Cot (256-467): sc-4337 WB



The Power to Question

BACKGROUND

The role of mitogen-activated protein kinases (MAPKs) in cell signaling pathways is well established. The rat gene Tpl-2, for tumor progression locus 2, and the human and mouse homologues c-Cot, for cancer osaka thyroid oncogene, encode a proto-oncogene serine/threonine protein kinase that was shown to play a role in the functional activation of the MAP kinase pathway. Overexpression of Cot induces MAP kinase activation in COS-1 and NIH/3T3 cells. Cot-mediated activation of MAP kinase is inhibited by both Ras N17, a dominant negative mutant of c-H-Ras, and Raf-1s621A, a dominant negative mutant of Raf-1, suggesting that Cot functions upstream of Ras and Raf-1. Other studies have shown that a kinase-negative, dominant negative mutant of Cot partially blocks Ras or Raf-1-induced MAP kinase activation, arguing that Cot functions downstream of Ras and Raf-1. To explain these contrasting findings, it has been suggested that Cot, Ras and Raf-1 may form a multimeric complex that phosphorylates MEK-1. Cot has also been shown to be implicated in T lymphocyte activation. Two forms of Cot, 58 and 52 kDa, are produced by alternative initiation of translation.

REFERENCES

- Haubruk, H. and McCormick, F. 1991. Ras p21: effects and regulation. Biochim. Biophys. Acta. 1072: 215-229.
- 2. Roberts, T.M. 1992. A signal chain of events. Nature 360: 534-535.
- Nishida, E. and Gotoh, Y. 1993. The MAP kinase cascade is essential for diverse signal transduction pathways. Trends Biochem. Sci. 18: 128-131.
- Fabian, J.R, Morrison, D., and Daar, I. 1993. Requirement for Raf and MAP kinase function during the meiotic maturation of *Xenopus* oocytes. J. Cell. Biol. 122: 645-652.
- 5. Aoki, M., Hamada, F., Sugimoto, T., Suminda, S., Akiyama, T., and Toyoshima, K. 1993. The human cot proto-oncogene encodes two protein serine/threonine kinases with different transforming activities by alternative initiation of translation. J. Biol. Chem. 268: 22723-22732.
- Patriotis, C., Makris, A., Chernoff, J., and Tsichlis, P.N. 1994. Tpl-2 acts in concert with Ras and Raf-1 to activate mitogen-activated protein kinase. Proc. Natl. Acad. Sci. USA 91: 9755-9759.
- 7. Ballester, A., Velasco, A., Tobena, R., and Alemany, S. 1998. Cot kinase activates tumor necrosis factor- α gene expression in a cyclosporin A-resistant manner. J. Biol. Chem. 273: 14099-14106.
- Channavajhala, P.L., Wu, L., Cuozzo, J.W., Hall, J.P., Liu, W., Lin, L.L., Zhang, Y. 2003. Identification of a novel human kinase supporter of Ras (hKSR-2) that functions as a negative regulator of Cot (Tpl2) signaling. J. Biol. Chem. 278: 47089-47097.

SOURCE

Cot (256-467) is expressed in *E. coli* as a 50 kDa tagged fusion protein corresponding to amino acids 256-467 of Cot of human origin.

PRODUCT

Cot (256-467) is purified from bacterial lysates (>98%) by column chromatography; supplied as 10 μg in 0.1 ml SDS-PAGE loading buffer.

APPLICATIONS

Cot (256-467) is suitable as a Western blotting control for sc-720 and sc-9167.

STORAGE

Store at -20° C; stable for one year from the date of shipment.

RESEARCH USE

For research use only, not for use in diagnostic procedures.

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