1	Hybrid Neuromuscular Training Promotes Musculoskeletal Adaptations in
2	Inactive Overweight and Obese Women: A Training-Detraining
3	Randomized Controlled Trial
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11 Abstract

12 This study investigated the effects of a 10-month high-intensity interval-type neuromuscular training 13 program on musculoskeletal fitness in overweight and obese women. Forty-nine inactive females 14 (36.4±4.4 yrs) were randomly assigned to either a control (N=21), a training (N=14, 10 months) or a 15 training-detraining group (N=14, 5 months training followed by 5 months detraining). Training used 16 progressive loaded fundamental movement patterns with prescribed work-to-rest intervals (1:2, 1:1, 2:1) 17 in a circuit fashion (2-3 rounds). Muscular strength and endurance, flexibility, passive range of motion 18 (PRoM), static balance, functional movement screen (FMS) and bone mass density (BMD) and content 19 (BMC) were measured at pre-, mid-, and post-intervention. Ten months of training induced greater 20 changes than the controls in (i) BMD (+1.9%, p<0.001) and BMC (+1.5%, p=0.023) ii) muscular 21 strength (25%-53%, p=0.001-0.005); iii) muscular endurance (103%-195%, p<0.001); and iv) mobility 22 (flexibility: 40%, p<0.001; PRoM [24%-53%, p=0.001-0.05;]; balance: 175%, p=0.058; FMS: +58%, 23 p < 0.001). The response rate to training was exceptionally high (86-100%). Five months of detraining 24 reduced but not abolished training-induced adaptations. These results suggest that a hybrid-type exercise 25 approach integrating endurance-based bodyweight drills with resistance-based alternative modes into a 26 real-world gym setting may promote musculoskeletal fitness in overweight or obese women. 27 Keywords: intermittent exercise; females; muscular strength; mobility; functional movement patterns,

- bone health.
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Word count: 4,361

31 Introduction

32 Obesity is a multifactorial chronic condition affecting one in three adults worldwide (29). 33 Individuals with obesity demonstrate low cardiorespiratory fitness, high metabolic risk and 34 physical limitations (11,32). This population demonstrates biomechanical deficits in activities 35 of daily living (ADL) and reduced passive range of motion (PRoM) in several joints (19) 36 compared to lean individuals. Such kinetic limitations due to restricted musculoskeletal fitness 37 and mobility levels predispose obese individuals to injuries and lead to impaired functionality 38 and reduced quality of life (42). Overweight or obese adults also exhibit a 15% and 48%, 39 respectively, higher risk of sustaining an injury (35) and these rates are even higher for industry 40 workers (17).

41 Musculoskeletal fitness is characterised by lean body mass (LBM), muscular strength and 42 endurance, balance and mobility, which contribute to physical performance, resting metabolic 43 rate improvement, osteoporosis prevention and body functionality in ADL (20). Low 44 musculoskeletal fitness levels may predispose obese adults to hip, foot, ankle, knee and 45 shoulder injuries and may cause soft tissue (cartilage, tendons and fascia) damage (42). 46 Although, the etiology is largely unclear, the overloading of the locomotor apparatus combined 47 with the poor musculoskeletal strength and mobility produce impaired mechanics during 48 movement that increases the stress within the soft tissue and bones (42). Obesity may also be 49 related to a lower bone mineral density (BMD) and content (BMC) (36). Although overweight 50 and obese adults tend to have higher absolute BMC values than adults of healthy weight, after 51 adjusting for total body mass, their BMC is markedly lower than that of their controls (36). In 52 other words, the elevated BMD usually measured in adults with obesity may not be adequate 53 to offset the greater forces developed during low- or high-velocity movements.

Regular exercise is an efficient tool for improving physical fitness, health and body composition in this population (1). Current exercise guidelines include progressive protocols of continuous endurance training (CET), resistance training (RT), or combined CET and RT 57 training (CT) to induce cardiorespiratory, neuromuscular and metabolic adaptations (1). 58 Structured RT is pivotal for preventing sarcopenia and osteoporosis in sedentary, 59 premenopausal women with obesity through improvements in LBM and physical performance 60 (21,40). High-intensity interval training (HIIT) is one of the most popular exercise modes (3) 61 requiring less exercise volume compared to CET, RT, or CT and demonstrating high 62 compliance rates when conducted under supervision in untrained individuals (34). HIIT includes repeated short-to-long bouts performed at an intensity that provokes a heart rate (HR) 63 \geq 80% of maximal hear rate (MHR) (39). The present, injury-free, hybrid-type exercise protocol 64 65 integrates progressive HIIT and functional resistance accessory training into a circuit training 66 format that has been shown to reduce body fat, increase LBM, RMR, endurance, exercise 67 behavioural regulation and vitality with exceptional adherence rates in previously inactive 68 women with obesity (4,6). Such an exercise approach incorporates endurance-based 69 bodyweight drills and resistance-based alternative modalities (18) performed in a circuit, 70 interval format and at moderate-to-high intensity adopting some of the principal characteristics 71 of multimodal integrative neuromuscular training (44).

72 Although there is evidence that overweight or obese adults need comprehensive exercise 73 strategies not only to reduce body mass and fat (1) but also to improve functional limitations 74 while avoiding physical training-related injuries (37), there is a paucity of longitudinal studies 75 to determine the efficacy of such a HIIT-type neuromuscular exercise approach on 76 musculoskeletal fitness, mobility and bone health. Additionally, it is important to investigate 77 potential changes in such physiological parameters, which result from the cessation of exercise, since exercise training is considered a fundamental component of every lifestyle intervention 78 79 for this population. It has been observed an unfavorable effect of detraining on neuromuscular 80 performance that was mainly influenced by the duration of training cessation, age, and training 81 status (45). Characteristically, when an 8-month multicomponent exercise program performed by older overweight women was interrupted for only three months, musculoskeletal
performance gains induced by previous training were abolished (49).

To our knowledge, there is no data concerning the effects of training cessation on musculoskeletal fitness parameters in sedentary overweight and obese individuals. We hypothesized that the training would induce favorable changes in musculoskeletal fitness. Therefore, this study aimed to determine the effects of the a HIIT-type, neuromuscular training protocol on (i) muscular strength and muscular endurance, (ii) joints' range of motion, (iii) balance, (iv) functional movement patterns, and (v) bone health of previously inactive, premenopausal Caucasian women with obesity.

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92 Materials and Methods

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94 Study design

This study is a part of a larger longitudinal research project whose purpose, methodology and primary outcomes are reported elsewhere (6). In this investigation, data upon musculoskeletal fitness are presented. This study was a randomised controlled trial based on a three-group, repeated-measures design in accordance with the Consolidated Standards of Reporting Trial (CONSORT) guidelines (Figure 1) and it was registered at clinicaltrials.gov as NCT03134781. Initially, 102 participants were assessed for eligibility, 66 were recruited and allocated to three groups, and 49 completed the trial as required (see Figure supplemental Digital Content 1).

Participants (36.4 \pm 4.4 years) were randomly assigned to a training (TR, *n*=14), a training detraining group (TRD, *n*=14) or a control (C, *n*=21) group. Following a 4-week adaptive and familiarization period as previously articulated (6), TR performed the 10-month exercise training protocol whereas TRD performed the same protocol for 5 months and then entered a 5-month detraining period. Abstinence from exercise in the detraining group was verified using accelerometry (GT3X-BT, ActiGraph, FL, USA). Accelerometry data were used in the analysis, only if participants had \geq 4 days and \geq 10 wear hours/day. Four vector magnitude data were used to calculate daily activity and sedentary time. Data were expressed as steps/day and time in sedentary, light, moderate, vigorous and moderate-to-vigorous physical activity as described (6). Assessments of musculoskeletal fitness were performed at pre-, mid-, and post-training

(Figure 1). All assessment procedures were completed with the same order (bone health,
flexibility, static balance, functional movement patterns, maximal strength, muscle endurance)
at pre-, mid- and post-training.

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FIGURE 1 ABOUT HERE

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118 **Participants**

119 Participants were medically cleared for strenuous exercise, were non-smokers and of low 120 regular PA or structured exercise for ≥ 6 months before the study. None of them were on 121 medication, diet or nutritional supplementation. Participants were excluded if they missed 122 \geq 20% of total exercise sessions, changed their eating habits and modified their PA levels during 123 the intervention. Participants during detraining need to have comparable PA levels with pre-124 training. Participants were informed about all risks, discomforts and benefits associated with 125 the study and provided a written consent. This investigation was carried out in accordance with 126 the guidelines contained in the 2013 Declaration of Helsinki and was approved by the 127 Institutional Ethics Committee. Participants' characteristics are shown in Table 1.

