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RESEARCH

Physiotherapy intervention in Parkinson's disease: systematic review and meta-analysis

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Abstract

Objective To assess the effectiveness of physiotherapy compared with no intervention in patients with Parkinson's disease.

Design Systematic review and meta-analysis of randomised controlled trials.

Data sources Literature databases, trial registries, journals, abstract books, and conference proceedings, and reference lists, searched up to the end of January 2012.

Review methods Randomised controlled trials comparing physiotherapy with no intervention in patients with Parkinson's disease were eligible. Two authors independently abstracted data from each trial. Standard meta-analysis methods were used to assess the effectiveness of physiotherapy compared with no intervention. Tests for heterogeneity were used to assess for differences in treatment effect across different physiotherapy interventions used. Outcome measures were gait, functional mobility and balance, falls, clinician rated impairment and disability measures, patient rated quality of life, adverse events, compliance, and economic analysis outcomes.

Results 39 trials of 1827 participants met the inclusion criteria, of which 29 trials provided data for the meta-analyses. Significant benefit from physiotherapy was reported for nine of 18 outcomes assessed. Outcomes which may be clinically significant were speed (0.04 m/s, 95% confidence

interval 0.02 to 0.06, P<0.001), Berg balance scale (3.71 points, 2.30 to 5.11, P<0.001), and scores on the unified Parkinson's disease rating scale (total score -6.15 points, -8.57 to -3.73, P<0.001; activities of daily living subscore -1.36, -2.41 to -0.30, P=0.01; motor subscore -5.01, -6.30 to -3.72, P<0.001). Indirect comparisons of the different physiotherapy interventions found no evidence that the treatment effect differed across the interventions for any outcomes assessed, apart from motor subscores on the unified Parkinson's disease rating scale (in which one trial was found to be the cause of the heterogeneity).

Conclusions Physiotherapy has short term benefits in Parkinson's disease. A wide range of physiotherapy techniques are currently used to treat Parkinson's disease, with little difference in treatment effects. Large, well designed, randomised controlled trials with improved methodology and reporting are needed to assess the efficacy and cost effectiveness of physiotherapy for treating Parkinson's disease in the longer term.

Introduction

Parkinson's disease is a complex neurodegenerative disorder¹ with wide reaching implications for patients and their families. The management of Parkinson's disease has traditionally centred on drug treatment,² but even with optimal medical management, patients still experience a deterioration of body function, daily

Extra material supplied by the author (see http://www.bmj.com/content/345/bmj. Web table 1: Characteristics of included studies Web table 2: Risk of bias in included studies Web figure 1: Two or six min walk test (m) Web figure 2: Freezing of gait questionnaire Web figure 3: 10 or 20 m walk test (s) Web figure 4: Unified Parkinson's disease rating scale – activities of daily living

Web figure 5: Unified Parkinson's disease rating scale - total score

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activities, participation,³ and decline in mobility.⁴ This can lead to increased dependence on others, inactivity,⁵ and social isolation,⁴ resulting in reduced quality of life.⁴ There has been increasing support for the inclusion of rehabilitation therapies as an adjuvant to pharmacological and neurosurgical treatment,⁶³ and a call for the move towards multidisciplinary management.¹⁷⁸ The physiotherapist is a member within this multidisciplinary team,¹⁹ with the purpose of maximising functional ability and minimising secondary complications through movement rehabilitation within a context of education and support for the whole person.^{10 11} Physiotherapy for Parkinson's disease focuses on transfers, posture, upper limb function, balance (and falls), gait, and physical capacity and (in)activity. It also uses cueing strategies, cognitive movement strategies, and exercise to maintain or increase independence, safety, and quality of life.4 12

Referral rates to physiotherapy for people with Parkinson's disease have historically been low, owing to a weak evidence base and poor availability of physiotherapy services.^{13 14} In recent years, supportive evidence for the inclusion of physiotherapy in the management of Parkinson's disease has grown, due to the increased number of trials particularly in the past five years.¹⁵ Recent management guidelines have supported physiotherapy, such as those from the United Kingdom National Institute for Health and Clinical Excellence (NICE)¹⁶ and the Royal Dutch Society of Physical Therapy.¹⁷ This has led to an increased number of referrals, with a survey by Parkinson's UK in 2008 reporting that 54% of the 13 000 members surveyed had seen a physiotherapist.¹⁸

To synthesise the latest trial reports with the older data, we have performed this systematic review and meta-analysis of all randomised controlled trials of physiotherapy in Parkinson's disease. This review includes trials assessing a variety of different physiotherapy methods as used to treat people with Parkinson's disease, to provide an overall assessment on the use of physiotherapy in this patient population. Previous reviews have focused on one type of physiotherapy (such as exercise or treadmill training).^{19 20} Detailed results have been published in the Cochrane Library²¹ updating the Cochrane review published in 2001.¹¹

Methods

Search strategy and selection criteria

A systematic search of the literature to the end of January 2012 was undertaken using a highly sensitive search strategy as recommended by the Cochrane Collaboration.²² We combined text and, where appropriate, Medical Subject Heading terms for physiotherapy, physical therapy, exercise, or rehabilitation; and Parkinson, Parkinson's disease, or parkinsonism. No language restrictions were applied. We identified relevant trials by electronic searches of general biomedical and science electronic databases (Medline, Embase, Cumulative Index to Nursing and Allied Health Literature, Web of Science), rehabilitation databases (Allied and Complimentary Medicine Database, REHABDATA, REHADAT, GEROLIT); English language databases of foreign language research and third world publications (Latin American and Caribbean Health Sciences Literature, MedCarib, Index medicus for the Eastern Mediterranean region); conference and grey literature databases (Conference Proceedings Citation Index, Dissertation Abstracts, Conference Papers Index, Index to Theses, Electronic Theses Online Service, ProQuest), and trial registries (Cochrane Central Register of Controlled Trials, CentreWatch Clinical Trials listing service, metaRegister of Controlled Trials, ClinicalTrials.gov,

Research Portfolio Online Reporting Tools, Physiotherapy Evidence Database, National Institute of disability and rehabilitation register, National research register). We also hand searched relevant general (for example, *Lancet*, *BMJ*, *JAMA*) and specific journals (for example, *Movement Disorders*, *Neurology*, *Archives of Physical Medicine and Rehabilitation*, *Clinical Rehabilitation*, *Physiotherapy*, *Physical Therapy*), abstract books, and conference proceedings (*International Congress of Parkinson's Disease and Movement Disorders*, *World Congress on Parkinson's Disease and Related Disorders*), as well as examined the reference lists of identified papers and other reviews.

