Clinical Rehabilitation

The effect of relaxation therapy on autonomic functioning, symptoms and daily functioning, in patients with chronic fatigue syndrome or fibromyalgia: a systematic review Mira Meeus, Jo Nijs, Tanja Vanderheiden, Isabel Baert, Filip Descheemaeker and Filip Struyf *Clin Rehabil* published online 8 September 2014 DOI: 10.1177/0269215514542635

The online version of this article can be found at: http://cre.sagepub.com/content/early/2014/09/04/0269215514542635

Published by: SAGE http://www.sagepublications.com

Additional services and information for Clinical Rehabilitation can be found at:

Email Alerts: http://cre.sagepub.com/cgi/alerts

Subscriptions: http://cre.sagepub.com/subscriptions

Reprints: http://www.sagepub.com/journalsReprints.nav

Permissions: http://www.sagepub.com/journalsPermissions.nav

>> OnlineFirst Version of Record - Sep 8, 2014

What is This?

Article

CLINICAL REHABILITATION

The effect of relaxation therapy on autonomic functioning, symptoms and daily functioning, in patients with chronic fatigue syndrome or fibromyalgia: a systematic review Clinical Rehabilitation I-13 © The Author(s) 2014 Reprints and permissions: sagepub.co.uk/journalsPermissions.nav DOI: 10.1177/0269215514542635 cre.sagepub.com



Mira Meeus¹⁻³, Jo Nijs^{3,4}, Tanja Vanderheiden¹, Isabel Baert^{1,3}, Filip Descheemaeker² and Filip Struyf^{1,3}

Abstract

Objective: To establish the effects of relaxation therapy on autonomic function, pain, fatigue and daily functioning in patients with chronic fatigue syndrome or fibromyalgia.

Method: A systematic literature study was performed. Using specific keywords related to fibromyalgia or chronic fatigue syndrome and relaxation therapy, the electronic databases PubMed and Web of Science were searched. Included articles were assessed for their risk of bias and relevant information regarding relaxation was extracted. The review was conducted and reported according to the PRISMA-statement. **Results:** Thirteen randomized clinical trials of sufficient quality were included, resulting in a total of 650 fibromyalgia patients (11 studies) and 88 chronic fatigue syndrome patients (3 studies). None of the studies reported effects on autonomic function. Six studies reported the effect of guided imagery on pain and daily functioning in fibromyalgia. The acute effect of a single session of guided imagery was studied in two studies and seems beneficial for pain relief. For other relaxation techniques (eg. muscle relaxation, autogenic training) no conclusive evidence was found for the effect on pain and functioning in fibromyalgia patients comparison to multimodal treatment programs. For fatigue a multimodal approach seemed better than relaxation, as shown in the sole three studies on chronic fatigue syndrome patients. **Conclusion:** There is moderate evidence for the acute effect of guided imagery on pain, although the content of the visualization is a matter of debate. Other relaxation formats and the effects on functionality and autonomic function require further study.

³Pain in Motion Research Group, Belgium

⁴Departments of Human Physiology and Rehabilitation Sciences, Faculty of Physical Education and Physiotherapy, Vrije Universiteit Brussel, Belgium

Corresponding author:

Mira Meeus, Rehabilitation Sciences and Physiotherapy Ghent Campus Heymans (UZ) 3 B3, De Pintelaan 185, Ghent, Belgium. Email: mira.meeus@ugent.be

Department of Rehabilitation Sciences and Physiotherapy, Faculty of Medicine and Health Sciences, University of Antwerp, Belgium

²Department of Rehabilitation Sciences and Physiotherapy, Ghent University and Artevelde University College, Ghent, Belgium

Keywords

Chronic fatigue syndrome, fibromyalgia, relaxation, visualization, imagery, autogenic training, pain, fatigue, autonomic functioning, daily functioning

Received: 13 September 2013; accepted: 14 June 2014

Introduction

Fibromyalgia and chronic fatigue syndrome fall into the spectrum of what might be termed functional disorders or stress-associated (or stress intolerance)^{1,2} syndromes by virtue of frequent onset after acute or chronic stressors and apparent exacerbation of symptoms during periods of physical or emotional stress.³

Consequently, much interest has been expressed in the possible role of the autonomic nervous system in the pathogenesis of fibromyalgia or chronic fatigue syndrome. Many of the common symptoms could be attributed to a dysfunction of the autonomic nervous system.⁴ In fact, many studies strongly support the notion that autonomic dysregulation is frequent in fibromyalgia or chronic fatigue syndrome.^{5–9}

This fits in the observations that both disorders appear to be preceded by (childhood) trauma, long periods of stress or a life event, suggesting that the stress may act by inducing a self-perpetuating vicious cycle.10 It seems that the illness onset might be facilitated by a shift within the stress system from chronic hyperfunction to hypofunction, implying an inability to adequately respond to new stressors and, eventually, giving rise to long-term disturbances in stress-regulating, pain-processing and immune mechanisms.11,12 Autonomic dysfunction may thus explain the diverse clinical manifesof chronic fatigue tations syndrome or fibromyalgia.4

In consequence, managing stress should be a rational therapy modality in the multidisciplinary and biopsychosocial approach of these patients.^{1,13} Obviously, relaxation therapy will be integrated in most rehabilitation approaches for fibromyalgia or chronic fatigue syndrome. In despite of its widespread use for the management of fibromyalgia or chronic fatigue syndrome, studies examining the

effectiveness for relaxation therapy in fibromyalgia or chronic fatigue syndrome have not been reviewed systematically. This lacuna makes it difficult for clinicians to apply evidence to practice, as many different kinds of relaxation therapy are available. In addition, it is unclear whether relaxation therapy influences autonomic function in patients with fibromyalgia or chronic fatigue syndrome. Is relaxation therapy capable of restoring the homeostasis of the stress response system, and will patients cope better with daily stressors by applying relaxation techniques? Does relaxation therapy avoids further or future overload of the stress response system in patients with chronic fatigue syndrome or fibromyalgia? And finally, does relaxation therapy leads to improvements in fatigue, pain and daily functioning in patients with chronic fatigue syndrome or fibromyalgia?

