

1 **The effects of supervised exercise training 12-24 months after bariatric surgery on physical**
2 **function and body composition: a randomised controlled trial**

3 Running title: The Motion Study

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37 **Conflicts of interest statement**

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49 **Abstract**

50 Background

51 Bariatric surgery is effective for the treatment of stage II and III obesity and its related diseases although
52 increasing evidence is showing weight regain approximately 12-24 months post-surgery. Weight regain
53 increases the risk of physical function decline which negatively affects an individual's ability to undertake
54 activities of daily living. The study assessed the effects of a 12-week supervised exercise intervention on
55 physical function and body composition in patients between 12-24 months post bariatric surgery.

56 Methods

57 Twenty-four inactive adult bariatric surgery patients whose body mass index remained $\geq 30\text{kg.m}^2$ 12-24 months
58 post-surgery, were randomised to an exercise intervention (n=12) or control group (n=12). Supervised exercise
59 consisted of three 60-minute gym sessions per week of moderate intensity aerobic and resistance training for 12-
60 weeks. Control participants received usual care. The incremental shuttle walk test (ISWT) was used to assess
61 functional walking performance after the 12-week exercise intervention, and at 24-weeks follow-up. Measures
62 of anthropometric, physical activity, cardiovascular, psychological, and biochemical outcomes were also
63 examined. Using an intention-to-treat protocol, independent t-tests were used to compare outcome measures
64 between groups.

65 Results

66 Significant improvements in the exercise group were observed for the ISWT, body composition, physical
67 function, cardiovascular, and self-efficacy measures from baseline to 12-weeks. A large baseline to 12-week
68 change was observed for the ISWT (exercise: $325.00 \pm 117.28\text{m}$; control: $355.00 \pm 80.62\text{m}$, $p < 0.001$). The
69 exercise group at 24-weeks recorded an overall mean improvement of 143.3 ± 86.6 metres and the control group
70 recorded a reduction of -32.50 ± 75.93 metres. Findings show a 5.6kg difference between groups in body mass
71 change from baseline to 24-months favouring the exercise group.

72 Conclusions

73 A 12-week supervised exercise intervention led to significant improvements in body mass and functional
74 walking ability post-intervention, with further improvements at the 6-month follow-up.

75

76 **Key words**

77 Obesity, exercise, bariatric surgery, physical activity, physical function

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81 Introduction

82 Bariatric surgery is an effective weight-loss intervention for morbidly obese patients and is successful in the
83 treatment of stage II and III obesity and its related diseases¹. However, there is increasing evidence of weight
84 regain in patients after bariatric surgery^{2, 3}, generally occurring between 12 and 24 months post-operatively^{2, 4}.
85 Weight regain increases the risk of physical function decline which negatively affects an individual's ability to
86 undertake activities of daily living⁵. Weight regain also increases the likelihood of obesity related co-morbidities
87 returning⁶, hence the importance of recommendations on diet and physical activity after bariatric surgery to
88 prevent weight regain⁷.

89 Post-operative lifestyle interventions that adopt a combined diet, exercise, and behaviour modification approach
90 have proven successful in aiding long-term weight maintenance and improving physical function⁸. NICE
91 recommend that the two year follow-up care package after bariatric surgery should incorporate physical activity
92 advice and support⁷.

93 At present, no quantifiable physical activity recommendations on post-operative exercise exist. There are a
94 limited number of exercise interventions in bariatric surgery patients, however these mainly target the initial
95 (first four) months after surgery. In a small non-randomised trial, Stegen *et al*⁹ observed that improvements in
96 physical fitness were not induced by the weight loss from surgery alone (in the absence of exercise). Castello *et*
97 *al*¹⁰, in a randomised controlled trial (RCT) that examined the effects of a 12-week aerobic exercise programme
98 after gastric band surgery, found a significant improvement in the six minute walk test (6MWT) distance in the
99 exercise group compared to those who received usual care¹⁰. More recently Coen *et al*¹¹ conducted a semi-
100 supervised moderately-intense exercise intervention in 128 patients who had undergone gastric bypass. This
101 RCT concluded that post-operative health education and exercise elicited similar improvements to weight and
102 body composition, but those who received the exercise intervention had additional fitness benefits. These
103 studies indicate the value of incorporating exercise into the early post-operative rehabilitation of patients after
104 bariatric surgery. Whether exercise therapy at a later stage after surgery, specifically attempting to address late
105 weight regain, is unknown.

106 The aim of this study was to examine the effects of a supervised 12-week exercise intervention on physical
107 function and body composition maintenance in patients who were between 12 and 24 months after bariatric
108 surgery.

