

**Regulation of skeletal muscle satellite cell proliferation and differentiation by transforming growth factor-beta, insulin-like growth factor I, and fibroblast growth factor.**

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Skeletal muscle satellite cells were cultured from mature rats and were treated in vitro with various combinations of transforming growth factor (TGF)-beta, fibroblast growth factor (FGF), and insulin-like growth factor I (IGF-I). In serum-free defined medium the following observations were made: TGF-beta depressed proliferation and inhibited differentiation; FGF stimulated proliferation and depressed differentiation; IGF-I stimulated proliferation to a small degree but demonstrated a more pronounced stimulation of differentiation. In evaluating combinations of these three factors, the differentiation inhibiting effect of TGF-beta could not be counteracted by any combination of IGF-I or FGF. The proliferation-depressing activity of TGF-beta, however, could not inhibit the mitogenic activity of FGF. Maximum stimulation of proliferation was observed in the presence of both FGF and IGF-I. The highest percentage fusion was also observed under these conditions, but differentiation with minimal proliferation resulted from treatment with IGF-I, alone. By altering the concentrations of TGF-beta, FGF, and IGF-I, satellite cells can be induced to proliferate, differentiate, or to remain quiescent.

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