

1 **Effect of exercise interventions in the early phase to improve physical**
2 **function after hip fracture – a systematic review and meta-analysis**

3

4 **Authors, professional and academic affiliations:**

5 Monica Beckmann, RPT, MSc ^{1,2}, Vigdis Bruun-Olsen, RPT, PhD ¹, Are Hugo Pripp, PhD ^{3,4},
6 Astrid Bergland, RPT, PhD ³, Toby Smith, PhD ⁵, Kristi E. Heiberg, RPT, PhD ^{1,3}

7

8 **Institutions:**

9

10 ¹Department of Medical Research, Bærum Hospital, Vestre Viken Hospital Trust

11 ²University of Oslo

12 ³OsloMet – Oslo Metropolitan University

13 ⁴Oslo Centre of Biostatistics and Epidemiology Research Support Services, Oslo University
14 Hospital

15 ⁵ Nuffield Department of Orthopaedics, Rheumatology and Musculoskeletal Sciences,
16 University of Oxford, UK.

17

18

19

20 **Corresponding author:**

21 **Monica Beckmann**

22 Department of Medical Research, Clinic of Bærum Hospital, Vestre Viken

23 3004 Drammen

24 Norway

25 **Phone:** 0047 03525

26 **E-mail:** mobeck@vestreviken.no

27 [Word count: 2413](#)

28 **Abstract**

29 **Background:** The efficacy of exercise interventions in the early recovery phase, i.e. started
30 within the first three months after hip fracture, has been poorly studied compared to prolonged
31 exercise interventions.

32 **Objective:** To examine the effect of exercise interventions to improve physical function in the
33 early phase after hip fracture.

34 **Data sources:** Seven databases including MEDLINE via Ovid, The Cochrane Library, Embase,
35 Cinahl, Pedro, AMED and Web of Science were comprehensively searched till December 2019.

36 **Eligibility criteria:** Randomised controlled trials (RCTs) of exercise interventions initiated
37 within the first three months after hip fracture to improve physical function, were eligible for
38 inclusion. Primary outcome was physical function assessed using walking ability, walking
39 speed, balance, muscle strength, mobility, and endurance.

40 **Data extraction and Data Synthesis:** We conducted subgroup analyses specifically to
41 investigate outcomes of these individual measurements. A meta-analysis was conducted to
42 examine the overall effect of early exercise interventions. A meta-regression was conducted to
43 examine the impact of study characteristic on exercise interventions. We used the PEDro score
44 to determine quality of the included studies.

45 **Results:** Nine studies (669 patients) were included. Despite high statistical heterogeneity, there
46 was high to moderate quality evidence that exercise provided benefit in improving physical
47 function (standardised mean difference (SMD) 1.07; 95% CI 0.44 – 1.70; $p < 0.001$). There was
48 no statistically significant difference in outcome, when measured by the individual physical
49 function outcome ($p > 0.05$). Meta-regression demonstrated no statistically significant
50 association between study characteristics and exercise interventions ($p > 0.05$).

51 **Conclusion:** Exercise in the early phase of hip fracture rehabilitation can improve physical
52 function. It remains unclear what type of exercise is superior in the early phase after hip fracture.

53 **Limitations:** This conclusion should be interpreted with caution given the high statistical
54 heterogeneity reported and non-significant subgroup analyses of specific physical function
55 measures, which were underpowered.

56 **Funding:** Vestre Viken HF, Norway.

57 **Keywords:** hip fracture, exercise, physical function, early phase, review, meta-analysis.

58 **Protocol Registration (PROSPERO):** CRD42018091135

59

60 **Introduction**

61 Hip fracture is the most serious fall-related fracture with a mortality rate of 5-10% during the
62 first month and 20-30% during the first year post-fracture [1]. In 2000 there was reported 1.6
63 million hip fractures worldwide, accounting for approximately 20% of all fractures in people
64 aged 50 years and older [2]. Age-adjusted rates of hip fractures are highest in Scandinavian and
65 North-American populations and lower in Southern Europe [3].

66 Hip fracture frequently leads to reduced physical function [4]. Physical function is the
67 capacity of an individual to perform their physical activities of daily living. It reflects motor
68 function and control, physical fitness, and habitual physical activity [5], and it is a strong
69 measure of biological age and a biomarker for health and quality of life in older people [6-9].
70 Physical function as an assessment domain is measured through various approaches including:
71 mobility, endurance, muscle strength, and balance [10]. Physical performance assessments as
72 outcome measures are vital when the effect of interventions of physical function are examined
73 [11].

74 Patients who experience a hip fracture are often vulnerable and fragile [12]. In the early
75 phase after hip fracture, reduced mobility and hospitalisation may cause severe decline in a
76 patient's muscle mass, muscle strength, and consequently, their physical function [13]. This
77 can lead to reduced independence with increased need of assistance in daily tasks [14].
78 Furthermore, these patients are at high risk of increased fear of falling, muscle weakness, and
79 post-operative pain through reduced mobility and loss of independence [12].

80 Exercise interventions after hip fracture aim to improve physical function and prevent
81 or reverse physical deconditioning [15]. However, a Cochrane systematic review and meta-
82 analysis on hip fracture recovery reported inconsistent effects of exercise interventions on
83 mobility [16]. A recent meta-analysis reported a small improvement in overall mobility

84 following a structured exercise intervention [4]. These meta-analyses included studies with
85 exercise interventions up to one year after the fracture [4, 16]. The efficacy of exercise
86 interventions in the early recovery phase, i.e. started within the first three months after hip
87 fracture, has been poorly studied [17].

88 The aim of this systematic review and meta-analysis is to determine the effectiveness of
89 exercise interventions on physical function in the early phase after hip fracture.

90

91 **Methods**

92 *Data Sources and Search Strategy for Identification of Studies*

93 This review was performed according to the Preferred Reporting Items for Systematic Reviews
94 and Meta-Analyses (PRISMA) guidelines [18]. The following electronic databases were
95 searched in May 2018 (updated December 2018): The Cochrane Database of Systematic
96 Reviews, Medline, Embase, Cinahl, PEDro, and AMED. Search strategies are shown in

97 **Supplementary File 1.**

98

99 *Eligibility Criteria*

100 Studies were included based on the following eligibility criteria: all randomised controlled
101 trials (RCTs) assessing the efficacy or effectiveness of exercise interventions commenced
102 within the first three months post-hip fracture surgery, for patients aged 65 years and older.
103 We define exercise interventions as interventions that include physical activity to improve or
104 maintain one or more components of physical function. Included studies reported at least one
105 performance based outcome measure of physical function, such as walking ability, walking
106 speed, strength, balance, mobility, and endurance. Studies investigating muscle stimulation,

107 passive management strategies, self-training interventions, and welfare technology studies
108 were excluded. We excluded studies that did not have a PEDro score of level 1 evidence (6 –
109 10 points).