109 Methods

110 Study design

111 A single-centre RCT with two parallel groups with balanced randomisation [1:1] was performed. Adult patients
112 who were 12-24 months post-bariatric surgery were randomised to either supervised exercise training for 12
113 weeks or to usual follow up care. Assessments were performed pre-intervention, post-intervention at 12-weeks,
114 and 24-weeks at Leicester Diabetes Centre, Leicester, UK. Ethical approval was received from the West
115 Midlands NHS research ethics committee (Reference: 13/WM/0445; ISRCTN 17240262).

116 Participants

117 Eligible participants were adult patients (≥ 18 years) who had undergone any type of bariatric surgery 12 to 24
118 months earlier, remained obese (BMI of $\geq 30\text{kg}\cdot\text{m}^2$ or $\geq 28\text{kg}\cdot\text{m}^2$ for South Asians, and were classified as inactive
119 (self-report ≤ 150 minutes MVPA per week). Participants completed a health assessment and treadmill exercise
120 test (Balke protocol) before being deemed healthy enough to participate in moderate intensity exercise by a
121 clinician. Volunteers with unstable diabetes, stage II hypertension, cardiovascular disease, pulmonary disease,
122 renal disease, orthopaedic limitations, motor neurone disease, or who were chair-bound, were excluded.

123 Randomisation and masking

124 Patients were recruited from the post-operative bariatric surgery lists from the NHS University Hospitals of
125 Leicester and Spire Leicester Private Hospital between January 2014 and February 2015. All patients who were
126 within 12-24 months of their surgery date were sent invitation letters with reply slips signed by their surgeon,
127 along with a participant information booklet. Upon the successful completion of consent, screening, and the

128 initial assessment, participants were randomly allocated into either the exercise or control group using random
129 number sequencing in concealed brown envelopes. The algorithm for randomisation was designed by a
130 statistician using the random permuted-block procedure (blocks of 4). The randomisation was performed by an
131 independent researcher, who had no other involvement in the study to ensure allocation concealment. Both the
132 participant and the researcher became aware of the study group allocation upon completion of the baseline
133 assessment.

134 **Study Groups**

135 **Exercise group**

136 The exercise intervention incorporated three 60 minute gym sessions per week for 12 weeks. Sessions were
137 hospital based and supervised by a qualified gym instructor with the appropriate immediate life support training.
138 All sessions consisted of a warm-up period, moderate intensity aerobic and resistance training ending with a
139 cool-down period. The aerobic exercise training element typically lasted 45 minutes; the first exercise
140 programme for participants consisted of 35 minutes with a longer warm-up period and was progressed to 45
141 minutes by the end of week two. Warm-ups were longer at the beginning of the 12 weeks, and reduced to 5
142 minutes as individuals' fitness improved. The exercise session was personalised for each individual and re-
143 assessed every two weeks within the 12 week programme. For the few participants who did not progress to 45
144 minutes within the first two weeks they were re-assessed weekly to reach this duration. Moderate intensity
145 aerobic exercise was expressed as a percentage of maximum heart rate; in the main exercise session this equated
146 to between 64 and 77% (RPE 12-14). Typically the resistance training element consisted of four core and lower
147 body resistance exercises (e.g. leg press, abdominal twists, leg extensions) per week. Moderate intensity for
148 resistance exercises was expressed as 60% of the participants' estimated one-repetition maximum (1-RM). At
149 the first session the 1-RM was identified which was equated by performing a weight where 17 repetitions were
150 possible¹². Two resistance exercises were performed per participant per gym session; these varied but only
151 included core and lower body exercises. Participants would perform three sets of 12 repetitions with 30-60
152 seconds rest between sets. The whole exercise session amounted to a total of 60 minutes per participant.

153 Programmes were personalised, specifying durations, resistances, inclines, sets and repetitions. Any limiting co-
154 morbidities were taken into consideration when designing the programmes. Due to the large variation in patients'
155 abilities, programmes were designed to meet the individuals' needs, and progression expectation varied but all
156 patients had the same target training element duration and all met this before the end of the 12 week training
157 programme. Programme progression was based on heart rate; ensuring patients were consistently working at
158 moderate intensity. The exercise programme was reviewed on fortnightly basis and the intensity was progressed
159 accordingly. Gym session attendance was monitored throughout the intervention.

160 Upon completion of the 12-week structured exercise training programme, the participants received a standard
161 lifestyle advice session lasting 30 to 60 minutes. This individualised advice session represented a typical
162 discharge advice session given to patients in follow-up care. Relevant topics such as physical activity
163 maintenance, overcoming barriers, and goal setting were discussed. In addition, a maintenance exercise
164 programme was provided (e.g. gym continuation or home-based alternatives). Finally, a diet information sheet
165 was provided based on standard post-operative advice given to patients after surgery.

