Effects of sex and joint action on voluntary activation

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The current study tested the hypothesis that a sex and joint-action interaction exists in the voluntary activation during maximal voluntary contraction (MVC). Twenty-eight healthy adults (14 of each sex) performed knee extensor MVC and plantar flexor MVC at extended and flexed knee positions. Voluntary activation during MVC was assessed using a twitch interpolation technique. A two-way analysis of variance demonstrated a significant interaction of sex and joint action. The voluntary activation during plantar flexor MVC at the extended knee position was significantly lower in women (88.3% ± 10.0%) than in men (96.2% ± 6.6%). In contrast, no significant sex differences were shown in the voluntary activation during knee extensor MVC and during plantar flexor MVC at the flexed knee position. The voluntary activation during knee extensor MVC was significantly higher than that during plantar flexor MVC at the extended knee position in women, whereas the corresponding difference was not observed in men. The results revealed that the sex difference in the voluntary activation during MVC depends on joint action.

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12 **Abstract**

13 The current study tested the hypothesis that a sex and joint-action interaction exists in the

14 voluntary activation during maximal voluntary contraction (MVC). Twenty-eight healthy adults

15 (14 of each sex) performed knee extensor MVC and plantar flexor MVC at extended and flexed

16 knee positions. Voluntary activation during MVC was assessed using a twitch interpolation

17 technique. A two-way analysis of variance demonstrated a significant interaction of sex and joint

18 action. The voluntary activation during plantar flexor MVC at the extended knee position was

19 significantly lower in women (88.3% ± 10.0%) than in men (96.2% ± 6.6%). In contrast, no

20 significant sex differences were shown in the voluntary activation during knee extensor MVC

21 and during plantar flexor MVC at the flexed knee position. The voluntary activation during knee

22 extensor MVC was significantly higher than that during plantar flexor MVC at the extended knee

23 position in women, whereas the corresponding difference was not observed in men. The results

24 revealed that the sex difference in the voluntary activation during MVC depends on joint action.

Introduction

26   The magnitude of muscle activation during maximal voluntary isometric contractions (MVCs) is

27   the major determinant of generated muscular force. Voluntary activation (VA%), determined by

28   the twitch interpolation technique (Shield & Zhou, 2004), is an index often used to represent the

29   magnitude of muscle activation.

30   It has been shown that the magnitude of VA% depends on joint action. For example,

31   male participants demonstrated a lower VA% during knee extensor MVC than during plantar

32   flexor MVC (Behm et al., 2002). As a possible reason for the difference, Behm et al. (2002)

33   proposed the difference in the muscle fiber type compositions of agonist muscle groups, i.e., a

34   difficulty in full recruitment of motor units during knee extensor MVC due to the relatively

35   higher proportion of type II fibers of the quadriceps femoris compared with the triceps surae. The

36   proportion of type II fibers of the vastus lateralis, the largest muscle among the quadriceps

37   femoris in both sexes (Ema et al., 2017), is lower in women than in men (Hunter, 2014). If the

38   proposal by Behm et al. (2002) is correct, women may show a higher magnitude of VA% during

39   knee extensor MVC than men. In contrast, the soleus, which has the largest physiological cross-

40   sectional area among the triceps surae (Fukunaga et al., 1992), comprises mainly type I fibers

41   even in men (Johnson et al., 1973), suggesting that any possible sex difference of the soleus fiber

42   type composition will be small. Moreover, given that knee flexion reduces the neural and

43   mechanical contribution of the gastrocnemius to plantar flexion strength (Wakahara et al., 2007),

44   it can be assumed that the sex difference in VA% during plantar flexor MVC, if any, will be

45   small, especially at a flexed knee position. A training program induced a similar extent of

46   strength gains between sexes despite a smaller magnitude of muscle hypertrophy in women

47   (Lemmer et al., 2000; Melnyk et al., 2009), which may indicate greater neural adaptation by the

48 training in women than in men. Considering that the magnitude of VA% before training

49 intervention (Gondin et al., 2005) and its training-induced change (Ema et al., 2018) were related

50 to strength gain, an investigation of the above notions should promote better understanding of the

51 sex dependency in training adaptation.

52   To the best of our knowledge, no studies have investigated the effects of sex and joint

53 action on VA% during MVC simultaneously. In the current study, we determined VA% during

54 knee extensor MVC and plantar flexor MVC at extended and flexed knee positions in both sexes.