110

111

112 *Article screening and selection*

113 Titles and abstracts from all individual citations were screened by one reviewer (MB) with
114 duplicates removed. One reviewer (MB) read all the included full-text articles and three
115 reviewers (KEH, VB-O, AB) read each a third of the selected full-text articles. Any
116 disagreements between the reviewers were addressed through discussion until consensus was
117 met.

118

119 *Data extraction*

120 For each included study, we extracted data using a specifically developed data extraction form
121 (**Supplementary File 2**). One reviewer extracted all data (MB). The extraction was verified by
122 one of the three other reviewers (KEH, VB-O, AB). Data extracted included: study origin,
123 interventions (exercise components, duration, frequency, and intensity), outcome measures, and
124 results. When differences in opinion occurred between the reviewers, consensus was met
125 through discussion.

126

127

128

129

130 *Quality assessment*

131 The PEDro score was used as each included study's methodological quality [19, 20]. A study
132 with a PEDro score of ≥ 6 was considered Level 1 evidence (6-8: good, 9-10: excellent) and
133 included in the review. [21]. One reviewer independently assessed the quality of the included
134 studies (MB). This was verified by one of three reviewers (KEH, VB-O, AB).

135

136 *Statistical Analysis*

137 Study heterogeneity was assessed by examining the data extraction tables. The reviewers
138 determined that the trial designs, population characteristics, and interventions were
139 satisfactorily homogeneous to permit a meta-analysis. Based on this, a random-effect model
140 meta-analysis was deemed appropriate. We calculated standardised mean differences (SMD)
141 with 95% confidence intervals (CIs) to assess pooled physical function outcomes. The
142 interpretations of the SMD as an effect size were based on Cohen's d (0.2 is small, 0.5 to 0.6 is
143 moderate, and 0.8 to 1.0 is large) [22]. I^2 statistics, p -values, Q statistics, and degree of freedom
144 were used to assess statistical heterogeneity [23, 24].

145 The primary analysis was the overall effect of exercise interventions on physical
146 function. When more than one physical function outcome was reported in a study, the outcome
147 measure considered most related to the intervention was selected, according to the principle of
148 specificity [25]. The estimates of SMD for the first post-intervention time point were extracted
149 for meta-analysis. Subgroup analyses were undertaken to assess the specific outcomes within
150 the domain of physical function i.e. walking speed, mobility, balance, and endurance.

151 We assessed for any association between study characteristics (study year, hospital or
152 community setting, days since surgery, sample size, mean age of participants, Pedro score of

153 the study, intensity of intervention, and follow-up weeks) and exercise intervention (strength
154 training versus non-strength training) through a meta-regression. Stata version 15 (Stata Corp,
155 Texas, USA) was used with the meta-analysis metan [26], metafunnel [27] and meta-regression
156 [28] packages.

157

158 **Results**

159 *Description of included studies*

160 The results of the search strategy are presented in Figure 1. In total, 2225 studies were identified
161 after duplicates were removed. From the 26 potentially eligible studies, nine met the eligibility
162 criteria and were included in the review.

163 The characteristics of the included studies are reported in Table 1. In total 669
164 participants with a mean age of 81 years (mean age range: 77 to 84 years) were included.
165 Included studies were published from 2002 to 2018. The sample sizes for the individual studies
166 varied from 20 to 160 participants. Three studies were undertaken in Australia, whereas single
167 studies originated from Italy, United States of America, Canada, Denmark, Netherlands, and
168 Germany.

169 The included studies examined combinations of exercise interventions (**Table 1**). Five
170 studies examined the effect of high-intensity exercise. Of these, two studies investigated the
171 effect of high-intensity physiotherapy [29, 30], three studies investigated the effect of high-
172 intensity progressive resistance training [31-33], two studies examined endurance training [34,
173 35], one study examined weight-bearing exercise [36], and one study examined balance task-
174 specific training [37]. Control groups for each study are presented in **Table 1**. Across the
175 studies, the number of sessions varied from 10-36 sessions. The overall duration of

176 interventions in the included studies varied from 1-12 weeks. In three studies, the interventions
177 were delivered to participants during their hospital stay [29, 32, 36] with intervention start at
178 mean nine days post-operatively (range 5-19 days after surgery). In three studies, the exercises
179 were delivered in a community setting [34, 35, 37] with intervention start at mean ten days post-
180 operative (range 5-14 days since surgery). Three studies delivered exercise interventions across
181 both hospital and community settings [30, 31, 33], commencing at a mean 45 days post-
182 operatively (range 14-90 days since surgery).

183

184 *Principal analysis*

185 We pooled data from four different measures of physical function (mobility, walking speed,
186 endurance, and balance). There was high to moderate quality evidence that exercise provided
187 benefit in improved physical function at three to six months post-operatively (SMD: 1.07; 95%
188 CI = 0.44 – 1.70; $p < 0.001$. $I^2 = 92.3\%$, $Q = 103.66$) (**Figure 2**). Because of the very large
189 effect in one study [37], we additionally performed a sensitivity meta-analysis excluding this
190 study. The meta-analysis still reported a statistically significant overall improvement in favour
191 of exercise (SMD: 0.36; 95% CI = 0.05 – 0.67; $p < 0.024$).

192

193 *Subgroup analyses*

194 Mobility as assessed with the Timed Up and Go (TUG) test was reported in five high-quality
195 studies ($n = 296$) [29, 31, 32, 34, 35]. The studies were of high quality according to the PEDro
196 score (**Table 2**). There was no statistically significant difference in TUG results between the
197 exercise and comparison groups (SMD: 0.48; 95% CI: -0.14 – 1.10; $p = 0.126$; $I^2 = 82\%$, $Q =$
198 21.76) (**Figure 3**).

199 Walking speed was assessed using the 6-meter and 10-meter walk test in four high-
200 quality studies (n = 334) [30, 31, 35, 36]. There was no statistically significant difference in
201 walk test results between the exercise and comparison groups (SMD: 0.35; 95% CI: -0.04 –
202 0.73; $p = 0.078$; $I^2 = 57%$, $Q = 7.01$) (**Figure 3**).

203 Endurance was measured using number of meters per minute, such as 2-minute walking
204 test, in two high-quality studies (n = 110) [33, 34]. There was no statistically significant
205 difference between the exercise and comparison groups (SMD: 1.52, 95% CI: -0.75 – 3.79; p
206 = 0.189; $I^2 = 92%$, $Q = 11.80$) (**Figure 3**).