166 **Control group**

167 During the 12-week exercise intervention period, participants in the control group continued with usual follow-
168 up care. After their 12-week assessment the control group also received the discharge advice session discussing
169 the same topics. All participants were given an example exercise programme and progression (e.g. home-based
170 exercise, walking, swimming), along with the diet information sheet.

171 **Outcomes**

172 All measurements were taken at the pre-intervention assessment (baseline), post-intervention assessment (12-
173 weeks) and at the follow-up assessment (24 weeks).

174 **Physical function measurements**

175 The primary measure of physical function was the ISWT. The ISWT reflects walking ability, an important
176 measure of daily living in these patients. This involved patients walking consecutive 10-metre shuttles in time

177 with an audible beep that became progressively faster, until they were no longer able to maintain that pace.
178 Patients performed a practice ISWT beforehand to minimise the influence of learning effects. Participants were
179 asked to walk for as long as possible until reaching test termination criteria while the assessor recorded the total
180 number of shuttles performed¹³. The ISWT has been validated against VO₂ max and VO₂ peak in clinical
181 populations¹⁴.

182 Grip strength was measured using the Takei A5001 Analogue Hand Grip Dynamometer (Takei Scientific
183 Instruments, Japan). The protocol was repeated three times on right and left sides. The five times sit to stand
184 (STS) test was used to measure functional lower limb muscle strength. Participants started seated with their
185 groups folded across their chest, they were then instructed to stand up and sit down five times as quickly as
186 possible.

187 Anthropometric measurements

188 Body composition outcomes (fat mass, fat-free mass, body fat percentage, body mass) were measured using
189 validated bioelectrical impedance (Tanita Scales BC-418-MA [Tanita Corporation, Japan])¹⁵. Body mass and
190 stretch stature were measured to calculate BMI. Waist (1cm above the iliac crest) and hip (widest area around
191 the gluteus maximus) circumferences were recorded.

192 Cardiovascular measurements

193 Blood pressure was obtained using the Omron M7 Digital Intellisense Upper Arm Cuff Blood Pressure Monitor
194 (Omron Corporation, Kyoto, Japan). Blood pressure was taken three times; the first measurement was discarded
195 and a mean of the following two measurements was reported. Resting heart rate was measured using the Contec
196 Full-Colour OLED USB Finger Pulse Oximeter & Heart Rate Monitor (CONTEC DTxInc, Melbourne, FL,
197 USA).

198

199 Physical activity measurements

200 Objective measures of physical activity were collected using an accelerometer (GT3X+, ActiGraph, Pensacola,
201 FL, USA). Participants wore the device on an elastic waist belt and positioned it in line with the axillary line of
202 the right iliac crest. Participants were instructed to wear the accelerometer for seven days from the moment they
203 woke up until they went to bed at night, only removing it for water-based activities such as showering and
204 swimming. Participants were asked to complete seven days following all three assessment visits. This is a
205 validated method of measuring physical activity with high inter-instrument reliability (0.97 ICC; $p < 0.001$)¹⁶.
206 The Freedson adult 1998 cut points were used to determine physical activity intensity¹⁷. The accelerometer
207 measured sedentary time (standing and sitting; <100 counts), light activity (100 to 1951), MVPA (>1951) and
208 step count. Data were included if it showed four valid days; a valid day was wear time of 10 waking hours.

209 Self-reported physical activity was measured using the short form international physical activity questionnaire
210 (IPAQ); a seven day recall measure assessing weekly physical activity and daily sitting time. The IPAQ-short
211 form is validated and has demonstrated moderate associations with accelerometer measurements¹⁸.

212 Self-efficacy for physical activity was assessed with the self-efficacy for regulating physical activity (SERPA)
213 scale. This 18-item questionnaire asks individuals to rate their degree of confidence to perform their exercise
214 routine regularly on a scale from 0 to 100. The results are reported as an average out of 100 to reflect the
215 individual's confidence¹⁹.

216 Dietary measurement

217 The 24-hour food recall was used to assess dietary behaviour. This is a validated method of assessing calorie
218 intake delivered via a structured interview²⁰. The investigator asked participants to recall all foods and drinks
219 they consumed the previous day whilst prompting for food quantities and portion sizes. All 24 hour food recalls
220 were analysed using NetWisp Version 4.0 (Tinuviel Software, Warrington, UK) software to estimate total daily
221 calorific intake in kilocalories (Kcal).