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131 Training

132 Three small-group (5-10 participants per session), supervised training sessions per week that133 used asynchronous music in the background were performed on non-consecutive days for TR

TABLE 1 ABOUT HERE

134 (10 months) and TRD (5 months) as previously reported (6). This hybrid-type protocol was organized in four progressive training phases (i.e., phase 1: weeks 1-7, phase 2: weeks 8-14, 135 136 phase 3: weeks 15-20 and phase 4: 21-40) (Figure 1). The mean weekly exercise volume was \sim 100 min, net exercise time was 6.5-24.0 min per session and total duration per session was 137 138 23-41 min (6). Exercises (see Table, Supplemental Digital Content 2) adapted basic movement 139 patterns (squat, hinge, lunge, push, pull, carry, rotation, plank) utilizing portable modalities 140 (suspension belts, balance balls, kettlebells, medicine balls, battle ropes, stability balls, speed 141 ladders, foam rollers, elastic bands) and bodyweight as resistance. Exercises (~10-12 per 142 session) were organized in a circuit format and performed in all planes of motion simulating 143 ADLs. The work-to-rest ratio was varied (1:2, 1:1, 2:1) using an interval duration of 20-40 sec 144 to provide progression (see Table, Supplemental Digital Content 2) (6). Verbal encouragement 145 was provided. Participants were instructed to execute as many repetitions as possible with the 146 correct form and with a controlled, moderate rhythm. A 10-min warm-up and a 5-min cool 147 down period was applied in all sessions (6). HR was monitored and recorded using telemetry 148 (Polar Team Solution, Polar Electro-Oy, Kempele, Finland) aiming to maintain an intensity 149 \geq 75% of MHR throughout each session. Rating (6-20) of perceived exertion (RPE) was 150 recorded at the end of each round in all sessions using the 6-20 Borg scale.

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152 Measurements

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154 Bone Health

Whole-body BMC and BMD were performed using a dual energy X-ray absorptiometry (DXA) scanner (GE Healthcare, Lunar DPX-NT) in the morning by the same experienced radiologist according to standard procedures (41). Briefly, participants were placed in a supine position with their body aligned along the central horizontal axis, their arms parallel to their body (without touching it), the forearms pronated, hands flat, legs fully extended, and feet secured using a velcro strap to prevent foot movement during the scan. The instrument was calibrated
daily using a calibration epoxy resin phantom. All analyses were performed using the 12.2 GE
enCORE software package.

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164 Flexibility, Static Balance and Mobility

A 3-min cycling warm-up preceded mobility testing (three consecutive measurements). Flexibility of lower back and hamstrings was measured under standardized conditions using the modified sit-and-reach test (2). Goniometry (Lafayette 01135, Lafayette Instrument Inc., Lafayette, IN, USA) was applied to assess the PRoM (in degrees) of ankle dorsiflexion, knee extension, hip extension, shoulder extension, and glenohumeral internal rotation (31). Both extremities were examined and the median of their measurements was reported as the value in goniometry.

172 Static balance was assessed using the modified Romberg test. Participants were asked to 173 stand without shoes on a firm surface, with eyes closed and arms crossed on the chest and the 174 dominant foot placed directly in front of the non-dominant foot. The time to failure was 175 measured manually by stopwatch in sec (30).

176 The Functional Movement Screening (FMS) with an ICC of >0.8 was used to evaluate 177 functional mobility, postural stability and movement behavior in different settings (23). Two 178 examiners (with an intra-rater reliability of 88.6%) performed this assessment. The FMS has 179 been reported as a simple, quick, non-invasive and suitable movement-based assessment tool 180 for middle-aged, overweight or obese aiming to evaluate their functional capacity levels (23). 181 In brief, the FMS assesses seven fundamental movement tasks (deep squat, hurdle step, in-line 182 lunge, active straight-leg raise, trunk stability push-up, rotary stability, shoulder mobility). Each 183 movement task was scored from 0 to 3 points (0=pain with pattern regardless of quality, 184 1=unable to perform pattern, 2=able to perform pattern with compensation/imperfection, 3=able to perform pattern as directed) and their sum provided the total score ranging from 0 to21 points (46).

- 187
- 188 Muscular Strength and Endurance

189 Maximal isotonic strength (one repetition maximal, 1RM) was assessed using standard 190 procedures for novice and untrained individuals following familiarization as previously 191 described (2) with an intra-class correlation coefficient (ICC) for test-retest trials of 0.88. Two 192 upper-body (vertical chest press, supinated closed-grip lat pulldown) and two lower-body 193 (seated leg extension, lying leg curl) exercises performed on traditional strength training 194 equipment (Panatta Sport, Apiro, Italy) were selected. Muscular endurance was assessed using 195 timed tests for the abdominal (partial curl-up), upper-body (modified kneeling push-up) and 196 lower-body (modified chair squat) musculature (2). All tests required the participants to 197 perform as many repetitions as possible within 60 sec using standard procedures and a 5-min 198 rest was provided between tests (2).

199

200 Statistical Analyses

201 A preliminary power analysis (effect size >0.55, probability error of 0.05, two-tailed alpha 202 level, power of 0.9) using the G*Power 3.0.10 program based on the study design suggested 203 that a sample of 36-40 participants was necessary to identify statistically meaningful trial 204 effects. For all dependent variables, differences (for both "between" and "within" groups) of 205 means (MD) and confidence intervals (CI) were calculated based on mixed models. Cohen's d 206 criteria were used to interpret the magnitude of MD as very small, small, medium, large, very 207 large and huge for values 0.01, 0.20, 0.50, 0.80, 1.20 and 2.0, respectively. No assumptions 208 were made for covariance matrices (unstructured) since repetitions were not that many to 209 significantly reduce the degrees of freedom. All estimations were corrected based on the 210 Bonferroni criterion for multiple comparisons. Results are presented as relative difference in

time (Δ %). Since the variability of the change score in the intervention groups was greater than that in C, the response rate was analysed using the number of differential responders relative to the ratio of variance in TR and C groups providing multiple differential responder groups (10). Statistical significance was set at *p*<0.05. Data were analysed using the SPSS 22.0 software (IBM Corp., Armonk, NY, USA).

216

217 Results

218 No injuries or other adverse effects occurred during the trial. Adherence rates for TR (10-month

219 intervention) and TRD (5-month intervention) were 93.5% and 82.6%, respectively, and a 4%

220 dropout rate was reported. Results are described in brackets as $\Delta\%/95\%$ CI/d/p levels.

221

222 Bone mineral density and bone mineral content

Changes in BMD and BMC are shown in Table 2. No changes were observed between TR and TRD. TR only improved BMD following training (Table 2) (+1.9%/0.010-0.035/2.61/p<0.001)and BMC (+1.5%/0.04-0.076/2.72/p=0.023) and its response rate to whole-body BMD was 100% (Figure 4).

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228 Flexibility, Static Balance and Mobility

Changes in flexibility and passive range of motion are shown in Table 2. At mid-training, TR and TRD demonstrated greater flexibility changes than C (TR vs. C: +38%/3.939-16.490/1.66/p=0.001; TRD vs. C: +34%/3.010-15.561/1.31/p=0.002). At post-training, TR and TRD elicited more favorable changes in flexibility than C (TR vs. C: +40%/4.465-17.130/1.83/p<0.001; TRD vs. C: +26%/0.572-13.238/1.02/p=0.028). No significant differences were observed between TR and TRD. In TR, the response rate to flexibility was 100% (Figure 4). 236 Changes in PRoM are shown in Table 2. At mid training, TR and TRD showed significant 237 differences than C in hip extension (TR vs. C: +43%/0.678-9.607/1.08/p=0.019; TRD vs. C: 238 +41%/0.393-9.322/1.08/p=0.029), glenohumeral internal rotation (TR vs. C: +24%/5.676-239 22.991/1.45/p<0.001; TRD vs. C: +17%/1.747-19.062/1.04/p=0.014). No changes were noted 240 between TR and TRD. At post-training, TR demonstrated a trend for a rise in PRoM compared 241 to C in ankle dorsiflexion (+44%/-0.543-10.924/0.86/p=0.088) and shoulder extension 242 (+24%/0.178-15.940/0.93/p=0.057). TR elicited greater PRoM changes than C and TRD in glenohumeral internal rotation (TR vs. C: +27%/7.814-24.519/1.59/p<0.001; TR vs. TRD: 243 244 +17%/1.636-19.935/1.83/p=0.016). In TR, the response rate to PRoM in the ankle, knee, hip, 245 shoulder, and glenohumeral joint was 93%, 100%, 93%, 86%, and 86%, respectively (Figure 246 4).

Changes in static balance are shown in Table 2. TR demonstrated a greater rise in static balance than C at post-training, which was close to being statistically significant (+143%/-0.784–64.436/0.80/p=0.058) and its response rate to static balance was 86% (Figure 4). No changes were found between TR and TRD at mid- and post-training.