This study investigates whether relaxation therapy is beneficial for patients with chronic fatigue syndrome or fibromyalgia.

Methods

This systematic review is reported following the PRISMA-guidelines (Preferred Reporting Items for Systematic reviews and Meta-Analyses).¹⁴

To identify relevant articles PubMed and Web Of Science were searched until May 2014. The search strategy was based on a combination of search terms related to "chronic fatigue syndrome" or "fibromyalgia" combined with terms related to "relaxation". The construct of the search strategy is presented in Supplementary Table 1.

To be included in the present systematic review, articles had to report the results of randomized controlled trials evaluating relaxation for patients with fibromyalgia or chronic fatigue syndrome on

	ב רמחובי					
Study	Sample	Design>	Experimental intervention (EI)	Control intervention (CI)	Outcome	Results
Guided imagery/visualization (FM)	ualization (FN	()				
Fors and Götestam, 2000 ²¹	61 ♀ FM (ACR) 45,7 ± 10y D0: 3.	T0 Ix I Tpost.	Ell: 1 × 30° audio tape guided imagery (visualization: pleasant nature images). El2: 1 × 30° audio tape patient education (visualization: natural pain-killing system).	l × 30' free talking about FM.	VAS pain.	EII & EI2: ∖ ≮ CI: =.
Fors et al., 2002 ²⁰	61 ♀ FM (ACR) 45,7 ± 10y D0: 6.	T0 4w I daily T.	EII: Daily × 30' audio tape guided imagery (visualization: pleasant nature images). E12: Daily × 30' audio tape patient education (visualization: natural pain-killing system).		VAS pain in daily diary.	EII:N 4w slope ≠. EI2 & CI:N 4w slope.
Castel et al., 2007 ²⁶	45 FM (ACR) 43,7 ± 8,6y 39 ♀ D0: /.	T0 I× I Tpost.	EII: Hypnosis (10') with relaxation (20': visualization pleasant beach). E12: Hypnosis (10') with analgesia suggestion (20': visualization blue liquid analgesic stream soothing pain).	Relaxation: 5' demo of relaxing body parts, 10' focus on diaphragmatic breathing, 5' suggestion of well-being.	VAS pain; MPQ.	Overall VAS, MPQ : EII = CI EI2 > EII, CI.
Menzies et al., 2006 ²⁵	48 FM 49,6 ± 10,53y 47 ♀ D0: /.	T0 6w I - T mid 4w I - T post.	 audio tapes with 20' guided imagery. basic script: familiarity with GI. pleasant scene imagery. pain related (2006) or journey in immune system (2014). w I-6: each tape used at least daily for 2w. 	Usual care.	MPQ FIQ.	MPQ = (El = Cl) FIQ' (El>Cl).
Menzies et al., 2014 ²⁷	72 ⊹ FM 46,9 ± 12,8y D0: 8.		w7-10: chosen tape at least daily.		BPI BFI.	BPI'N (El > CI) BFI'N (⇔CI'N).
Verkaik et al., 2014	70 FM (ACR) 47,4 ± 11,4y 69 ♀ D0: 17.	T 0 4w I T mid FU 6w.	2x 1,5h (4w interval) group guided imagery session (instructions and discussion guided imagery and exercises; pleasant and pain related) + CD with guided imagery exercise 2x/d during 4 w.	2x 1,5h (4w interval) group discussion 2x 1,5h (4w interval) group discussion.	Daily VAS FIQ.	FIQ, VAS = (EI =CI).

(Continued)

	(הפר					
Study	Sample	Design>	Experimental intervention (EI)	Control intervention (CI)	Outcome	Results
Ost applied relaxation (CFS)	tion (CFS)					
Deale et al., 1997 ¹⁹	60 CFS (CDC) 31,9 ± 9y 41 ♀ DO: 7.	T0 4-6m l T mid, Post FU l, 3,	13 sessions CBT (graded activity, pacing, sleep hygiene, cognitive strategies, etc).	Ost applied relaxation: PMR and visualisation, fast relaxation; taught during 13 treatment sessions, and practiced 2x/d.	SF-36: physical functioning. Fatigue questionnaire.	6m: SF-36N El = Cl FatigueN > + ourcome
						- outcome (SF-36) El > CI.
Deale et al., 2001 ¹⁸	60 CFS (CDC) 41 4 +	T0 4-6m FI J 6m				6m: + outcome (SF-36N &
	10,4y 36 ⊹0 D0: 7.	5y.				fatigueN): El > Cl 5v: El = Cl.
Thomas et al., 2008 ²⁸	40CFS (CDC) 28 년	T0 10w l T ₂₀₀₅ t	10x 1h individual session. Multiconvergent therapy (CBT + GET, pacing, mindfulness, etc.) (n=17, 48 ± 8,03y).	CII: 10x 1h Ost applied relaxation techniques) $(n=14, 45 \pm 12,56y)$. C12: control $(n=9, 46 + 11,04x)$	Karnofsky performance, Improvement	mproved patients El >
	DO: 5.	FU 6m.			in fatigue and disability.	(T post & FU 6m); Means =
(Progressive) Muscle Relaxation (PMR) (FM)	e Relaxation	(PMR) (FM	()			
Field et al.,2002 ²³	20 FM (ACR) 50,9y D0:/.	T0 5w I T post.	2x/w 30' massage therapy (Swedish and Shiatsu).	2x/w 30' PMR instructions.	10-point Likert pain and fatigue; Tender points and pain	Overall⊻, but El>CI.
					assessment (algometry).	
Buckelew et al.,I 998 ²²	I 19 FM (Yunus criteria)	T0 6w l T post	Phase 1: 6w individual training (1,5-3h/w + 2x/w home work) Phase 2: 2y group maintenance (1h/m)	<u>ase 2:</u> 2y group maintenance (1h/m)	Tender point index (TPI). Mvalgic score	6w, 3m : TPI
	43,98 ± 9,6y	2y I FU 3m,			(algometry). VAS pain.	3m, 2y : VAS 凶 in El
	108 DO:18.	ly, 2y.				(no group ≠) 2 y: Myalgic
						score ⊿ III El2,3 (no group ≠).