222

223 Statistical analysis

224 A sample size calculation indicated that a total of 28 participants were required to detect a difference of 50
225 metres in the ISWT (primary outcome) between the two groups at the 12-week assessment point, with 80%
226 power, and a two-sided 0.05 significance level, and a standard deviation of 45 metres. A difference of 50 metres

227 has been identified as clinically meaningful for another clinical population²¹. In order to allow for a 20% drop
228 out rate, 34 participants was the recruitment target.

229 Data analysis used an intention to treat (ITT) protocol to include all participants who were randomised, using
230 the last-observation-carried-forward method for missing data. The baseline to 12 week change and baseline to
231 24 week change were compared between groups using an independent t-test. Change differences for objectively
232 measured physical activity between each group were determined using ANCOVA controlling for accelerometer
233 wear time. The magnitude of an effect has been reported using the Cohen's d statistic.

234

235 **Results**

236 **Participant characteristics**

237 Of 115 patients initially invited, 57 responded, and 47 expressed interest, and were screened for trial eligibility.
238 A total of 24 patients met study criteria and consented to randomisation. Three discontinued before the end of
239 the trial. Figure 1 details the flow of participants throughout the trial.

240 Mean time since surgery was 19.3 ± 5.4 months. Participants had a mean age of 48.4 ± 8.9 years, and mean pre-
241 operative body mass of 136.3 ± 18.7 kg. At baseline assessment, mean body mass was 106.8 ± 16.7 kg (BMI of
242 39.0 ± 5.2 kg·m²). Of a possible 36 gym sessions, those randomised to the intervention group of the study
243 attended a mean of 34.2 ± 2.5 sessions (95% adherence). No adverse events or injuries were recorded
244 throughout the exercise intervention. Table 1 presents baseline characteristics.

245 **Functional outcomes**

246 Table 2 displays the physical function change data between baseline and 12 and 24 weeks by trial group. For the
247 primary outcome (ISWT), significant differences favouring the exercise group were recorded for the changes
248 recorded at both 12-weeks ($t_{(22)}= 5.820$, $p<0.001$, $d=2.38$), and 24-weeks ($t_{(22)}= 5.289$, $p<0.001$, $d=2.16$). The
249 overall mean improvement in distance walked at 24-weeks was 143.3 ± 86.6 metres for the exercise group,
250 while there was a reduction of 32.50 ± 75.93 metres for the control group (Figure 2). Similarly for the STS test,
251 performance had improved by 4.2 ± 4.0 seconds in the exercise group at 6 months, and slowed by 0.2 ± 2.1
252 seconds in the control group ($t_{(22)}=-3.411$, $p=0.003$, $d=1.39$).

253 **Anthropometric outcomes**

254 Table 3 presents the anthropometric outcome change data between baseline and 12 and 24 weeks by trial group.
255 The exercise groups body mass change data was significantly different to the control group at both 12-weeks
256 ($t_{(22)}=-3.278$, $p=0.003$, $d=1.34$) and 24-weeks ($t_{(22)}=-3.179$, $p=0.004$, $d=1.30$), amounting to a 5.6 kg difference
257 between groups by 24-weeks. Fat mass change was also significantly different between groups at both 12-
258 weeks ($t_{(22)}=-3.573$, $p=0.002$, $d=1.46$) and 24-weeks ($t_{(22)}=-2.843$, $p=0.009$, $d=1.16$). By 24-weeks, total fat mass
259 was 4.0kg lower among the exercise group compared with the control group.

260 **Physical activity outcomes**

261 Table 4 displays the physical activity and self-efficacy change data between baseline and 12 and 24 weeks by
262 trial group. Objectively measured MVPA change from baseline to 12-weeks showed a significant difference
263 between groups ($f_{(2,19)}=4.788$, $p=0.043$, $d=0.98$), with the exercise group increasing by a mean 10.5 minutes per
264 day. No other significant group differences were observed in objective or self-reported outcomes. By 24-weeks,
265 mean daily MVPA in the exercise group was 7.5 minutes greater than baseline, and not significantly different
266 from the change in the control group. From 12 to 24 weeks, there were declines in all physical activity measures
267 in both the exercise and control groups. The exercise group's self-efficacy was highest at 12-weeks showing a
268 mean increase of 20.44 ± 18.90 points whereas the control group remained the same -0.42 ± 7.91 points,
269 showing a statistically significant difference between groups ($t_{(22)}=3.527$, $p=0.002$, $d=1.44$). When focusing on
270 baseline to 24-week mean change, the exercise group sustained an increase from baseline, however this was
271 lower than the 12-week change (6.05 ± 23.32 points). The control group also displayed a mean improvement of
272 9.04 ± 17.06 points at 24-weeks. There was no significant difference between the groups in the self-efficacy
273 change at 24-weeks.