55 We tested the hypothesis that a sex and joint-action interaction exists in VA% during MVC.

56

57 **Methods**

58 *Participants*

59 A sample size estimation (G*Power 3.1.7, Kiel University, Germany) was performed to detect a

60 within-between interaction for VA%. The expected effect size, α, power, and correlation among

61 repeated measures were set at 0.25, 0.05, 0.80, and 0.5, respectively. The estimation showed that

62 28 participants are required. It was proposed that physical activity and the existence of practice

63 for strength testing affect the magnitude of VA% (Hunter et al., 2016). Therefore, we recruited

64 untrained healthy young adults, and all participants visited our laboratory in advance for

65 familiarization and practice in performing MVCs with the experimental setting for the right leg.

66 A total of 28 adults (14 of each sex) with no habitual resistance exercises, knee or ankle injuries

67 participated in the study (Table 1). We confirmed no significant sex difference in the magnitude

68 of habitual physical activity, assessed with the long version of the International Physical Activity

69 Questionnaire (Craig et al., 2003). The strength testing for knee extension and plantar flexion

70 was performed on different days in random order among the participants. This study was

71   approved by the Ethics Committee of the Shibaura Institute of Technology (Acceptance number:

72   16-008). All participants were informed of potential risks and the study's purpose, and they

73   provided written informed consent before participation.

74

75   ***Evoked twitch responses***

76   To provide insights into the effects of muscle fiber type composition on VA%, we investigated

77   the twitch contractile properties, because the properties have been reported to be associated with

78   the composition (Harridge et al., 1996; Hamada et al., 2000). Participants sat (for knee

79   extensions) or lay supine (for plantar flexions) on the bench of an isokinetic dynamometer

80   (CON-TREX MJ, PHYSIOMED, Germany) while being secured at the pelvis and torso to the

81   dynamometer with nonelastic straps. The knee and hip joint angles were set at 90° and 80°,

82   respectively, for knee extension (anatomical position = 0°). For plantar flexion, the knee joint

83   angle was 0° (K0) or 90° (K90) and the ankle joint angle was 0° (Kennedy & Cresswell, 2001).

84   The centers of rotation of the dynamometer and the right knee/ankle joints were visually adjusted.

85   Using a constant current variable voltage stimulator (DS7A, Digitimer Ltd, UK), the quadriceps

86   femoris and triceps surae twitch responses were obtained with rectangular pulses of 1 ms. For the

87   quadriceps femoris, to percutaneously stimulate the femoral nerve, a cathode (2 × 2 cm) was

88   placed in the femoral triangle, and an anode (4 × 5 cm) was placed in midway between the

89   superior aspect of the greater trochanter and the inferior border of the iliac crest. For the triceps

90   surae, the tibial nerve was stimulated percutaneously in the popliteal fossa with the cathode and

91   over the ventral aspect of the thigh with the anode. The supramaximal stimulus intensity was

92   determined by increasing the current intensity until plateaus in the twitch torque occurred.

93   Thereafter, five supramaximal twitch responses at a higher current (≥20%) were obtained every

94   10 seconds. Torque signals were recorded at 4 kHz and stored in a personal computer after A/D

95   conversion (PowerLab16/35, ADInstruments, Australia). After low-pass filtering the signal at

96   500 Hz, contraction onset was manually identified as described previously (Ema et al., 2018). A

97   previous study used the time to peak twitch torque (TPT), i.e., the duration from torque onset to

98   peak twitch torque, as an index of estimated muscle fiber type composition (Kubo & Ikebukuro,

99   2010). Moreover, TPT was associated with muscle fiber type composition (Hamada et al., 2000).

100   However, TPT is possible to depend on the magnitude of the peak value of twitch torque, making

101   it unsuitable for comparisons between sexes and between different joint actions. Therefore, we

102   determined the twitch torque at 50 ms from torque onset relative to the peak value of twitch

103   torque (normalized $Twitch_{0-50}$) (Balshaw et al., 2016), and used this metric as the index of

104   estimated muscle fiber type composition. The data were averaged across five contractions.

