

1 Sex Comparison of Knee Extensor Size,
2 Strength And Fatigue Adaptation to Sprint
3 Interval Training

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22 **Keywords:** Sprint Interval Training, Exercise, Skeletal Muscle, Fatigue, Torque-Velocity

23 Relationship

24 **Abstract**

25 **Background:** Regular sprint interval training (SIT) improves whole-body aerobic capacity and
26 muscle oxidative potential, but very little is known about knee extensor anabolic or fatigue
27 resistance adaptations, or whether effects are similar for males and females. The purpose of
28 this study was to compare sex-related differences in knee extensor size, torque-velocity
29 relationship and fatigability adaptations to 12 weeks SIT.

30 **Methods:** Sixteen males and fifteen females (mean (SEM) age: 41 (± 2.5) yrs) completed
31 measurements of total body composition assessed by DXA, quadriceps muscle cross-sectional
32 area (CSA_Q) assessed by MRI, the knee extensor torque-velocity relationship (covering 0 –
33 240°·sec⁻¹) and fatigue resistance, which was measured as the decline in torque from the first
34 to the last of 60 repeated concentric knee extensions performed at 180°·sec⁻¹. SIT consisted
35 of 4 x 20 second sprints on a cycle ergometer set at an initial power output of 175% of power
36 at VO₂max, three times per week for 12 weeks.

37 **Results:** CSA_Q increased by 5% (p=0.023) and fatigue resistance improved 4.8% (p=0.048),
38 with no sex differences in these adaptations (sex comparisons: p=0.140 and p=0.282,
39 respectively). Knee extensor isometric and concentric torque was unaffected by SIT in both
40 males and females (p>0.05 for all velocities).

41 **Conclusions:** 12 weeks SIT, totalling 4 minutes very intense cycling per week, significantly
42 increased fatigue resistance and CSA_Q similarly in males and females, but did not significantly
43 increase torque in males or females. These results suggest that SIT is a time-effective training
44 modality for males and females to increase leg muscle size and fatigue resistance.

45 INTRODUCTION

46 In recent years, there has been a resurgence of research interest in high intensity interval
47 training and sprint interval training (SIT). Studies usually set out to understand the cellular
48 regulation of training adaptations and to investigate the implementation of training practices
49 to improve health status of various populations (6). However, the majority of research in
50 these areas included only young, male participants, with females and middle-aged people
51 being under-represented.

52 It should not be taken as a certainty that the findings from studies including only males will
53 apply equally to females. Males have higher maximal skeletal muscle strength and power
54 compared with females (30) and higher maximal power output during sprinting (16, 19).
55 However, the superior performance does not lie entirely with males, since males fatigue more
56 quickly than females during controlled isometric contractions of single muscle groups (25, 35)
57 and during sprinting, while females recover faster during short rest periods between repeated
58 sprinting bouts (16, 19).

59 Male advantage when producing maximal muscle force and power is in part due to the higher
60 relative muscle mass (8), larger muscles (28) and larger fibre cross-sectional areas in males
61 than females (18). Any sex-related comparisons for muscle force and power should therefore
62 normalise values to muscle size ('normalised' force and power), sometimes termed as
63 "muscle quality" in the literature (22, 30). However, it is not only muscle mass that exhibits a
64 sex-related difference, but also the contractile and metabolic characteristics. For instance,
65 males have been reported to have higher concentrations of glycolytic enzymes and faster
66 rates of contraction and relaxation than females (35). Males may use relatively more
67 carbohydrates during sub-maximal aerobic exercise than females (45), shift to anaerobic

68 metabolism at lower relative intensity during incremental exercise (39) and during maximal
69 sprinting higher glycolytic contributions were reported in males compared with females (17).
70 Skeletal muscle characteristics such as these that affect energetics may also confer sex
71 differences in fatigability and may influence adaptations to SIT.

72 Few studies directly examine sex-related differences in adaptations to SIT. We recently
73 reported sex differences for the changes to maximal rate of oxygen uptake (VO_{2max}) and
74 body fat after 12 weeks SIT (5). Focussing on peak power output, Esbjörnsson-Liljedahl et al.
75 (18) showed that females increased their power output more than males after 4 weeks SIT,
76 but another study looking at adaptation to just 6 SIT sessions reported similar gains in peak
77 power for males and females (4).