274 **Cardiovascular and dietary outcomes**

275 Table 5 presents the cardiovascular and dietary measurements change data between baseline and 12 and 24
276 weeks by trial group. Systolic blood pressure was significantly lower in the exercise group at 12 ($t_{(22)}=-2.738$,
277 $p=0.012$, $d=1.12$) and 24-weeks ($t_{(22)}=-2.738$, $p=0.012$, $d=1.12$).

278 Resting heart rate decreased to a greater extent in the exercise group (11.25 ± 9.04 bpm) compared with the
279 control group (2.83 ± 7.52 bpm) between baseline and 12-weeks ($t_{(22)}=-2.480$, $p=0.021$, $d=1.01$). The mean
280 change at 24-weeks remained lower in both groups but was not significantly different. No significant differences
281 in dietary intake were noted between groups at 12 or 24 weeks.

282 Discussion

283 This is the first RCT to introduce supervised structured exercise for post-bariatric surgery patients when weight
284 regain is most likely. Significant improvements in physical function, anthropometric, cardiovascular, self-
285 efficacy and physical activity outcomes were observed directly after 12 weeks of exercise training compared
286 with routine care. After a further 12-week follow up period, the exercise group had maintained an advantage
287 over the control participants in physical function, anthropometric, and cardiovascular outcomes.

288 Physical functioning relates to the ability to perform basic activities of daily living such as walking, stair
289 climbing, and transitioning from sitting to standing. These functional abilities are often limited in obese
290 individuals, hence exercise training that improves physical function is important⁵. The ISWT is a valid field
291 based test of functional capacity and aerobic fitness as it strongly relates to VO_2 max²². Improvements displayed
292 in blood pressure and resting heart rate in the exercise group alongside an increased walking distance could also
293 indicate enhanced fitness. The increased mean ISWD in the exercise group after 24-weeks was 143 metres,
294 representing a very large effect size ($d=2.2$). Although, minimum clinically important differences (MCID) for
295 the ISWT in bariatric surgery patients have not been established, this is considerably higher than the 47.5 metres
296 identified as a MCID in performance for patients with chronic obstructive pulmonary disease²¹, hence is likely
297 to translate into meaningful improvements in functional ability.

298 The largest functional improvements in the exercise group occurred from baseline to 12-weeks (during the
299 supervised training). Nonetheless, although the changes were smaller, walking performance, sit-to-stand speed,
300 and grip strength all continued to improve in the 12 weeks after completing the supervised training programme
301 (follow up phase). The degree to which the sustained levels of physical function are directly attributable to the
302 maintenance of physical activity after the intervention is difficult to determine. Objectively recorded MVPA was
303 7.5 minutes higher per day than at baseline in the exercise group, and step counts and self-reported activity also
304 remained higher than at baseline. However, increases in some of these outcomes were also observed in the
305 control group after the advice session received at the end of the intervention period where an example exercise
306 programme and diet sheet were provided, although no significant differences existed between groups. The
307 increases seen in moderate intensity physical activity are noteworthy since general adult activity guidelines are
308 based on moderate intensity (≥ 150 minutes weekly²³) and moderately intense exercise is currently recommended
309 for interventions in obese populations for retention and motivation purposes²⁴. The exercise group recorded
310 positive changes from baseline to 6-months in MVPA, this equated to 52.4 minutes more MVPA weekly
311 whereas MVPA in the control group decreased. Previous research has suggested that 89% of bariatric surgery
312 patients were not meeting the minimum MVPA guideline of ≥ 150 minutes weekly 12 months after surgery²⁵.
313 Despite the encouraging changes in physical activity, conclusions are nonetheless limited by discrepancies
314 between self-reported and objectively measured behaviour at baseline.

315 Despite physical activity being an important method for optimising surgical outcomes, it can sometimes lead to
316 a compensatory response of increased calorific intake²⁶. The American Society for Metabolic and Bariatric
317 Surgery (ASMBS) has reported that exercise changes body composition, with increased fat-free mass resulting
318 in slower loss of overall body mass. The frequency and intensity of exercise may also affect metabolic rate
319 contributing to weight loss plateaus²⁶. Based on this and the results of previous trials of post-operative exercise
320 interventions^{9-11, 27, 28}, total body mass change was not expected in this trial. However body mass in the exercise
321 group decreased progressively at every assessment with a loss of 2.7kg by 24-weeks. In contrast, the control
322 group had gained 2.9 kg over the study period. A similar change in body mass was achieved in a 10-month
323 running intervention initiated between one and three years after bariatric surgery for seven patients, with a mean
324 BMI loss of $2.2\text{kg}\cdot\text{m}^2$ reported compared with control participants, and a 1% reduction in body fat²⁹.

