

# Effectiveness of exercise in patients with osteoarthritis of hip or knee: nine months' follow up

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## Abstract

**Objective**—To determine whether the effects of an exercise programme in patients with osteoarthritis of hip or knee are sustained at six and nine months' follow up.

**Methods**—A randomised, single blind, clinical trial was conducted in a primary care setting. Patients with osteoarthritis of hip or knee (ACR criteria) were selected. Two intervention groups were compared. Both groups received treatment from their general practitioner, including patient education and drug treatment if necessary. The experimental group also received exercise treatment from a physiotherapist in primary care. The treatment period was 12 weeks, with an ensuing 24 week follow up. The main outcome measures were pain, drug use (non-steroidal anti-inflammatory drugs), and observed disability.

**Results**—201 patients were randomly allocated to the exercise or control group, and 183 patients completed the trial. At 24 weeks exercise treatment was associated with a small to moderate effect on pain during the past week (difference in change between the two groups  $-11.5$  (95% CI  $-19.7$  to  $-3.3$ ). At 36 weeks no differences were found between the groups.

**Conclusions**—Beneficial effects of exercise decline over time and finally disappear.

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In treatment guidelines for osteoarthritis (OA) of the hip or knee the use of exercise has been advocated.<sup>1,2</sup> In a recent update, exercise again is recommended as an important non-pharmacological treatment in patients with OA of the hip and knee.<sup>3</sup>

Exercise treatment in OA aims at reducing pain and disability.<sup>4</sup> This is achieved through improvement of muscle strength, stability of joints, range of motion, and aerobic fitness. These functions are often impaired in patients with OA, presumably contributing to pain and disability.<sup>5</sup> Improving these functions is assumed to reduce pain and disability. In addition, exercise aims directly at reducing disability—for example, through correction of the walking pattern.<sup>6,7</sup>

In a systematic review, based on 11 randomised controlled trials (RCTs), we concluded that there is evidence for the beneficial effects of exercise on pain, disability, and

patient's global assessment.<sup>8</sup> Since the publication of this review several new RCTs on exercise in patients with OA have been published. These newly published RCTs confirm this conclusion.<sup>9–13</sup>

Most randomised clinical trials, however, were confined to post-treatment results, ignoring residual effectiveness. Randomised controlled trials on the long term effects of exercise in patients with OA of the hip or knee are few. In four trials the residual effects of exercise were studied after completion of the programme.<sup>9–15</sup> Beneficial effects at one year were found in four trials, on pain,<sup>9,14</sup> disability,<sup>14</sup> and muscle strength.<sup>9,13</sup> In one trial no beneficial effects could be shown.<sup>16</sup> However, several methodological shortcomings hamper the interpretation of these results. These include the absence of an intention to treat analysis,<sup>13,14</sup> a high loss to follow up,<sup>15</sup> combined data presentation for different patient groups (OA and rheumatoid arthritis)<sup>14</sup> or for different assessments,<sup>9</sup> and a low power owing to small numbers of patients at randomisation.<sup>9</sup>

One trial studied the effectiveness of exercise consisting of a three month centre based phase followed by a 15 month home based phase.<sup>16,17</sup> The effects of exercise after completion of the exercise programmes were not studied. When data from follow ups at 3, 9, and 18 months were combined, beneficial effects were reported for pain, disability, muscle strength, and aerobic capacity. Separate follow up data were presented for a subset of outcome measures. These data indicated a maximal beneficial effect at three months (after completion of the supervised programme) for self reported disability and pain, and at nine months for walking speed. Most effects showed a slight decline afterwards, even when the home based exercise programme was continued.

In uncontrolled studies too, scarcely any attention has been paid to the residual effects of exercise in OA. Fisher *et al* reported beneficial follow up effects (eight and 12 months) of a four month muscle rehabilitation programme.<sup>18</sup> Beneficial effects on pain and observed disability (including walking) remained at post-treatment levels, at both eight and 12 months. Beneficial effects on muscle strength and endurance had declined at 12 months, but remained significantly higher than baseline levels. In a case study, Marks presented effects of quadriceps exercises on muscle strength.<sup>19</sup> After a six week supervised

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training period quadriceps torque had improved by 40%, with a 10% reduction after a six week non-training period.

In conclusion, the residual effects of exercise in OA of the hip or knee after completion of treatment have been investigated in only a few studies. It remains unclear how long beneficial effects are sustained after completion of the exercise treatment.