251 Changes in FMS are shown in Table 2. TR and TRD induced a greater rise of the FMS total 252 score than C at mid-training (TR vs. C: +49%/3.860-5.855/6.42/p<0.001; TRD vs. C: 253 +45%/3.431-5.426/5.0/p<0.001) and post-training (TR vs. C: +58%/4.559-254 6.774/7.71/p<0.001; TRD vs. C: +38%/2.631-4.845/4.0/p<0.001). No changes were reported 255 between TR and TRD at mid-training, but TR had more favorable changes in the FMS total 256 score relative to TRD (+14%/0.716-3.142/2.83/p=0.001) at post-training. In TR, the response 257 rate to the FMS total score was 100% (Figure 4).

258

259 Muscular strength

260 Changes in muscular strength are shown in Figures 2A-D. At mid-training, TR exhibited 261 superior changes than C in chest press (+22%/0.244-12.566/0.82/p=0.039), lat pulldown

(+17%/2.767-12.233/1.29/p=0.001), leg extension (+41%/5.986-17.252/2.04/p<0.001), and 262 263 leg curl (+27%/4.871-12.796/2.62/p<0.001). TRD demonstrated greater changes than C in 264 chest press (+33%/3.530-15.851/1.59/p=0.001),lat pulldown (+22%/5.053-14.519/2.16/p < 0.001), leg extension (+39%/5.379-16.645/1.87/p < 0.001), and leg curl 265 266 (+23%/3.656-11.582/2.05/p<0.001). TR and TRD induced comparable strength gains at mid-267 training.

268 At post-training, TR showed greater changes than C in chest press (+29%/2.167-14.666/1.23/p=0.005), lat pulldown (+25%/6.475-15.977/2.03/p<0.001), leg extension 269 270 (+53%/9.133-20.462/2.79/p<0.001), and leg curl (+38%/7.826-16.698/2.71/p<0.001). In leg 271 curl, TR presented greater changes than TRD (+12%/0.105-9.823/0.91/p=0.044). TRD 272 exhibited superior changes than C in chest press (+30%/2.296-14.795/1.50/p=0.004), lat 273 pulldown (+23%/5.511-15.013/2.03/p<0.001),leg extension (+41%/5.633-274 16.962/2.23/p < 0.001), and leg curl (+23%/2.862 - 11.733/1.79/p = 0.001). In TR, the response 275 rate to all 1RM measures was 100% (Figure 4).

276

277 Muscular endurance

278 Changes in muscular endurance are shown in Figures 2E-G. At mid-training, TR resulted in 279 greater changes of muscular endurance than C (curl-up: +75%/11.024– 9.595/3.39/p<0.001; 280 push-up: (+143%/3.402-11.026/2.04/p<0.001;bodyweight squat: +97%/13.060-281 18.559/6.03/p < 0.001) and TRD demonstrated superior changes in muscular endurance than C 282 (curl-up: +87%/13.238-21.810/3.85/p<0.001; push-up: +168%/3.590-11.267/2.56/p<0.001; 283 bodyweight squat: 102%/11.961-18.658/5.33/p<0.001). TR and TRD resulted in similar 284 muscular endurance gains.

At post-training, TR exhibited greater changes of muscular endurance than C (curl-up: +103%/16.185–25.196/5.0/p<0.001; push-up: +195%/7.876–15.553/3.14/p<0.001; bodyweight squat: +136%/18.176–24.872/7.33/p<0.001) and TRD showed superior changes in

288	muscular endurance than C (curl-up: $+83\%/12.185-21.196/3.27/p<0.001$; push-up:
289	+123%/3.590-11.267/1.99/p<0.001; bodyweight squat: $+97%/11.961-8.658/4.66/p<0.001$).
290	TR had more favorable changes in muscular endurance relative to TRD (push-up: $+32\%/0.081-$
291	8.491/0.80/p=0.044; bodyweight squat: +20%/2.547–9.882/1.69/p<0.001). In TR, the response
292	rate to all muscular endurance measures was 100% (Figure 3).
293	
294	FIGURE 2 ABOUT HERE
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296	TABLE 2 ABOUT HERE
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298	FIGURE 3 ABOUT HERE
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300	Discussion

301 This 10-month study revealed that the implementation of a HIIT-type, integrated 302 neuromuscular exercise program performed in a real-world gym setting using portable 303 equipment induces considerable improvement of musculoskeletal fitness in previously inactive, 304 overweight or obese women. These adaptations were reduced but not lost after prolonged 305 detraining.

306 This trial focused on physically inactive, middle-aged, overweight or obese who are 307 characterized by increased cardiometabolic risk (11), poor functional capacity (32), a higher 308 risk for musculoskeletal disorders (42) and physical limitations (19) compared to normoweight 309 women. Overweight and obese adults are more prone to sustain injuries and exhibit knee 310 osteoarthritis than individuals of a normal BMI (35). Hence, progressive, injury-free and 311 effective exercise protocols are critical to reduce functional deficits that are responsible for a 312 smaller PRoM in several joints (19), impaired quality of life and a rising prevalence of injuries 313 in this population (17). In this study, a 10-month training program designed for inactive, 314 overweight or obese women was injury-free and reported high adherence and low dropout rates.
315 This outcome may support the necessity of prescribing progressive and supervised exercise
316 regimens for this population aiming to promote a safe exercise experience that may be pivotal
317 for behavioural regulation in exercise (4).

318

319 Bone adaptations

320 The implementation of high-impact training for inactive, middle-aged, overweight or obese 321 women is critical for preventing osteopenia, osteoporosis and injuries (20). Training improved 322 whole-body BMD (+1.9%) and BMC (+1.5%) only in TR at post-training indicating that this 323 type of program may meet the essential features of a high-impact, weight-bearing training 324 program capable of activating bone cell mechanisms and hormonal factors. It is worth 325 mentioning that exercise-induced weight loss in this cohort was not accompanied by a decline 326 in BMD as it was seen in overweight or obese elderly (40), which is important for bone health 327 and injury prevention.

328

329 Flexibility, Static Balance and Mobility

330 Due to insufficient use of joints in inactive, overweight or obese individuals, the 331 functional length of muscles' that cross these joints is reduced resulting in decreased PRoM 332 (19). Hamstring and lower back flexibility improved by 40%, whereas PRoM in ankle, hip, 333 shoulder, glenohumeral joints improved by 24-44% after 10 months of implementation. These 334 adaptations were maintained after 5 months of detraining. These results coincide with a 10-38% 335 increase in flexibility of inactive, overweight or obese older adults following long-term RT 336 (13,14,15). These outcomes may be attributed to the features of the protocol, i.e. the 337 incorporation of whole-body multiplanar movements mimicking ADLs. RT may promote 338 flexibility if exercises are performed through a full range of motion to adequately activate both 339 the agonist and antagonist muscle groups (15). Resistive exercises may not only increase muscle mass and contractility but they also improve the strength of tendons and ligaments thereby augmenting joints' PRoM (43). Studies employing compensatory overload models have shown a simultaneous elevation of muscle's strength and tendon's active fibroblast numbers, collagen synthesis and turnover rate (43). The strength of the junction between bone and ligament is also enhanced by this type of training (38). The association between body fat and flexibility performance changes in response to training supports the evidence that body composition may play some role in flexibility and mobility performance in overweight or obese adults (19).

347 Although this study did not examine a stretching intervention, it appears that improved 348 PRoM in overweight and obese adults demonstrates exceptional trainability to a hybrid-type 349 exercise training protocol and it may be linked to the improved functionality seen in response 350 to this type of training. The role of flexibility as a major fitness component has been questioned 351 (47). Although the goal of this study was not to determine the value of flexibility as a main 352 fitness component in the overweight and obese population, it appears that PRoM may be pivotal 353 for adequate levels of functionality and quality of life. DoIT seems not to induce negative 354 adaptations to motor control, physical performance and injury rate in a population commonly 355 characterized by reduced mobility and functionality despite the lack of static stretching (47). 356 This is an interesting outcome that highlights the rationale for integrated neuromuscular training 357 methodology adapted for overweight and obese individuals although that static stretching has 358 been classified as a major component of exercise prescription for this population (1).

359 Static balance improved by 150%, an adaptation not lost following detraining. This 360 finding complements the marked (+58%) improvement seen in the FMS score suggesting a 361 noticeable improvement in neuromuscular functional status. The large increase in static balance 362 may be related to the low ceiling effect and the relative insensitivity to change of the assessment 363 used, especially in younger individuals without clinical neurological conditions or balance 364 impairments (27). However, sedentary populations with obesity are likely to demonstrate 365 significantly impaired components of motor skills related to fitness such as balance and 366 coordination (19). Thus, a 10-month intervention incorporating various neuromotor exercises
367 into a structured training regimen with a frequency of 3 times per week may reasonably promote
368 a large improvement in this cohort.

