

3050 Spruce Street Saint Louis, Missouri 63103 USA Telephone (800) 325-5832 (314) 771-5765 Fax (314) 286-7828 email: techserv@sial.com sigma-aldrich.com

ProductInformation

ANTI-CAM KINASE IV (CAMKIV), Developed in Rabbit IgG Fraction of Antiserum

Product Number C 9973

Product Description

Anti-CaM Kinase IV (CaMKIV) is developed in rabbit using a synthetic peptide (K-GSSAVGFEVPQQDVIL-PEY) corresponding to the C-terminal region of human CaM Kinase IV (amino acids 455-473 with N-terminally added lysine) conjugated to KLH as immunogen. This sequence has limited homology (~50% identity) with rat and mouse CaM Kinase IV. Whole antiserum is fractionated and then further purified by ion-exchange chromatography to provide the IgG fraction of antiserum that is essentially free of other rabbit serum proteins.

Anti-CaM Kinase IV recognizes human CaMKIV (65 kDa) by immunoblotting. The antibody also reacts with rat CaMKIV. By immunoblotting, staining of CaMKIV is specifically inhibited with the CaMKIV immunizing peptide (human CaMKIV, amino acids 455-473 with N-terminally added lysine).

Ca²⁺ /Calmodulin dependent protein kinases (CaM kinases, CaMKs) are a family of Ser/Thr protein kinases including CaMKI, CaMKII, CaMKIII and CaMKIV.¹ In response to Ca²⁺ signaling, CaMKs are considered to play a central role in many cellular functions, including synthesis and secretion of neurotransmitters, axonal transport, long term potentiation (LTP), receptor function, modification of the cytoskeleton and regulation of gene expression. CaM kinase IV (65 kDa) is abundantly expressed in the brain $^{2-5}$ (with the highest levels found in the cerebellum), in the thymus,⁶ and is localized to both the cytoplasm and nucleus.^{5,7} CaMKIV is thought to play a central role in controlling a wide range of Ca²⁺ mediated cellular functions in the central nervous system (CNS) and the immune system. In the brain it phosphorylates a wide range of substrates including synapsin I, microtubule associated protein (MAP2), tau and tyrosine hydroxylase. In Jurkat T lymphocytes, CaMKIV is activated through CD3mediated signaling pathway.⁸ CaMKIV is a monomeric enzyme containing a catalytic and autoinhibitory subunit. CaMKIV is transiently phosphorylated and

activated in response to rise in intracellular Ca²⁺, by CaM kinase kinase (CaMKK), upstream in the CaMK signaling cascade.^{8,9} CaMKIV translocates to the nucleus, where it phosphorylates cAMP- response element-binding protein (CREB) at Ser¹³³, a key regulatory site controlling transcriptional activity.¹⁰ CaMKIV phosphorylation and activity is regulated by the serine/threonine phosphatase, PP2A, that dephosphorylates and inactivates CaMKIV. CaMKIV and PP2A have been shown to associate and form a stable signaling complex.¹¹

Reagents

The product is provided as IgG fraction of antiserum in 0.01 M phosphate buffered saline, pH 7.4, containing 15 mM sodium azide as a preservative.

Precautions and Disclaimer

Due to the sodium azide content a material safety sheet (MSDS) for this product has been sent to the attention of the safety officer of your institution.

Consult the MSDS for information regarding hazardous and safe handling practices.

Storage/Stability

For continuous use, store at 2-8 °C for up to one month. For extended storage, freeze in working aliquots. Repeated freezing and thawing is not recommended. Storage in "frost-free" freezers is not recommended. If slight turbidity occurs upon prolonged storage, clarify the solution by centrifugation before use. Working dilution samples should be discarded if not used within 12 hours.

Product Profile

A minimum working dilution of 1:10,000 is determined by immunoblotting using a whole cell extract of the human T-cell leukemia Jurkat cell line.

A minimum working dilution of 1:2,500 is determined by immunoblotting using a cytosolic fraction of rat brain extract. Note: In order to obtain best results and assay sensitivity in different techniques and preparations we recommend determining optimal working dilutions by titration.

References

- Hanson, P.I. and Schulman, H., Ann. Rev. Biochem. 61, 559 (1992).
- 2. Miyano, O., et al., J. Biol. Chem. 267, 1198 (1992).
- 3. Okuno, S., et al., J. Biochem. 116, 923 (1994).
- 4. Okuno, S., et al., J. Biochem. 119, 1176 (1996).
- 5. Nakamura, Y., et al., Neuroscience **68**, 181 (1995).
- Frangakis, M.V., et al., J. Biol. Chem. 266, 17592 (1991).
- 7. Jensen, K.F., et al., Proc. Natl. Acad. Sci. USA 88 2850 (1991).
- 8. Park, I-K, and Soderling, T.R. J. Biol. Chem. **270**, 51 30464 (1995).
- Tokumitsu, H., et al., J. Biol. Chem. 270, 51 19320 (1995).
- 10. Sheng, M., et al., Science 252, 1427 (1991).
- 11. Westphal, R.S., et al., Science **280**, 1258 (1998).

lpg 7/99

Sigma brand products are sold through Sigma-Aldrich, Inc.

Sigma-Aldrich, Inc. warrants that its products conform to the information contained in this and other Sigma-Aldrich publications. Purchaser must determine the suitability of the product(s) for their particular use. Additional terms and conditions may apply. Please see reverse side of the invoice or packing slip.