Study	Sample	Design>	Experimental intervention (EI)	Control intervention (CI)	Outcome	Results
Other relaxation			EII: Biofeedback/relaxation (cognitive and muscular relaxation strategies with biofeedback for trapezius tension). EI2: Exercise group (ROM, aerobic, strength, posture) EI3: EII+EI2.	Educational information on diagnosis, treatment and general health.		
Hammond et al., 2006 ²⁴	183 FM (ACR) ±48,5 y ±90% _や D0: 82.	T0 FU 4, 8m.	10x 2h group educational exercise therapy (education, exercise therapy, pacing, sleep hygiene, relaxation, pain- fatigue-stress management) + activity program 5x 30/w.	10x1 h group relaxation lessons (visualisation, breathing exercises) .	FIQ (pain, physical function and total).	Overall S El > Cl: total (4m); El=Cl: pain and physical function (4, 8m),
Keel et al., 1998 ²⁹	32 FM (ACR) ±49 y 24 ♀ DO: 5.	T0 I 5w I T post FU 3m.	15 × 105-120' group therapy sessions (information, self-control and acceptance, exercise therapy, relaxation, group discussions) (n =16n=14, incl 12 $+$; 48y).	I5 × 45-60' group relaxation sessions (autogenic training) (n=16n=13, incl 12 ♀; 50y).	Pain diary.	FU: EI ≠ CI.
FM = Fibromyalgia, /	ACR = 1990 A w = week m	American Col = month v	FM = Fibromyalgia, ACR = 1990 American College of Rheumatology criteria for FM, CFS = Chronic fatigue syndrome, CDC = 1994 Centre of Disease Control criteria for CFS, DO = deconcuter H = houre w = workh w = wore two from the secondary for end of the secondary fill = following 1 = foreworking (CRT = Committie Bahavioura)	syndrome, CDC = 1994 Centre of Disease Co ant accessment FI1 = following 1 = Interventio	ontrol criteria for C	FS, DO = Behavioural

Table I. (Continued)

drop-out. H = hour, w = week, m = month, y = year, T0 = pre-treatment assessment, T post = post-treatment assessment, FU = follow-up, I = Intervention. CBT = Cognitive Behavioural Therapy; PMR = Progressive Muscle Relaxation; GI= Guided Imagery; GET = Graded Exercise Therapy; ROM = Range of Motion; VAS = Visual Analogue Scale, MPQ = McGill Pain Questionnaire, FIQ = Fibromyagia Impact Questionnaire, BPI = Brief Pain Inventory; BFI = Brief Fatigue Inventory; SF-36 = Short Form 36 Health Survey. & indicate a reduction or increase for a certain outcome; >, <, =, ≠, indicate whether these changes are larger, smaller, equal, different or opposite in the Intervention Group (IG) and Control Group (CG). Eligibility assessment of the search results was performed according to following eligibility criteria:

- Study subjects were adult (> 18 years) chronic fatigue syndrome or fibromyalgia patients.

- All kinds of relaxation (as stand-alone therapy) were allowed and its efficacy was studied.

- Relaxation as part of a multimodal program or combined with movement therapies (eg. Yoga, stretching, etc.) were not allowed.

- Only randomized controlled trials published in full text record in English, French, Dutch or German were included.

First, all search results were independently searched and screened by two of the researchers (MM and TV), based on title and abstract. The full text article was retrieved if the citation was considered potentially eligible and relevant. In the second phase, each full text article was once again independently evaluated by the two researchers whether it fulfilled the inclusion criteria. If any of the eligibility criteria were not fulfilled, then the article was excluded from the literature review. In case of disagreement the last researcher was consulted (FS).

Information was extracted from each included trial on: (1) characteristics of trial participants; (2) type/format of intervention; (3) type/format of control intervention; (4) outcome measure and therapy effects regarding pain, fatigue, daily functioning and autonomic function. Data were extracted from included studies by TV and MM. When extracted data did not match, the study was discussed in order to find a consensus.