Study selection

Studies eligible for this review were randomised controlled trials (including the first phase of crossover trials) of patients with Parkinson's disease comparing a physiotherapy intervention with no intervention or placebo control. Physiotherapy encompasses a wide range of techniques, so we were inclusive in our definition of physiotherapy intervention, including trials of general physiotherapy, exercise, treadmill training, cueing, dance, and martial arts versus no intervention.

General physiotherapy was a broad category, including a variety of techniques traditionally used by physiotherapists to treat people with Parkinson's disease. Trials in this category may include multifaceted interventions using both active participation in treatment by the patient (such as exercise and practising of functional activities) and hands-on techniques delivered by the therapist (for example, massage, passive stretching, the Bobath technique). Exercise interventions were those that included only active exercise participation techniques targeting a variety of symptoms, such as balance, falls prevention, and walking speed. An existing Cochrane review analysed trials of treadmill training for Parkinson's disease, and thus these trials were considered separately to other exercise interventions. Martial arts and dance interventions included participation in universally recognised activities not specifically designed for treating disease, but which appeared in trials of Parkinson's disease evaluating relevant physiotherapy outcome measures. We excluded trials of multidisciplinary team interventions because it was difficult to ascertain the amount of physiotherapy input. Ultimately, all trials were meta-analysed to give an overall picture of the effect of delivery of a physiotherapy intervention versus no physiotherapy intervention.

Data extraction and quality assessment

All articles were read by two independent review authors (CLT, SP, CM, or CPH) and data extracted according to predefined criteria, with any discrepancies resolved by discussion. Publications were assessed for methodological quality by recording specified eligibility criteria, method of randomisation and blinding, concealment of allocation, similarity of patients in treatment groups at baseline, variation in cointerventions received by patients throughout the trial period, whether an intention to treat analysis was performed, and the number of patients lost to follow-up.

Quantitative data synthesis

Outcome data included gait outcomes (such as the two or six min walk test, 10 or 20 m walk test, speed, cadence, stride and step length, freezing of gait questionnaire), functional mobility and balance outcomes (such as the timed up and go test, functional reach test, Berg balance scale, activities specific balance confidence scale), falls data (such as number of falls, falls efficacy scale), clinician rated disability scales (such as the

unified Parkinson's disease rating scale (UPDRS)), patient rated

economics where available.

quality of life (such as Parkinson's disease questionnaire 39), adverse events, compliance or withdrawals, and health n=4, table $1 \downarrow$). Results of each trial were combined using standard meta-analytic methods to estimate an overall effect for physiotherapy versus no intervention. Since all outcomes were continuous variables, weighted mean difference methods were used.²³ Briefly, for each trial, this involved calculating the mean change (and standard deviation) from baseline to the time point after intervention, for both the intervention and no intervention groups. From these numbers, the mean difference and its variance between arms for each trial could be calculated and then combined using a fixed effects model. The primary analysis was a comparison of physiotherapy with

no intervention (control) using change from baseline to the first assessment after treatment (which, in most cases, was immediately after intervention). This comparison was chosen as the primary analysis, because in most trials it was the main data analysis reported, and few trials reported data at assessment points in the longer term (that is, after six months). Some trials also allowed patients in the control group to receive physiotherapy intervention after this point, so this primary analysis allowed a clean comparison of physiotherapy intervention versus no intervention.

Since the different trials implemented various types of physiotherapy, trials were divided according to the type of intervention (general physiotherapy, exercise, treadmill, cueing, dance, or martial arts). If any trials with three or more treatment arms were identified, we made two assumptions for the analysis. Firstly, if the trial was comparing two or more physiotherapy methods in the same category of intervention (as described above) versus control, then the data for those physiotherapy arms were combined to give one comparison of physiotherapy intervention versus control for that trial.

Secondly, if the trial was comparing two or more physiotherapy methods that were in different categories (as described above) versus control, then the data for those physiotherapy arms were kept separate, and the data for that trial were included in the appropriate physiotherapy categories. Therefore, in some cases, the control arms for some trials were included twice in the analysis. However, this related to only a small number of trials and patients, and it was judged that this double inclusion would not overly influence the analysis. We used tests of heterogeneity to make indirect comparisons to investigate whether the treatment effect differed across the different intervention categories.24

Results

Of 76 potentially relevant studies identified, 31 were excluded (for example, studies were not properly randomised, or crossover trials did not report data for the first intervention period) and six were ongoing trials for which no data were available (fig 1 \Downarrow). Therefore, we included 39 randomised controlled trials of 1827 patients in the systematic review (fig 1, web table 1).²⁵⁻⁶³ There were nine trials with multiple arms. $^{\rm 27\ 31\ 37\ 38\ 40\ 45\ 52\ 55\ 60}$ In five trials, two intervention arms were in the same physiotherapy category; therefore, these arms were combined to give one physiotherapy comparison versus no intervention.37 40 52 55 60 In five trials, two intervention arms were in different physiotherapy categories, so the trial contributed data to two physiotherapy comparisons.^{27 31 38 45 60} This meant that these five trials were included multiple times in the analysis, and the control arms

from these trials were counted more than once. Therefore, 39 trials contributed data for 44 comparisons within the six different physiotherapy interventions (physiotherapy n=7, exercise n=14, treadmill training n=8, cueing n=9, dance n=2, and martial arts

Methodological quality

The amount of methodological detail reported in the trials was variable, with several quality indicators not fully discussed in many publications (fig $2\downarrow$, web table 2). Only six (15%) studies^{26 32 33 36 57 59} reported a sample size calculation in the trial report. Less than half of the trials described the randomisation method used, and information on concealment of treatment allocation was also poorly reported (14 (36%)). Blinded assessors were used in 24 (62%) studies (although in one study, the assessors correctly guessed the treatment allocation in nearly 30% of patients³³). Finally, only nine trials stated intention to treat as the primary method of analysis,^{29 32 33 36 39 40 47 57 63} three trials stated per protocol as the primary method of analysis, $^{\rm 34\ 52\ 55}$ and the remaining trials did not describe the method of analysis.