We here report a follow up study to determine whether the effects of exercise in patients with OA of the hip or knee are sustained over time. The effects of exercise at 12 and 24 weeks after completion of a supervised exercise programme are presented. In addition, the prognostic value of patient characteristics for outcome after completion of treatment is explored. The short term effects of exercise in our trial have been reported in detail elsewhere.<sup>20</sup>

### Patients and methods

#### STUDY GROUP

Patients were selected by GPs in the period May 1994 to February 1996. The GPs were situated in four cities and surrounding villages in the eastern part of the Netherlands. Inclusion criteria for patients were OA of the hip or knee according to the clinical criteria of the American College of Rheumatology.<sup>21 22</sup> Exclusion criteria were another disease which might explain the complaints; complaints in fewer than 10 of 30 days; treatment for these complaints with exercise in the preceding six months; age under 40 or over 85; indication for hip or knee replacement; contraindication for exercise treatment; contraindications for analgesics or non-steroidal anti-inflammatory drugs (NSAIDs); and inability to understand the Dutch language. After having given oral consent, patients were registered and their names forwarded to the research team. Radiographs were obtained and evaluated by one radiologist using grading scales (0–3) for individual radiographic features.<sup>23</sup> All patients were visited and their eligibility was checked by a GP research fellow (DB). All eligible patients were asked to give written informed consent. Afterwards, patients were randomly allocated equally to either the exercise or the control group, using sequentially numbered, opaque, sealed envelopes containing the treatment assigned.

Patients were split into groups according to their pain in the past week (VAS 0–30 v 31–100 mm) and location of OA (hip or knee) in order to achieve comparability in these prognostic factors.

We aimed at having 200 patients participating in the study, as this number of patients leads to a power of 0.80 to detect small to medium sized effects with an  $\alpha$  of 0.05.<sup>24</sup>

The study protocol was approved by the ethics committee of the Maastricht University Hospital (Maastricht, the Netherlands).

#### INTERVENTIONS

Two interventions were compared. The patients in the exercise group were given exercise treatment individually by a physiotherapist in

primary care. In addition, their GP provided patient education (including a brochure) and drug treatment, if necessary. Treatment of the control group was restricted to that given by their GP, as described above (patient education and drugs, if necessary). The treatment period was 12 weeks, with 24 weeks of follow up.

Exercise treatment was given according to a written protocol<sup>25</sup> and included exercises for muscle functions (strength and length), mobility, and coordination, and exercises for elementary movement abilities and locomotion abilities. Also, instructions for the adaptation of activities of daily living and home exercises were given. Content and intensity of treatment were described by treatment goals and corresponding exercises. Content, intensity, and frequency of treatment were tailored to the patient's needs. Depending on the physiotherapist's diagnostic findings, specific treatment goals with corresponding exercises were chosen. A number of sessions a week were prescribed, ranging from one to three times a week, depending on the pain level. A physiotherapy session in primary care lasted approximately 30 minutes. Exercise treatment could be discontinued within the 12 week period if, according to the physiotherapists, treatment goals had been achieved. Physiotherapists were trained to use the protocol.

A protocol was also used for the prescription of drugs. The GP prescribed preferably paracetamol; prescription of NSAIDs was restricted to naproxen, diclofenac sodium, and ibuprofen. The patient was instructed to use as few drugs as possible. The GP also provided a brochure for patient education, covering diagnosis, prognosis, advice about rest, daily activities and diet, the use of aids and medical treatment. No advice about exercise was included. GPs were instructed to minimise treatment in the follow up period. Physiotherapists and GPs recorded detailed information about the actual treatments on standardised forms, including any deviation from the protocol.

#### OUTCOME ASSESSMENT

##### Primary outcome measures

Primary outcome measures were pain, use of NSAIDs, and observed disability. Patients rated their pain in the past week on a visual analogue scale (0 mm = no pain to 100 mm = very severe pain) in answer to the question: How severe was your pain during the past week? The amount of NSAIDs used was based on prescription data and a count of remaining drugs during evaluation sessions. Observed disability was determined by studying videos of the patients' performance of a series of standardised tasks, using an adaptation of the method described by Keefe *et al.*<sup>26 27</sup> The tasks included walking, sitting down, bending, and reclining. Movement times and quality of performance were both assessed. The inter-observer reliability of this method is good.<sup>27–29</sup> A total score was calculated based on five measures: 5 m walking time, stand to sit time, stand to recline time, and the levels of caution and rigidity during the performance of the

tasks. Standardised scores (Z scores) of separate measurements were first calculated and then summed to obtain an overall score. To enhance comparability the resulting overall score was standardised to produce a score with a mean of zero and a standard deviation of one.<sup>30</sup> The internal consistency of the constructed overall score was good ( $\alpha=0.84$ ).<sup>31</sup>