207 Balance measured using Berg Balance scale was reported in three high-quality studies
208 (n = 162) [33, 34, 37]. There was no statistically significant difference in balance function
209 between the exercise and comparison groups (SMD: 2.84, 95% CI: -0.25 – 5.93; $p = 0.071$; I^2
210 = 97%, $Q = 69.09$) (**Figure 3**).

211

212 *Meta-regression*

213 The meta-regression indicated no statistically significant association between study
214 intervention characteristics and exercise interventions ($p > 0.05$).

215

216 **Discussion**

217 The overall results suggest that exercise interventions prescribed in the early phase of recovery
218 after hip fracture, improve physical function. However, within the subgroup analyses of
219 mobility, walking speed, endurance, and balance no effects were demonstrated, compared to
220 control groups. The meta-analyses presented high statistical heterogeneity and should therefore

221 be interpreted with caution. The meta-regression could not identify an exercise intervention that
222 was superior in association with physical function.

223 To our knowledge, this is the first systematic review and meta-analysis to examine the
224 effectiveness of exercise interventions in the early phase after hip fracture. The overall
225 improvement in physical function compared to the lack of improvement in the subgroup
226 analyses may be explained through various reasons. Firstly, the sub-group analyses were
227 underpowered. The non-statistical difference may therefore reflect a type-2 statistical error.
228 With an assumed clinically significant effect of 0.5 SMD, high heterogeneity between studies
229 and an average study size of 50 participants in each group, we estimated that five to six studies
230 would be needed to obtain around 80% statistical power [38]. This may have been magnified
231 by a potential reduced between-group difference in physical function since both experimental
232 and control interventions offered an exercise intervention. As the evidence-base developed,
233 further exploration of the specific instruments used to assess physical function is desirable, to
234 better interpret the results reported in this analysis.

235 The results of this review are in agreement with a recent meta-analysis reporting
236 outcomes at a later-stage in hip fracture recovery [39]. The meta-analysis reported that balance
237 training from early to the chronic phase after hip fracture improved overall physical function,
238 such as balance, walking, lower limb strength, ADLs, performance task score, and health related
239 quality of life scores [39]. The authors highlight that balance exercises were particularly
240 effective at this later-stage, but it was not possible to ascertain whether a specific exercise
241 programme offered superior outcomes over another [39]. However, we can surmise that
242 exercise in this phase of recovery is beneficial. Further research to reveal whether there are
243 differences in outcome by specific exercise interventions are warranted.

244 Timing may be a confounding factor on the effect of exercise interventions after hip
245 fracture. Binder et al. (2004) and Hauer et al. (2002) showed that progressive resistance

246 exercise was an effective intervention for improving physical function after hip fracture [31,
247 33]. In the study by Binder et al. (2004) the intervention started at mean 90 days after the hip
248 fracture surgery [33]. Hauer et al. (2002) started their intervention 6-8 weeks after hip fracture
249 surgery. Physiological timing may be assumed to be a potential confounding factor where
250 pain and healing after surgery may impact on the patient's ability to engage with exercise and
251 therefore outcomes.

252 A systematic review by Cadore et al. (2013) reported that a multicomponent exercise
253 program consisting of strength training, endurance, and balance appeared to be the best
254 strategy to improve physical function and maintain functional capacity in older adults [40].
255 Given the results from Cadore et al. and the summary of results above, one may argue that a
256 wide scope of exercise can be beneficial for these elderly patients with hip fracture. The
257 limited association reported in our review may have been attributed to type-2 errors. Further
258 research should be undertaken to better reveal the potential impact of timing of different
259 exercise interventions and whether they influence outcomes in the early phase following hip
260 fracture.

261

262 *Limitations*

263 This systematic review and meta-analysis is presented with some limitations that should be
264 considered. Firstly, only a single assessor (MB) screened all citations. Therefore it may be a
265 higher risk that the single assessor, instead of three assessors, missed out of relevant studies.
266 Secondly, given the low number of studies identified, it was not possible to assess for the risk
267 of publication bias through a funnel plot or statistical means. Thirdly, there was sufficient
268 homogeneity to pool the data in a meta-analysis. However, we acknowledge high statistical

269 heterogeneity across the meta-analyses indicating unknown between-trial variability. The
270 results of the meta-analysis should therefore be viewed with caution.

271

272 **Conclusion**

273 In this systematic review and meta-analysis evidence from early phase exercise after hip
274 fracture was evaluated. Based on moderate to high-quality evidence, exercise interventions
275 could have the potential to improve the patients' physical function after hip fracture. The results
276 should be interpreted with caution due to high statistical heterogeneity and underpowered
277 subgroup analyses. The clinical implications from our results suggested that different types of
278 exercise could be beneficial in the early phase after hip fracture.

279

280 **Key messages:**

- 281 • Exercise has the potential to improve the patients' physical function in the early
282 phase after hip fracture
- 283 • It remains unclear what type of exercise is superior for this population in this
284 phase
- 285 • More research is warranted to elaborate on these conclusions