Data available for analysis

Of 13 trials reported in abstract form, five had data available for meta-analysis.³⁴ ³⁷ ³⁹ ⁵¹ ⁵⁵ From the studies with full publications, one trial had relevant data that could not be extracted because it was only available in graph form,⁵⁶ and another trial published only median and interquartile range data, which could not be meta-analysed in this format.³⁰ Therefore, data were not available from ten trials, and data available for meta-analysis was provided by 29 trials.

Effects of intervention

Gait outcomes

Speed was significantly increased with physiotherapy compared with no intervention (mean difference 0.04 m/s, 95% confidence interval 0.02 to 0.06; P<0.001, fig 3U). There were also benefits of borderline significance for the two or six minute walk test and the freezing of gait questionnaire. We saw a greater increase in the distance walked in two or six mins (13.37 m, 0.55 to 26.20; P=0.04, web figure 1) and an improvement in score for the freezing of gait questionnaire (-1.41, -2.63 to -0.19;P=0.02, web figure 2) after physiotherapy. By contrast, we saw borderline significance in favour of no intervention for the time taken to walk 10 or 20 m (0.40 s, 0.00 to 0.80; P=0.05, web figure 3). There was no significant difference between physiotherapy and no intervention for cadence (-1.57 steps/min,-3.81 to 0.67; P=0.17), stride length (0.03 m, -0.02 to 0.08; P=0.24), and step length (0.02 m, 0.00 to 0.04; P=0.06), (table 21).

Functional mobility and balance outcomes

We found significant improvements with physiotherapy for the timed up and go test (-0.63 s, 95% confidence interval -1.05 to -0.21; P=0.003), functional reach test (2.16 cm, 0.89 to 3.43; P<0.001), and Berg balance scale (3.71 points, 2.30 to 5.11; P<0.001); (table 2, figs $4-6\downarrow\downarrow\downarrow\downarrow\downarrow$). There was no difference with physiotherapy compared with no intervention for activity specific balance confidence scale (2.40 points, -2.78 to 7.57;P=0.36; table 2).

In the analysis for the timed up and go test, one trial was heavily weighted in the analysis owing to small standard deviations compared with other studies (fig 4).⁶⁰ Furthermore, in the trial publication, a non-significant effect of martial arts intervention

was reported (P=0.093), but when the data as reported in the paper were included in our analysis, a significant difference was found (P=0.003). We contacted the authors of this study to check whether the data reported in the paper were in fact standard errors, but they were confirmed as standard deviations. We therefore performed a sensitivity analysis, removing this study, and found that the overall result became not significant (-0.38 s, 95% confidence interval -0.96 to 0.21; P=0.21); thus, this result should be interpreted with caution.

Falls

Seven trials collected data for falls using a falls diary, reporting either the number of patients falling or the number of falls per patient.^{33 36 39 50 57 61 62} For both outcomes, there was a decrease in falls after physiotherapy. However, only three studies compared the two treatment groups, with two reporting no difference between the arms,^{50 57} and one reporting a significant difference in the falls efficacy scale between the two treatment arms (-1.91 points, 95% confidence interval -4.76 to 0.94; P=0.19; table 2).

Clinician rated disability on UPDRS

The UPDRS motor score improved with physiotherapy compared with no intervention (-5.01 points, 95% confidence interval -6.30 to -3.72; P<0.001, fig 7 \Downarrow). We also saw significant improvements in the UPDRS subscore for activities of daily living (-1.36 points, -2.41 to -0.30; P=0.01; web figure 4) and total scores with physiotherapy (-6.15, -8.57 to -3.73; P<0.001; web figure 5), but no difference in mental subscore (-0.44, -0.98 to 0.09; P=0.10; table 2).

Patient rated quality of life using Parkinson's disease questionnaire 39

Only data for the mobility domain and summary index of the Parkinson's disease questionnaire 39 were available for meta-analysis. We saw no difference between treatment arms for either overall patient rated quality of life using the summary index (-0.38 points, 95% confidence interval -2.58 to 1.81; P=0.73) or the mobility domain (-1.43, -8.03 to 5.18; P=0.67).

Treatment compliance, adverse events, and health economics

Only 14 trials discussed patient compliance, with 12^{26 29 32 36 37 39:41 47 49 59 63} quantifying it in some form. No trials reported data for health economics, and only one commented on adverse events, stating that none had occurred during treatment sessions.³⁶

Subgroup analysis

Only one outcome, the UPDRS motor subscore, showed significant heterogeneity between the treatment effects of the different classes of intervention. In all other cases, there was no evidence of any differences (table 2). One outlying trial was the cause of this heterogeneity in the motor score;³⁴ when this trial was excluded from the analysis, the result remained significant (-3.77 points, 95% confidence interval -5.15 to -2.39; P<0.001), but the test for between trial and between subgroup heterogeneity was no longer significant (P=0.44 and P=0.08, respectively).

Discussion

A variety of physiotherapy methods are used to treat people with Parkinson's disease. Previous reviews have focused on one type of physiotherapy (for example, exercise, treadmill training).^{19 20} This review brings together all the evidence from the numerous trials evaluating the various physiotherapy methods into one review to assess the overall effect of physiotherapy versus no physiotherapy, and it also allows an indirect comparison of the different physiotherapy methods used.

This review provides evidence on the efficacy of physiotherapy in the short term (mean follow-up <three months), in the treatment of Parkinson's disease. Significant benefit of physiotherapy was reported for nine outcomes: gait speed, two or six min walk test, freezing of gait questionnaire, timed up and go test, functional reach test, Berg balance scale, and UPDRS (total scores, activities of daily living and motor subscores). Although most of the observed differences between the two treatment groups were small, the improvements seen for speed, Berg balance scale, and UPDRS scores were at levels that may be of clinical importance.⁶⁴⁻⁶⁷ By contrast, the only patient rated quality of life outcome measure (Parkinson's disease questionnaire 39) showed no significant benefit from treatment with physiotherapy, and was only reported in eight trials.

The review also highlights the wide range of physiotherapy techniques being used in the treatment of Parkinson's disease. Indirect comparisons provided no evidence of differences in the treatment effect between different types of physiotherapy. With all this in mind and considering the low methodological quality, small size, and short duration of many of the included trials, this evidence supporting the use of physiotherapy for people with Parkinson's disease must be balanced against the lack of long term evidence currently available.