78 Increased power output could in theory be due to changes to neural activation, but the results
79 from Esbjörnsson-Liljedahl et al. (18) allude to proportionally larger gains in vastus lateralis
80 fibre cross sectional area (particularly Type IIx) in females than males as a mechanism. This
81 would suggest that females have a greater hypertrophic adaptation to SIT than males, but
82 there is little evidence to this effect because muscle-specific hypertrophy has been largely
83 overlooked in studies of SIT. Hypertrophy follows a net increase in anabolic signalling over
84 time, and in this regard Fuentes et al. (2012) found no sex-related differences in their anabolic
85 responses to a single SIT session (20). Another study, however, reported approximately 150%
86 higher rates of muscle protein synthesis 48 hours following 3 weeks of SIT in males than
87 females (40). This is in conflict with the previous report of higher increases in Type IIx muscle
88 fibre CSA in females than males (18), although it should be noted that type IIx fibres typically
89 account for approximately 10% of vastus lateralis fibres (44). Irrespective of the conflicting
90 reports, neither of these cross-sectional studies of acute training adaptation examined
91 changes to muscle size after several weeks SIT. Measurements of total body lean mass by

92 DEXA are contradictory in this area. Heydari et al. observed a 2% significant increase after 12
93 weeks HIIT in young, overweight males (24), whereas Trapp et al. (utilising the same protocol
94 but over 15 weeks) saw no change in total body lean mass in young females (21). These
95 studies taken together suggest that the muscle hypertrophic response to SIT may be sex-
96 specific, but the contradictory results highlight the need for further evidence.

97 Thus, the aim of the present study was to compare sex-related differences in muscle size,
98 knee extensor torque-velocity relationship and fatigability (occurring after 60 maximal
99 voluntary concentric knee extensions), and their adaptation to 12 weeks SIT. It was
100 hypothesised that 12 weeks cycling SIT would promote knee extensor hypertrophy, increased
101 torque and fatigue resistance. Based on the limited available data indicating larger gains in
102 type IIx fibre CSA in females than males, females were hypothesised to have a greater
103 hypertrophic adaptation compared with males. Assuming no change to the muscle quality, it
104 was further hypothesised that females would increase maximal torque more than males in
105 line with the greater hypertrophic adaptation.

106 **METHODS**

107 **Experimental Approach to the Problem**

108 To determine the effect of SIT on knee extensor hypertrophy, torque production and fatigue
109 resistance, 16 males and 15 females were recruited from the general population through
110 advertisement in local and national newspaper articles, a local gym and campus
111 advertisement. The participants reported to the laboratory and completed measurements of
112 knee extensor cross sectional area (Magnetic Resonance Imaging, MRI), total body
113 composition (Dual Energy X-ray Absorptiometry, DXA) and knee extensor torque and fatigue

114 (Unilateral Knee Extension Dynamometry). The participants then completed 12 weeks of SIT
115 in a local gym or in the laboratory before returning to repeat the measurements.

116 **Subjects**

117 The study conformed to the latest revisions of the Declaration of Helsinki (47) and was
118 approved by the Ethics Committee at Manchester Metropolitan University. Volunteers
119 provided written, informed consent prior to participation. Those with a history of
120 cardiovascular, neuromuscular or metabolic disease were excluded as well as people whom
121 had suffered a leg fracture within the past two years. Participants who were involved in
122 competitive sports or cycled for more than 15 minutes per day for three or more days per
123 week were also excluded. The included participants were from the same group as previously
124 reported in (5). Included participants ranged from 20 to 69 years. Although not reported in
125 the present manuscript, the mean VO_2 max values (43 and 34 mL.kg.min⁻¹ for males and
126 females, respectively) previously reported for the study group (5) were at around the 60th
127 percentile of population-based values (2, 10, 46). A total of 31 participants completed the 12-
128 week SIT intervention and the primary outcome measurements for this study were changes
129 to: quadriceps cross-sectional area (CSA_Q), knee extensor maximal isometric and concentric
130 torque, and fatigue resistance. Participant characteristics are shown in *Table 1*.

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140 **Table 1. Participant characteristics and muscle size in males (n=16) and females (n=15) before and after 12**
 141 **weeks SIT.**