325 Typically, when patients undergo bariatric surgery, rapid weight loss occurs losing both fat mass and fat-free
326 mass which negatively impacts basal metabolic rate³⁰. Fat-free mass loss typically accounts for between 33%
327 and 50% of total body mass loss³¹. Exercise interventions implemented during this period of rapid weight loss
328 have not found any significant differences in body composition through the addition of exercise^{9-11, 27, 28}.
329 However in the current trial with exercise training introduced later, a reduction of 2.1kg of fat mass was
330 observed after 12 weeks of training, which was largely maintained 12 weeks later.

331 Conversely, fat-free mass decreased in the intervention group and increased in the control group; this is not
332 surprising because of the control group's overall increase in body mass. Loss of fat-free mass at the end of the
333 12 week exercise intervention amounted to 13% (0.32kg) of the total body mass reduction. This is lower than
334 observed in the trials initiated earlier (23-39%)^{9, 10, 27}, and did not negatively impact strength outcomes (grip
335 strength and STS performance), which continued to improve during the trial.

336 Previous research has found that in stage II and III obesity, exercise education alone is insufficient for
337 preventing declines in health related fitness³². The supervised exercise approach in the current trial provided
338 regular professional support, on-going counselling and an increased knowledge and understanding of exercise.
339 The benefits of this are reflected in the self-efficacy levels of the exercise group, with low baseline scores
340 increasing markedly after the completion of the exercise intervention. Notably self-efficacy was increased
341 among control participants following the advice session. Due to the variation of abilities in this patient
342 population it is not possible to develop a standardised universal exercise prescription, therefore individualised
343 exercise prescription is necessary. However, the development of guidelines is possible and is needed for
344 optimising post-operative physical function and body composition outcomes. Despite the exercise intervention
345 having a substantial effect on physical function, little effect on objectively measured physical activity and
346 sedentary time was observed. This supports previous research in this clinical population which indicates that
347 physical function improvements do not directly translate into increased physical activity and ultimately
348 undermines the argument that limited exercise capacity is a result of inactivity in this patient population^{8,33}. This
349 therefore indicates the need for a combined approach of supervised exercise with a behavioural component to
350 promote exercise uptake and long term physical activity maintenance to fit into patients' lifestyles. The control
351 participants in the current study demonstrated that an example home based programme was insufficient to
352 improve physical function or MVPA as they declined further highlighting the importance of the need for further
353 support.

354 The strengths of the trial include its rigorous design. This is the first RCT initiated at the point of weight regain,
355 delivered at 12 to 24 months post-bariatric surgery. It is also the first intervention to report follow-up outcomes
356 12 weeks after completion of the exercise intervention, and to include objective measurement of physical
357 activity. The study obtained dietary information to allow potential confounding by dietary changes to be
358 detected. Finally in comparison with previous research, there were low drop-out rates and high session
359 attendance indicating a good level of adherence with the intervention. Of the 115 patients invited by postal
360 invitation, 41% were assessed for eligibility. previous exercise and behavioural interventions have
361 predominantly recruited directly in clinics via clinicians or via telephone and report higher percentages of
362 patients interested. A good example is the recent Bari-Active trial, a behavioural intervention to increase home-
363 based walking exercise, which screened 85% of patients referred for eligibility³⁴, previous exercise training
364 studies to our knowledge do not report response rates; therefore we cannot make a direct comparison with other
365 exercise trials. However, as with all trials this raises the possibility of a selection bias.

366 Despite a thorough recruitment and screening process, the recruited sample was slightly smaller than intended,
367 contributing to some minor differences between intervention and control groups at baseline. However, none of
368 these were statistically significant, and analysis of change data indicated large and significant inter-group
369 differences in the primary outcome measure (ISWT), and many other outcomes. Nonetheless it is difficult to
370 know if baseline differences in physical activity have influenced some of the outcomes. In addition, not all
371 participants had undertaken the same bariatric procedure and no formal record of post-operative weight loss
372 patterns was available; however patients typically self-reported weight plateau or gain. The recruited sample
373 was predominantly female, with only four men randomised, reflecting the general gender bias in the
374 characteristics of bariatric surgery patients [at a ratio of 3:1].

375 Exercise intervention research after bariatric surgery is still in its infancy. This RCT has provided a foundation
376 for future research for the use of physical activity to optimise long term post bariatric surgery outcomes. In
377 particular, larger scale RCTs and longer follow-up periods to determine maintenance of outcomes from such a
378 programme are needed. The combination of pre and post-operative counselling targeting physical activity
379 behaviour change before initiating supervised exercise requires investigation, along with cost effectiveness and
380 feasibility of implementation in routine care.