286

287

288 *Conflict of interest:* There are no conflicts of interest

289

290 **References**

- 292 1. Sogaard AJ, Holvik K, Meyer HE, Tell GS, Gjesdal CG, Emaus N, et al. Continued decline in hip
293 fracture incidence in Norway: a NOREPOS study. *Osteoporosis int.* 2016;27(7):2217-22.
- 294 2. Johnell O, Kanis JA. An estimate of the worldwide prevalence and disability associated with
295 osteoporotic fractures. *Osteoporosis int.* 2006;17(12):1726-33.
- 296 3. Sambrook P, Cooper C. Osteoporosis. *The Lancet.* 2006;367(9527):2010-8.
- 297 4. Diong J, Allen N, Sherrington C. Structured exercise improves mobility after hip fracture: a
298 meta-analysis with meta-regression. *Br J Sports Med.* 2016;50(6):346-55.
- 299 5. Garber CE, Blissmer B, Deschenes MR, Franklin BA, Lamonte MJ, Lee IM, et al. Quantity and
300 quality of exercise for developing and maintaining cardiorespiratory, musculoskeletal, and
301 neuromotor fitness in apparently healthy adults: guidance for prescribing exercise. *Med Sci Sports
302 Exerc.* 2011;43(7):1334-59.
- 303 6. Pavasini R, Guralnik J, Brown JC, di Bari M, Cesari M, Landi F, et al. Short Physical
304 Performance Battery and all-cause mortality: systematic review and meta-analysis. *BMC Med.*
305 2016;14(1):215.
- 306 7. Vestergaard S, Patel KV, Bandinelli S, Ferrucci L, Guralnik JM. Characteristics of 400-meter
307 walk test performance and subsequent mortality in older adults. *Rejuvenation Res.* 2009;12(3):177-
308 84.
- 309 8. Cesari M, Kritchevsky SB, Newman AB, Simonsick EM, Harris TB, Penninx BW, et al. Added
310 value of physical performance measures in predicting adverse health-related events: results from the
311 Health, Aging And Body Composition Study. *J Am Geriatr Soc.* 2009;57(2):251-9.
- 312 9. Halaweh H, Willen C, Grimby-Ekman A, Svantesson U. Physical Activity and Health-Related
313 Quality of Life Among Community Dwelling Elderly. *J Clin Med Res.* 2015;7(11):845-52.
- 314 10. Brovold T, Skelton DA, Bergland A. Older adults recently discharged from the hospital: effect
315 of aerobic interval exercise on health-related quality of life, physical fitness, and physical activity. *J
316 Am Geriatr Soc.* 2013;61(9):1580-5.
- 317 11. Guralnik JM, Simonsick EM, Ferrucci L, Glynn RJ, Berkman LF, Blazer DG, et al. A short physical
318 performance battery assessing lower extremity function: association with self-reported disability and
319 prediction of mortality and nursing home admission. *J Gerontol.* 1994;49(2):M85-94.
- 320 12. Auais MA, Eilayyan O, Mayo NE. Extended exercise rehabilitation after hip fracture improves
321 patients' physical function: a systematic review and meta-analysis. *Phys Ther.* 2012;92(11):1437-51.
- 322 13. Suetta C, Magnusson SP, Rosted A, Aagaard P, Jakobsen AK, Larsen LH, et al. Resistance
323 training in the early postoperative phase reduces hospitalization and leads to muscle hypertrophy in
324 elderly hip surgery patients--a controlled, randomized study. *J Am Geriatr Soc.* 2004;52(12):2016-22.
- 325 14. Taraldsen K, Sletvold O, Thingstad P, Saltvedt I, Granat MH, Lydersen S, et al. Physical
326 behavior and function early after hip fracture surgery in patients receiving comprehensive geriatric
327 care or orthopedic care--a randomized controlled trial. *J Gerontol A Bio Sci Med Sci.* 2014;69(3):338-
328 45.
- 329 15. Ftouh S, Morga A, Swift C. Management of hip fracture in adults: summary of NICE guidance.
330 *BMJ (Clinical research ed).* 2011;342:d3304.
- 331 16. Handoll HH, Sherrington C, Mak JC. Interventions for improving mobility after hip fracture
332 surgery in adults. *Cochrane Database Syst Rev.* 2011(3):Cd001704.
- 333 17. Asplin G, Carlsson G, Ziden L, Kjellby-Wendt G. Early coordinated rehabilitation in acute
334 phase after hip fracture - a model for increased patient participation. *BMC Geriatrics.*
335 2017;17(1):240.
- 336 18. Moher D, Shamseer L, Clarke M, Ghersi D, Liberati A, Petticrew M, et al. Preferred reporting
337 items for systematic review and meta-analysis protocols (PRISMA-P) 2015 statement. *Systematic
338 reviews.* 2015;4:1.
- 339 19. de Morton NA. The PEDro scale is a valid measure of the methodological quality of clinical
340 trials: a demographic study. *Aust J Physiother.* 2009;55(2):129-33.

