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video / ebook /.Raf-1 is involved in growth factor induction of MMP-2 and cell invasion by TPA in human osteosarcoma cells. Raf-1 is activated by the tyrosine kinase receptors (TKR) and is also a downstream signal molecule of the Ras/MAPK signaling cascade.

Expression of activated Raf-1 is frequently observed in human osteosarcoma tissues. In this study, we investigated the possible role of Raf-1 in growth factors and TPA (12-O-tetradecanoylphorbol-13-acetate)-induced migration and invasion in osteosarcoma

cells. We found that expression of active Raf-1 was elevated after stimulation of osteosarcoma cells with epidermal growth factor (EGF), insulin-like growth factor-1 (IGF-1), platelet-derived growth factor (PDGF), or nerve growth factor (NGF).

Treatment of osteosarcoma cells with EGF, IGF-1, PDGF, or NGF caused a modest induction of MMP-2, but in combination with TPA, produced a strong induction of MMP-2 activity, cell invasion, and inactivation of MAPK. The effect of TPA was suppressed by inhibitors of

Raf-1, protein kinase C, or MAPK. The combination of EGF and TPA increased Raf-1 activity in a time-dependent manner, and EGF treatment caused a dramatic increase in phosphorylation of Raf-1 and MAPK. Moreover, EGF increased Raf-1 protein kinase activity, which caused

a significant increase in MEK activity. Thus, our results suggest that Raf-1 is involved in the MAPK activation cascade. Raf-1 may be a potential novel therapeutic target for the treatment of human osteosarcoma.// // Generated by class-dump 3.5 (64 bit)

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