369 The FMS testing battery was used as an assessment tool only since its internal and 370 external validity as a predictive tool for injury has been questioned (23). Although there is no 371 data on the effects of various exercise training modalities on the FMS score in untrained, 372 overweight or obese adults, this score (<15) for sedentary middle-aged women is considered 373 moderate-to-low (33). Considering that individuals with obesity demonstrate biomechanical 374 deficiencies in ADLs (19), neuromuscular-type protocols may aid to reduce these limitations 375 by using progressive integrated neuromuscular exercises characterized by multiple angles and 376 planes of motion such as bending, lifting, pushing, pulling, carrying, and rotating (22) using 377 non-traditional portable modalities (18). Such training introduces increased cognitive and motor 378 processing demands that ultimately favor not only strength but also body and joint stability, 379 coordination, balance and PRoM. These outcomes are aligned with recent evidence suggesting 380 that the functionality and mobility of overweight or obese women be improved through 381 neuromotor training programs (37).

382 Improvements of knee flexor and extensor strength in response to neuromuscular exercise 383 training, as in this study, are associated with increased balance and gait (48) that ultimately 384 improves functional status and reduces falls (16). A potential explanation for these adaptations 385 may be related to the increase in LBM and strength that are seen in response to similar protocols 386 (7). Another explanation for static balance adaptations may be linked to the activation of the 387 vermis of the cerebellum, which is the principal part of a central coordinating mechanism (27). 388 Additionally, postural sensibility to convey information concerning position may play an 389 important role for improving the function of sensory pathways and proprioception (27). These 390 findings highlight the need to integrate multicomponent neuromuscular exercise interventions 391 of sufficient power in the muscular system.

393 Muscular performance adaptations

394 Strength improved in both upper- (+25-29%) and lower-body (+38-53%) in response to 395 training. Interestingly, detraining did not reverse these gains in upper- (+23-30% vs. pre-396 training) and lower-body (+23-41% vs. pre-training) musculature. Likewise, muscular 397 endurance increased in upper-body, lower-body and abdominal musculature by 195%, 136% 398 and 103%, respectively. These improvements were also maintained following detraining in 399 upper-body (+104%), lower-body (+73%) and abdominal musculature (+58%) compared to 400 baseline levels. These findings are aligned with previous reports in lean women involved in 401 either a short-term conventional CT or circuit-based whole-body RT (28), suggesting a similar 402 trainability. These results corroborate a previous report of improved musculoskeletal fitness 403 and body composition in response to short-term HIIT-type programs that use a whole-body RT 404 approach (6,25). RT is highly recommended as a fundamental component of an exercise 405 program targeted to preventing, managing and treating obesity while eliciting neuromuscular 406 adaptations in individuals (1). The increase in muscular strength and endurance may be 407 attributed to neuromuscular adaptations (25,28,38) and a rise in DXA-assessed LBM (6).

408

409 Detraining

410 Detraining is a serious issue for overweight or obese individuals participating in exercise 411 interventions (26). There are no data for the impact of detraining on the adaptations obtained 412 from hybrid, HIIT-type programs. In this study, training gains were reduced but not eliminated 413 following a 5-month detraining period. This outcome corroborates previous findings suggesting 414 that musculoskeletal fitness may be maintained above pre-training levels ever after a training 415 cessation of 5 months or longer if previous training was of sufficient intensity (14). 416 Additionally, it has been documented that RT status may limit the type of neural adaptations 417 that are responsible for the increase in muscular strength and probably the speed of reversibility (45). As such, previously untrained individuals are likely to rapidly lose the adaptations induced
by short-term (8-12 weeks) RT programs during a detraining period (45). Detraining-induced
loss of musculoskeletal fitness seems to be intensity-dependent (14) and may be associated with
an attenuation of muscle fiber size and motor unit recruitment efficiency (24).

422

423 Practical applications

424 The outcomes of this study coincide with studies using HIIT-type protocols (34,36) 425 suggesting that ~100 min of training per week without changes in eating patterns and habitual 426 PA may be an effective long-term approach for musculoskeletal fitness improvement in inactive 427 overweight or obese women. Interestingly, prolonged detraining did not abolish the musculoskeletal fitness adaptations obtained from this fully-supervised longitudinal exercise 428 429 intervention. These findings underline a safe, time-efficient and motivating (4) exercise 430 approach to promote musculoskeletal health in overweight or obese women that may be a valuable addition to current exercise recommendations for this population (1). However, further 431 432 research is needed in this area investigating the efficacy of such an exercise protocol in males 433 and other age and race groups as previously described (5).

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- 436 and impressive commitment of the participants.
- 437

438 **Disclosure of interest**

439 The authors report no conflict of interest.

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Figure 1.

CONSORT diagram of the study.

Figure 2.

Experimental flowchart.

TR, training group (5 months); TRD, training (5 months) - detraining (5 months) group; DoIT, exercise protocol; PRoM, passive range of motion; FMS, functional movement screen; ¹for all groups (4-week adaptive period); ²only for TR and TRD; ³for all groups.

Figure 3.

Changes in muscular strength and endurance throughout the experimental period.

C, control group; TR, training group; TRD, training-detraining group; 1-RM, one repetition maximum; reps, repetitions; † different from Pre, p < 0.05; ‡ different from Mid, p < 0.05; § different from C, p < 0.05; # different from TR, p < 0.05.

Figure 4.

Multiple differential responder groups to exercise in TR following a 10-month intervention. 1-RM, one repetition maximum; PRoM, passive range of motion; FMS, functional movement screen. Supplemental Digital Content 1. The exercises of the 10-month DoIT protocol.

1	1	Hybrid Neuromuscular Training Promotes Musculoskeletal Adaptations in
2 3 4	2	Inactive Overweight and Obese Women: A Training-Detraining
5 6 7	3	Randomized Controlled Trial
8 9 10	4	
11 12 13	5	Alexios Batrakoulis ^a , Panagiotis Tsimeas ^a , Chariklia K. Deli ^a , Dimitrios
14 15 16	6	Vlachopoulos ^b , Esther Ubago-Guisado ^c , Athanasios Poulios ^a , Athanasios
17 18 19	7	Chatzinikolaou ^d , Dimitrios Draganidis ^a , Konstantinos Papanikolaou ^a , Kalliopi
20 21 22	8	Georgakouli ^a , Dimitrios Batsilas ^a , Luis Gracia-Marco ^e , Athanasios Z. Jamurtas ^a ,
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1 2	26	Hybrid Neuromuscular Training Promotes Musculoskeletal Adaptations in
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6 7	28	Randomized Controlled Trial
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This study investigated the effects of a 10-month high-intensity interval-type neuromuscular training program on musculoskeletal fitness in overweight and obese women. Forty-nine inactive females (36.4±4.4 yrs) were randomly assigned to either a control (N=21), a training (N=14, 10 months) or a training-detraining group (N=14, 5 months training followed by 5 months detraining). Training used progressive loaded fundamental movement patterns with prescribed work-to-rest intervals (1:2, 1:1, 2:1) in a circuit fashion (2-3 rounds). Muscular strength and endurance, flexibility, passive range of motion (PRoM), static balance, functional movement screen (FMS) and bone mass density (BMD) and content (BMC) were measured at pre-, mid-, and post-intervention. Ten months of training induced greater changes than the controls in (i) BMD (+1.9%, p<0.001) and BMC (+1.5%, p=0.023) ii) muscular strength (25%-53%, p=0.001-0.005); iii) muscular endurance (103%-195%, p<0.001); and iv) mobility (flexibility: 40%, p<0.001; PRoM [24%-53%, p=0.001-0.05;]; balance: 175%, p=0.058; FMS: +58%, p < 0.001). The response rate to training was exceptionally high (86-100%). Five months of detraining reduced but not abolished training-induced adaptations. These results suggest that a hybrid-type exercise approach integrating endurance-based bodyweight drills with resistance-based alternative modes into a real-world gym setting may promote musculoskeletal fitness in overweight or obese women. Keywords: intermittent exercise; females; muscular strength; mobility; functional movement patterns,

47 bone health.

Word count: 4,361

Obesity is a multifactorial chronic condition affecting one in three adults worldwide (29). Individuals with obesity demonstrate low cardiorespiratory fitness, high metabolic risk and physical limitations (11,32). This population demonstrates biomechanical deficits in activities of daily living (ADL) and reduced passive range of motion (PRoM) in several joints (19) compared to lean individuals. Such kinetic limitations due to restricted musculoskeletal fitness and mobility levels predispose obese individuals to injuries and lead to impaired functionality and reduced quality of life (42). Overweight or obese adults also exhibit a 15% and 48%, respectively, higher risk of sustaining an injury (35) and these rates are even higher for industry workers (17).