Methodological quality of the different studies was assessed with the specific checklist for randomized controlled trials of the Dutch Institute for Healthcare Improvement CBO provided by the Dutch Cochrane Centre (http://dcc.cochrane.org/ beoordelingsformulieren-en-andere-downloads). This checklist assesses 9 items: 1: Are patients randomized to a group; 2: Is it a blind allocation?; 3: Are patients blinded?; 4: Are therapists blinded?; 5: Are assessors blinded?; 6: Were groups comparable?; 7: Is there a sufficient portion of patients included in the follow-up of?; 8: Are all patients analysed the group to which they were randomized?; 9: Are groups treated equally? All items are score with "yes", "no" or "lack of information". Since blinding of patients was not possible, we checked whether patients were naïve to the intervention. If patients remained naïve, studies obtained one point for this item. If studies only relied on self-reports, the item regarding the blinding of the assessor was ignored. Item 7 was positively appreciated if the drop-out was less than 10% and similar in the different groups. Based on the methodological quality of the randomized controlled trials, studies could reach a level of evidence A2 (good quality, sufficient sample size and double-blinded) or B (if previous criteria not fulfilled) (www.cbo.nl).

Methodological quality was assessed independently by two researchers (TV and MM), who were blinded from each other's' quality assessment. After rating the selected articles, the results of both researchers were compared and differences were analysed. In case of disagreement, the reviewers screened the article a second time to obtain a consensus. When consensus could not be reached a third opinion was provided by the last author (FS).

After pooling the results, the overall quality of evidence for each outcome was rated with the Grades of Recommendation, Assessment, Development and Evaluation (GRADE) approach.¹⁵ GRADING the evidence was done by the first author and final author. For every type of relaxation a GRADE summary statement is provided under the respective paragraph in boxes in italics.

Results

Figure 1 shows the process of study selection. In the second screening phase most studies were excluded based on the intervention used. After reading the full text, relaxation therapy was often not used as sole treatment component. Finally, 13 randomized controlled trials that met the inclusion criteria were included in the systematic review

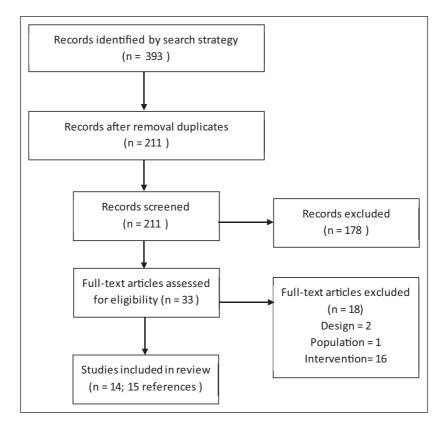


Figure 1. Flow chart study selection.

(reported in 14 articles given the fact that the study of Verkaik was reported in a Dutch paper in 2011¹⁶ and in an English paper in 2014)¹⁷. The two papers by Deale et al. are counted as 2 studies, even though the 2001 study¹⁸ was in fact the long-term followup report of the 1997 study.¹⁹ Also the study by Fors et al.²⁰ is a 4 week extension of the study of Fors and Götestam.²¹ In total the review contained 650 fibromyalgia patients and 88 chronic fatigue syndrome patients.

The risk of bias and the level of evidence of the different studies are reported in Supplementary Table 2.

In most cases (94 % or 110 of the 117 items) the two researchers agreed. After a comparison of the 7 differences, the reviewers reached a consensus for 1 item. The remaining 6 points of discussion were solved after a third opinion. The final score of each study is presented in Supplementary Table 2, with the explanation for the loss of points.

Methodological quality varied between 2/8 to 6/8. Only in 2 studies 9 items were evaluated.^{22,23} The other 10 studies relied on self-reports. This means that assessor and patient are one person. Therefore blinding was only scored once (blinding of the patient). None of the studies reported the therapist to be blinded, which is obvious given the nature of the therapy.

Also blinding of the patients was impossible, in some studies patients were kept naïve for the different interventions,^{21,24} These studies obtained 1 point if the format of the interventions was quite similar (item 9), in this that patients would not assume to be in a control or experimental session. All studies scored well on randomization and comparability of groups.

Because none of the studies was doubleblinded, all studies obtained a GRADE level of evidence B.

For each study, the characteristics for which data were are presented in Table 1.

Regarding intervention, 6 studies evaluated the effects of guided imagery,^{16,17,20,21,25-27} 3 studied (the sole studies in chronic fatigue syndrome patients) the effects of Ost applied relaxation techniques (explained later on),^{18,19,28} 2 used muscle relaxation techniques,^{22,23} 1 autogenic training,²⁹ and finally the last article studied the effect of a more undefined relaxation session.²⁴ The latter might be due to the fact that relaxation therapy served as control intervention, as it did in 6 other studies.^{18,19,23,26,28,29}

Autogenic training refers to a series of mental exercises involving relaxation and autosuggestion, which focus the mind on the body's experience of relaxation,³⁰ Progressive Muscle Relaxation is a technique for learning to control the state of tension in ones muscles.³¹ In Ost applied relaxation techniques, it is aimed at learning to relax rapidly as soon as signs of anxiety are recognised. The client is learned to watch for early signs of anxiety (worrying thoughts, somatic symptoms e.g. palpitations, abdominal discomfort, muscle tension) as cues to immediately start Progressive Muscle Relaxation.³² All varying techniques share the fact that they enhance self-efficacy.

Interventions that were compared to relaxation therapy were diverse: hypnosis, massage, exercise therapy, usual care, Cognitive Behavioural Therapy, etc.

Two studies reported the effect of one session of relaxation,^{21,26} while the others used a treatment program of at least four weeks.

Only three studies reported the effects in chronic fatigue syndrome patients,^{18,19,28} all fulfilling the 1994 Centre of Disease Control criteria³³ for chronic fatigue syndrome. Besides Menzies et al.²⁵ and Buckelew et al.,²² fibromyalgia studies (n=10) used the 1990 American College of Rheumatology criteria³⁴ for diagnosing fibromyalgia. The majority of the participants was female with a mean age varying between 31.9 years¹⁹ and 50.9 years.23 Most outcome measures concerned self-report measures and questionnaires. For evaluating pain, Visual Analogue Scales and the McGill Pain Questionnaire were most often used. Regarding functionality the Fibromyalgia Impact Questionnaire was the most prevalent in the selected studies.