	Males (Pre)	Males (Post)	Females (Pre)	Females (Post)	Time effect (p- value)	Sex effect (p-value)	Sex x Time interaction (p-value)
Age (yrs)	40.8 (3.2)		40.9 (3.9)				
SIT Sessions		32 (2)		34 (2)		0.602	
VO ₂ max (L·min ⁻¹)	3.5 (0.2)	3.6 (0.2)	2.0 (0.1)	2.4 (0.1)	0.032	<0.001	0.595
VO ₂ max (mL·kg·min ⁻¹)	43.7 (2.1)	45.8 (1.7)	34.3 (2.3)	40.8 (2.4)	0.001	0.013	0.815
Total body mass (kg)	80.0 (2.2)	79.2 (2.1)	62.5 (2.6)	62.1 (2.5)	0.039	<0.001	0.207
Height (m)	1.75 (0.02)		1.66 (0.01)				
Body Mass Index (kg·m ²)	25.9 (0.9)	25.8 (0.9)	21.8 (0.7)	21.8 (0.7)	0.055	0.001	0.495
Body Fat (%)	22.5 (1.4)	21.3 (1.4)	30.4 (1.4)	29.9 (1.3)	0.807	<0.001	0.133
Total body lean mass (kg)	60.4 (1.6)	61.1 (1.6)	39.1 (0.9)	39.2 (1.0)	0.147	<0.001	0.067
Leg lean mass (kg)	21.6 (0.7)	21.0 (0.7)	13.6 (0.3)	13.0 (0.4)	0.992	<0.001	0.990
Right thigh lean mass (kg)	5.0 (0.2)	5.1 (0.2)	3.4 (0.1)	3.5 (0.2)	0.547	<0.001	0.880
CSA _Q (cm ²)	86.2 (3.3)	89.7 (3.1)	56.7 (1.9)	60.0 (1.6)	0.023	<0.001	0.140

142 *Data are shown as mean (SEM). Knee Extensor CSA (cm²) is measured in 15 males and 10 females due to*
 143 *equipment maintenance.*

144 Procedures

145 Total body and leg lean mass were assessed by dual-energy x-ray absorptiometry (DXA: Lunar
146 Prodigy Advance; GE Medical; EnCore version 10.50.086) using the same procedures as
147 previously reported by our group (5), and thigh lean mass was also recorded (33). The test-
148 retest variation in measurement for this equipment has been determined in our laboratory
149 as 1% (*unpublished*).

150 Magnetic resonance imaging (MRI) was used to measure peak quadriceps cross-sectional area
151 (CSA_Q) using a T1-weighted turbo 3D sequence (256x256 matrix, Repetition Time 40ms, Echo
152 Time 16ms) on a 0.25-T scanner (G-scan, Esaote, Genoa, Italy) with the participant supine and
153 hips and knees fully extended. The scanning coil was positioned over the thigh of the
154 dominant leg and contiguous transverse-plane slices of 6 mm thickness were collected with
155 no gap between slices. Images were analysed using OsiriX imaging software (OsiriX medical
156 imaging, OsiriX, Atlanta, USA) by manually tracing the quadriceps muscles and avoiding any
157 visible fat deposits in the muscle. Slices at 24mm apart were analysed and the slice with the
158 highest quadriceps anatomical cross-sectional area was recorded. Analyses were carried out
159 by the same investigator. Using the same equipment and measurement techniques, our
160 laboratory previously reported a co-efficient of variation of 0.43, 0.35, 0.30 and 0.31% for
161 repeated measurements of *Vastus Lateralis*, *Rectus Femoris*, *Vastus Medialis* and *Vastus*
162 *Intermedius* muscles, respectively (14). The scanning procedure was repeated after 12 weeks
163 of SIT.

164 The knee extensor torque-velocity relationship and fatigue resistance were assessed in 16
165 males and 15 females using unilateral extensions on a Cybex Norm Dynamometer (Cybex,
166 division of Lumex Inc, Ronkonkoma, New York, USA). Participants were seated upright (hip

167 angle of 85°) with straps secured firmly around the upper body and the hips to limit
168 extraneous body movements. The torque lever was strapped 2 cm above the ankle malleolus
169 of the dominant leg (as determined by the participant) and the centre of knee rotation was
170 aligned with the point of rotation of the dynamometer lever arm. A brief warm up included
171 six isokinetic contractions at 180°·sec⁻¹ using approximately 60-70% of maximal effort. The
172 maximal voluntary isometric torque (MVC) was assessed three times with 60 seconds of rest
173 between efforts at a knee angle of 90°. The highest torque value was recorded. Following a 3
174 minute rest, isokinetic torque was assessed over two efforts separated by 60 seconds rest
175 between efforts at any velocity at 60, 120, 180, 240°·sec⁻¹ in a random order, blinded from
176 the participant, with a 60 second rest between velocities. Each trial started with the leg flexed
177 as far as possible and participants made two maximal efforts at each velocity through the full
178 range of movement until the leg reached full extension. The peak torque occurring at any
179 point during the concentric contraction was recorded. A rest of 30 seconds was given between
180 maximal efforts and strong verbal encouragement was given throughout. The data obtained
181 from MRI scanning was then combined with the data from the isokinetic torque production
182 to measure 'normalized torque', that is, torque produced per cm² of CSA_Q, in order to give an
183 indication of muscle quality before and after SIT:

184
$$\text{Normalized Torque} = \text{Isokinetic Torque (Nm) at a given velocity (}^\circ\cdot\text{sec}^{-1}\text{)} \div \text{CSA}_Q \text{ (cm}^2\text{)}$$

185 Isokinetic torque as a percentage of isometric torque produced was assessed by:

186
$$\% \text{Isometric Torque} = \text{Isokinetic Torque (Nm) at a given velocity (}^\circ\cdot\text{sec}^{-1}\text{)} \div \text{Isometric Torque}$$

187
$$\text{measured at } 90^\circ \text{ (Nm)}$$

188 Knee extensor fatigue resistance was assessed after a 3 minute rest. The test started with the
189 knee flexed as far as possible and participants performed 60 maximal-effort isokinetic

190 contractions over 2 minutes (one every 2 seconds, as timed by a metronome), moving through
191 the full range of knee extension at a velocity of $120^{\circ}\cdot\text{sec}^{-1}$ and returning to the fully flexed
192 knee angle between contractions. The highest torque produced during the first 3 contractions
193 was recorded as the highest contraction torque during the test and in all cases the lowest
194 torque was produced during the final contraction and this was recorded as the lowest torque.
195 The fatigue index was calculated using the formula:

$$196 \quad \text{Fatigue Index} = (\text{Torque Produced in final contractions} \div \text{Torque Produced in First} \\ 197 \quad \text{Contractions}) \times 100$$

198 In this instance, a higher value indicates greater fatigue resistance, *i.e.* the percentage of
199 muscle torque output maintained after 60 contractions relative to the first contraction.

200 The isometric MVC ICC using these techniques is 0.854 with isokinetic variation ICC of 0.819
201 at $60^{\circ}\cdot\text{sec}^{-1}$, 0.810 at $120^{\circ}\cdot\text{sec}^{-1}$, 0.850 at $180^{\circ}\cdot\text{sec}^{-1}$, 0.836 at $240^{\circ}\cdot\text{sec}^{-1}$.

202 *Sprint Interval Training*

203 Participants completed an incremental cycling test to establish the workload at the maximal
204 rate of oxygen uptake (VO_2max), as previously described (5) and this was used to determine
205 the SIT workload. SIT was completed on cycle ergometers (Cateye, Japan). The training
206 consisted of 2 minutes of warm-up at a self-selected moderate intensity. This was followed
207 by four bouts of 20 seconds maximal effort sprints at a workload (in Watts) that was set at a
208 power output corresponding to 175% of the workload attained in the VO_2max test at the
209 initial laboratory visit. This target workload was increased every two weeks by 5%, reaching
210 200% of the workload attained in the VO_2max test at the initial laboratory visit after 12 weeks
211 (a standard incremental cycling test was used to determine the VO_2max , as described
212 previously (5). Each of these bouts was separated by 2 minutes of very low intensity cycling

213 (a workload of 20% of that attained in the initial VO₂max test). This training protocol was
214 chosen due to its brevity, as well as previous studies yielding significant changes in
215 physiological measurements in short time periods (11, 36). Thus, each training session lasted
216 less than 10 minutes and only 80 seconds was completed at an intensity that would be
217 expected to influence the primary outcome variables: knee extensor size, maximal torque and
218 fatigue resistance.

219 The first training session for each participant was supervised by the research team in the
220 research laboratory and participants received clear instructions on the use of the cycle
221 ergometers and the training regimen. Participants were then instructed to train three times
222 per week for 12 weeks (36 sessions in total) using the ergonomic cycles (Cateye, Japan) that
223 we provided in a local gym or at our laboratory. Participants completed on average 33 (± 2)
224 sessions over 12 weeks, with males and females completing similar numbers of sessions
225 (Table 2). Exercise instructors at the local gym were fully informed of the research and training
226 protocols, they were available to provide a safe training environment and to assist
227 participants if needed during training sessions. The exercise instructors were not involved in
228 the data collection process or in the interpretation of data. Participants maintained a training
229 logbook to record workloads during training sessions and were otherwise asked to maintain
230 their usual dietary and exercise habits throughout the intervention period.

