

Inhibition of β -adrenergic signalling impairs functional repair of rat skeletal muscle after injury

F. Beitzel,¹ M.N. Sillence² and G.S. Lynch,¹ ¹Basic and Clinical Myology Laboratory, Department of Physiology, University of Melbourne, Victoria 3010, Australia and ²School of Agriculture, Charles Sturt University, Wagga Wagga, New South Wales 2678, Australia.

β -adrenoceptor (β -AR) mediated signalling plays an important role in muscle regeneration, by altering the β -AR population and adenylate cyclase activity, to enhance cAMP production that promotes protein accretion. The aim was to block β -AR after muscle injury, to assess the importance of β -AR signalling in regeneration. Adult rats were anaesthetised (ketamine 100 mg/kg and xylazine 10 mg/kg, i.p.), and the EDL and soleus muscles of the right hindlimb injected with bupivacaine to cause complete destruction of all fibres. Rats received twice daily injections of the β_2 -selective antagonist, ICI-118551 (12.5 mg/kg, i.p.), the nonspecific β_1 - and β_2 -antagonist, propranolol (12.5mg/kg, i.p.), or saline, commencing 2 days prior to injury and continuing for 14 days post-injury. Rats were anaesthetised (as described above) for excision of isolated muscles and then killed by cardiac excision. Maximum force (P_o) of injured EDL muscles from saline-treated rats was restored to 59% of control values, compared to only 50% and 53% of controls for ICI- and propranolol-treated rats, respectively. For injured soleus muscles from saline-treated rats, P_o was restored to 43% of control values, and only 33% and 38% of controls for ICI- and propranolol-treated rats, respectively. These findings indicate a reduced rate of functional restoration in regenerating muscles with β -antagonist administration, particularly ICI, where P_o of injured EDL and soleus muscles was 15% and 23% lower than for injured muscles from saline-treated rats. The impairment in functional recovery with β -AR blockade highlights the role of β -AR signalling in successful muscle repair.

Supported by the National Health & Medical Research Council.