Musculoskeletal fitness is characterised by lean body mass (LBM), muscular strength and endurance, balance and mobility, which contribute to physical performance, resting metabolic rate improvement, osteoporosis prevention and body functionality in ADL (20). Low musculoskeletal fitness levels may predispose obese adults to hip, foot, ankle, knee and shoulder injuries and may cause soft tissue (cartilage, tendons and fascia) damage (42). Although, the etiology is largely unclear, the overloading of the locomotor apparatus combined with the poor musculoskeletal strength and mobility produce impaired mechanics during movement that increases the stress within the soft tissue and bones (42). Obesity may also be related to a lower bone mineral density (BMD) and content (BMC) (36). Although overweight and obese adults tend to have higher absolute BMC values than adults of healthy weight, after adjusting for total body mass, their BMC is markedly lower than that of their controls (36). In other words, the elevated BMD usually measured in adults with obesity may not be adequate to offset the greater forces developed during low- or high-velocity movements.

Regular exercise is an efficient tool for improving physical fitness, health and body
 composition in this population (1). Current exercise guidelines include progressive protocols of
 continuous endurance training (CET), resistance training (RT), or combined CET and RT

training (CT) to induce cardiorespiratory, neuromuscular and metabolic adaptations (1). Structured RT is pivotal for preventing sarcopenia and osteoporosis in sedentary, premenopausal women with obesity through improvements in LBM and physical performance (21,40). High-intensity interval training (HIIT) is one of the most popular exercise modes (3)

requiring less exercise volume compared to CET, RT, or CT and demonstrating high compliance rates when conducted under supervision in untrained individuals (34). HIIT includes repeated short-to-long bouts performed at an intensity that provokes a heart rate (HR) \geq 80% of maximal hear rate (MHR) (39). The present, injury-free, hybrid-type exercise protocol integrates progressive HIIT and functional resistance accessory training into a circuit training format that has been shown to reduce body fat, increase LBM, RMR, endurance, exercise behavioural regulation and vitality with exceptional adherence rates in previously inactive women with obesity (4,6). Such an exercise approach incorporates endurance-based bodyweight drills and resistance-based alternative modalities (18) performed in a circuit, interval format and at moderate-to-high intensity adopting some of the principal characteristics of multimodal integrative neuromuscular training (44).

Although there is evidence that overweight or obese adults need comprehensive exercise strategies not only to reduce body mass and fat (1) but also to improve functional limitations while avoiding physical training-related injuries (37), there is a paucity of longitudinal studies to determine the efficacy of such a HIIT-type neuromuscular exercise approach on musculoskeletal fitness, mobility and bone health. Additionally, it is important to investigate potential changes in such physiological parameters, which result from the cessation of exercise, since exercise training is considered a fundamental component of every lifestyle intervention for this population. It has been observed an unfavorable effect of detraining on neuromuscular performance that was mainly influenced by the duration of training cessation, age, and training status (45). Characteristically, when an 8-month multicomponent exercise program performed by older overweight women was interrupted for only three months, musculoskeletalperformance gains induced by previous training were abolished (49).

To our knowledge, there is no data concerning the effects of training cessation on musculoskeletal fitness parameters in sedentary overweight and obese individuals. We hypothesized that the training would induce favorable changes in musculoskeletal fitness. Therefore, this study aimed to determine the effects of the a HIIT-type, neuromuscular training protocol on (i) muscular strength and muscular endurance, (ii) joints' range of motion, (iii) balance, (iv) functional movement patterns, and (v) bone health of previously inactive, premenopausal Caucasian women with obesity.

111 Materials and Methods

113 Study design

This study is a part of a larger longitudinal research project whose purpose, methodology and primary outcomes are reported elsewhere (6). In this investigation, data upon musculoskeletal fitness are presented. This study was a randomised controlled trial based on a three-group, repeated-measures design in accordance with the Consolidated Standards of Reporting Trial (CONSORT) guidelines (Figure 1) and it was registered at clinicaltrials.gov as NCT03134781. Initially, 102 participants were assessed for eligibility, 66 were recruited and allocated to three groups, and 49 completed the trial as required (see Figure supplemental Digital Content 1).

Participants (36.4 \pm 4.4 years) were randomly assigned to a training (TR, *n*=14), a training detraining group (TRD, *n*=14) or a control (C, *n*=21) group. Following a 4-week adaptive and familiarization period as previously articulated (6), TR performed the 10-month exercise training protocol whereas TRD performed the same protocol for 5 months and then entered a 5-month detraining period. Abstinence from exercise in the detraining group was verified using accelerometry (GT3X-BT, ActiGraph, FL, USA). Accelerometry data were used in the analysis,

only if participants had ≥4 days and ≥10 wear hours/day. Four vector magnitude data were used to calculate daily activity and sedentary time. Data were expressed as steps/day and time in sedentary, light, moderate, vigorous and moderate-to-vigorous physical activity as described (6). Assessments of musculoskeletal fitness were performed at pre-, mid-, and post-training (Figure 1). All assessment procedures were completed with the same order (bone health, flexibility, static balance, functional movement patterns, maximal strength, muscle endurance) at pre-, mid- and post-training.

FIGURE 1 ABOUT HERE

Participants

Participants were medically cleared for strenuous exercise, were non-smokers and of low regular PA or structured exercise for ≥ 6 months before the study. None of them were on medication, diet or nutritional supplementation. Participants were excluded if they missed $\geq 20\%$ of total exercise sessions, changed their eating habits and modified their PA levels during the intervention. Participants during detraining need to have comparable PA levels with pretraining. Participants were informed about all risks, discomforts and benefits associated with the study and provided a written consent. This investigation was carried out in accordance with the guidelines contained in the 2013 Declaration of Helsinki and was approved by the Institutional Ethics Committee. Participants' characteristics are shown in Table 1.

TABLE 1 ABOUT HERE

150 Training

Three small-group (5-10 participants per session), supervised training sessions per week that used asynchronous music in the background were performed on non-consecutive days for TR

(10 months) and TRD (5 months) as previously reported (6). This hybrid-type protocol was organized in four progressive training phases (i.e., phase 1: weeks 1-7, phase 2: weeks 8-14, phase 3: weeks 15-20 and phase 4: 21-40) (Figure 1). The mean weekly exercise volume was \sim 100 min, net exercise time was 6.5-24.0 min per session and total duration per session was 23-41 min (6). Exercises (see Table, Supplemental Digital Content 2) adapted basic movement patterns (squat, hinge, lunge, push, pull, carry, rotation, plank) utilizing portable modalities (suspension belts, balance balls, kettlebells, medicine balls, battle ropes, stability balls, speed ladders, foam rollers, elastic bands) and bodyweight as resistance. Exercises (~10-12 per session) were organized in a circuit format and performed in all planes of motion simulating ADLs. The work-to-rest ratio was varied (1:2, 1:1, 2:1) using an interval duration of 20-40 sec to provide progression (see Table, Supplemental Digital Content 2) (6). Verbal encouragement was provided. Participants were instructed to execute as many repetitions as possible with the correct form and with a controlled, moderate rhythm. A 10-min warm-up and a 5-min cool down period was applied in all sessions (6). HR was monitored and recorded using telemetry (Polar Team Solution, Polar Electro-Oy, Kempele, Finland) aiming to maintain an intensity \geq 75% of MHR throughout each session. Rating (6-20) of perceived exertion (RPE) was recorded at the end of each round in all sessions using the 6-20 Borg scale.

171 Measurements

173 Bone Health

Whole-body BMC and BMD were performed using a dual energy X-ray absorptiometry (DXA) scanner (GE Healthcare, Lunar DPX-NT) in the morning by the same experienced radiologist according to standard procedures (41). Briefly, participants were placed in a supine position with their body aligned along the central horizontal axis, their arms parallel to their body (without touching it), the forearms pronated, hands flat, legs fully extended, and feet secured

a instrument was cal

using a velcro strap to prevent foot movement during the scan. The instrument was calibrated
daily using a calibration epoxy resin phantom. All analyses were performed using the 12.2 GE
enCORE software package.

3 Flexibility, Static Balance and Mobility

A 3-min cycling warm-up preceded mobility testing (three consecutive measurements). Flexibility of lower back and hamstrings was measured under standardized conditions using the modified sit-and-reach test (2). Goniometry (Lafayette 01135, Lafayette Instrument Inc., Lafayette, IN, USA) was applied to assess the PRoM (in degrees) of ankle dorsiflexion, knee extension, hip extension, shoulder extension, and glenohumeral internal rotation (31). Both extremities were examined and the median of their measurements was reported as the value in goniometry.

Static balance was assessed using the modified Romberg test. Participants were asked to stand without shoes on a firm surface, with eyes closed and arms crossed on the chest and the dominant foot placed directly in front of the non-dominant foot. The time to failure was measured manually by stopwatch in sec (30).