231 **Statistical Analyses**

232 The primary outcome measurements were: peak quadriceps muscle cross sectional area
233 (CSA_Q); torque measured at the different velocities; and fatigue index after 60 maximal effort
234 concentric contractions. A secondary outcome measurement was lean mass measured by
235 DXA. The differences between pre- and post- 12 weeks SIT were calculated and sex-related

236 differences in adaptation were compared. All data were tested for normality of distribution
237 using the Kolmogorov-Smirnov test. Independent samples t-test was used to examine sex-
238 related differences in the number of training sessions completed (*Sex effect, Table 1*) and
239 baseline sex-related differences in all recorded measurements. If no sex difference was found
240 at baseline, a two-factor repeated measures ANOVA was used to assess sex differences in
241 training adaptation and between isokinetic velocities. If a baseline sex difference was found,
242 a two-factor repeated measures ANCOVA was used with baseline values as a co-variate. In
243 examining sex-related differences over the training intervention, the sex x time interaction
244 effect refers to sex-related differences as a result of the training intervention (pre- to post-
245 training intervention). Three-Factor repeated measures ANOVA was used to assess sex- and
246 time related differences in the force-velocity profile. Relationships between measurement
247 outcomes and participant age were examined using partial correlation coefficients controlling
248 for sex. The data were analysed using SPSS (v.20 IBM) and statistical significance was accepted
249 at $p < 0.05$. Data are presented as mean \pm standard error of mean (SEM).

250 Results

251 No significant correlations were found between participant age and the adaptations to SIT
252 for any of the outcome variables (all $p > 0.200$).

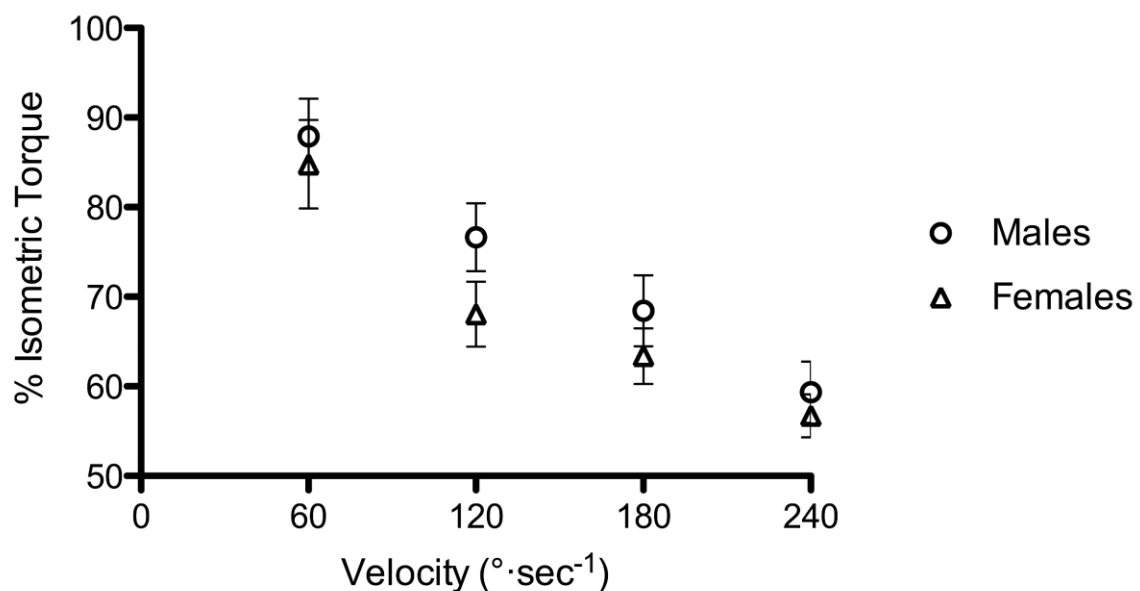
253 Body composition and knee extensor muscle size

254 *Table 1* shows variables relating to body composition and skeletal muscle size in males and
255 females. Total body lean mass was unchanged after training ($p = 0.147$), with no sex difference
256 in this adaptation (sex x time interaction: $p = 0.067$). Analysis of leg lean mass and thigh lean
257 mass from DXA scans showed no significant changes after training. However the more
258 detailed analysis of CSA_Q from MRI showed a significant increase of 3.25 cm² after SIT

259 (p=0.023), with both males and females increasing CSA_Q similarly after training (sex x time
260 interaction: p=0.140).