Most studies evaluated the effect on both pain and functionality and four additionally studied the effect on fatigue. Fatigue was measured with different tools in five studies. No studies were found assessing the effect of relaxation on autonomic parameters.

Effects of guided imagery/visualization (fibromyalgia)

All of the six studies that investigated the effects of guided imagery/visualisation in fibromyalgia patients reported effects on pain. Two studies found an acute reduction in pain after one session of guided imagery on pain.^{21,26} The effect depended on the content of the visualization. In the study of Fors and Götestam,²¹ the visualization of a pleasant environment led to a decrease in pain, while the visualization of the human analgesic system did not. The study by Castel,²⁶ found that analgesia suggestion was however more pain relieving than the pleasant visualization. After a prolonged guided imagery program, pain significantly ameliorated in those visualizing pleasant things, while worsening in those visualizing the human analgesic system or subjected to a single 30' free talking session about fibromyalgia.20 In the studies of Verkaik^{16,17} and Menzies,^{25,27} pain did not change after a prolonged guided imagery program including both pleasant visualization and pain or immune related visualization.

For functionality, measured with the Fibromyalgia Impact Questionnaire, Verkaik^{16,17} did not found an effect, while Menzies²⁵ reported a larger improvement in the relaxation group compared to the usual care group.

In the latest study of Menzies²⁷ the relaxation group showed an improvement in *fatigue*, while the control group reported worsened fatigue.

There is conflicting evidence for the isolated effect of guided imagery or visualization on pain and functionality in patients with fibromyalgia. The acute effect of a single guided imagery session seems beneficial for pain reduction in fibromyalgia (moderate evidence). The content is a matter of debate.

Ost applied relaxation (chronic fatigue syndrome)

In both of the studies by Deale^{18,19} and in the study of Thomas,²⁸ the Ost applied relaxation therapy (aimed at learning to relax rapidly as soon as signs of anxiety are recognised by watching for early signs of anxiety) was used as a control intervention. Immediately after, until six months after the treatment programs, a multimodal program (consisting of Cognitive Behavioural Therapy, pacing, mindfulness, etc.) resulted in less fatigue compared to the relaxation therapy alone.¹⁹

Regarding functionality, results may be confusing: at six months follow-up the increases in the subscale physical functioning and the Karnofsky performance scale were equal between the multimodal groups and the relaxation group,^{19,28} while the amount of patients reaching a defined improvement is higher in the multimodal groups.^{19,28} At five years follow-up, no differences were found between the relaxation and Cognitive Behavioural Therapy groups.¹⁸

In short term (up to six months) there is preliminary evidence for a less beneficial outcome, especially regarding fatigue, in chronic fatigue syndrome patients only receiving Ost relaxation compared to more comprehensive rehabilitation programs.

Muscle relaxation (fibromyalgia)

Muscle relaxation programs resulted in decreased pain and fatigue, both on self-report measures as on

algometry measures, but changes were not always significantly different compared to the control intervention.^{22,23} Massage therapy however was more efficacious than Progressive Muscle Relaxation instructions in reducing self-reported pain and fatigue and pain assessments.²³ Muscle relaxation with biofeedback and/or exercise were more efficacious in reducing tender point index compared to educational information, but it seems that adding exercise may result in better outcome.²²

There is very limited evidence for the isolated pain relieving effects of muscle relaxation in fibromyalgia, possibly other modalities (massage, exercise, biofeedback) are more beneficial or need to be added to generate a synergistic effect.

Other relaxation therapy formats (fibromyalgia)

Regarding pain, autogenic training (patients repeating a set of visualisations themselves) in group sessions was equally effective as integrated group therapy sessions (including exercise therapy, relaxation, acceptance, discussion etc.) in symptom reduction, immediately post-intervention. However, at follow-up, there was an increase in pain in the relaxation group and a decrease in the experimental group, a difference that was statistically significant.²⁹

Ten sessions of group relaxation, including breathing exercises and visualisation, resulted in quite similar improvements at 4 and 8 months on the Fibromyalgia Impact Questionnaire, compared to a multimodal approach consisting of exercise therapy, activity program, pacing, stress-management, relaxation etc. Only for the total score of the Fibromyalgia Impact Questionnaire (including subscales like anxiety, depression etc.), the comprehensive approach was better at four months follow-up.²⁴

Discussion

Most studies investigated the effect on pain and functioning and the results are conflicting.

This may come as no surprise given the huge variance in used protocol, formats, control therapies etc.

Nevertheless, it seems that a single session of guided imagery has beneficial acute effects on pain in patients with fibromyalgia.^{21,26} For prolonged treatment programs of guided imagery more study is warranted, because the studies of Verkaik^{16,17} and Menzies²⁵ could not confirm the findings of Fors et al.²⁰ that the regular use of pleasant guided imagery pain has alleviating effects during a four week period and has consequently clinical utility. The fact that results were not univocal may be due to the content of the guided imagery, which is indeed a matter of debate. Fors et al.20 compared two types of imagery: one pleasant visualization not referring to pain or other negative aspects and one visualizing the human endogenous system. The imagery exercise of Verkaik^{16,17} and Menzies^{25,27} incorporated both pleasant imagery and pain or immune system related imagery. It seems thus beneficial to distract attention from the pain/body. From these findings it seems better to distract and guide the patient towards a decreased health anxiety,²¹ and away from the pain.