The Functional Movement Screening (FMS) with an ICC of >0.8 was used to evaluate functional mobility, postural stability and movement behavior in different settings (23). Two examiners (with an intra-rater reliability of 88.6%) performed this assessment. The FMS has been reported as a simple, quick, non-invasive and suitable movement-based assessment tool for middle-aged, overweight or obese aiming to evaluate their functional capacity levels (23). In brief, the FMS assesses seven fundamental movement tasks (deep squat, hurdle step, in-line lunge, active straight-leg raise, trunk stability push-up, rotary stability, shoulder mobility). Each movement task was scored from 0 to 3 points (0=pain with pattern regardless of quality, 1=unable to perform pattern, 2=able to perform pattern with compensation/imperfection,

=able to perform pattern as directed) and their sum provided the total score ranging from 0 to 21 points (46).

Muscular Strength and Endurance

Maximal isotonic strength (one repetition maximal, 1RM) was assessed using standard procedures for novice and untrained individuals following familiarization as previously described (2) with an intra-class correlation coefficient (ICC) for test-retest trials of 0.88. Two upper-body (vertical chest press, supinated closed-grip lat pulldown) and two lower-body (seated leg extension, lying leg curl) exercises performed on traditional strength training equipment (Panatta Sport, Apiro, Italy) were selected. Muscular endurance was assessed using timed tests for the abdominal (partial curl-up), upper-body (modified kneeling push-up) and lower-body (modified chair squat) musculature (2). All tests required the participants to perform as many repetitions as possible within 60 sec using standard procedures and a 5-min rest was provided between tests (2).

Statistical Analyses

A preliminary power analysis (effect size >0.55, probability error of 0.05, two-tailed alpha level, power of 0.9) using the G*Power 3.0.10 program based on the study design suggested that a sample of 36-40 participants was necessary to identify statistically meaningful trial effects. For all dependent variables, differences (for both "between" and "within" groups) of means (MD) and confidence intervals (CI) were calculated based on mixed models. Cohen's d criteria were used to interpret the magnitude of MD as very small, small, medium, large, very large and huge for values 0.01, 0.20, 0.50, 0.80, 1.20 and 2.0, respectively. No assumptions were made for covariance matrices (unstructured) since repetitions were not that many to significantly reduce the degrees of freedom. All estimations were corrected based on the Bonferroni criterion for multiple comparisons. Results are presented as relative difference in

time (Δ %). Since the variability of the change score in the intervention groups was greater than that in C, the response rate was analysed using the number of differential responders relative to the ratio of variance in TR and C groups providing multiple differential responder groups (10). Statistical significance was set at *p*<0.05. Data were analysed using the SPSS 22.0 software (IBM Corp., Armonk, NY, USA).

Results

No injuries or other adverse effects occurred during the trial. Adherence rates for TR (10-month intervention) and TRD (5-month intervention) were 93.5% and 82.6%, respectively, and a 4% dropout rate was reported. Results are described in brackets as Δ %/95% CI/*d/p* levels.

Bone mineral density and bone mineral content

Changes in BMD and BMC are shown in Table 2. No changes were observed between TR and TRD. TR only improved BMD following training (Table 2) (+1.9%/0.010-0.035/2.61/p<0.001) and BMC (+1.5%/0.04-0.076/2.72/p=0.023) and its response rate to whole-body BMD was 100% (Figure 4).

247 Flexibility, Static Balance and Mobility

Changes in flexibility and passive range of motion are shown in Table 2. At mid-training, TR and TRD demonstrated greater flexibility changes than C (TR vs. C: +38%/3.939-16.490/1.66/p=0.001; TRD vs. C: +34%/3.010-15.561/1.31/p=0.002). At post-training, TR and TRD elicited more favorable changes in flexibility than C (TR vs. C: +40%/4.465-17.130/1.83/p<0.001; TRD vs. C: +26%/0.572-13.238/1.02/p=0.028). No significant differences were observed between TR and TRD. In TR, the response rate to flexibility was 100% (Figure 4).

Changes in PRoM are shown in Table 2. At mid training, TR and TRD showed significant differences than C in hip extension (TR vs. C: +43%/0.678-9.607/1.08/p=0.019; TRD vs. C: +41%/0.393-9.322/1.08/p=0.029), glenohumeral internal rotation (TR vs. C: +24%/5.676-22.991/1.45/p<0.001; TRD vs. C: +17%/1.747-19.062/1.04/p=0.014). No changes were noted between TR and TRD. At post-training, TR demonstrated a trend for a rise in PRoM compared to C in ankle dorsiflexion (+44%/-0.543-10.924/0.86/p=0.088) and shoulder extension (+24%/0.178-15.940/0.93/p=0.057). TR elicited greater PRoM changes than C and TRD in glenohumeral internal rotation (TR vs. C: +27%/7.814-24.519/1.59/p<0.001; TR vs. TRD: +17%/1.636-19.935/1.83/p=0.016). In TR, the response rate to PRoM in the ankle, knee, hip, shoulder, and glenohumeral joint was 93%, 100%, 93%, 86%, and 86%, respectively (Figure 4).

Changes in static balance are shown in Table 2. TR demonstrated a greater rise in static balance than C at post-training, which was close to being statistically significant (+143%/-0.784-64.436/0.80/p=0.058) and its response rate to static balance was 86% (Figure 4). No changes were found between TR and TRD at mid- and post-training.

Changes in FMS are shown in Table 2. TR and TRD induced a greater rise of the FMS total score than C at mid-training (TR vs. C: +49%/3.860-5.855/6.42/p<0.001; TRD vs. C: +45%/3.431-5.426/5.0/p<0.001) and post-training (TR vs. C: +58%/4.559-6.774/7.71/p<0.001; TRD vs. C: +38%/2.631-4.845/4.0/p<0.001). No changes were reported between TR and TRD at mid-training, but TR had more favorable changes in the FMS total score relative to TRD (+14%/0.716-3.142/2.83/p=0.001) at post-training. In TR, the response rate to the FMS total score was 100% (Figure 4).

Muscular strength

Changes in muscular strength are shown in Figures 2A-D. At mid-training, TR exhibited superior changes than C in chest press (+22%/0.244-12.566/0.82/p=0.039), lat pulldown

(+17%/2.767-12.233/1.29/p=0.001), leg extension (+41%/5.986-17.252/2.04/p<0.001), and 281 leg curl (+27%/4.871-12.796/2.62/p<0.001). TRD demonstrated greater changes than C in 282 283 chest (+33%/3.530-15.851/1.59/p=0.001),lat pulldown (+22%/5.053press 14.519/2.16/p < 0.001), leg extension (+39%/5.379-16.645/1.87/p<0.001), and leg curl 284 (+23%/3.656-11.582/2.05/p<0.001). TR and TRD induced comparable strength gains at midtraining.

At post-training, TR showed greater changes than C in chest press (+29%/2.167-14.666/1.23/p=0.005), lat pulldown (+25%/6.475-15.977/2.03/p<0.001), leg extension (+53%/9.133-20.462/2.79/p<0.001), and leg curl (+38%/7.826-16.698/2.71/p<0.001). In leg curl, TR presented greater changes than TRD (+12%/0.105-9.823/0.91/p=0.044). TRD exhibited superior changes than C in chest press (+30%/2.296-14.795/1.50/p=0.004), lat pulldown (+23%/5.511-15.013/2.03/p<0.001), leg extension (+41%/5.633 -16.962/2.23/p < 0.001), and leg curl (+23%/2.862 - 11.733/1.79/p = 0.001). In TR, the response rate to all 1RM measures was 100% (Figure 4).

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Muscular endurance

Changes in muscular endurance are shown in Figures 2E-G. At mid-training, TR resulted in greater changes of muscular endurance than C (curl-up: +75%/11.024– 9.595/3.39/p<0.001; push-up: (+143%/3.402-11.026/2.04/p<0.001;bodyweight squat: +97%/13.060-18.559/6.03/p<0.001) and TRD demonstrated superior changes in muscular endurance than C (curl-up: +87%/13.238-21.810/3.85/p<0.001; push-up: +168%/3.590-11.267/2.56/p<0.001; bodyweight squat: 102%/11.961-18.658/5.33/p<0.001). TR and TRD resulted in similar muscular endurance gains.

At post-training, TR exhibited greater changes of muscular endurance than C (curl-up: +103%/16.185-25.196/5.0/p<0.001;+195%/7.876-15.553/3.14/p<0.001;push-up: bodyweight squat: +136%/18.176-24.872/7.33/p < 0.001) and TRD showed superior changes in 307 muscular endurance than C (curl-up: +83%/12.185-21.196/3.27/p<0.001; push-up: 308 +123%/3.590-11.267/1.99/p<0.001; bodyweight squat: +97%/11.961-8.658/4.66/p<0.001). 309 TR had more favorable changes in muscular endurance relative to TRD (push-up: +32%/0.081-310 8.491/0.80/p=0.044; bodyweight squat: +20%/2.547-9.882/1.69/p<0.001). In TR, the response 311 rate to all muscular endurance measures was 100% (Figure 3).

