Please cite this article in press as: Kidgell DJ et al. Increased cross-education of muscle strength and reduced corticospinal inhibition following eccentric strength training. Neuroscience (2015), http://dx.doi.org/10.1016/j.neuroscience.2015.05.057

Neuroscience xxx (2015) xxx-xxx

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- INCREASED CROSS-EDUCATION OF MUSCLE STRENGTH AND REDUCED CORTICOSPINAL INHIBITION FOLLOWING ECCENTRIC STRENGTH TRAINING
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- 18 Abstract-Aim: Strength training of one limb results in a substantial increase in the strength of the untrained limb, however, it remains unknown what the corticospinal responses are following either eccentric or concentric strength training and how this relates to the crosseducation of strength. The aim of this study was to determine if eccentric or concentric unilateral strength training differentially modulates corticospinal excitability, inhibition and the cross-transfer of strength. Methods: Changes in contralateral (left limb) concentric strength, eccentric strength, motor-evoked potentials, short-interval intracortical inhibition and silent period durations were analyzed in groups of young adults who exercised the right wrist flexors with either eccentric (N = 9) or concentric (N = 9) contractions for 12 sessions over 4 weeks. Control subjects (N = 9) did not train. Results: Following training, both groups exhibited a significant strength gain in the trained limb (concentric group increased concentric strength by 64% and eccentric group increased eccentric strength by 62%) and the extent of the cross-transfer of strength was 28% and 47% for the concentric and eccentric group, respectively, which was different between groups (P = 0.031). Transcranial magnetic stimulation revealed that eccentric training reduced intracortical inhibition (37%), silent period duration (15-27%) and increased corticospinal excitability (51%) compared to concentric training for the

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untrained limb (P = 0.033). There was no change in the control group. *Conclusion:* The results show that eccentric training uniquely modulates corticospinal excitability and inhibition of the untrained limb to a greater extent than concentric training. These findings suggest that unilateral eccentric contractions provide a greater stimulus in crosseducation paradigms and should be an integral part of the rehabilitative process following unilateral injury to maximize the response. © 2015 Published by Elsevier Ltd. on behalf of IBRO.

Key words: corticospinal inhibition, cross-activation, crosstransfer, ipsilateral motor cortex, strength, recovery.

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INTRODUCTION

The potential to increase muscle strength following resistance training is well-documented and overloading skeletal muscle with eccentric strength training has been shown to be superior compared to concentric strength training for increasing muscle strength (Hortobágyi et al., 1996). An interesting observation, originally described by Scripture et al. (1894), is the phenomena of crosseducation, whereby strength training of a single limb was found to increase the strength of the untrained limb. Since this initial report, several studies have provided evidence to support the existence of cross-education using concentric, eccentric and isometric strength training (Cannon and Cafarelli, 1987; Brown et al., 1990; Munn et al., 2005; Lee et al., 2009). Interestingly eccentric training produces the largest changes in strength compared to concentric and isometric (Enoka, 1996; Hortobágyi et al., 1997, 1999). However, the mechanism that modulates the greater cross-education effect following eccentric training remains unknown and untested.

Given lack of muscle hypertrophy in the untrained limb 40 (Farthing et al., 2003), along with reports of increased 41 corticospinal excitability (Kidgell et al., 2011; Goodwill 42 et al., 2012), reduced corticospinal inhibition (Latella 43 et al., 2012), reduced interhemispheric inhibition (IHI) 44 (Hortobágyi et al., 2011) and increased voluntary activa-45 tion (Lee et al., 2009), cross-education is believed to 46 occur as a result of neural adaptations. While direct evi-47 dence to substantiate such claims is increasing, the exact 48 mechanisms and specific locus of adaptation for the 49 cross-education of strength remain unresolved (Carroll 50 et al., 2006; Ruddy and Carson, 2013). 51

Abbreviations: ANOVA, analysis of variance; FCR, flexor carpi radialis; GABA_A, γ -aminobutyric acid; ICI, intracortical inhibition; IHI, interhemsipheric inhibition; iM1, ipsilateral primary motor cortex; ISI, inter-stimulus interval; MEP amplitude, motor-evoked potential; M_{MAX} , maximum compound wave; MVIC, maximal voluntary isometric contraction force; rmsEMG, root mean square electromyography; RMT/AMT, resting and active motor thresholds; sEMG, surface electromyography; SICI, short-interval intracortical inhibition; SP, silent period; TMS, transcranial magnetic stimulation.