Clinical implications

We saw no evidence of an improvement in patient rated quality of life after physiotherapy, and the observed differences in the nine significant outcomes were relatively small. Their relevance and benefit to patients with Parkinson's disease must be put into context, in terms of what is considered a recommended minimally important difference. Little good quality evidence is available for minimal clinically important differences in these outcome measures. Data for the minimally important difference of walking speed is lacking for people with Parkinson's disease, but a small cohort study of patients with stroke has been reported. Perry and colleagues⁶⁴ reported that an increase in speed of just 0.03 m/s could translate into a change from a limited to an unlimited household walker. The improvement in speed of 0.04 m/s with physiotherapy reported here accords with the findings of the Perry study.⁶⁴

Data for minimally important differences for the two or six min walk test and the freezing of gait questionnaire are also lacking. Therefore, while a 13 m increase in distance walked would probably be considered clinically important, the importance of a 1.4 point improvement in the freezing of gait questionnaire is less clear. Five points has been reported by a small test-retest study as the minimal detectable change on the Berg balance scale.⁶⁵ In this review, we recorded a four point improvement after physiotherapy for this outcome. The minimally important difference for the timed up and go test in Parkinson's disease patients is 11 s,⁶⁵ which is much larger than the 0.6 s improvement observed within this review. Similarly, the 2 cm improvement in the functional reach test seen in this review

was lower than the minimally important difference of 9 cm and 7 cm for forward and backward functional reach tests, respectively.⁶⁵ The small changes observed in this review may, for some outcomes, translate into clinically relevant improvements in a person's functional mobility. A greater evidence base is required to support or refute the clinical significance of these results.

We also observed significant improvements after physiotherapy intervention for clinician rated UPDRS scores (that is, total score, and activities of daily living and motor subscores). The total score improved by six points, the activities of daily living subscore by one point, and motor subscore by five points (table 2). The minimally important differences for the UPDRS have been reported in two studies. One analysed data from two independent randomised controlled trials and concluded that the minimally important difference was eight points for the total score, between two and three points for the activities of daily living subscore, and five points for the motor score.⁶⁶ The second study performed a cross sectional analysis on 653 patients with Parkinson's disease, and reported a minimally important difference of 2.3 to 2.7 points for motor subscore and 4.1 to 4.5 points for total score.⁶⁷ Taking into account the recommendations of both Schrag and colleagues⁶⁶ and Shulman and colleagues,⁶⁷ the improvements observed within this review are approaching or are at these minimally important differences. This similarity suggests that a physiotherapy intervention is beneficial in improving clinician rated motor symptoms and may have a positive effect on activities of daily living. However, we found no effect on patient rated quality of life (measured using the Parkinson's disease questionnaire 39).

Over the past decade, steps have been taken to provide best practice consensus in the form of Dutch guidelines for physical therapy in patients with Parkinson's disease (Koninklijk Nederlands Genootschap voor Fysiotherapie).¹² However, this publication provides a guidance framework rather than a recipe for treatment. Additionally, Parkinson's disease is recognised as a complex condition with an individualised presentation.⁶⁸ For this reason, Morris and colleagues⁶⁹ recognised the importance of the physiotherapist understanding the specific experience of Parkinson's disease in each patient, and advocated that treatment be tailored to fit a person's complaints, lifestyle, and personal interests, as opposed to a "one size fits all" approach. We found no evidence of any differences in the treatment effect between the different physiotherapy interventions. However, data within each physiotherapy intervention were limited, and these comparisons were based on indirect comparisons, which should be interpreted with caution. Therefore, physiotherapy interventions should be compared against each other within rigorous trial designs to determine which, if any, are effective. This analysis could provide therapists with a menu of treatment strategies from which they can devise individualised interventions.

Limitations of the review

The methodological quality and reporting of the majority of trials was variable, and often inadequate. Of 39 trials, only 18 provided information on the randomisation method and only five used a central randomisation procedure to ensure concealment of treatment allocation. Blinded assessors were used in 24 studies, and only nine reported using intention to treat analysis. The lack of information in many reports may not necessarily indicate poor implementation within the trial, but without this information, the level of bias within each trial is difficult to assess. The need for further improvement in the methodological quality of trials in physiotherapy for Parkinson's

disease was noted in another recent systematic review.⁷⁰ Future trials therefore need to ensure that their designs fulfil the requirements of a methodologically sound, large randomised controlled trial, and that the reporting follows the CONSORT guidelines.⁷¹

Furthermore, the trials included in the review were relatively small, and most compared the effect of physiotherapy intervention with no intervention over a short period of time (<three months). Three trials followed up their patients for three months or longer, but all used different follow-up periods, the longest being six months.^{33 37 63} With such limited data, no meaningful meta-analysis could be performed on these longer term data to assess the duration of any improvement after therapy. It is also important to consider the results alongside the possibility of a so-called honeymoon effect⁷² in the period during or just after physiotherapy, which could inflate the treatment effect in favour of physiotherapy. For a long term disease such as Parkinson's disease, the effect of therapy over a much longer period is needed.

Studies included in the review all used standard physiotherapy and Parkinson's disease outcomes. However, Parkinson's disease is a multidimensional disease, and many important outcomes were either poorly or not reported. This included data for quality of life, the number of falls, depression and anxiety, adverse events, and the health of the carer supporting the person with Parkinson's disease. We saw little focus on patient orientated outcomes, without which studies cannot necessarily capture the difficulties experienced by patients in everyday life or their opinions on treatment acceptability and personal improvements. Patient reported outcomes such as the walk 12G scale, which has been shown to have validity in Parkinson's disease,73 should be more commonly used in trials of physiotherapy to increase the effect of the studies themselves and of subsequent meta-analyses. Furthermore, none of the reports included a health economics analysis of the physiotherapy intervention studied, and therefore, little is known about the cost effectiveness and economic value of these various therapies. Implementation of a community based professional network of physiotherapists working according to evidence based recommendations has been shown to reduce costs of provision of healthcare compared with usual care.⁷⁴ Unfortunately, the evidence base, required to inform the types of techniques which should be recommended for use in these networks, is inconclusive.

Outcome reporting bias may have created a deceptively positive impression of the effectiveness of the studied interventions. Unfortunately, the proportion of outcomes that went unreported could not be assessed here, owing to a lack of information on trial protocol.