FIGURE 2 ABOUT HERE

TABLE 2 ABOUT HERE

FIGURE 3 ABOUT HERE

19 Discussion

This 10-month study revealed that the implementation of a HIIT-type, integrated neuromuscular exercise program performed in a real-world gym setting using portable equipment induces considerable improvement of musculoskeletal fitness in previously inactive, overweight or obese women. These adaptations were reduced but not lost after prolonged detraining.

This trial focused on physically inactive, middle-aged, overweight or obese who are characterized by increased cardiometabolic risk (11), poor functional capacity (32), a higher risk for musculoskeletal disorders (42) and physical limitations (19) compared to normoweight women. Overweight and obese adults are more prone to sustain injuries and exhibit knee osteoarthritis than individuals of a normal BMI (35). Hence, progressive, injury-free and effective exercise protocols are critical to reduce functional deficits that are responsible for a smaller PRoM in several joints (19), impaired quality of life and a rising prevalence of injuries in this population (17). In this study, a 10-month training program designed for inactive, overweight or obese women was injury-free and reported high adherence and low dropout rates. This outcome may support the necessity of prescribing progressive and supervised exercise regimens for this population aiming to promote a safe exercise experience that may be pivotal

for behavioural regulation in exercise (4).

Bone adaptations

The implementation of high-impact training for inactive, middle-aged, overweight or obese women is critical for preventing osteopenia, osteoporosis and injuries (20). Training improved whole-body BMD (+1.9%) and BMC (+1.5%) only in TR at post-training indicating that this type of program may meet the essential features of a high-impact, weight-bearing training program capable of activating bone cell mechanisms and hormonal factors. It is worth mentioning that exercise-induced weight loss in this cohort was not accompanied by a decline in BMD as it was seen in overweight or obese elderly (40), which is important for bone health and injury prevention.

Flexibility, Static Balance and Mobility

Due to insufficient use of joints in inactive, overweight or obese individuals, the functional length of muscles' that cross these joints is reduced resulting in decreased PRoM (19). Hamstring and lower back flexibility improved by 40%, whereas PRoM in ankle, hip, shoulder, glenohumeral joints improved by 24-44% after 10 months of implementation. These adaptations were maintained after 5 months of detraining. These results coincide with a 10-38% increase in flexibility of inactive, overweight or obese older adults following long-term RT (13,14,15). These outcomes may be attributed to the features of the protocol, i.e. the incorporation of whole-body multiplanar movements mimicking ADLs. RT may promote flexibility if exercises are performed through a full range of motion to adequately activate both the agonist and antagonist muscle groups (15). Resistive exercises may not only increase muscle mass and contractility but they also improve the strength of tendons and ligaments thereby augmenting joints' PRoM (43). Studies employing compensatory overload models have shown a simultaneous elevation of muscle's strength and tendon's active fibroblast numbers, collagen synthesis and turnover rate (43). The strength of the junction between bone and ligament is also enhanced by this type of training (38). The association between body fat and flexibility performance changes in response to training supports the evidence that body composition may play some role in flexibility and mobility performance in overweight or obese adults (19).

Although this study did not examine a stretching intervention, it appears that improved PRoM in overweight and obese adults demonstrates exceptional trainability to a hybrid-type exercise training protocol and it may be linked to the improved functionality seen in response to this type of training. The role of flexibility as a major fitness component has been questioned (47). Although the goal of this study was not to determine the value of flexibility as a main fitness component in the overweight and obese population, it appears that PRoM may be pivotal for adequate levels of functionality and quality of life. DoIT seems not to induce negative adaptations to motor control, physical performance and injury rate in a population commonly characterized by reduced mobility and functionality despite the lack of static stretching (47). This is an interesting outcome that highlights the rationale for integrated neuromuscular training methodology adapted for overweight and obese individuals although that static stretching has been classified as a major component of exercise prescription for this population (1).

Static balance improved by 150%, an adaptation not lost following detraining. This finding complements the marked (+58%) improvement seen in the FMS score suggesting a noticeable improvement in neuromuscular functional status. The large increase in static balance may be related to the low ceiling effect and the relative insensitivity to change of the assessment used, especially in younger individuals without clinical neurological conditions or balance impairments (27). However, sedentary populations with obesity are likely to demonstrate significantly impaired components of motor skills related to fitness such as balance and coordination (19). Thus, a 10-month intervention incorporating various neuromotor exercises into a structured training regimen with a frequency of 3 times per week may reasonably promote a large improvement in this cohort.

The FMS testing battery was used as an assessment tool only since its internal and external validity as a predictive tool for injury has been questioned (23). Although there is no data on the effects of various exercise training modalities on the FMS score in untrained, overweight or obese adults, this score (<15) for sedentary middle-aged women is considered moderate-to-low (33). Considering that individuals with obesity demonstrate biomechanical deficiencies in ADLs (19), neuromuscular-type protocols may aid to reduce these limitations by using progressive integrated neuromuscular exercises characterized by multiple angles and planes of motion such as bending, lifting, pushing, pulling, carrying, and rotating (22) using non-traditional portable modalities (18). Such training introduces increased cognitive and motor processing demands that ultimately favor not only strength but also body and joint stability, coordination, balance and PRoM. These outcomes are aligned with recent evidence suggesting that the functionality and mobility of overweight or obese women be improved through neuromotor training programs (37).

Improvements of knee flexor and extensor strength in response to neuromuscular exercise training, as in this study, are associated with increased balance and gait (48) that ultimately improves functional status and reduces falls (16). A potential explanation for these adaptations may be related to the increase in LBM and strength that are seen in response to similar protocols (7). Another explanation for static balance adaptations may be linked to the activation of the vermis of the cerebellum, which is the principal part of a central coordinating mechanism (27). Additionally, postural sensibility to convey information concerning position may play an important role for improving the function of sensory pathways and proprioception (27). These findings highlight the need to integrate multicomponent neuromuscular exercise interventions of sufficient power in the muscular system.

Strength improved in both upper- (+25-29%) and lower-body (+38-53%) in response to training. Interestingly, detraining did not reverse these gains in upper- (+23-30% vs. pretraining) and lower-body (+23-41% vs. pre-training) musculature. Likewise, muscular endurance increased in upper-body, lower-body and abdominal musculature by 195%, 136% and 103%, respectively. These improvements were also maintained following detraining in upper-body (+104%), lower-body (+73%) and abdominal musculature (+58%) compared to baseline levels. These findings are aligned with previous reports in lean women involved in either a short-term conventional CT or circuit-based whole-body RT (28), suggesting a similar trainability. These results corroborate a previous report of improved musculoskeletal fitness and body composition in response to short-term HIIT-type programs that use a whole-body RT approach (6,25). RT is highly recommended as a fundamental component of an exercise program targeted to preventing, managing and treating obesity while eliciting neuromuscular adaptations in individuals (1). The increase in muscular strength and endurance may be attributed to neuromuscular adaptations (25,28,38) and a rise in DXA-assessed LBM (6).

Detraining

Muscular performance adaptations

Detraining is a serious issue for overweight or obese individuals participating in exercise interventions (26). There are no data for the impact of detraining on the adaptations obtained from hybrid, HIIT-type programs. In this study, training gains were reduced but not eliminated following a 5-month detraining period. This outcome corroborates previous findings suggesting that musculoskeletal fitness may be maintained above pre-training levels ever after a training cessation of 5 months or longer if previous training was of sufficient intensity (14). Additionally, it has been documented that RT status may limit the type of neural adaptations that are responsible for the increase in muscular strength and probably the speed of reversibility 437 (45). As such, previously untrained individuals are likely to rapidly lose the adaptations induced
438 by short-term (8-12 weeks) RT programs during a detraining period (45). Detraining-induced
439 loss of musculoskeletal fitness seems to be intensity-dependent (14) and may be associated with
440 an attenuation of muscle fiber size and motor unit recruitment efficiency (24).

Practical applications

The outcomes of this study coincide with studies using HIIT-type protocols (34,36) suggesting that ~100 min of training per week without changes in eating patterns and habitual PA may be an effective long-term approach for musculoskeletal fitness improvement in inactive overweight or obese women. Interestingly, prolonged detraining did not abolish the musculoskeletal fitness adaptations obtained from this fully-supervised longitudinal exercise intervention. These findings underline a safe, time-efficient and motivating (4) exercise approach to promote musculoskeletal health in overweight or obese women that may be a valuable addition to current exercise recommendations for this population (1). However, further research is needed in this area investigating the efficacy of such an exercise protocol in males and other age and race groups as previously described (5).