#### Secondary outcome measures

A set of 13 secondary outcome measures were included in the trial, to study the working mechanisms of exercise treatment in OA. In this article, a subset of these secondary outcome measures is included: use of paracetamol assessed in the same way as the use of NSAIDs; global perceived effect as assessed by the patients themselves on an eight point scale (1 = vastly worsened; 8 = completely recovered)<sup>32</sup>; self reported disability with the IRGL questionnaire (Influence of Rheumatic disease on General health and Lifestyle)<sup>33</sup>; muscle strength measured with a hand held dynamometer<sup>34</sup>; joint motion measured with a goniometer<sup>35</sup>; and the level of physical activity according to the Zutphen Physical Activity Questionnaire (excluding sedentary hobbies).<sup>36</sup> Overall scores were compiled for muscle strength of hip and knee and joint range of motion of hip and knee. The procedure was similar to data reduction in observed disability (mean = 0, SD = 1, see van Baar *et al*<sup>30</sup>).

#### Prognostic factors

The following prognostic factors were studied:

- Demographic characteristics: age and sex
- Clinical features: location of OA of the hip (yes/no) and the knee (yes/no), OA of the hand according to Altman *et al*<sup>37</sup> (yes/no), radiological OA as measured on grading scales for individual radiographic features (yes/no) (for details see van Baar *et al*<sup>30</sup>), obesity (body mass index  $<30/\geq 30$ )<sup>38</sup>, duration of complaints and comorbidity (yes/no)
- Baseline levels of outcome measures: pain during the past week, drug use (NSAIDs, paracetamol), observed and self reported disability, muscle strength, and range of joint motion, both for hip and knee, and level of physical activity
- Pain coping strategies (six dimensions) as assessed with the Pain Coping Inventory.<sup>39</sup> In addition, the extent to which patients believe physical activity to affect their pain was assessed using an adaptation for patients with OA of the Fear-Avoidance Beliefs Questionnaire<sup>40</sup>
- Psychological wellbeing: anxiety, depression, and cheerfulness as assessed with the IRGL questionnaire<sup>33</sup>
- Compliance with exercise as reported by both the patient and the physiotherapist. Patients were phoned by a research fellow (MvB) at weeks 6 and 12 and asked, using standardised questions, whether they had done their home exercises as often as prescribed. Possible answers were: (almost) never, sometimes, regularly, often, and very often. This contact was not used to reinforce compliance with the exercise programme.

Physiotherapists, using a standardised form, were asked to estimate whether the patient complied with the home exercise instructions in the last four weeks of treatment. Possible answers were: certainly not, probably not, probably, and certainly. A patient was considered compliant (a) when the patient reported exercising (very) often or (b) when the physiotherapist estimated that the patient had certainly exercised at home. Patients were evaluated by a “blinded” research assistant at baseline and at 12 weeks (the end of treatment), 24 weeks, and 36 weeks. The research assistants had been trained to perform the measurements in a standardised manner. The evaluations took place in local healthcare centres.

#### STATISTICAL ANALYSIS

Analyses were performed according to the intention to treat principle.<sup>41</sup> Patient data were analysed in the intervention groups to which they had initially been assigned. This included patients who had withdrawn and patients not treated according to the assigned treatment. In addition, a per-protocol analysis was performed, excluding patients with deviations from the treatment protocol and late ineligible patients.