In the study by Castel et al.²⁶ the hypnosis plus analgesia suggestion was however more efficacious in reducing pain, compared to relaxation or to hypnosis with relaxation suggestion. But there might be a difference between hypnosis with analgesia or relaxation suggestion. A deeper hypnosis was used in the hypnosis plus analgesia suggestion group compared to the relaxation suggestion group. Different from imagery studies using the visualization of the endogenous pain inhibitory system, the participants were asked to imagine a liquid or blue analgesic stream that filtered through their skin and reached different parts of their body (muscles, joints, bones, internal organs). It was suggested that the liquid soothed the pain in the most affected areas, eliminated the tension, and created feelings of wellbeing. ²⁶ This means that an external source of pain inhibition was visualized.

For functionality results are more inconclusive. Some results indicate improved functionality following guided imagery,²⁵ others report similar improvements in the relaxation therapy group comparison to often more comprehensive treatment programs including exercise therapy and activity programs²⁴ and still others report ambiguous results.^{19,28} The effect on this outcome measure certainly deserves further attention, but it is remarkable that none of the studies provide firm evidence for multicomponent therapy programs being more efficacious in improving functional status than relaxation alone. The restoring effect of relaxation therapy is mirrored e.g. in a reduction in the number of visits to a doctor by almost a third, suggesting that the effect of support from both the group leader and peers may have led to less health care seeking. Half of the group in the study of Hammond et al. considered relaxation therapy as beneficial for their fibromyalgia.²⁴ Taken together these findings suggest that the positive effects of relaxation therapy in patients with fibromyalgia are (partly) due to improvements in self-efficacy. A systematic literature review identified self-efficacy as one of the major factors responsible for a positive rehabilitation outcome in patients with chronic pain.35

But, as previously mentioned, the reporting of study findings is sometimes confusing and further study should account for that. In the studies of Deale^{18,19} it is reported that Cognitive Behavioural Therapy was more efficacious than relaxation therapy in improving functional status in chronic fatigue syndrome patients, because substantial self-reported improvement occurred in 70% (63% with the drop-outs) of the Cognitive Behavioural Therapy patients, compared with 19% (17% with the drop-outs) of the patients in the relaxation sessions. But on the other hand there was no significant interaction effect for the change in the subscale physical functioning of the Short form 36 health survey (SF-36). This may suggest that the results of "general improvement" may be influenced by the patients' perceptions. It could be that patients' expectancies are higher regarding comprehensive treatment approaches compared to relaxation therapy alone. Global improvement was rated on a 7-point scale from "very much better" through "unchanged" to "very much worse." Consequently, the score of improvement may rather reflect therapy satisfaction than improvement, as changes on the SF-36 were not significantly better. On the other hand, improvement was sometimes defined

as a 50% change on the SF-36 or a defined cut-off (83%) on the SF-36.¹⁹ The same goes for the study by Thomas,²⁸ in which the amount of patients presenting a 10% increase or a 80% score on the Karnofsky performance scores post-treatment is significant higher in the Multiconvergent therapy group, but the change in performance score was not statistically different.²⁸

Fatigue was only studied in three chronic fatigue syndrome studies^{18,19,28} and in the fibromyalgia studies by Field²³ and Menzies.²⁷ Besides the latter who found guided imagery to be better than usual care,²⁷ all these studies used relaxation therapy as control intervention and found a larger reduction in fatigue in the massage therapy group,²³ Cognitive Behavioural Therapy group.²⁸

No studies directly evaluating the effect of relaxation on autonomic parameters in patients with chronic fatigue syndrome or fibromyalgia were revealed. This may be warranted, since different studies reported autonomic aberrances in these populations.^{7,9,36,37}

Inconsistencies in some of the finding may be due to the variety in protocols and the different relaxation formats. Some of the methods are performed alone; some require the help of another person (often a trained professional or an audio tape); some involve movement, some focus on stillness; while other methods involve different elements.

All these techniques can be performed alone, with some techniques relying on mental exercises (eg. autogenic training), others requiring contracting and relaxing of the muscles (Ost relaxation and progressive muscle relaxation). The guided imagery, on the other hand, consists of suggestions given to a client by a trained practitioner and is thus induced with the assistance of a therapist or tape.

Based on the present review the findings of the studies using guided imagery are the most promising for treating patients with fibromyalgia. This may suggest that fibromyalgia patients need an external cue to be able to relax and to benefit from the relaxation therapy. It seems the most beneficial to visualize pleasant things and guide the brain away from the pain, unless the suggestion of administering an analgesic is given. But the content of the guided imagery demands for further research.

The present review is based on randomized controlled trials which is in favour of the level of evidence. Unfortunately, risk of bias was possible in all studies. Due to the nature of the intervention, it was hard to blind patients for the type of intervention. Only in the studies evaluating different interventions provided by audiotape, blinding or assuring naïve patients was possible. Furthermore, most of the studies relied on self-report measures, prohibiting blinding of the assessors as well. The combination of both self-report measures and the lack of blinding of patients, may be an important shortcoming.

Secondly, it is a pity that no studies were found evaluating the effect of relaxation on autonomic function in these patients. This may be due to the very stringent eligibility criteria and studying only two databases. Only studies evaluating the effect of relaxation therapy as a stand-alone therapy were included, to allow studying the isolated effect of relaxation therapy. In the excluded studies relaxation therapy was often combined with stretching or flexibility exercises,38,39 yoga40 etc. To avoid bias through studies incorporating more "movement oriented modalities", these studies were excluded. This was done in order to study the primary aim of the literature study. Probably, combining relaxation therapy with other treatment modalities like patient education, counselling, activity management and exercise imparts synergistic effects.