Implications for research

A larger and better quality body of evidence is required before a recommendation for change in practice can be made. The majority of the studies in this review were small and had a short follow-up period. Larger randomised controlled trials are needed, particularly those focusing on improving trial methodology and reporting. Rigorous methods of randomisation should be used and the allocation of treatment be adequately concealed. Data should be analysed according to intention to treat principles, and trials should be reported according to CONSORT guidelines.⁷¹ This review also illustrates the need for the universal use of relevant, reliable, and sensitive outcome measures. Additionally, only three trials looked at the benefit of physiotherapy intervention in the longer term. To assess whether or how long any improvement owing to physiotherapy intervention may last, long term follow-up should be performed without crossover from control to active intervention. Moreover, this review highlights the variety of physiotherapy interventions being used in the treatment of Parkinson's disease. More specific trials with improved treatment strategies are needed to underpin the most appropriate choice of physiotherapy intervention.

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This paper is based on a Cochrane review by the same authors. Cochrane reviews are regularly updated as new evidence emerges and in response to comments and criticisms. This work has been undertaken to update the physiotherapy versus placebo or no intervention in Parkinson's disease first published in 2001. The Cochrane Library should be consulted for the most recent version of the review.

Contributors: CEC, RS, CS, KW, and NI contributed to the design of the protocol. CLT, RS, LS, and CPH designed and implemented the searches. CLT, RS, CM, SP, and CPH selected the studies. CLT, SP, CM, and CPH undertook data extraction and assessment of risk of bias. CM, who throughout the review and analysis period was a research physiotherapist, provided expertise on technical aspects of the project as necessary. CLT, SP, NI, and CPH were involved in the data analysis. All authors were involved in interpretation of the review. NI is the study guarantor.

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Data sharing: No additional data available.

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What is already known this on this topic

Referral rates for physiotherapy in Parkinson's disease are historically low in the UK

Evidence from published trials and guidelines have suggested potential benefits of physiotherapy for patients with Parkinson's disease

What this study adds

A variety of physiotherapy methods currently exist for treating Parkinson's disease

Physiotherapy could provide clinically meaningful benefits in the short term for patients, although many relevant trials have been of low methodological quality, small size, and short duration. Indirect comparisons indicate little difference in treatment effect between interventions

It is uncertain whether physiotherapy is beneficial in the longer term, and if so, which type of physiotherapy is best to deliver

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Tables

					Mean			
Physiotherapy intervention	Number of trials	Total number of participants (% male)	Mean age (years)	Mean stage on Hoehn and Yahr scale	duration of Parkinson's disease (years)	Duration of treatment sessions (min)	Duration of trial period	Examples of types of therapy
Physiotherapy	7 ^{2526,27,28,29,30,31}	244 (69)	65	2.4	4	30-60	4-12 months	Bobath training; gait and balance exercises; hands-on techniques; education and advice on transfer, posture, physical fitness
Exercise	14 ^{32,33,24,35,36,37,38,39,40,41,42,43,44,45}	769 (60)	69	2.6	6	30-120	3-24 weeks	Strengthening and balance training; walking; falls prevention; neuromuscular facilitation; resistance exercise; aerobic training; education; relaxing techniques
Treadmill	8 ^{27,31,46,47,48,49,50,51}	179 (61)	68	2.4	5	30-60	4-8 weeks	Walking on treadmill with speed or incline adjustments; body weight supported treadmill training; step and gait training
Cueing	g ^{38,45,52,53,54,55,56,57,58}	371 (59)	67	2.6	7	4-30	Single session of 13 weeks	Three types of cueing: audio (music, spoken instructions); visual (computer images); sensory (vibration)
Dance	2 ^{59,60}	120 (63)	69	2.3	7	60	12-13 weeks	Tango; waltz; foxtrot
Martial arts	4 ^{60,61,62,63}	143 (74)	65	2.1	6	60	12-24 weeks	Tai chi; gigong

For one trial,³¹ 60 patients split between physiotherapy and treadmill categories not included in table because group split not given. For multiple arm trials, which are included in more than one intervention type, control arm patients (n=59) were counted twice. Web table 1 provides detailed information on individual trials.

Outcomes	No of trials; no of comparisons	Physiotherapy interventions	No of participants	Mean difference (95% CI)	Heterogeneity	Subarour
Gait	compansons	r nysiotherapy interventions	participants	Mean unerence (35 % Ci)	Theterogeneity	Subgroup
Speed (m/s)	15; 19 ^{25,26,27,32,34,38,40,45,47,50,52,53,55,57,60}	General physiotherapy, exercise, treadmill, cueing, dance, martial arts	814	0.04 (0.02 to 0.06), P<0.001	P=0.55	P=0.25
2 or 6 min walk test (m)	6; 7 ^{39,41,42,47,59,60}	Exercise, treadmill, dance, martial arts	242	13.37 (0.55 to 26.20), P=0.04	P=0.44	P=0.19
10 or 20 m walk test (s)	4; 4 ^{39,41,43,49}	Exercise, treadmill	169	0.40 (0.00 to 0.80), P=0.05	P=0.19	P=0.51
Freezing of gait questionnaire	4 ; 4 ^{32,57,59,60}	Exercise, cueing, dance	298	–1.41 (–2.63 to –0.19), P=0.02	P=0.74	P=0.55
Cadence (steps/min)	7; 9 ^{27,40,45,50,53,55,57}	General physiotherapy, exercise, treadmill, cueing	350	-1.57 (-3.81 to 0.67), P=0.17	P=0.73	P=0.97
Stride length (m)	6; 9 ^{27,45,50,53,55,60}	General physiotherapy, exercise, treadmill, cueing, dance, martial arts	225	0.03 (-0.02 to 0.08), P=0.24	P=0.33	P=0.23
Step length (m)	5; 6 ^{27,34,40,52,57}	General physiotherapy, exercise, treadmill, cueing	383	0.02 (0.00 to 0.04), P=0.06	P=0.71	P=0.47
Functional mobility a	nd balance					
Timed up and go test (s)	9; 10 ^{34,36,37,40,42,43,52,57,60}	Exercise, cueing, dance, martial arts	639	-0.63 (-1.05 to -0.21), P=0.003	P=0.12	P=0.33
Functional reach test (cm)	4; 4 ^{33,41,43,57}	Exercise, cueing	393	2.16 (0.89 to 3.43), P<0.001	P=0.15	P=0.48
Berg balance scale	5; 6 ^{33,36,44,46,60}	Exercise, treadmill, dance, martial arts	385	3.71 (2.30 to 5.11), P<0.001	P=0.06	P=0.47
Activity specific balance confidence	3 ; 3 ^{37,42,58}	Exercise, cueing	66	2.40 (-2.78 to 7.57), P=0.36	P=0.61	P=0.32
Falls						
Falls efficacy scale	4;4 ^{32,36,46,57}	Exercise, cueing	353	-1.91 (-4.76 to 0.94), P=0.19	P=0.44	P=0.28
Clinician rated disability	(UPDRS)					
Total	3; 4 ^{26,27,34}	General physiotherapy, exercise, treadmill	207	-6.15 (-8.57 to -3.73), P<0.001	P=0.03	P=0.01
Subscores						
Mental	2; 3 ^{26,27}	General physiotherapy, treadmill	105	-0.44 (-0.98 to 0.09), P=0.10	P=0.80	P=0.82
Activities of daily living	3; 4 ^{26,27,59}	General physiotherapy, treadmill, dance	157	–1.36 (–2.41 to –0.30), P=0.01	P=0.28	P=0.19
Motor	12; 14 ^{25,26,27,34,40,47,52,53,58,59,60,63}	General physiotherapy, exercise, treadmill, cueing, dance, martial arts	593	–5.01 (–6.30 to –3.72), P<0.001	P<0.001	P<0.001
Patient rated quality of	life (Parkinson's disease ques	tionnaire 39)				
Summary index	7; 8 ^{25,32,37,39,47,57,60}	General physiotherapy, exercise, treadmill, cueing, dance, martial arts	405	-0.38 (-2.58 to 1.81), P=0.73	P=0.89	P=0.87
Mobility subscore	2; 3 ^{29,60}	General physiotherapy, dance, martial arts	105	-1.43 (-8.03 to 5.18), P=0.67	P=0.11	P=0.11