To analyse the effects, change scores were calculated (follow up score minus baseline score). For drug use and global perceived effect, follow up scores were compared because no change scores could be calculated. The scores for global perceived effect and physical activity were dichotomised in order to study the number of patients who had improved. In the analysis, the change (or follow up) scores were compared between the interventions by analysis of covariance. Adjusted analyses were performed. The baseline level of each outcome measure was included to enhance the precision of the effect estimates. In addition, drug use and fear avoidance beliefs, the measures on which the groups differed at baseline (see “Results”), were included as covariates in order to control for baseline differences. Group differences and 95% confidence intervals were calculated for all outcome measures. In addition, effect sizes were calculated by taking the difference between the change scores of the intervention groups and dividing it by the standard deviation of the change score of the total group. An effect size of 0.2 is considered to be a small beneficial effect of exercise, 0.5 a medium effect, and 0.8 a large effect.<sup>24</sup>

The analysis for studying the prognostic value of patient characteristics for long term outcome of exercise was restricted to patients who had received exercise treatment. These patients were classified as (a) being stable or improved or (b) having deteriorated, based on their results for the primary outcome measures during follow up (week 12 to week 36). Patients pain during the past week or observed disability was classified as stable or improved if their scores in week 36 were equal or lower than in week 12. Patients were classified as stable or improved for NSAIDs if they did not change drug use (or non-use) or if they had reduced or

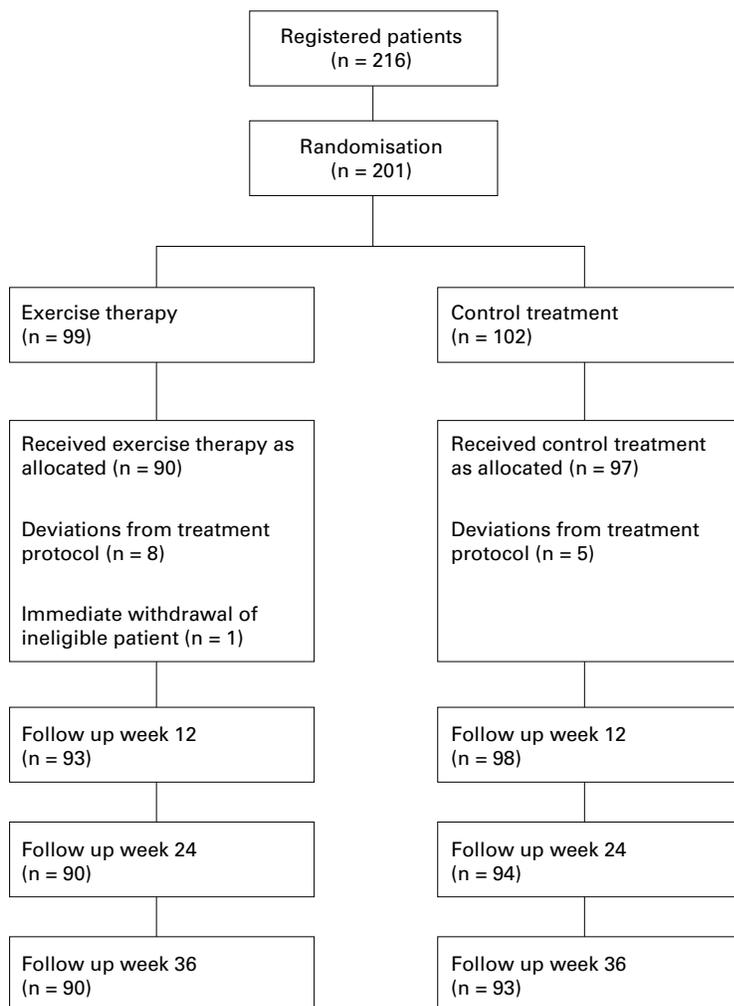


Figure 1 Participant flow and follow up.

Table 1 Comparability of intervention groups at baseline\*. Results are given as number (%) except where otherwise stated

	Exercise therapy (n=98)	Control (n=102)
Sex (female)	76 (78)	81 (79)
Age (years), mean (SD)	68.3 (8.4)	67.7 (9.2)
Comorbidity	63 (64)	62 (62)
Location of OA		
Knee	58 (59)	61 (60)
Hip	36 (37)	35 (34)
Both	4 (4)	6 (6)
Duration of complaints (years)		
≤0.5	34 (35)	42 (41)
0.5–≤1	13 (13)	12 (12)
1–≤5	29 (30)	25 (25)
>5	22 (22)	23 (23)
Radiological OA (score ≥ 1)		
Joint space narrowing	62 (69)	55 (60)
Osteophytes	65 (72)	65 (71)
Previous medical treatment	59 (61)	64 (64)
Pain in past week (VAS† 0–100), mean (SD)	46.9 (27.7)	43.1 (26.8)
At assessment, † mean (SD)	34.0 (27.2)	28.7 (26.0)
Drug use in past week (taking)		
Paracetamol	51 (52)	39 (38)
NSAIDs‡	34 (35)	23 (23)
Observed disability		
5 m walking time (s), median (IQR†)	4.9 (4.3–6.0)	5.0 (4.3–6.0)
Sit time (s), median (IQR)	3.4 (2.9–4.1)	3.4 (2.9–4.0)
Recline time (s), median (IQR)	6.9 (5.4–9.0)	6.1 (5.1–7.9)
Guarding (0–1), mean (SD)	0.58 (0.48)	0.53 (0.50)
Rigidity (0–1), mean (SD)	0.34 (0.38)	0.33 (0.37)