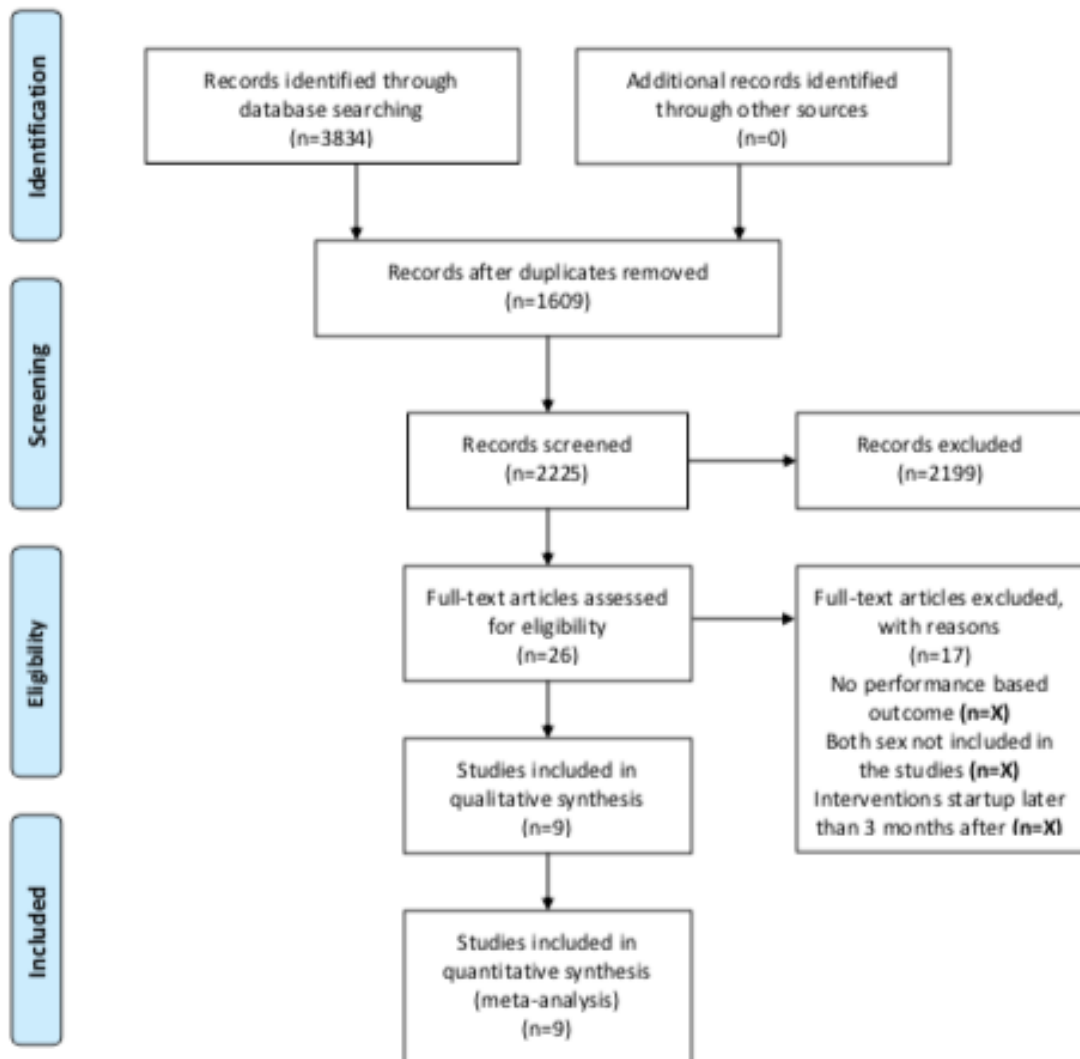
- 341 20. Maher CG, Sherrington C, Herbert RD, Moseley AM, Elkins M. Reliability of the PEDro scale
342 for rating quality of randomized controlled trials. *Phys Ther.* 2003;83(8):713-21.
- 343 21. Foley NC, Teasell RW, Bhogal SK, Speechley MR. Stroke Rehabilitation Evidence-Based
344 Review: methodology. *Top Stroke Rehabil.* 2003;10(1):1-7.
- 345 22. Cohen J. *Statistical power analysis for the behavioral sciences.* Rev. ed. ed. New York:
346 Academic Press; 1977.
- 347 23. Borenstein M HL, Higgins JPT, et al. *Introduction to meta-analysis.* Wiltshire, UK: John Wiley
348 & Sons Ltd. 2009.
- 349 24. Higgins JP, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analyses.
350 *BMJ (Clinical research ed).* 2003;327(7414):557-60.
- 351 25. Hawley JA. Specificity of training adaptation: time for a rethink? *Journal Physiol.*
352 2008;586(1):1-2.
- 353 26. Harris R, Bradburn M, Deeks J, Harbord R, Altman D, Steichen T, et al. METAN: Stata Module
354 for Fixed and Random Effects Meta-analysis. Boston College Department of Economics, Statistical
355 Software Components series. 2007. p. 3-28 .
- 356 27. Sterne J. METAFUNNEL: Stata module to produce funnel plots for meta-analysis. Statistical
357 Software Components S434101. Boston College Department of Economics. 2003.
- 358 28. Harbord R, Higgins J. METAREG: Stata module to perform meta-analysis regression.
359 Statistical Software Components S446201. Boston College Department of Economics. 2004.
- 360 29. Kimmel LA, Liew SM, Sayer JM, Holland AE. HIP4Hips (High Intensity Physiotherapy for Hip
361 fractures in the acute hospital setting): a randomised controlled trial. *Med Aust.* 2016;205(2):73-8.
- 362 30. Moseley AM, Sherrington C, Lord SR, Barracough E, St George RJ, Cameron ID. Mobility
363 training after hip fracture: a randomised controlled trial. *Age Ageing.* 2009;38(1):74-80.
- 364 31. Hauer K, Specht N, Schuler M, Bartsch P, Oster P. Intensive physical training in geriatric
365 patients after severe falls and hip surgery. *Age Ageing.* 2002;31(1):49-57.
- 366 32. Kronborg L, Bandholm T, Palm H, Kehlet H, Kristensen MT. Effectiveness of acute in-hospital
367 physiotherapy with knee-extension strength training in reducing strength deficits in patients with a
368 hip fracture: A randomised controlled trial. *PloS One.* 2017;12(6):e0179867.
- 369 33. Binder EF, Brown M, Sinacore DR, Steger-May K, Yarasheski KE, Schechtman KB. Effects of
370 extended outpatient rehabilitation after hip fracture: a randomized controlled trial. *JAMA.*
371 2004;292(7):837-46.
- 372 34. Mendelsohn ME, Overend TJ, Connelly DM, Petrella RJ. Improvement in aerobic fitness
373 during rehabilitation after hip fracture. *Arch Phys Med Rehabil.* 2008;89(4):609-17.
- 374 35. van Ooijen MW, Roerdink M, Trekop M, Janssen TW, Beek PJ. The efficacy of treadmill
375 training with and without projected visual context for improving walking ability and reducing fall
376 incidence and fear of falling in older adults with fall-related hip fracture: a randomized controlled
377 trial. *BMC Geriatrics.* 2016;16(1):215.
- 378 36. Sherrington C, Lord SR, Herbert RD. A randomised trial of weight-bearing versus non-weight-
379 bearing exercise for improving physical ability in inpatients after hip fracture. *Aust J Phys.*
380 2003;49(1):15-22.
- 381 37. Monticone M, Ambrosini E, Brunati R, Capone A, Pagliari G, Secci C, et al. How balance task-
382 specific training contributes to improving physical function in older subjects undergoing
383 rehabilitation following hip fracture: a randomized controlled trial. *Clin Rehabil.* 2018;32(3):340-51.
- 384 38. Valentine JC, Pigott TD, Rothstein HR. How Many Studies Do You Need?: A Primer on
385 Statistical Power for Meta-Analysis. *J Educational Behav Stat.* 2010;35(2):215-47.
- 386 39. Lee SY, Jung SH, Lee SU, Ha YC, Lim JY. Effect of Balance Training After Hip Fracture Surgery: A
387 Systematic Review and Meta-analysis of Randomized Controlled Studies. *JGeront A, Biol Sci Med Sci.*
388 2019;74(10):1679-85.
- 389 40. Cadore EL, Rodriguez-Manas L, Sinclair A, Izquierdo M. Effects of different exercise
390 interventions on risk of falls, gait ability, and balance in physically frail older adults: a systematic
391 review. *Rejuvenation Res.* 2013;16(2):105-14.