Table 2| Summary of results

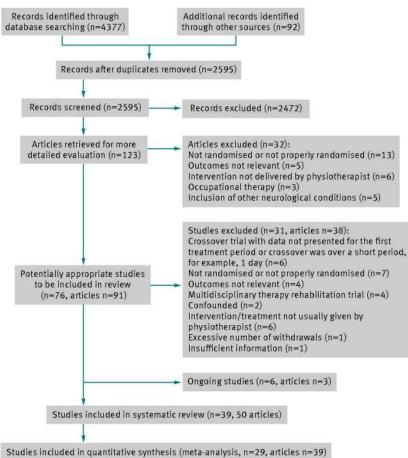


Fig 1 Trial flow diagram to summarise the stages of systematic review

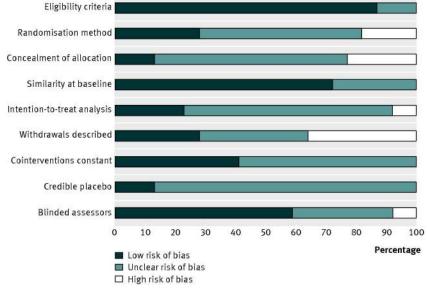


Fig 2 Review authors' judgments about each risk of bias item, presented as percentage across all included studies

	Interve	ntion		No inter	vention			
Study	Mean	Standard deviation	Total	Mean	Standard deviation	Total	Weight (%)	Mean difference (95% Cl)
General physiothe	erapy v co	ntrol						
Chandler 1999	0.13	0.36	26	0.10	0.30	26	1.5	0.03 (-0.15 to 0.21)
Ellis 2005	0.16	0.22	32	0.01	0.21	33	4.5	0.15 (0.05 to 0.25)
Fisher 2008	0.02	0.18	10	0.02	0.17	10	2.1	0.00 (-0.15 to 0.15)
Subtotal			68			69	8.0	0.09 (0.01 to 0.17)
Heterogeneity	x ² =3	.01, df=2 (P=	0.22); l ² =3	4%				
Test for overall eff	ect Z=2.	23 (P=0.03)						
Exercise v control								
Allen 2010	0.02	0.27	21	0.02	0.29	24	1.8	0.00 (-0.16 to 0.16)
Boehm 2011	0.01	0.24	50	-0.002	0.23	52	5.8	0.01 (-0.08 to 0.10)
Mak 2008	0.02	0.08	19	0	0.06	14	21.4	0.02 (-0.03 to 0.07)
Sage 2009a	0.06	0.20	31	-0.004	0.22	15	2.8	0.06 (-0.07 to 0.20)
Thaut 1996	0.07	0.18	11	-0.05	0.27	11	1.3	0.12 (-0.07 to 0.31)
Subtotal			132			116	33.2	0.03 (-0.01 to 0.06)
Heterogeneity	x ² =1	.49, df=4 (P=	0.83); l ² =0	1%				
Test for overall eff			anterstiller – Sil					
Treadmill v contro								
Canning 2008	0.09	0.12	9	0.06	0.09	9	5.1	0.03 (-0.07 to 0.13)
Fisher 2008	0.06	0.20	10	0.00	0.17	10	1.8	0.04 (-0.12 to 0.20)
Protas 2005	0.17	0.35	9	0.01	0.23	9	0.7	0.16 (-0.11 to 0.43)
Subtotal			28			28	7.6	0.04 (-0.04 to 0.12)
Heterogeneity	x2=0	.77, df=2 (P=		1%		10.00	100000	
Test for overall eff		것이야지? 이 다시겠어? 이						
Cueing v control		,,						
and The contract	0.06	0.19	28	0.01	0.27	14	2.0	0.05 (-0.11 to 0.21)
Almeida 2012	0.08	0.19	11	-0.02	0.27	14	1.8	0.05 (-0.11 to 0.21)
de Bruín 2010a	-0.05	0.22	17	-0.02 0.048	0.17	6	1.8	
Haase 2011	0.05	0.26	17	0.048	0.20	14	28.4	-0.10 (-0.30 to 0.10) 0.05 (0.01 to 0.09)
Mak 2008	0.05	0.16	76	0.02	0.08	77	12.4	0.06 (0.00 to 0.12)
Nieuwboer 2007 Thaut 1996	0.08	0.10	15	-0.05	0.23	11	1.3	0.21 (0.02 to 0.40)
Subtotal	0.10	0.22	166	-0.05	0.27	133	47.0	0.05 (0.02 to 0.09)
Heterogeneity	v2_A	.71, df=5 (P=		1%			47.0	0.02 (0.02 (0 0.03)
Test for overall eff				0.0				
	LLL <u>2</u> =3.	24 (F = 0.001)						
Dance v control								
Hackney 2009	0.05	0.20	31	0.02	0.38	17	1.3	0.03 (-0.16 to 0.22)
Subtotal	100000000000000000000000000000000000000		31			17	1.3	0.03 (-0.16 to 0.22)
Heterogeneity		pplicable						
Test for overall eff	ect Z=0.	30 (P=0.76)						
Martial arts v con	trol							
Hackney 2009	0.01	0.21	13	0.10	0.11	13	2.9	-0.09 (-0.22 to 0.04)
Subtotal			13			13	2.9	-0.09 (-0.22 to 0.04)
Heterogeneity	Not a	pplicable						
Test for overall eff	ect Z=1.	37 (P =0.1 7)						
Total			438			376	100	0.04 (0.02 to 0.06)
Heterogeneity	x ² =1	6.63, df=18 (P=0.55);	2=0%				
Test for overall eff								
Test for subgroup				=0.25); 12=2	24.7%			