\*Missing values: comorbidity (1 in exercise therapy, 2 in controls); duration of complaints (2 in exercise therapy, 6 in controls); radiological OA (7 in exercise therapy, 12 in controls); previous medical treatment (1 in exercise therapy, 2 in controls).

†VAS = visual analogue scale; NSAIDs = non-steroidal anti-inflammatory drugs; IQR = interquartile range.

stopped using NSAIDs in the period week 12 to week 36. A  $\chi^2$  test and  $t$  test for independent samples were used to test for baseline differences or differences in compliance between these two groups.

The analyses were carried out using SPSS/PC+ 5.0.

## Results

### PARTICIPANT FLOW

Figure 1 shows the patient flow and follow up between registration and week 36. In total 201 patients were included in the study, 99 patients were allocated to exercise therapy, 102 patients to the control treatment. One patient allocated to exercise treatment was excluded because of additional medical information (erythrocyte sedimentation rate >45 mm/1st h) obtained immediately after random allocation to the group.

The week 12 assessment was completed by 191 patients; 183 patients completed the trial up to 36 weeks. Reasons for withdrawal and loss to follow up were comorbidity (n=8), loss of motivation (n=6), family circumstances (n=2), and adverse effects of exercise (n=1). The number of patients withdrawing and lost to follow up was relatively low. In addition, numbers and reasons were equally divided across the intervention groups.

Table 1 presents the baseline characteristics in the two intervention groups. Patients were generally similar, and also with respect to OA and the contralateral leg (data not shown). There were two exceptions: patients allocated to the exercise treatment group reported a higher use of drugs in the seven days before participation in the study and scored higher on fear avoidance beliefs concerning physical activity.

### TREATMENT

Patients receiving exercise treatment visited their GP 1.6 (SD 0.7) times in the 12 week treatment period, control patients 1.8 (0.9) times (p=0.03). The GP prescribed drugs for 54% and 66% of the patients in the two groups, respectively (p>0.05).

In the exercise group the mean number of physical therapy sessions was 16.8 (SD 7.0). The median number of sessions was 14 (range 3–30). The median duration of treatment was 11 weeks; 38% of the patients received 12 weeks of treatment (for details, Baar *et al*<sup>9</sup>).

Compliance with home exercises was moderate to good: 66% of the patients reported exercising often or very often. The physical therapist estimated good compliance (that is, certainly did the home exercises) for 53% of the patients, and moderate compliance (that is, probably did the home exercises) for 33%.

### PRIMARY OUTCOME MEASURES

Table 2 presents follow up results for the primary outcome measures. At 24 weeks (12 weeks after completion of treatment), a beneficial effect was seen for pain during the past week. Compared with the post-treatment level (week 12) the effect size had declined to 0.36, indicating a small to moderate effect. No

Table 2 Primary outcome measures: improvements and differences between intervention groups\*

	Exercise therapy		Control		Difference (95% CI) exercise therapy - controls	Effect size
	n	Mean change†	n	Mean change†		
Pain past week						
Week 12	93	-22.8	98	-5.7	-17.0 (-23.6 to -10.4)	0.58
Week 24	88	-17.2	91	-5.6	-11.5 (-19.7 to -3.3)	0.36
Week 36	90	-15.5	92	-8.9	-6.6 (-14.7 to 1.6)	0.20
Drug use: NSAIDs						
Week 12	93	42%	98	36%	6% (-8% to 20%)	0.12
Week 24	90	39%	94	48%	-9% (-22 to 4%)	0.18
Week 36	90	40%	93	51%	-10% (-23% to 3%)	0.22
Observed disability						
Week 12	91	-0.21	98	-0.02	-0.19 (-0.38 to -0.01)	0.28
Week 24	88	-0.13	91	-0.04	-0.09 (-0.30 to 0.12)	0.12
Week 36	88	-0.16	91	-0.06	-0.10 (-0.31 to 0.11)	0.13

\*Analyses are adjusted for baseline differences (fear avoidance beliefs towards physical activity, use of paracetamol, use of NSAIDs) and baseline score on specific outcome parameter.