Finally, many of the included studies used relaxation therapy as a control intervention and this may have influenced the format of the relaxation, the description of the formats and the reporting of the results. It might have even introduced therapist bias into the trial (i.e. that the therapist, being aware of the study hypothesis, becomes biased towards a less favourable outcome of the control intervention). Furthermore, since relaxation is often used as control intervention in studies, it might be that we have missed some of these studies as they might have been identified by other MeSh terms, more related to their experimental intervention.

Consequently, it is clear that more research is warranted into the isolated effects of relaxation

therapy in patients with fibromyalgia and especially in chronic fatigue syndrome. Especially since studies on physiological effects are lacking and the current effects on functioning are heading for clearer comparisons with the more time and money consuming multicomponent programs. However, we do not propose relaxation therapy as a stand-alone therapy, but it should be considered which components are the most valuable in a multicomponent program, as functional outcome does not seem very different to that of relaxation on its own, based on the present knowledge. Using physiological (i.e. autonomic and HPA-axis responsiveness) outcomes is warranted to examine the mechanism of action of relaxation therapy for patients with fibromyalgia or chronic fatigue syndrome.

Six of the included studies concerned guided imagery, but studies on other popular relaxation techniques, as Progressive Muscle Relaxation and autogenic training are scarce. These are however frequently used techniques and deserve further study. It would be interesting for instance to study the autonomic response to different kinds of relaxation to monitor which formats can induce the best relaxation effect.

Clinical messages

- Although firm evidence for the isolated effects of relaxation therapy is lacking, the acute effect of a single guided imagery session seems beneficial for pain reduction in fibromyalgia. The content is a matter of debate, but probably pleasant visualization (away from the pain) is the best.
- Muscle relaxation alone seems less beneficial for pain relief in fibromyalgia than a in a combination with other modalities.
- Longer relaxation therapy programs seem beneficial for improving daily functioning, competing with more multi-modal programs.

Conflict of interest

The authors declare that there is no conflict of interest.

Funding

This research received no specific grant from any funding agency in the public, commercial, or not-for-profit sectors.

References

- Van Houdenhove B, Verheyen L, Pardaens K, Luyten P and Van Wambeke P. Rehabilitation of decreased motor performance in patients with chronic fatigue syndrome: should we treat low effort capacity or reduced effort tolerance? *Clinical rehabilitation* 2007; 21: 1121–1142.
- Van Houdenhove B, Egle U and Luyten P. The role of life stress in fibromyalgia. *Current rheumatology reports* 2005; 7: 365–370.
- Crofford LJ and Demitrack MA. Evidence that abnormalities of central neurohormonal systems are key to understanding fibromyalgia and chronic fatigue syndrome. *Rheumatic diseases clinics of North America* 1996; 22: 267–284.
- Martinez-Lavin M and Hermosillo AG. Autonomic nervous system dysfunction may explain the multisystem features of fibromyalgia. *Seminars in arthritis and rheumatism* 2000; 29: 197–199.
- Frith J, Zalewski P, Klawe JJ, Pairman J, Bitner A, Tafil-Klawe M, et al. Impaired blood pressure variability in chronic fatigue syndrome – a potential biomarker. *Qjm* 2012; 105: 831–830.
- Jones DE, Hollingsworth KG, Taylor R, Blamire AM and Newton JL. Abnormalities in pH handling by peripheral muscle and potential regulation by the autonomic nervous system in chronic fatigue syndrome. *Journal of internal medicine* 2010; 267: 394–401.
- Newton JL, Okonkwo O, Sutcliffe K, Seth A, Shin J and Jones DE. Symptoms of autonomic dysfunction in chronic fatigue syndrome. *QJM* 2007; 100: 519–526.
- Newton JL, Pairman J, Hallsworth K, Moore S, Plotz T and Trenell MI. Physical activity intensity but not sedentary activity is reduced in chronic fatigue syndrome and is associated with autonomic regulation. *QJM* 2011; 104: 681–687.
- De Becker P, Dendale P, De Meirleir K, Campine I, Vandenborne K and Hagers Y. Autonomic testing in patients with chronic fatigue syndrome. *The American journal of medicine* 1998; 105: 22S–26S.
- Pall ML. Common etiology of posttraumatic stress disorder, fibromyalgia, chronic fatigue syndrome and multiple chemical sensitivity via elevated nitric oxide/peroxynitrite. *Medical hypotheses* 2001; 57: 139–145.
- 11. Clauw DJ and Chrousos GP. Chronic pain and fatigue syndromes: overlapping clinical and neuroendocrine features and potential pathogenic mechanisms. *Neuroimmunomodulation* 1997; 4: 134–153.
- Van Houdenhove B and Egle UT. Fibromyalgia: a stress disorder? Piecing the biopsychosocial puzzle together. *Psychotherapy and psychosomatics* 2004; 73: 267–275.
- 13. Nijs J, Mannerkorpi K, Descheemaeker F and Van Houdenhove B. Primary care physical therapy in people

with fibromyalgia: opportunities and boundaries within a monodisciplinary setting. *Physical therapy* 2010; 90: 1815–1822.