Table 1. Characteristics of included trials

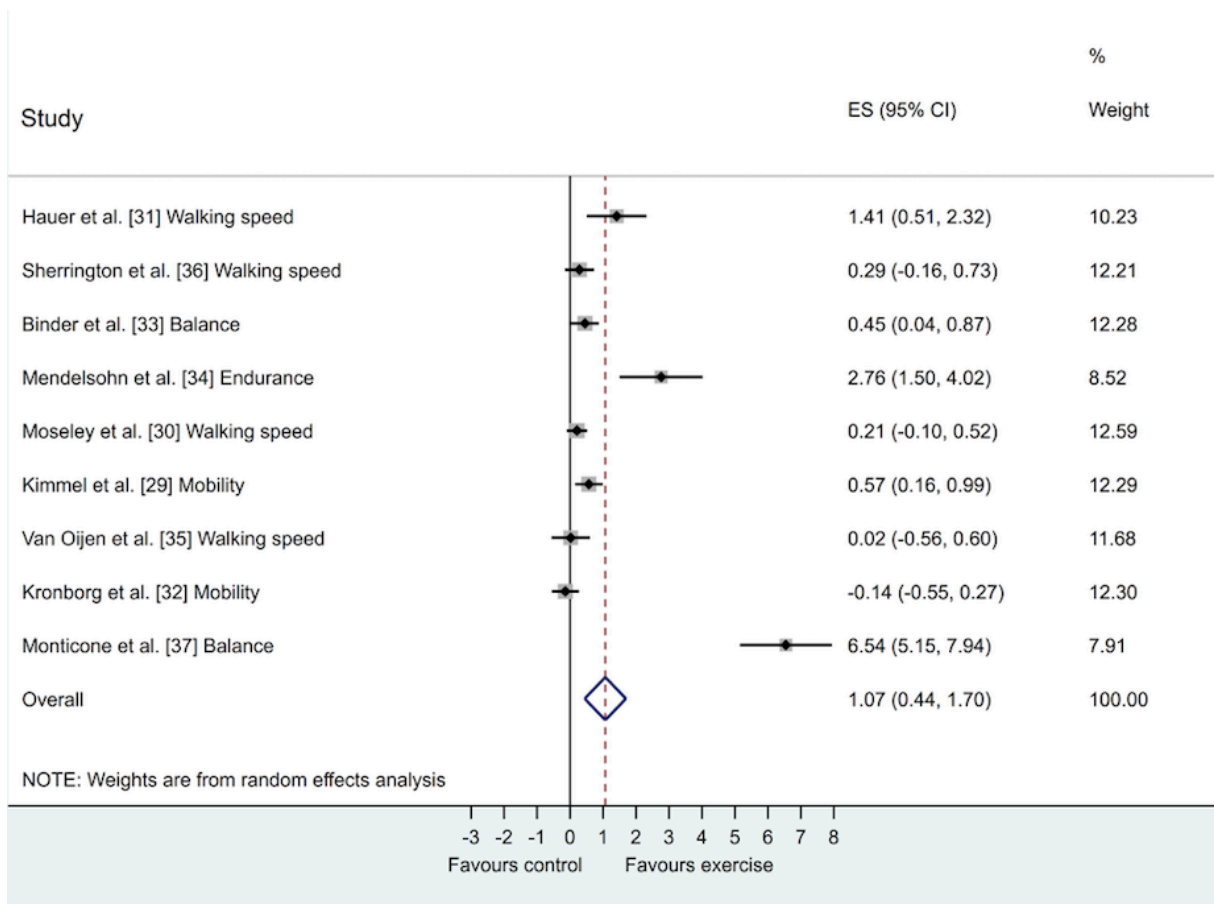
| | Setting | Start of intervention/Days since surgery | Sample size | Mean Age (years) | Pedro Score | Outcome | Characteristics of intervention | Comparator | Number of sessions | Duration of intervention (weeks) |
|--------------------------------------|----------------------|--|-------------|------------------|-------------|---|--|---|--------------------|----------------------------------|
| Binder et al 2004 ²⁵ | Hospital & Community | 90 | 90 | 81 | 7 | (S) Walking endurance m/min (S) Bergs balance scale | High intensive progressive resistance training of lower limb | Low intensity non-progressive strength training of lower limb | 36 sessions | 12 |
| Hauer et al 2002 ²³ | Hospital & Community | 42 - 56 | 24 | 81 | 7 | (S) Timed Up & Go (S) Max walking speed m/s | High intensity progressive resistance training of lower limb | Placebo motor activities such as calisthenics, games and memory tasks whilst seated | 36 sessions | 12 |
| Kimmel et al 2016 ²¹ | Hospital | 5 | 92 | 81 | 8 | (S) Timed Up & Go | Intensive physiotherapy consisting of strength exercises and gait re-training three times daily | Physiotherapy consisting of strength exercises and gait re-training one time daily | 21 sessions | 1 |
| Kronborg et al 2017 ²⁴ | Hospital | 3 | 90 | 79 | 8 | (S) Timed Up & Go | Progressive knee-extension strength training in addition to basic mobility and exercise therapy aimed at lower extremities | Basic mobility and exercise therapy aimed at lower extremities | 14 sessions | 2 |
| Mendelsohn et al 2008 ²⁶ | Community | 5 | 20 | 81 | 7 | (S) Timed Up & Go (S) Walking endurance m/min (S) Bergs balance scale | Endurance training by using an arm crank ergometer in addition to physical and occupational therapy 5 times a week | Physical and occupational therapy consisting of balance, strength, gait, flexibility and ADLs 5 times a week | 12 sessions | 4 |
| Monticone et al 2018 ²⁹ | Community | 10 | 52 | 77 | 8 | (S) Bergs balance scale | Balance task-specific training | General physiotherapy including open kinetic chain exercises and walking | 15 sessions | 3 |
| Moseley et al 2009 ²⁷ | Hospital & Community | 14 | 160 | 84 | 8 | (P) Walking speed m/s | Weight bearing exercises twice daily for a total of 60 minutes per day | Exercises sitting or lying in addition to a small amount of walking for a total of 30 minutes per day | 28 sessions | 4 |
| Sherrington et al 2003 ²⁸ | Hospital | 19 | 80 | 81 | 7 | (S) Walking speed m/s | Weight bearing exercises in addition to usual physiotherapy such as walking, bed mobility sit to stand and stair climbing | Non weight bearing exercises in addition to usual physiotherapy such as walking, bed mobility sit to stand and stair climbing | 10 sessions | 2 |
| Van Dijen et al 2016 ²⁷ | Community | 14 | 70 | 83 | 7 | (P) Timed Up & Go (S) Walking speed m/s | Endurance training, adaptability treadmill training | Conventional treadmill training | 30 sessions | 6 |