381 **Conclusion**

382 These findings suggest that the implementation of a supervised exercise intervention at the typical point of
383 weight regain is effective for improving physical function and body composition in this population. However,
384 since physical activity declined after the end of the supervised intervention, patients may need on-going support
385 to develop independence in order to sustain these improvements long-term.

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390 Leadership in Applied Health Research and Care - East Midlands (NIHR CLAHRC - EM), and the Leicester
391 Clinical Trials Unit. The views expressed are those of the author(s) and not necessarily those of the NHS, the
392 NIHR or the Department of Health.

393 **Conflict of interest**

394 Professor Melanie Davies has acted as consultant, advisory board member and speaker for Novo Nordisk,
395 Sanofi-Aventis, Lilly, Merck Sharp & Dohme, Boehringer Ingelheim, AstraZeneca and Janssen and as a speaker
396 for Mitsubishi Tanabe Pharma Corporation. She has received grants in support of investigator and investigator
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398 Professor Stuart Biddle reports that Funding has been received since 2013 for consultancy work from Fitness
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405 All other authors have no conflicts of interest to report.

References

1. Colquitt JL, Picot J, Loveman E, Clegg AJ. Surgery for obesity. *Cochrane Database of Systematic Reviews* 2009; **15**(2): 1-243.
2. Sjöström L, Lindroos AK, Peltonen M, Torgerson J, Bouchard C, Carlsson B *et al.* Lifestyle, diabetes, and cardiovascular risk factors 10 years after bariatric surgery. *The New England Journal of Medicine* 2004; **351**(26): 2683-2693.
3. Courcoulas AP, Christian NJ, Belle SH, Berk PD, Flum DR, Garcia L *et al.* Weight change and health outcomes at 3 years after bariatric surgery among individuals with severe obesity. *Journal of the American Medical Association* 2013; **310**(22): 2416-2425.
4. Magro DO, Geloneze B, Delfini R, Pareja BC, Callejas F, Pareja JC. Long-term weight regain after gastric bypass: a 5-year prospective study. *Obesity Surgery* 2008; **18**(6): 648-51.
5. Steele T, Cuthbertson DJ, Wilding JP. Impact of bariatric surgery on physical functioning in obese adults. *Obesity Reviews* 2015; **16**(3): 248-58.
6. Shah M, Simha V, Garg A. Long-Term Impact of Bariatric Surgery on Body Weight, Comorbidities, and Nutritional Status. *The Journal of Clinical Endocrinology and Metabolism* 2006; **91**(11): 4223-4231.
7. National Institute for Health and Care Excellence (NICE). Obesity: identification, assessment and management of overweight and obesity in children, young people and adults. In. NICE clinical guideline 189 ed, 2014.
8. Josbeno DA, Jakicic JM, Hergenroeder A, Eid GM. Physical activity and physical function changes in obese individuals after gastric bypass surgery. *Surgery for Obesity and Related Diseases* 2010; **6**(4): 361-366.
9. Stegen S, Derave W, Calders P, Van Laethem C, Pattyn P. Physical fitness in morbidly obese patients: effect of gastric bypass surgery and exercise training. *Obesity Surgery* 2011; **21**(1): 61-70.
10. Castello V, Simões RP, Bassi D, Catai AM, Arena R, Borghi-Silva A. Impact of Aerobic Exercise Training on Heart Rate Variability and Functional Capacity in Obese Women After Gastric Bypass Surgery. *Obesity Surgery* 2010; **21**(11): 1739-1749.
11. Coen PM, Tanner CJ, Helbling NL, Dubis GS, Hames KC, Xie H *et al.* Clinical trial demonstrates exercise following bariatric surgery improves insulin sensitivity. *The Journal of clinical investigation* 2015; **125**(1): 248-57.
12. Williams MA, Haskell WL, Ades PA, Amsterdam EA, Bittner V, Franklin BA *et al.* Resistance Exercise in Individuals With and Without Cardiovascular Disease: 2007 Update. *A Scientific Statement From the American Heart Association Council on Clinical Cardiology and Council on Nutrition, Physical Activity, and Metabolism* 2007; **116**(5): 572-584.
13. Lewis ME, Newall C, Townend JN, Hill SL, Bonser RS. Incremental shuttle walk test in the assessment of patients for heart transplantation. *Heart* 2001; **86**: 183-7.
14. MacSween A, Brydson G, Creed G, Capell HA. A Preliminary Validation of the 10-metre Incremental Shuttle Walk Test as a Measure of Aerobic Capacity in Women with Rheumatoid Arthritis. *Physiotherapy*; **87**(1): 38-44.