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13 457	Disclosure of interest
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Figure 1.

CONSORT diagram of the study.

Figure 2.

Experimental flowchart.

TR, training group (5 months); TRD, training (5 months) - detraining (5 months) group; DoIT, exercise protocol; PRoM, passive range of motion; FMS, functional movement screen; ¹for all groups (4-week adaptive period); ²only for TR and TRD; ³for all groups.

Figure 3.

Changes in muscular strength and endurance throughout the experimental period.

C, control group; TR, training group; TRD, training-detraining group; 1-RM, one repetition maximum; reps, repetitions; † different from Pre, p < 0.05; ‡ different from Mid, p < 0.05; § different from C, p < 0.05; # different from TR, p < 0.05.

Figure 4.

Multiple differential responder groups to exercise in TR following a 10-month intervention.

1-RM, one repetition maximum; PRoM, passive range of motion; FMS, functional movement screen.

Table 1.

Participants' baseline characteristics (range of values are shown in parentheses).

	C (n = 21)	TR (n = 14)	TRD (n = 14)
Age (yrs)	36.0 ± 4.2	36.4 ± 5.0	36.9 ± 4.3
	(30.0 - 44.0)	(30.0 - 45.0)	(30.0 - 45.0)
Body mass (kg)	80.2 ± 8.9	78.0 ± 9.9	78.7 ± 7.9
	(69.0 – 103.0)	(64.0 – 97.5)	(68.0 - 91.0)
Body height (m)	1.65 ± 0.5	1.66 ± 0.5	1.64 ± 0.6
	(1.55 – 1.75)	(1.60 – 1.77)	(1.55 – 1.76)
BMI (kg⋅m ⁻²)	29.6 ± 3.0	28.2 ± 2.8	29.1 ± 3.0
	(27.0 – 33.6)	(25.9 – 34.3)	(26.9 – 31.5)
PA (steps·day ⁻¹)	$6,400 \pm 1,851$	6,331 ± 1,042	$6,870 \pm 2,031$
	(2,694 – 9,025)	(4,358 – 8,676)	(2,865 - 9,452)
Body fat (%)	46.7 ± 6.5	47.5 ± 3.2	46.2 ± 3.9
	(35.7 – 58.3)	(41.4 – 53.2)	(38.0 - 52.5)

C, control group; TR, training group; TRD, training-detraining group; BMI, body mass index; PA, physical activity; Data are means \pm SD.

	С		TR			TRD			
Variables	Pre	Mid	Post	Pre	Mid	Post	Pre	Mid	Post
Modified sit and reach (cm)	27.3 ± 7.0	27.1 ± 6.7	27.0 ± 6.7	28.1 ± 6.5	$37.3\pm6.6^{~\dagger,\$}$	$37.8 \pm 6.9^{+, \ddagger, \$}$	27.7 ± 8.1	$36.4\pm8.8^{~\dagger,\S}$	$33.9 \pm 8.6^{+, \ddagger, \$}$
Ankle dorsiflexion (deg.)	12.0 ± 7.0	12.1 ± 6.9	11.8 ± 7.0	11.3 ± 7.7	$16.1\pm8.6~^\dagger$	$17.0\pm7.6~^\dagger$	11.7 ± 6.4	14.9 ± 5.4	13.8 ± 4.9
Knee extension (deg.)	41.0 ± 30.9	41.4 ± 31.1	41.9 ± 30.8	34.5 ± 9.3	$21.6\pm8.3~^\dagger$	$21.6\pm8.2~^\dagger$	43.0 ± 37.7	$35.6\pm40.3~^\dagger$	37.4 ± 38.7 ^{†,‡}
Hip extension (deg.)	12.3 ± 6.3	12.1 ± 5.7	11.8 ± 5.6	12.7 ± 6.8	$17.3\pm4.5~^{\dagger,\$}$	$18.1\pm4.6^{~\dagger,\S}$	11.1 ± 5.1	$17.0\pm5.0^{~\dagger,\S}$	$15.4\pm4.7~^\dagger$
Shoulder extension (deg.)	33.1 ± 10.8	32.8 ± 10.0	32.5 ± 9.9	33.5 ± 11.5	$39.5\pm9.1~^\dagger$	$40.4\pm8.3^{\dagger,\ddagger,\$}$	31.6 ± 9.6	$39.8\pm10.0~^{\dagger}$	$37.3\pm9.7^{~\dagger,\ddagger}$
Glenohumeral rotation (deg.)	60.3 ± 12.8	59.8 ± 12.3	59.5 ± 12.5	67.1 ± 11.2	$74.1\pm7.8~^{\dagger,\$}$	$75.6\pm6.9^{~\dagger,\$}$	58.6 ± 10.8	$70.2\pm8.1~^{\dagger,\$}$	$64.9 \pm 7.0^{+,*,\#}$
Sharpened Romberg (sec)	24.1 ± 16.3	22.8 ± 14.5	22.3 ± 13.9	21.7 ± 26.3	37.4 ± 36.8	$54.2\pm61.2~^\dagger$	23.8 ± 18.4	40.2 ± 36.7	33.9 ±32.7
FMS (total score)	10.00 ± 1.10	9.86 ± 1.28	9.76 ± 1.09	10.14 ± 1.51	$14.71\pm0.73^{~\dagger,\$}$	$15.43 \pm 0.65^{+1.5}$	10.29 ± 1.14	$14.29\pm1.13^{~\dagger,\$}$	$13.50 \pm 1.91^{~\dagger, \ddagger, \$, \#}$
Whole-body BMD (g/cm ²)	1.192 ± 0.063	1.194 ± 0.064	1.193 ± 0.063	1.180 ± 0.060	1.187 ± 0.056	$1.202 \pm 0.058^{+,+}$	1.196 ± 0.066	1.201 ± 0.066	1.195 ± 0.066
Whole-body BMC (g/cm ²)	2.576 ± 0.24	2.574 ± 0.24	2.573 ± 0.24	2.599 ± 0.21	2.609 ± 0.23	$2.639 \pm 0.22^{+}$	2.587 ± 0.24	2.598 ±0.26	2.589 ± 0.25

Table 2. Changes in flexibility, passive range of motion, static balance, FMS and bone health throughout the experimental period.

C, control group; TR, training group; TRD, training-detraining group; FMS, functional movement screen; BMD, body mineral density; BMC, body mineral content; [†] different from Pre, p < 0.05; [‡] different from Mid, p < 0.05; [§] different from C, p < 0.05; [#] different from TR, p < 0.05.



Figure











CONSORT 2010 Flow Diagram



Supplemental Digital Content 1. The exercises of the training protocol.

Exercises					
Adjunct Modalities	Phase 1	Phase 2	Phase 3	Phase 4	
1 – Balance Ball	Over dome ankle touch	Straddle jump	Split jack	Over dome hand touch	
2 – Suspension Exercise Device	Neutral grip row	Wide grip row	Y deltoid raise	Chest press	
3 – Kettlebell	Sumo deadlift	Sumo deadlift high pull	Two-arm swing	Two-arm snatch	
4 – Bodyweight	Straight-arm plank	Forearm plank	Straight-arm reverse plank	Side plank rotation	
5 – Speed Ladder	Low knee skip	Lateral shuffle	Heel flick	High knee skip	
6 – Battling Rope	Bilateral wave	Alternating wave	Side-to-side wave	Slam	
7 – Medicine Ball	Alternating static lunge	Forward lunge with press	Lunge to chest pass	Twisting chop	
8 – Foam Roller	Forearm plank	Forearm plank with leg lift	Shifting Plank	Forearm plank with leg lift	
9 – Bodyweight	Jumping jack	Split jack	Ice skater	Burpee	
10 – Resistance Band with Stick	Squat to overhead press	Lateral shuffle press	Hockey slap shot	Axe chop	
11 – Resistance Band		Squat row	Reverse fly with lunge	Squat to overhead press	
12 – Medicine Ball			Squat throw	Swing	

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	Phase 1	Phase 2	Phase 3	Phase 4
Training Parameters	(Week 1-7)	(Week 8-14)	(Week 15-20)	(Week 21-40)
Session duration (min)	23.0	38.0	41.0	41.0
Effort time (min) ^a	6.66	16.5	24.0	24.0
Recovery time (min) ^b	16.34	21.5	17.0	17.0
Work-to-rest ratio	1:2	1:1	2:1	2:1
Work interval (sec)	20.0	30.0	40.0	40.0
Rest interval (sec)	40.0	30.0	20.0	20.0
Exercises amount	10	11	12	12
Rounds	2	3	3	3
Rest time/round (min)	3.0	2.5	2.5	2.0
Movement number ^c	Maximal	Maximal	Maximal	Maximal

Supplemental Digital Content 1. The characteristics of the training protocol.

^aEffort time = session duration – recovery time.

^bRecovery time = session duration – effort time.

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