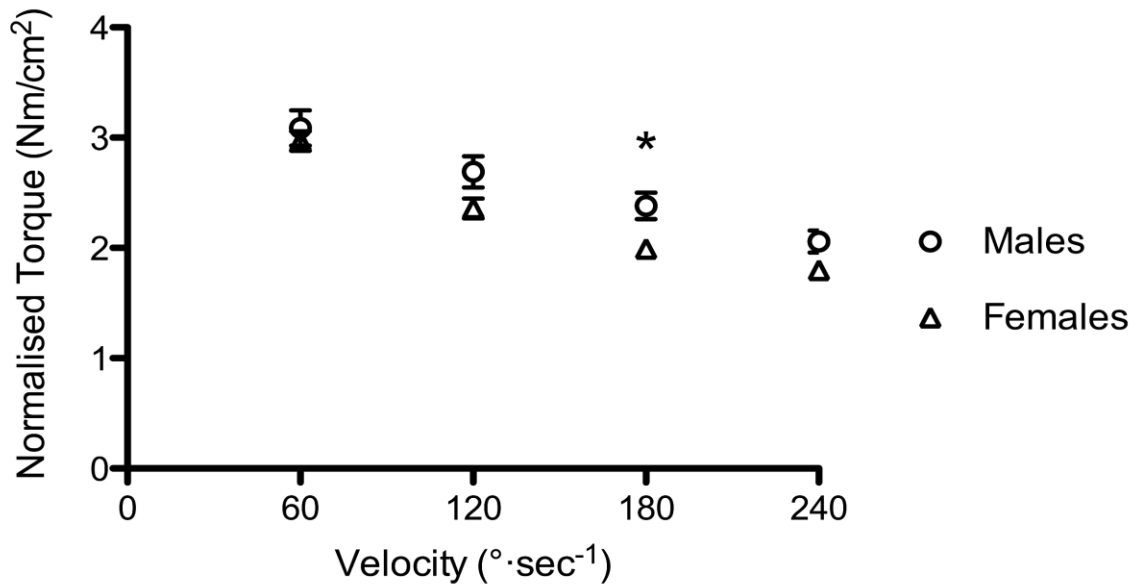
261 Knee extensor maximal torque and fatigue resistance

262 Males had higher isometric torque than females at baseline (*Table 2*), but both sexes generally
263 showed similar percentage decline in torque as contraction velocity increased, with torque at
264 all velocities ($^{\circ}\cdot\text{sec}^{-1}$) relative to isometric MVC being similar for males and females (sex x
265 velocity interaction; p=0.374) (*Figure 1a*). When normalising torque to CSA_Q, males had higher
266 values than females at $180^{\circ}\cdot\text{sec}^{-1}$ (*Figure 1b*), with sex-related difference in the decrease in
267 torque per cross-sectional area of muscle approaching significance (sex x velocity interaction;
268 p=0.051). There were no significant sex-related differences in the torque-velocity profile after
269 SIT (sex x time x velocity interaction; p=0.425).



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271 *Figure 1a: Isokinetic knee extensor torque relative to isometric MVC plotted as a function of contraction*
272 *velocity at baseline (Pre-SIT). Data are mean \pm SEM and plotted separately for males (n=16, circles) and*
273 *females (n=15, triangles). *indicates statistically significant sex-related difference (p<0.05)*



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275 *Figure 1b: Isokinetic knee extensor torque normalised to knee extensor cross sectional area plotted as a*
 276 *function of contraction velocity at baseline (Pre-SIT). Data are mean ± SEM and plotted separately for males*
 277 *(n=16, circles) and females (n=15, triangles). *indicates statistically significant sex-related difference*
 278 *(p<0.05)*

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280 The fatigue index (proportion of torque that remained after 60 maximal effort concentric
 281 contractions) showed no sex-related differences at baseline (*Table 2*). Fatigue index improved
 282 overall by 4.8% after training (p=0.048), with no sex differences in this adaptation (sex x time
 283 interaction; p=0.127) (*Table 2*).

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291 **Table 2. Knee extensor torque in males and females before and after 12 weeks SIT**

	Males (Pre)	Males (Post)	Females (Pre)	Females (Post)	Time effect	Sex effect	Sex x Time interaction
Isometric MVC (Nm)	314.32 (21.86)	322.98 (23.10)	193.33 (9.43)	204.20 (9.65)	0.079	<0.001	0.385
Isometric MVC (Nm·cm ²)	3.65 (0.21)	3.57 (0.24)	3.23 (0.09)	3.23 (0.11)	0.712	0.248	0.705
60°·sec ⁻¹	260.25 (13.78)	265.89 (16.45)	161.23 (9.64)	175.66 (6.45)	0.080	<0.001	0.548
120°·sec ⁻¹	227.01 (12.27)	231.29 (13.42)	128.98 (6.34)	145.22 (4.03)	0.066	<0.001	0.806
180°·sec ⁻¹	199.55 (8.70)	201.50 (11.03)	120.11 (5.05)	120.54 (3.04)	0.495	<0.001	0.522
240°·sec ⁻¹	173.93 (8.22)	172.73 (9.91)	107.57 (3.93)	107.19 (3.25)	0.552	<0.001	0.659
Fatigue Index (%)	52.40 (2.52)	56.42 (3.27)	55.92 (3.56)	64.80 (2.26)	0.048	0.282	0.127