- Moher D, Liberati A, Tetzlaff J and Altman DG. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *Annals of internal medicine* 2009; 151: 264–269.
- Balshem H, Helfand M, Schunemann HJ, Oxman AD, Kunz R, Brozek J, et al. GRADE guidelines: 3. Rating the quality of evidence. *Journal of clinical epidemiology* 2011; 64: 401–406.
- Verkaik R, Busch M, Koeneman T, van den Berg R, Spreeuwenberg PMM and Francke AL. Guided imagery in people with fibromyalgia: Effects on pain, self-efficacy and functional status. *Psychologie & Gezondheid* 2011; 39: 282–291.
- Verkaik R, Busch M, Koeneman T, van den Berg R, Spreeuwenberg P and Francke AL. Guided imagery in people with fibromyalgia: a randomized controlled trial of effects on pain, functional status and self-efficacy. J Health Psychol 2014; 19: 678–688.
- Deale A, Husain K, Chalder T and Wessely S. Long-term outcome of cognitive behavior therapy versus relaxation therapy for chronic fatigue syndrome: a 5-year followup study. *The American journal of psychiatry* 2001; 158: 2038–2042.
- Deale A, Chalder T, Marks I and Wessely S. Cognitive behavior therapy for chronic fatigue syndrome: a randomized controlled trial. *The American journal of psychiatry* 1997; 154: 408–414.
- Fors EA, Sexton H and Gotestam KG. The effect of guided imagery and amitriptyline on daily fibromyalgia pain: a prospective, randomized, controlled trial. *J Psychiatr Res* 2002; 36: 179–187.
- Fors EA and Gotestam KG. Patient education, guided imagery and pain related talk in fibromyalgia coping. *European Journal of Psychiatry* 2000; 14: 233–240.
- Buckelew SP, Conway R, Parker J, Deuser WE, Read J, Witty TE, et al. Biofeedback/relaxation training and exercise interventions for fibromyalgia: a prospective trial. *Arthritis care and research* 1998; 11: 196–209.
- Field T, Diego M, Cullen C, Hernandez-Reif M, Sunshine W and Douglas S. Fibromyalgia pain and substance P decrease and sleep improves after massage therapy. *Jcr-Journal of Clinical Rheumatology* 2002; 8: 72–76.
- Hammond A and Freeman K. Community patient education and exercise for people with fibromyalgia: a parallel group randomized controlled trial. *Clinical rehabilitation* 2006; 20: 835–846.
- Menzies V, Taylor AG and Bourguignon C. Effects of guided imagery on outcomes of pain, functional status, and self-efficacy in persons diagnosed with fibromyalgia. *Journal of alternative and complementary medicine* 2006; 12: 23–30.
- Castel A, Perez M, Sala J, Padrol A and Rull M. Effect of hypnotic suggestion on fibromyalgic pain: Comparison

between hypnosis and relaxation. *European Journal of Pain* 2007; 11: 463–468.

- Menzies V, Lyon DE, Elswick RK, Jr, McCain NL and Gray DP. Effects of guided imagery on biobehavioral factors in women with fibromyalgia. *Journal of behavioral medicine*. 2014; 37: 70–80.
- Thomas MA, Sadlier MJ and Smith AP. A multiconvergent approach to the rehabilitation of patients with chronic fatigue syndrome: a comparative study. *Physiotherapy* 2008; 94: 35–42.
- Keel PJ, Bodoky C, Gerhard U and Muller W. Comparison of integrated group therapy and group relaxation training for fibromyalgia. *The Clinical journal of pain* 1998; 14: 232–238.
- Schultz JH. Autogenic training in general practice. Medizinische Klinik 1950; 45: 945–949.
- Jacobson E. *Progressive relaxation*. Chicago: Press UoC, 1938.
- Ost LG. Applied relaxation: description of a coping technique and review of controlled studies. *Behav Res Ther* 1987; 25: 397–409.
- 33. Fukuda K, Straus SE, Hickie I, Sharpe MC, Dobbins JG and Komaroff A. The chronic fatigue syndrome: a comprehensive approach to its definition and study. International Chronic Fatigue Syndrome Study Group. *Annals of internal medicine* 1994; 121: 953–959.
- Wolfe F, Smythe HA, Yunus MB, Bennett RM, Bombardier C, Goldenberg DL, et al. The American College of Rheumatology 1990 Criteria for the Classification of Fibromyalgia. Report of the Multicenter Criteria Committee. *Arthritis and rheumatism* 1990; 33: 160–172.
- 35. Miles CL, Pincus T, Carnes D, Homer KE, Taylor SJ, Bremner SA, et al. Can we identify how programmes aimed at promoting self-management in musculoskeletal pain work and who benefits? A systematic review of subgroup analysis within RCTs. *European journal of pain* 2011; 15: 775.
- Reyes del Paso GA, Garrido S, Pulgar A and Duschek S. Autonomic cardiovascular control and responses to experimental pain stimulation in fibromyalgia syndrome. J *Psychosom Res* 2011; 70: 125–134.
- Martinez-Lavin M, Hermosillo AG, Rosas M and Soto ME. Circadian studies of autonomic nervous balance in patients with fibromyalgia: a heart rate variability analysis. *Arthritis Rheum* 1998; 41: 1966–1971.
- Wallman KE, Morton AR, Goodman C, Grove R and Guilfoyle AM. Randomised controlled trial of graded exercise in chronic fatigue syndrome. *Medical Journal of Australia* 2004; 180: 444–448.
- Fulcher KY and White PD. Randomised controlled trial of graded exercise in patients with the chronic fatigue syndrome. *British medical journal* 1997; 314: 1647–1652.
- Martin L, Nutting A, MacIntosh BR, Edworthy SM, Butterwick D and Cook J. An exercise program in the treatment of fibromyalgia. *Journal of Rheumatology* 1996; 23: 1050–1053.