(P) = Primary, (S) = Secondary

Figure 1: Flow chart of the screening and selection process



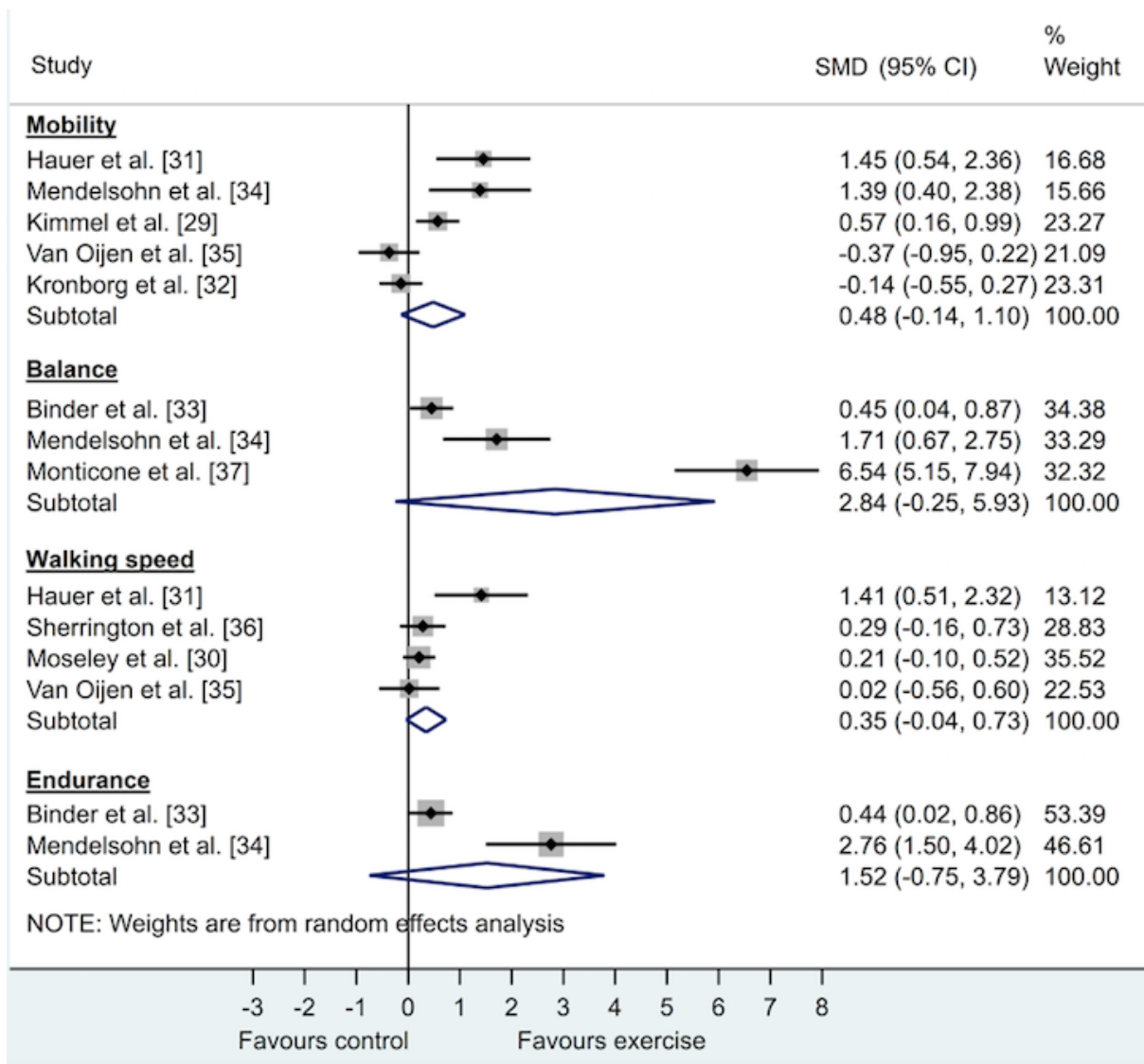
398 Figure 2



399

400

401 Figure 3



402

Search history

Database: Ovid MEDLINE(R) Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R) 1946 to Present

Date: 18.05.2018, updated November 2018

Hits: 654

| # | Searches | Results |
|----|---|---------|
| 1 | Hip Fractures/ | 14000 |
| 2 | Femoral Fractures/ | 15310 |
| 3 | Femoral Neck Fractures/ | 8240 |
| 4 | ((hip or femur or femoral or trochanter* or subtrochanter* or intertrochanter* or pertrochanter* or acetabulum) adj3 fracture*).tw,kw,kf. | 36008 |
| 5 | or/1-4 | 46927 |
| 6 | physical therapy modalities/ | 33689 |
| 7 | movement/ | 68193 |
| 8 | gait/ | 24164 |
| 9 | locomotion/ | 22939 |
| 10 | Exercise Therapy/ | 34888 |
| 11 | exercise/ | 91826 |
| 12 | motor activity/ | 92338 |
| 13 | Walking/ | 28384 |
| 14 | Early Ambulation/ | 2549 |
| 15 | rehabilitation/ | 17613 |
| 16 | muscle strength/ | 15566 |
| 17 | postural balance/ | 19931 |
| 18 | resistance training/ | 6429 |
| 19 | (exercise* or walking or training or retraining or mobil* or locomotion or gait or balanc* or physiotherap* or physio therap* or physical therap* or weight bearing or physical activit*).tw,kw,kf. | 1171214 |
| 20 | or/6-19 | 1366259 |
| 21 | 5 and 20 | 6421 |
| 22 | AGED/ | 2778152 |
| 23 | Frail Elderly/ | 9484 |
| 24 | "AGED, 80 AND OVER"/ | 798955 |
| 25 | (aged or elder or elders or elderly or old or older or olds or geriatric patient*).tw,kw,kf. | 1772821 |
| 26 | or/22-25 | 4055039 |
| 27 | 21 and 26 | 4604 |

| | | |
|-----|--|---------|
| S12 | (MH "Motor Activity") | 4,969 |
| S13 | (MH "Walking") | 13,112 |
| S14 | (MH "Early Ambulation") | 656 |
| S15 | (MH "Rehabilitation") | 12,314 |
| S16 | (MH "Muscle Strength") | 12,172 |
| S17 | (MH "Muscle Strengthening") | 9,991 |
| S18 | (MH "Resistance Training") | 2,009 |
| S19 | (MH "Weight-Bearing") | 3,384 |
| S20 | (MH "Balance Training, Physical") | 825 |
| S21 | T1 ((exercise* or walking or training or retraining or mobili* or locomotion or gait or balanc* or physiotherap* or "physio therap*" or "physical therap*" or "weight bearing" or "physical activit*")) OR AB ((exercise* or walking or training or retraining or mobili* or locomotion or gait or balanc* or physiotherap* or "physio therap*" or "physical therap*" or "weight bearing" or "physical activit*")) | 256,723 |
| S22 | S5 OR S6 OR S7 OR S8 OR S9 OR S10 OR S11 OR S12 OR S13 OR S14 OR S15 OR S16 OR S17 OR S18 OR S19 OR S20 OR S21 | 313,079 |
| S23 | (MH "Aged") OR (MH "Aged, 80 and Over") | 429,221 |
| S24 | T1 ((aged or elder or elders or elderly or old or older or olds or geriatric patient*)) OR AB ((aged or elder or elders or elderly or old or older or olds or geriatric patient*)) | 243,459 |
| S25 | S23 OR S24 | 549,052 |
| S26 | S4 AND S22 AND S25 | 1,104 |
| S27 | (MH "Randomized Controlled Trials") | 42,132 |
| S28 | T1 ((random* or rct* or metaanaly* or "meta analy*" or quasiexperiment* or "quasi experiment*")) OR AB ((random* or rct* or metaanaly* or "meta analy*" or quasiexperiment* or "quasi experiment*")) | 183,089 |
| S29 | S27 OR S28 | 193,436 |
| S30 | S26 AND S29 | 214 |

