; Pain Management [\*methods; psychology]; Psychotherapy [\*methods]; Randomized Controlled Trials as Topic; Recurrence

### MeSH check words

Adolescent; Child; Humans