105

106   ***VA% evaluations***

107   After several warm-up contractions involving two maximal MVCs, participants performed knee

108   extensor/plantar flexor MVCs two times. Verbal encouragement was provided during the

109   contractions. Supramaximal triplet stimulations at 100 Hz were interpolated during and after

110   MVC. If the difference in the peak value of torque before stimulation was above 10%, an

111   additional contraction was requested with sufficient rest between contractions. The VA% was

112   calculated as follows: (1 - [superimposed triplet torque/potentiated resting triplet torque]) × 100

113   (Miyamoto et al., 2012). The mean of the two trials was used for subsequent analyses.

114

115   ***Statistical analyses***

116   Statistical analyses were performed using SPSS version 22 (IBM, USA). All data are shown as

117   means ± standard deviation. A two-way analysis of variance (ANOVA) with repeated measures

118   was conducted to examine the effects of sex (men and women) and joint action (knee extension,

119   plantar flexion in K0 and K90) on dependent variables. When a significant interaction was

120   shown, follow-up ANOVAs with Bonferroni multiple-comparisons were used. Partial η^2 was

121   calculated as an index of effect size for ANOVA. The significance level was set at $P < 0.05$.

122

123   **Results**

124   Figure 1 shows VA% during MVC. There was a significant sex × joint-action interaction (F [2,

125   52] = 3.221, $P = 0.048$, partial $\eta^2 = 0.110$) and significant main effect of sex (F [1, 26] = 4.27, P

126   = 0.049, partial $\eta^2 = 0.141$) but not of joint action (F [2, 52] = 1.198, $P = 0.310$, partial η^2 =

127   0.044). Regarding the sex difference, VA% during plantar flexor MVC in K0 was significantly

128   higher in men than in women ($P = 0.020$), whereas no significant sex difference was found for

129   VA% during knee extensor MVC ($P = 0.501$) or plantar flexor MVC in K90 ($P = 0.086$). For

130   joint action dependency, VA% during knee extensor MVC was significantly higher than that

131   during plantar flexor MVC in K0 in women ($P = 0.001$) but not in men ($P = 0.421$). No

132   corresponding differences were shown between other joint actions in each sex ($P = 0.152$–0.772).

133   The normalized Twitch_{0-50} results are described in Figure 2. A significant main effect of

134   joint action (F [2, 52] = 90.321, $P < 0.001$, partial $\eta^2 = 0.776$) without a main effect of sex (F [1,

135   26] = 0.376, $P = 0.545$, partial $\eta^2 = 0.014$) or an interaction of the two factors (F [2, 52] = 0.607,

136   $P = 0.549$, partial $\eta^2 = 0.023$) was shown. The normalized Twitch_{0-50} of knee extension was

137   significantly greater than those of plantar flexions for both extended ($P < 0.001$) and flexed knee

138   positions ($P < 0.001$).

139

Discussion

The main finding of the current study was that the sex difference in VA% was shown only during plantar flexor MVC at the extended knee position. Compared with men, women showed lower VA% during plantar flexor MVC at the extended knee position. In contrast, corresponding sex difference in VA% was not observed during knee extensor MVC or during plantar flexor MVC at the flexed knee position. In addition, in women but not in men, VA% during plantar flexor MVC at the extended knee position was significantly different from that during knee extensor MVC. These results indicate that the sex difference in VA% during MVC depended on joint action.

The only sex difference in VA% was shown during plantar flexor MVC in K0. The normalized Twitch$_{0-50}$ was not significantly different between sexes for any joint action (Figure 2), suggesting that the muscle fiber type composition is not a major factor for the observed sex and joint-action dependency in VA%. Co-contraction of the antagonist tibialis anterior during plantar flexor MVC might be the explainable factor for the current result. Compared with males, females exhibited higher magnitude of the tibialis anterior activation relative to medial gastrocnemius activation during the push-off phase of countermovement jumping (Márquez et al., 2017). During countermovement jumping, just before take-off, the fascicles of the medial gastrocnemius contracted quasi-isometrically (Kurokawa et al., 2003); therefore, the contraction type of the triceps surae may be partly similar between the current (i.e., isometric contraction) and previous (Márquez et al., 2017) studies. Because of reciprocal inhibition (Crone et al., 1987), co-contraction of the tibialis anterior during plantar flexor MVC could diminish the magnitude of triceps surae activation during plantar flexion. In contrast, the absence of a sex difference in

162 VA% during knee extensor MVC (Figure 1) is in line with previous studies (Krishnan &

163 Williams, 2009; Lee et al., 2017), and the magnitude of hamstring activation as antagonists

164 during knee extensor MVC was not different between sexes (Krishnan & Williams, 2009). Such

165 notions may be also related to the higher VA% during knee extensor MVC than during plantar

166 flexor MVC in K0 only in women. Taken together, a possible sex difference in the antagonist

167 activation may account for the current sex- and joint-action differences in VA%.