†For drug use follow up scores are presented; percentage of patients taking drugs.

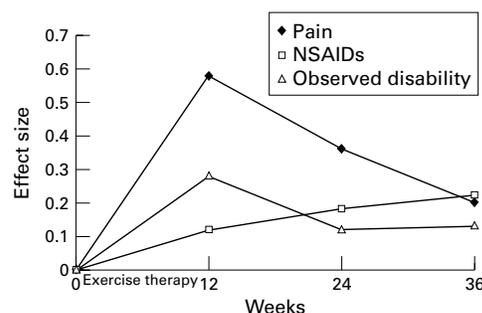


Figure 2 Effect sizes of primary outcome measures in the course of the trial.

effects were found for NSAIDs and observed disability. At 36 weeks (24 weeks after completing treatment), no differences were found between the intervention groups.

Figure 2 gives an overview of effect sizes over the course of the trial for the primary outcome measures. Beneficial effects for pain and disability were maximal at week 12, just after completion of treatment. In the follow up period the size of the effects declined to non-significant levels in week 36, except for the results for the use of NSAIDs, which remained stable during follow up.

#### SECONDARY OUTCOME MEASURES

Table 3 presents follow up results on the secondary outcome measures. At 24 weeks, beneficial effects of exercise were found for use of paracetamol and patient's global perceived effect. No effects were found for self reported disability, muscle strength, and range of motion. Similar effects were found at week 36. As with the primary outcome measures, the size of the effects declined in the follow up period (week 12 to week 36). Again, an exception is found in the use of drugs where the effects of the use of paracetamol remained stable during follow up.

Exploratory analyses were performed to study the effects of exercise on muscle strength and joint range of motion only in patients with hip or knee OA. Two of 24 tests reached significance. At week 12 patients with knee OA in the group receiving exercise treatment showed improved muscle strength of the hip.

At week 24 patients with hip OA in the control group showed improved muscle strength of the hip.

#### OUTCOME IN PER-PROTOCOL ANALYSIS

A per-protocol analysis was conducted, excluding all patients who were classified as late ineligible patients owing to a changed medical diagnosis (n=4) or owing to treatment not according to the protocol (n=21). For 13 patients (eight of whom had been allocated to the exercise treatment group) treatment in the intervention period deviated from the protocol. Treatment in the follow up period had to be restricted to a minimum. However, in eight patients (five receiving exercise treatment and three controls) a medical specialist intervened, by performing a total hip replacement (n=4), a total knee replacement (n=1), an arthroscopy (n=1), a meniscus operation (n=1), or a corticosteroid injection (n=1).

The results of the per-protocol analysis were similar to the results of the intention to treat analysis. The effect sizes were somewhat higher, as can be explained by the nature of the analysis. This applies to both primary and secondary outcome measures (data not presented).

#### PROGNOSTIC VALUE OF PATIENT CHARACTERISTICS FOR OUTCOME AFTER COMPLETION OF SUPERVISED TREATMENT

Exploratory analyses were performed to study the prognostic value of demographic characteristics, clinical features, kinesiological characteristics, lifestyle, psychological wellbeing, and compliance for the residual effectiveness of exercise treatment. A study examined whether patients who were stable or improved in the follow up period (week 12 to week 36) differed from patients who deteriorated in this period, in their baseline characteristics or their compliance with treatment. Patients were classified as being stable/improved versus having deteriorated, based on their results for two primary outcome measures (pain during the past week and observed disability) after completion of treatment (week 12 to week 36) (see statistical analysis). A classification based on NSAIDs did not result in a useful division of patients (table 4).

Sixty two tests (31 patient characteristics and two outcome measures) were performed to study differences between patients who were stable/improved and those who had deteriorated. These patients groups differed in two aspects of their baseline characteristics. Patients who were stable or showed improvement (in an observed disability) in the follow up period were more often overweight (body mass index  $\geq 30$ ) at baseline (46% v 19%, p=0.01). Patients whose pain was stable or improved in the follow up period reported lower levels on the pain coping strategy "transforming of pain" (8.3 v 10.4, p=0.001).