292 *Data are shown as mean (SEM). Males (n=16), Females (n=15)*

293 Discussion

294 This SIT programme included only 4 minutes per week of very high intensity exercise and led
 295 to significant increases to fatigue resistance and CSA_Q, but no change to knee extensor
 296 concentric torque. The training effects were similar for males and females. These findings
 297 advance previous studies of physiological adaptation to SIT which typically focussed on
 298 aerobic and metabolic adaptation in young adult males and females separately.

299 Muscle size

300 There are two contradictory reports in the literature concerning the possible sex differences
 301 in hypertrophic responses to sprint interval or high intensity interval training. Esbjornsson-
 302 Liljedhal et al., reported that females increased Type IIx fibre CSA more than males after 4

303 weeks of SIT (18), while another study reported lower skeletal muscle anabolic response to
304 training in females compared with males (40), but this latter work was based only on acute
305 responses and did not follow up after a period of training. In the present study, we found that
306 males and females showed similar increases in CSA_Q, which are agonist muscles during cycling
307 (7). This could be expected since sprint interval exercise activates Type I and IIx fibres similarly
308 in males and females (17, 41), suggesting motor unit recruitment and therefore the training
309 stimulus received by the IIx fibres, is similar for males and females.

310 A review of the literature indicated that the extent of hypertrophy may depend on the length
311 of the training programme: training 6-weeks or less did not cause significant changes to fibre
312 cross-sectional areas (38). However, longer term training of 7-weeks or more generally
313 increased fibre cross-sectional areas, although it should be noted that studies in this area
314 included only small sample sizes ($n = 8$ to 13) and none of them compared sex-related
315 differences in adaptation (38). For example, a six-week SIT protocol caused a non-significant
316 increase of fibre CSA by 6-12% in 11 untrained males (mean age= 23 ± 5 years) (1). However, 8
317 months SIT increased fibre CSA by 8-16% in 13 athletes (8 males and 5 females, mean age=
318 17 ± 1 years), although sex comparisons were not possible due to small sample sizes and the
319 females in the study were trained in sprinting prior to the intervention whilst the males were
320 not (12).

321 Total body lean mass measured by DXA was unchanged by SIT, with no sex difference in this
322 adaptation (*Table 1*). The leg lean mass and thigh lean mass measured using DXA, also showed
323 no significant changes after training. It is not clear why the DXA detected no significant
324 changes with training, while the MRI clearly showed significant hypertrophy of the quadriceps
325 muscles. The problem is not due to lack of consistency of repeated scans, since Kiebzak et al.

326 (29) observed a 1% variance in lean mass with repeated DXA scans on the same participants
327 over consecutive days and in previous pilot work we determined the test-retest variation to
328 be 1% from our scanner over 4 weeks (*unpublished*). Disparity between MRI and DXA for
329 measuring muscle size has been reported previously, with DXA also failing to reveal the full
330 extent of muscle loss with ageing (32, 33). So it is possible that the DXA is not suitable for
331 detecting these relatively small changes to muscle tissue.

332 Muscle maximal torque

333 The higher values in males compared with females across the concentric torque-velocity
334 relationship are mainly due to the larger muscle mass of males, but there was a clear trend
335 for the torque per muscle cross-sectional area to be higher in males than in females and this
336 was significant at $180^{\circ}\cdot\text{sec}^{-1}$. Torque decreased with increasing knee extension velocity (*Figure*
337 *1b*). These findings fit with previous estimations of torque per muscle cross sectional area,
338 sometimes described as “muscle quality” in the literature. Lindle et al. (30) observed in 346
339 males and 308 females aged 20-90 that males have a 9% higher concentric peak torque per
340 muscle cross sectional area than females. Similarly, Goodpaster et al. (22) observed
341 approximately 17% higher muscle quality in older males compared with older females (mean
342 age= 73 ± 3 years) during isokinetic contractions, suggesting that this sex difference is
343 maintained throughout the lifespan. Previous studies examining sex differences after SIT in
344 muscle strength characteristics have been significantly shorter, not lasting more than 4 weeks
345 (4, 18).