Database: Web of Science (Core collection: Indexes=SCI-EXPANDED, ESCI)

Date: 22.05.2018, updated november 2018

Hits: 1265

| Set | Searches | Results |
|-----|---|-----------|
| # 5 | #4 AND #3 AND #2 AND #1 Indexes=SCI-EXPANDED, ESCI Timespan=All years | 1,265 |
| # 4 | TS=((random* or rct* or metaanaly* or "meta analy*" or quasiexperiment* or "quasi experiment*")) Indexes=SCI-EXPANDED, ESCI Timespan=All years | 1,593,406 |
| # 3 | TS=((aged or elder or elders or elderly or old or older or "geriatric patient*")) Indexes=SCI-EXPANDED, ESCI Timespan=All years | 2,961,620 |

406

| | | |
|-----|--|-----------|
| # 2 | TS=((exercise* or walking or training or retraining or mobili* or locomotion or gait or balanc* or physiotherap* or "physio therap*" or "physical therap*" or "weight bearing" or "physical activit*" or "motor activit*")) Indexes=SCI-EXPANDED, ESCI Timespan=All years | 1,683,444 |
| # 1 | TS((((hip or femur or femoral or trochanter* or subtrochanter* or intertrochanter* or pertrochanter* or acetabulum) NEAR/2 fracture*)) Indexes=SCI-EXPANDED, ESCI Timespan=All years | 35,884 |

Database: Pedro

Date: 23.05.2018, updated November 2018

Hits: 53 + 157 + 14 + 50 + 4 + 25 (many duplicates)

Hip fracture AND systematic review: 53

Hip fracture AND clinical trial: 157

Femoral fracture AND systematic review: 14

Femoral fracture AND clinical trial: 50

Femur fracture AND systematic review: 4

Femur fracture AND clinical trial: 25

407

408

409 Supplementary File 2

Supplementary file 2. Extraction form

| Reference (year), Country | Study design and study duration | Sample size | Intervention (IG) / Control group (CG) | Outcomes of interest (a) | Results (b) | Pedro score |
|------------------------------|---|-------------|---|---|---|-------------|
| Binder et al 2004 USA | RCT 6 months Baseline 3-4 months after hip fracture | 90 | Supervised physical therapy and exercise training, progressive resistance training (IG) Home exercise, low-intensity (CG) | Modified physical performance test (PPT), Berg Balance instrument | IG can improve physical function and quality of life and reduce disability compared with CG | 7/10 |
| Kronborg et al 2017 Denmark | RCT In hospital, 10 days+ | 90 | In hospital physiotherapy with additional Progressive knee-extension strength training of the fractured limb (IG) Physiotherapy without strength training (CG) | Maximal isometric knee-extension strength in the fracture limb TUG | Improvements in favor of IG on primary outcome | 8/10 |
| Moseley et al 2009 Australia | RCT 16 weeks Baseline after hospital discharge | 160 | IG received a higher dose (60 min/day) exercise programme conducted while standing and the CG received a lower dose exercise program (30 min/day) primarily conducted whilst seated /supine | Knee extensor muscle strength in fractured leg, walking speed | No difference between groups with respect to the primary outcome measures | 8/10 |
| Mendelsohn et al 2008 Canada | RCT 4 weeks | 20 | Training group IG and control group CG. Both groups attended physical and occupational therapy sessions 5 times a week. Training group also used an arm crank ergometer 3 times/week for 4 weeks | TUG, Bergs Balance Scale, 2MWT, 10MWT | IG performed significantly better in TUG, 2MWT, 10 MWT and Bergs than CG. | 7/10 |

410

| | | | | | | |
|----------------------------------|--|----|---|--|---|------|
| Sherrington et al 2003 Australia | RCT 2 weeks program In hospital | 80 | Two weeks program of either weight-bearing (IG) or non-weight bearing (CG) exercise prescribed by a physiotherapist Exercises for improving strength, balance, gait and functional performance | strength, balance, gait and functional performance | Weight bearing and non-weight bearing exercise programs produce similar effects on strength, balance, gait and functional performance in early phase. | 7/10 |
| Hauer et al 2002 Germany | RCT, 6-8 weeks after hip surgery | 28 | High intensity progressive resistance training and progressive functional training 3 days week for 12 weeks versus motor placebo activities | 1 RM leg press, handgrip strength, max gait speed, TUG, stair climbing, | IG improved strength, functional performance and balance, control group showed no improvement | 7/10 |
| Kimmel et al 2016 Australia | RCT In hospital | 92 | usual care physiotherapy (daily; control group) or intensive physiotherapy (three times daily; intervention group). | post-operative Day 5, at discharge, and at 6 months. modified Iowa Level of Assistance (mILOA) score, TUG, hospital length of stay (LOS). | Intensive acute hospital physiotherapy is safe and reduces hospital LOS after an isolated hip fracture | 9/10 |
| Monticone et al 2018 Italy | RCT 90 minutes 5 times a week for three weeks | 52 | The experimental group (IG) underwent a rehabilitation programme, based on balance task-specific training. The control group (CG) underwent general physiotherapy, including open kinetic chain exercises and walking training. | a Pain Numerical Rating Scale, the Berg Balance Scale, the Functional Independence Measure and the 36-item Short-Form Health Survey. The participants were evaluated before and after training, and after 12 months. | Significant effects of were found for all outcome measures in favour of the experimental group. | 8/10 |

411

| | | | | | | |
|-----------------------------------|--|----|---|----------------------------------|---|------|
| Van Ooijen et al 2016 Netherlands | RCT Inpatient rehabilitation 6 weeks intervention | 70 | six weeks inpatient adaptability treadmill training (n = 24), conventional treadmill training (n = 23) or usual physical therapy (n = 23) | Walking ability, fear of falling | Overall, adaptability treadmill training, conventional treadmill training and usual physical therapy, resulted in similar effects on walking ability, fear of falling and fall incidence. | 7/10 |
|-----------------------------------|--|----|---|----------------------------------|---|------|

412