Test for subgroup differences x'=6.64, df=5 (P=0.25); |'=24.7%

Fig 3 Comparison of physiotherapy interventions in relation to speed (m/s). Studies denoted as a or b distinguishes those published by the same first author and in the same year

	Interve	ntion		No inte	rvention				
Study	Mean	Standard deviation	Total	Mean	Standard deviation	Total	Weight (%)	Mean difference (95% Cl)	
Exercise v control									
Boehm 2011	-0.40	5.20	50	-0.20	3.90	52	5.5	-0.20 (-1.99 to 1.59)	
Goodwin 2009	0.13	11.09	61	-0.48	13.92	62	0.9	0.61 (-3.83 to 5.05)	
Klassen 2007	-1.30	2.50	17	-0.20	1.85	6	4.9	-1.10 (-3.00 to 0.80)	
Sage 2009a	-0.60	2.21	31	0	2.33	15	8,8	-0.60 (-2.01 to 0.81)	
Schilling 2008	-0.10	0.70	8	-0.75	1.20	7	17.2	0.65 (-0.36 to 1.66)	
Stozek 2003	-2.36	2.63	30	1.10	7.15	31	2.4	-3.46 (-6.15 to -0.77)	
Subtotal			197			173	39.7	-0.21 (-0.88 to 0.45)	•
Heterogeneity	x ² =9	.66, df=5 (P=	0.09); ² =4	8%					
Test for overall eff	ect Z=0.	63 (P=0.53)							
Cueing v control									
Almeida 2012	-1.45	3.20	28	0.10	3.16	14	4.2	-1.55 (-3.59 to 0.49)	
Nieuwboer 2007	-1.59	4.59	76	-1.34	5.78	77	6.4	-0.25 (-1.90 to 1.40)	
Subtotal			104			91	10.7	-0.77 (-2.05 to 0.52)	
Heterogeneity	x2=0	.94, df=1 (P=	0.33); ²=0	1%					
Test for overall eff	ect Z=1.	17 (P=0.24)							
Dance v control									
Hackney 2009	-1.10	4.31	31	2.00	9.28	17	0.8	-3.10 (-7.76 to 1.56)	
Subtotal			31			17	0.8	-3.10 (-7.76 to 1.56)	
Heterogeneity	Not a	pplicable							
Test for overall eff	ect Z=1.	30 (P=0.19)							
Martial arts v con	trol								
Hackney 2009	-1.00	0.10	13	-0.10	1.10	13	48.8	-0.90 (-1.50 to -0.30)	
Subtotal			13			13	48.8	-0.90 (-1.50 to -0.30)	•
Heterogeneity	Not a	pplicable							•
Test for overall eff		94 (P=0.003)							
Total			345			294	100	-0.63 (-1.05 to -0.21)	•
Heterogeneity	x ² =1	4.00, df=9 (P		=36%		075,500	767176		
Test for overall eff		18 83							10 -5 0 5
Test for subgroup				=0.33); '=	11.6%				Favours intervention Favours no interv

Fig 4 Comparison of physiotherapy interventions with controls in relation to the timed up and go test (s). Studies denoted as a or b distinguishes those published by the same first author and in the same year

	Interve	ntion		No inte	rvention								
Study	Mean	Standard deviation	Total	Меап	Standard deviation	Total	Weight (%)	Mean difference (95% Cl)					
Exercise v control													
Ashburn 2007	0.40	6.56	67	-1.00	7.00	66	30.3	1.40 (-0.91 to 3.71)				_	
Schenkman 1998	1.57	4.45	23	-0.28	4.17	23	26.0	1.85 (-0.64 to 4.34)					
Stozek 2003	5.80	6.01	30	-0.03	7.38	31	14.2	5.83 (2.46 to 9.20)					
Subtotal			120			120	70.5	2.46 (0.94 to 3.97)					
Heterogeneity	x ² =4	.88, df=2 (P=	0.09); l ² =5	9%									
Test for overall effe	ct Z=3.	18 (P=0.001)											
Cueing v control													
Nieuwboer 2007	1.80	5.28	76	0.34	9.02	77	29.5	1.46 (-0.88 to 3.80)				_	
Subtotal			76			77	29.5	1.46 (-0.88 to 3.80)				-	
Heterogeneity	Not a	applicable											
Test for overall effe	ct Z=1.	22 (P=0.22)											
Total			196			197	100.0	2.16 (0.89 to 3.43)					
Heterogeneity	x ² =5	.37, df=3 (P=	0.15); l ² =4	4%					-10	-5	o	5	10
Test for overall effe	ct Z=3.	34 (P=0.0008	1)						Favour	s no interv	ention Fav	ours intervo	ention
Test for subgroup o	ifference	x2=0 /	9 df=1 (P=	=0 (8) · 12=0	1%								

Test for subgroup differences x²=0.49, df=1 (P=0.48); l²=0%

Fig 5 Comparison of physiotherapy interventions with controls in relation to the functional teach test (cm)

