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The description is invalid or has not been completed. Effect of amino acids on the insulin-like growth factor-II ligand in chick embryo bone marrow stromal cells. Insulin-like growth factor II (IGF-II) is a potent peptide mitogen for adult fibroblasts and also induces the differentiation of adult and embryonic stem cells to bone. Chicken bone marrow stromal cells respond to IGF-II by expressing a receptor (IGF-IIR) and, after a lag period, responding to addition of IGF-II by entering the S phase of the cell cycle. It has been suggested that the mechanism of cell entry into the S phase is by stimulation of phosphoinositide hydrolysis. We used a transfection assay to determine the role of IGF-IIR and second messenger systems in bone marrow stromal cell response to IGF-II.

Coexpression of IGF-IIR and the phospholipase C-inducing G protein (Gq) was necessary and sufficient for activation of phospholipase C and subsequent S phase entry. However, IGF-II stimulated other signalling pathways, including cyclic AMP-mediated inhibition of proliferation and activation of ERK. To determine whether IGF-II mediates its effects through either or both Gq- and Gs-mediated pathways, we varied the concentration of amino acids in the culture medium. Increasing the supply of amino acids or decreasing their removal inhibited proliferation stimulated by IGF-II, increased the lag in S phase entry, and enhanced ERK activation. We conclude that the response of bone marrow stromal cells to IGF-II is dependent on IGF-IIR- and G protein (Gq)-mediated phospholipase C

activation, and that the amino acid-mediated inhibition of cell proliferation is mediated by a mechanism that involves inhibition of S phase entry. The

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