No significant differences were found for compliance in the supervised treatment period. Non-compliance, as reported by the patient, occurred as often in stable/improved patients as it occurred in patients who had deteriorated.

Table 3 Secondary outcome measures: improvements and differences between intervention groups\*

	Exercise therapy		Control		Difference (95% CI) exercise therapy – controls	Effect size
	n	Mean change†	n	Mean change†		
Drug use: paracetamol						
Week 12	93	35%	98	51%	-16% (-29% to -3%)	0.33
Week 24	90	39%	94	55%	-17% (-30% to -3%)	0.32
Week 36	90	41%	93	57%	-17% (-30% to -3%)	0.32
Global perceived effect: improved‡						
Week 12	91	47%	98	18%	28% (15% to 42%)	0.64
Week 24	88	49%	91	25%	24% (10% to 38%)	0.50
Week 36	90	46%	93	29%	17% (3% to 31%)	0.35
Self reported disability§						
Week 12	93	-1.1	98	-0.0	-1.1 (-2.3 to 0.1)	0.26
Week 36	90	-1.2	92	-0.3	-0.8 (-2.2 to 0.5)	0.18
Muscle strength hip						
Week 12	91	0.22	94	0.04	0.17 (0.02 to 0.33)	0.32
Week 24	85	0.16	87	0.13	0.04 (-0.14 to 0.21)	0.06
Week 36	85	0.26	87	0.22	0.04 (-0.12 to 0.21)	0.08
Muscle strength knee						
Week 12	92	0.19	98	0.06	0.13 (-0.04 to 0.29)	0.22
Week 24	88	0.14	91	0.14	0.00 (-0.17 to 0.17)	0.00
Week 36	86	0.19	90	0.19	0.00 (-0.18 to 0.17)	0.01
Joint range of motion hip						
Week 12	92	0.21	98	0.06	0.15 (-0.03 to 0.32)	0.23
Week 24	87	0.36	90	0.27	0.09 (-0.13 to 0.31)	0.11
Week 36	87	0.19	89	0.19	0.00 (-0.21 to 0.22)	0.01
Joint range of motion knee						
Week 12	92	0.17	98	0.09	0.08 (-0.09 to 0.25)	0.13
Week 24	90	0.11	94	0.15	-0.04 (-0.24 to 0.17)	0.05
Week 36	87	0.11	90	0.05	0.05 (-0.15 to 0.25)	0.10
Physical activity: improved¶						
Week 12	87	46%	97	48%	-2% (-17% to 13%)	0.04
Week 36	89	42%	91	49%	-8% (-22% to 7%)	0.15

\*Analyses are adjusted for baseline differences (fear avoidance beliefs towards physical activity, use of paracetamol, use of NSAIDs) and baseline score on specific outcome parameter.

†For drug use follow up scores are presented; percentage of patients taking drugs.

‡Results on an eight point scale are dichotomised as improved (completely recovered, very much improved, and much improved) and not improved (slightly improved, not changed, slightly worsened, much worsened, and vastly worsened).

§Not assessed at 24 week follow up.

¶Results are dichotomised as improved (increase in physical activity level, change = 1 min/week) versus not improved (stabilisation or decrease in physical activity, change = 0 min/week).

Table 4 Residual effects in patients receiving exercise treatment: results at 36 weeks, compared with results at week 12 (n=90)

	Stable or improved patients* (n)	Patients who deteriorated (n)
Pain in past week	46	44
Drug use: NSAIDs	84	6
Observed disability†	39	48

\*Results are dichotomised as stable or improved (reduction or stabilisation of pain, drug use, or disability) versus deteriorated (increase of pain or disability, start taking NSAIDs, or increase in use).

†Missing values: 3.

Table 5 Compliance with exercise treatment and its effect on the residual effect of exercise (results week 36 compared with week 12), as assessed by pain in the past week and observed disability

	Yes	No	p Value
Compliance according to the patient (n)	61	28	
Pain in past week: stable or improved (%)	54.1	46.4	0.50
Observed disability: stable or improved (%)	45.9	40.0	0.62
Compliance according to the physiotherapist (n)	73	11	
Pain in past week: stable or improved (%)	56.2	27.3	0.07
Observed disability: stable or improved (%)	46.6	22.2	0.17

However, though not significant, non-compliance, as reported by the physiotherapist, seems to occur more often in patients who deteriorated during follow up than in stable/improved patients (table 5).