168   Another possible factor for the joint-action dependency of sex difference in VA% is the

169 corresponding differences in motor unit firing rates. It was shown that VA% during knee

170 extensor MVC was associated with motor unit firing rates of the vastus lateralis (Knight &

171 Kamen, 2008). A previous study failed to find a significant sex difference in the motor unit firing

172 rates of the vastus medialis, one of the muscles of the quadriceps femoris (Tenan et al., 2013). In

173 contrast, in an animal study, males showed more rapid firing rates than females (English &

174 Widmer, 2003). Therefore, if the aforementioned findings are applicable to other quadriceps

175 femoris muscles and human trieps surae, they may relate to the present VA% results.

176   The lack of significant difference in VA% between joint actions in male participants is

177 not consistent with the previous finding of a higher value of VA% during plantar flexor MVC

178 than during knee extensor MVC in men (Behm et al., 2002). This discrepancy may be related to

179 the difference in participant backgrounds. We recruited untrained participants because resistance

180 training can affect the magnitude of VA% (Ema et al., 2018). In contrast, Behm et al. (2002)

181 examined subjects who participanted in habitual resistance exercises or competitive sport

182 activities. Although the kind of resistance training that the subjects had performed was not

183 mentioned, it is possible that the effects of training on VA% during MVC differed between joint

184 actions, likely resulting in the higher VA% during plantar flexor MVC than during knee extensor

185 MVC (Behm et al., 2002).

186   Previous studies demonstrated that greater muscle hypertrophy in men than in women

187 after resistance training were accompanied by a lack of sex difference in knee extension strength

188 gains (Lemmer et al., 2000; Melnyk et al., 2009). The previous results imply a greater neural

189 adaptation in women than in men, because there was a negative correlation between VA% before

190 training and the magnitude of strength improvement (Gondin et al., 2005). However, no

191 significant sex difference in VA% during knee extensor MVC was found in the present study;

192 therefore, it is difficult to explain the aforementioned results in terms of a sex difference in

193 neural adaptations. In contrast, our data might suggest that the training-induced increase in

194 plantar flexion strength can be expected to be greater in women than in men; future attempts are

195 required to clarify this subject.

196

Conclusion

197

198 The significant sex difference in VA% was limited during plantar flexor MVC at the extended

199 knee position, and only women showed joint action dependency in VA%. These results revealed

200 that there is an interaction of sex and joint action for VA% during MVC.

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268 **Figure Captions**

269 **Figure 1**

270 Scatterplots showing individual data of voluntary activation (VA%) during maximal voluntary

271 contractions of knee extension, plantar flexion at extended (K0) and at flexed (K90) knee

272 positions. Solid line shows the mean value. *Indicates a significant difference between joint

273 actions. †Shows a significant difference between sexes.

274

275 **Figure 2**

276 Scatterplots showing individual data of rate of torque development of twitch torque (RTD_{twitch})

277 of knee extension, plantar flexion at extended (K0) and at flexed (K90) knee positions Solid line

278 shows the mean value. *Indicates a significant difference between joint actions.

Physical characteristics of participants.

MET, metabolic equivalent. Data are shown as mean ± standard deviation.

Table 1 Physical characteristics of participants

		Men (n = 14)				Women (n = 14)		
Age	years	23	±	4		22	±	1
Height	cm	169.7	±	4.4		157.6	±	4.1
Body mass	kg	62.0	±	6.2		51.5	±	6.6
Physical activity	MET min/wk	3108	±	2177		2836	±	2542

MET, metabolic equivalent. Data are shown as mean ± standard deviation.

Figure 1

Scatterplots showing individual data of voluntary activation (VA%) during maximal voluntary contractions of knee extension, plantar flexion at extended (K0) and at flexed (K90) knee positions.

Solid line shows the mean value. *Indicates a significant difference between joint actions.

†Shows a significant difference between sexes.

Figure 2

Scatterplots showing individual data of rate of torque development of twitch torque (RTD_{twitch}) of knee extension, plantar flexion at extended (K0) and at flexed (K90) knee positions.

Solid line shows the mean value. *Indicates a significant difference between joint actions.