346 The increase in quadriceps muscle size in the present study did not lead to gains in knee
347 extension maximal torque in either males or females. A previous study that measured torque
348 before and after SIT (repeated Wingate tests over 3 weeks) also found no significant changes

349 to knee extensor MVC in 11 males or 9 females (3). It is possible that the mode of exercise
350 training (cycling) might not have trained the neural control needed for isolated knee
351 extensions, as was suggested when the converse was observed when knee extension training
352 did not increase cycling power output (15).

353 **Fatigue resistance**

354 The majority of studies into sex differences in muscle fatigue during controlled exercise of
355 individual muscle groups utilised isometric contractions (25, 26) and the results from such
356 studies generally indicate that females have superior fatigue resistance compared with males
357 (35, 42). Some studies involving concentric contractions also suggest superior fatigue
358 resistance of females compared with males (25, 37, 48). However, fatigability, measured in
359 the present study as the decline in torque after 60 maximal-effort unilateral moderate-
360 velocity concentric knee extensions, was similar in males and females at baseline, with torque
361 during the final contractions dropping to 55% of the first 3 contractions. A possible
362 explanation for why our findings differ from other previous studies is that other studies
363 tended to include young adults or older adults, whereas we included a range of young and
364 middle aged adults. Differences between studies in the velocity of contraction might also
365 influence the fatigability and it is interesting to note that a study utilising maximal velocity
366 contractions at a load equal to 20% of the participant's MVC also reported similar fatigue in
367 males and females during knee extension (42).

368 There are reports that fatigue characteristics of individual muscle groups are unaffected by
369 sprint training in males (23), but no previous studies compared chronic training adaptation of
370 males and females. Fatigue resistance improved after 12-weeks SIT in the present study.
371 When examining isokinetic contractions, previous work suggests a sex dimorphism in muscle

372 fatigue, with males fatiguing significantly faster than females when velocity is controlled.
373 However, when examining as a product of exercise intensity (i.e., a percentage of maximal
374 power or 1-repetition maximum, not a controlled velocity), a sex difference is no longer
375 observed (34). When matched for initial sprinting mechanical work, males and females see
376 similar decline in muscle power after repeated sprint exercise (9, 43), similar to isokinetic
377 contractions which are matched to initial power output, suggesting that the mechanism for
378 increased fatigue resistance in females seen when measuring power output in repeated sprint
379 exercise is an initial higher power output in males (25). In the present study, where training
380 was normalised to cycling power at VO_2Max , it was observed that males and females
381 increased their resistance to fatigue similarly after SIT when measured as a product of
382 contraction velocity (sex x time interaction: $p=0.127$). Taken together, this could suggest that
383 differing prescription of exercise training (i.e., normalised to initial power output versus
384 velocity or body mass etc.) may have sex-related differences in muscle fatigue. The
385 physiological mechanisms underlying the training-induced improvement to fatigue cannot be
386 identified from the present study, but are likely to be associated with increases in
387 mitochondrial concentrations (and therefore improvements in skeletal muscle metabolism)
388 (31) and capillary density (13) that have been found after SIT and collectively improve muscle
389 oxidative energy recovery during the brief rest intervals between contractions.

390 Limitations

391 The design of the training programme, being performed in a local gym or the research
392 laboratory, gave exercise volunteers more control and although this is the case in real-life
393 situations, it may confer less commitment or obligation to training compared with typical fully
394 supervised laboratory-based programmes and we were not able to directly record the power

395 output completed by the study volunteers during training. It was not possible to control for
396 physical activities outside of the training programme and dietary intake was not monitored
397 throughout the training programme. Instead, participants were asked to maintain their usual
398 patterns of food and drink consumption. Finally, there was no control for menstrual cycle
399 variations, which potentially limits the interpretation of some aspects of the data relating to
400 sex-related differences in adaptation. In this regard however, all female participants should
401 equally have completed 3 full menstrual cycles before returning to the laboratory for post-
402 training measurements. Furthermore, evidence points toward there being no change in the
403 muscle parameters measured in this study (27).

404 **Practical Applications**

405 The novel aspects of this research were that we examined maximal knee extensor torque, size
406 and fatigue resistance in males and females ranging from young through to middle-aged
407 adults before and after a period of cycling SIT. Although knee extension torque across a wide
408 range of velocities did not change with training, fatigue resistance and CSA_Q were improved
409 in males and females. Practitioners can use these findings as evidence that SIT is a time-
410 effective option to increase muscle size and resistance to fatigue.

411

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