http://dx.doi.org/10.1016/j.neuroscience.2015.05.057

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52 Two theories for the potential mechanism underpinning cross-education have been presented 53 (Ruddy and Carson, 2013 for detailed review). Firstly, 54 the 'bilateral-access' hypothesis involves the develop-55 ment of motor engrams (i.e. reorganization of movement 56 representations within the M1) following unilateral prac-57 tice, that can be accessed not only by the trained limb, 58 but also by the untrained limb. The second 'cross-59 activation' hypothesis is based on the concept of unilat-60 eral contractions being driven by bilateral cortical activity 61 in both the contralateral M1 and the ipsilateral primary 62 motor cortex (iM1), producing lasting neuroplastic 63 changes in both cortices. Certainly, transcranial magnetic 64 65 stimulation (TMS) studies have shown that bilateral corticospinal excitability is facilitated by high-force contrac-66 tions, with the scale of ipsilateral corticospinal effects 67 being relative to the level of force gradation (Dettmers 68 et al., 1995; Muellbacher et al., 2000; Hortobágyi et al., 69 2003). However, the neural adaptation underpinning 70 71 cross-education following unilateral eccentric training remains unknown. 72

The corticospinal control of eccentric contractions are 73 organized differently to concentric contractions (Enoka, 74 1996; Sekiguchi et al., 2003; Gruber et al., 2009; 75 76 Howatson et al., 2011), with the duration of the silent per-77 iod (SP) being reduced during eccentric contractions 78 compared to concentric (Sekiguchi et al., 2003), intracor-79 tical inhibition (ICI) is significantly reduced, while intracortical facilitation (ICF) is increased during forceful eccentric 80 contractions, but not during concentric contractions 81 (Howatson et al., 2011). Cortical excitability of the iM1 is 82 facilitated during eccentric contractions of the right wrist 83 flexors compared to concentric contractions (Howatson 84 et al., 2011). Taken together, compared to concentric 85 contractions, cortical excitability is facilitated in both con-86 tralateral and iM1's during eccentric contractions and the 87 neural networks involved in ICI and IHI appear to be influ-88 89 enced by the type of contraction. On this basis ICI and IHI might be the primary mechanism underpinning the cross-90 91 education effects following eccentric training compared to 92 concentric, however, this remains to be tested.

Cross-education has gained scientific and clinical interest, primarily for the potential to minimise strength loss and enhance recovery in patients that are unable to perform training due to single limb injury or impairment (Farthing et al., 2009). Unilateral training of the free limb has been found to maintain strength, attenuate muscle atrophy and function of the untrained limb following periods of immobilization (Farthing et al., 2009; Magnus 100 et al., 2010) and fracture (Magnus et al., 2013). Given 101 the clinical relevance of cross-education, the purpose of 102 the present study was to determine whether the TMS 103 responses following eccentric or concentric cross-104 education training are different and whether this differ-105 ence may explain the change in strength of the untrained 106 limb. We hypothesized that unilateral eccentric training 107 would provide a greater stimulus to the non-exercising 108 limb than concentric training and these behavioral 109 changes in strength would be accompanied with modula-110 tion in neurophysiological indices associated with cross-111 education. 112

EXPERIMENTAL PROCEDURES

Participants

Twenty-seven participants (15 males aged 25 ± 1 years 115 and 12 females aged 27 \pm 2 years) were selected on a 116 voluntary basis. All volunteers provided written informed 117 consent prior to participation in the study, which was 118 approved by the Human Research Ethics Committee in 119 accordance to the standards established by the 120 Declaration of Helsinki. All participants were right-hand 121 dominant as determined by the Edinburgh Handedness 122 Inventory (Oldfield, 1971), had not participated in strength 123 training for at least 12 months, and were free from any 124 known history of peripheral or neurological impairment. 125 Prior to the experiment, all participants completed the 126 adult safety screening questionnaire to determine their 127 suitability for TMS (Keel et al., 2001). 128

Experimental approach

Fig. 1 outlines the organization of the study. Participants 130 were required to attend a familiarization session that 131 involved performing five eccentric, concentric and 132 isometric contractions of the right and left wrist flexors 133 along with exposure to single-pulse TMS. Following the 134 familiarization session, participants were randomly 135 (based strength) allocated to a control, eccentric training 136 or concentric training group. All participants underwent 137 TMS, ultrasonography, and maximum strength testing 138 (isometric, eccentric and concentric) before and after a 139 4-week supervised strength-training program; however 140 control participants only undertook pre- and post-testing. 141 Post-testing was carried out between 36 and 48 h after 142 the final training session. 143



Fig. 1. Schematic representation of the experimental design with measures obtained pre and following 4 weeks of maximal unilateral eccentric or concentric strength training of right wrist flexors. Pre and post measures included assessment of peripheral muscle excitability (M-waves), corticospinal excitability and inhibition recruitment curves, short interval intracortical inhibition (SICI) and muscle strength.

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