15. Kushner RF, Kunigk A, Alspaugh M, Andronis PT, Leitch CA, Schoeller DA. Validation of bioelectrical-impedance analysis as a measurement of change in body composition in obesity. *American Journal of Clinical Nutrition* 1990; **52**(2): 219-23.
16. Santos-Lozano A, Marin PJ, Torres-Luque G, Ruiz JR, Lucia A, Garatachea N. Technical variability of the GT3X accelerometer. *Medical Engineering and Physics* 2012; **34**(6): 787-90.
17. Freedson PS, Melanson E, Sirard J. Calibration of the Computer Science and Applications, Inc. accelerometer. *Medicine and Science in Sports and Exercise* 1998; **30**(5): 777-81.
18. Bond DS, Phelan S, Wolfe LG, Evans RK, Meador JG, Kellum JM *et al.* Becoming physically active after bariatric surgery is associated with improved weight loss and health-related quality of life. *Obesity* 2008; **17**(1): 78-83.
19. Bandura A. *Self-Efficacy Beliefs of Adolescents* Information Age Publishing, Inc, 2006.
20. Gersovitz M, Madden JP, Smiciklas-Wright H. Validity of the 24-hr. dietary recall and seven-day record for group comparisons. *Journal of the American Dietetic Association* 1978; **73**(1): 48-55.
21. Singh SJ, Jones PW, Evans R, Morgan MD. Minimum clinically important improvement for the incremental shuttle walking test. *Thorax* 2008; **63**(9): 775-7.
22. Singh SJ, Morgan MD, Hardman AE, Rowe C, Bardsley PA. Comparison of oxygen uptake during a conventional treadmill test and the shuttle walking test in chronic airflow limitation. *The European Respiratory Journal* 1994; **7**(11): 2016-20.
23. World Health Organisation (WHO). Global Strategy on Diet, Physical Activity and Health In: *Physical Activity and Adults - Recommended levels of physical activity for adults aged 18 - 64 years*: WHO, 2015.
24. De Feo P. Is high-intensity exercise better than moderate-intensity exercise for weight loss? *Nutrition, Metabolism and Cardiovascular Diseases* 2013; **23**(11): 1037-42.
25. King WC, Hsu JY, Belle SH, Courcoulas AP, Eid GM, Flum DR *et al.* Pre- to postoperative changes in physical activity: report from the longitudinal assessment of bariatric surgery-2 (LABS-2). *Surgery for Obesity and Related Diseases* 2012; **8**(5): 522-532.
26. American Society of Metabolic and Bariatric Surgery. *ASMBS Public and Professional Education Committee Bariatric Surgery: Postoperative Concerns.*, 2008. Report no.: http://s3.amazonaws.com/publicASMBS/GuidelinesStatements/Guidelines/asbs_bspc.pdf.
27. Huck CJ. Effects of Supervised Resistance Training on Fitness And Functional Strength in Patients Succeeding Bariatric Surgery. *Journal of Strength and Conditioning Research* 2014; **29**(3): 589-595.
28. Castello V, Simões PR, Beltrame T, Bassi D, Maria Catai A, Arena R *et al.* Effects of aerobic exercise training on variability and heart rate kinetic during submaximal exercise after gastric bypass surgery - a randomized controlled trial. *Disability and Rehabilitation* 2013; **35**(4): 334-342.
29. Marchesi F, De Sario G, Reggiani V, Tartamella F, Giammaresi A, Cecchini S *et al.* Road Running After Gastric Bypass for Morbid Obesity: Rationale and Results of a New Protocol. *Obesity Surgery* 2015; **25**(7): 1162-70.

30. Das SK, Roberts SB, McCrory MA, Hsu LKG, Shikora SA, Kehayias JJ *et al.* Long-term changes in energy expenditure and body composition after massive weight loss induced by gastric bypass surgery. *American Journal of Clinical Nutrition* 2003; **78**(1): 22-30.
31. O'Brien PE, McPhail T, Chaston TB, Dixon JB. Systematic review of medium-term weight loss after bariatric operations. *Obesity Surgery* 2006; **16**(8): 1032-40.
32. Herring LY, Wagstaff C, Scott A. The efficacy of 12 weeks supervised exercise in obesity management. *Clinical Obesity* 2014; **4**(4): 220-227.
33. Bond DS, Evans RK, DeMaria E, Wolfe L, Meador J, Kellum J *et al.* Physical activity and quality of life improvements before obesity surgery. *American Journal of Health Behavior* 2006; **30**(4): 422-434.
34. Bond DS, Vithiananthan S, Thomas JG, Trautvetter J, Unick JL, Jakicic JM *et al.* Bari-Active: a randomized controlled trial of a preoperative intervention to increase physical activity in bariatric surgery patients. *Surgery for Obesity Related Diseases* 2015; **11**(1): 169-77.