## Discussion

Several randomised clinical trials have shown the beneficial effects of exercise in patients with OA immediately after completion of treatment.<sup>9-14</sup> Only a few, inconclusive studies have investigated whether effects are sustained

over time. To rectify this lack of knowledge we studied the effectiveness of exercise in patients with OA at 12 and 24 weeks after completion of treatment. Our results indicate a slow decline of the beneficial effects of exercise treatment, resulting in a similar clinical status for patients receiving exercise treatment and controls.

Our results are in line with the few earlier studies on this topic, in the sense that all studies reported some decline in the post-treatment effects.<sup>9 13-19</sup> However, in most earlier work the decline seems less pronounced both in controlled trials<sup>9 13 14 16 17</sup> and in one uncontrolled study.<sup>18</sup> Several explanations are possible. The trials of Minor<sup>14</sup> and Deyle<sup>13</sup> lacked an intention to treat analysis on follow up data. This probably resulted in an overestimation of treatment effects.<sup>41</sup> In the FAST trial,<sup>16 17</sup> treatment continued, albeit at home, until the last assessment. Therefore, no “non-training” effect could be studied. Interpretation of the results of Røgind *et al* is hampered by the high number of outcome measures, in combination with the small number of patients.<sup>9</sup> In the uncontrolled study,<sup>18</sup> four of 15 patients dropped out during the intervention. There, the remaining patients were probably a select and highly motivated group of patients, resulting in beneficial and sustaining effects.

Our results are in line with a prospective cohort study into the effects of stopping exercise programmes in healthy older adults.<sup>42</sup> The beneficial effects on cardiovascular capacity and muscle strength after a 16 week cardiovascular or resistance exercise programme decreased after a 10 week non-training period.

Exploratory analyses were made on the prognostic value of patient characteristics for long term outcome. A beneficial long term outcome was found in patients who were overweight at baseline and in patients with a relatively low level of the specific pain coping strategy (transformation of pain). However, given the high number of tests performed (n=62), it is entirely possible that these results are due to chance. A more thorough analysis of the factors affecting beneficial long term outcome is outside the scope of our study. However, future research should focus on these prognostic factors. A thorough analysis of the underlying mechanisms is needed, including analyses of interaction of these factors. We did not find a relation between compliance and beneficial outcome after completion of treatment. However, we did not have data on long term compliance—that is, on compliance after completion of the exercise treatment. Therefore, we have no insight into the highly relevant relation between long term compliance and outcome after completion of treatment. In future trials long term compliance should be studied.

The clinical implications of our results for the application of exercise treatment in patients with OA are clear. The evidence available clearly indicates the usefulness of exercise in patients with OA of the hip or knee.<sup>8-13</sup> Small to moderate beneficial effects on pain and disability can be produced, which are the main symptoms of OA. However, the effects slowly

decline, indicating that measures must be taken to maintain the beneficial effects of exercise. One measure is to introduce some form of retraining or prolonged training. Both intermittent exercise treatment and follow up booster sessions can be applied. There are some indications that the content and timing of an exercise programme affects results. In the FAST trial the relation between compliance and outcome was studied.<sup>43</sup> The most beneficial outcome was found in patients with OA who did aerobic exercises three days a week for about 35 minutes each session. Patients spending more time exercising during a session had outcomes similar to those of non-exercising patients. These results are based on a secondary analysis of the dose-response relation between compliance and outcome. However, to our knowledge, no clinical studies on this topic are available as yet. The optimisation of content and timing of exercise treatment deserves further study.

Another measure is to maximise the patient's compliance with exercise, even after having completed a supervised programme. An important component of an exercise programme should be to facilitate acceptance and practice of exercises. Sluijs and Knibbe distinguished between short term and long term compliance and, subsequently, different approaches to be applied by healthcare providers.<sup>44</sup> To improve long term compliance, a self regulated approach seems useful. In future trials, explicit attention should be paid to improvement of both short term and long term compliance.

In conclusion, exercise is effective in patients with OA of the knee or hip. However, these effects decline over time and finally disappear. Future research should focus on finding ways of optimally maintaining beneficial effects over time. Important issues are optimal content and timing of treatment and compliance of the patient with exercise therapy.

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