	Interve	ntion		No inte	rvention					
Study	Mean	Standard deviation	Total	Mean	Standard deviation	Total	Weight (%)	Mean difference (95% Cl)		
Exercise v control	L									
Ashbum 2007	1.50	9.51	67	1.60	10.21	66	17.5	-0.10 (-3.45 to 3.25)		
Goodwin 2009	3.10	11.07	61	-0.55	9.88	62	14.3	3.65 (-0.06 to 7.36)		
Taheri 2011	8.40	9.70	12	-1.08	3.70	12	5.7	9.48 (3.61 to 15.35)		
Subtotal			140			140	37.6	2.79 (0.50 to 5.08)	•	
Heterogeneity	x2=8	.04, df=2 (P=0	0.02); 12=7	5%						
Test for overall eff	ect Z=2.	38 (P=0.02)								
Treadmill v contro	ol									
Cakit 2007	7.09	8.50	21	-1.20	10.07	10	3.8	8.29 (1.07 to 15.51)		
Subtotal			21			10	3.8	8.29 (1.07 to 15.51)		
Heterogeneity	Not a	pplicable								
Test for overall eff	ect Z=2.	25 (P=0.02)								
Dance v control										
Hackney 2009	3.95	4.70	31	-1.20	9.32	17	8.8	5.15 (0.42 to 9.88)		÷.
Subtotal			31			17	8.8	5.15 (0.42 to 9.88)		
Heterogeneity	Not a	pplicable								
Test for overall eff										
Martial arts v con	trol									
Hackney 2009	3.30	3.00	13	-0.50	2.10	13	49.8	3.80 (1.81 to 5.79)		
Subtotal	(T. (T. (T.)		13	10.71	10000	13	49.8	3.80 (1.81 to 5.79)		
Heterogeneity	Not a	pplicable								
Test for overall eff)							
Total			205			180	100.0	3.71 (2.30 to 5.11)		
Heterogeneity	x ² =1	0.57, df=5 (P=		53%				80 - 1 - 1 - 1 - 1 - 1 - 1 - 1 - 1 - 1 - 1	-20 -10 0	10 20
Test for overall eff			- 32	<i>,,,,</i> ,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,					Favours no intervention Favours i	727-1 X727
Test for subgroup		0-00 -	2012 St. 51	=0.47); 2=0	0%					

Fig 6 Comparison of physiotherapy interventions with controls in relation to the Berg balance scale. Studies denoted as a or b distinguishes those published by the same first author and in the same year

	Interver	ntion		No inte	vention				
Study	Mean	Standard deviation	Total	Mean	Standard deviation	Total	Weight (%)	Mean difference (95% Cl)	
General physiothe	erapy v co	ntrol							
Chandler 1999	-1.00	7.00	26	3.00	6.24	26	12.7	-4.00 (-7.60 to -0.40)	
Ellis 2005	-3.00	6.60	32	-0.20	5.30	33	19.5	-2.80 (-5.72 to 0.12)	
Fisher 2008	-3.80	8.17	10	-2.70	8.15	10	3.2	-1.10 (-8.25 to 6.05)	
Subtotal			68			69	35.5	-3.08 (-5.24 to -0.92)	•
Heterogeneity	x ² =0.	58, df=2 (P=	0.75); l²=0	%					
est for overall eff	ect Z=2.	79 (P=0.005)							
xercise v control									
Boehm 2011	-7.00	8.60	50	6.00	9.50	52	13.4	-13.00 (-16.51 to -9.49)	
5age 2009a	-3.70	6.72	31	1.20	7.81	15	7.8	-4.90 (-9.51 to -0.29)	
Subtotal			81			67	21.2	-10.02 (-12.81 to -7.23)	•
leterogeneity	x ² =7.	51, df-1 (P-	0.006); 12-	87%					
Test for overall eff									
readmill v contro	ł		1.1252						
Canning 2008	2.40	5.10	9	2.30	4.90	9	7.8	0.10 (-4.52 to 4.72)	
Fisher 2008	-2.80	9.72	10	-2.70	8.15	10	2.7	-0.10 (-7.96 to 7.76)	
Subtotal			19			19	10.4	0.05 (-3.93 to 4.03)	-
leterogeneity	x ² =0.	00, df=1 (P=	0.97): l ² =0	%					
est for overall eff				2220					
Cueing v control									
Almeida 2012	1.40	8.57	28	2.10	9.28	14	4.9	-0.70 (-6.51 to 5.11)	
de Bruin 2010a	-5.60	9.17	11	-1.80	6.53	11	3.7	-3.80 (-10.45 to 2.85)	
Shankar 2008	-4.00	8.29	14	1.21	7.75	14	4.7	-5.21 (-11.15 to 0.73)	
Subtotal		101000	53		1000000	39	13.3	-3.15 (-6.68 to 0.37)	-
leterogeneity	x ² =1	18, df=2 (P=	$0.55 \cdot l^2 = 0$	%					
est for overall eff			0.55), 1 0						
Dance v control									
Duncan 2012	-12.70	12.10	26	-3.00	9.40	26	4.8	-9.70 (-15.59 to -3.81)	
Hackney 2009	-2.10	10.96	31	5.00	10.33	17	4.2	-7.10 (-13.34 to -0.86)	
Subtotal	2.10	0.000000	57	3.00	10.55	43	9.0	-8.48 (-12.76 to -4.19)	
Heterogeneity	v2- 0	.35, df = 1 (P		- 0%			9.0	0.40(12.7010 4.17)	
Fest for overall eff				-076					
Martial arts v cont		.00 (7 - 0.00	•1)						
	-1.50	6.60	13	4.30	5.60	13	7.5	-5.80 (-10.51 to -1.09)	
Hackney 2009 Schmitz-Hubsch 200		10.90	31	5.54	14.77	21	3.0	-5.86 (-13.25 to 1.53)	
Schmitz-Hubsch 200 Subtotal	0.32	10.90	31 44	2.24	14.//	34	3.0 10.5	-5.86 (-13.25 to 1.53)	
	~ ~	00 df 1 /D		0/		24	10.5	-3.62 (-9.79 10 -1.85)	
Heterogeneity		00, df=1 (P=	S10930	70					
est for overall eff	ect Z=2.0	57 (P =0.0 04)	322			271	100.0	E 01 (6 20 to 2 72)	
Total						2/1	100.0	-5.01 (-6.30 to -3.72)	•
Heterogeneity		4.99, df=13 (;1′=63%					-20 -10 0 10
ect for overall eff.	ect Z=7.0	53 (P(0.0000)	1)						Favours intervention Favours no intervention

Fig 7 Comparison of physiotherapy interventions with controls in relation to the UPDRS motor subscale. Studies denoted as a or b distinguishes those published by the same first author and in the same year