



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

Product Information

Anti-Tuberin (VV-18)

Developed in Rabbit
Affinity Isolated Antibody

Product Number **T 5075**

Product Description

Anti-Tuberin (VV-18) is developed in rabbit using as immunogen a synthetic peptide corresponding to amino acid residues 1790-1807 of human Tuberlin with N-terminal added cysteine, conjugated to KLH. The corresponding sequence differs by two amino acids in rat and mouse tuberlin. The antibody is affinity-purified using the immunizing peptide immobilized on agarose.

Anti-Tuberin (VV-18) recognizes human, rat, and mouse tuberlin. Applications include the detection of tuberlin by immunoblotting (180-200 kDa) and immunoprecipitation. Detection of the tuberlin band by immunoblotting is specifically inhibited with the immunizing peptide.

Tuberlin is the protein product of the tumor suppressor gene TSC2.¹ Hamartin, the product of the TSC1 tumor suppressor gene, contains two coiled-coil regions that have been shown to mediate its binding to tuberlin.^{2,3} Hamartin and tuberlin are involved in the regulation of cell cycle, cell growth, cell differentiation, cell adhesion and vesicular trafficking.⁴ Mutations in either the TSC1 or the TSC2 gene are responsible for tuberous sclerosis complex (TSC), an autosomal dominant hereditary disease characterized by mental retardation, seizures, and benign tumors (hamartomas) in multiple organs including the kidney, brain, heart and skin.⁵

Tuberlin is widely expressed in mammalian cell lines and tissues. It co-localizes with hamartin in most tissues and cell types.⁶ Tuberlin is found in cytosolic, microsomal, cytoskeletal, and vesicular fractions, and in certain cells it is also localized to the nucleus.^{7,8}

Tuberlin has a GTPase activating protein homology (GAP) domain that has been reported to be involved in the regulation of the small GTPase Rheb both *in vitro* and *in vivo*.⁹ Tuberlin forms together with hamartin a functional cytoplasmic complex that inhibits growth by inhibiting phosphorylation of S6K and 4EBP, probably through their upstream modulator mammalian target of rapamycin (mTOR).¹⁰

Tuberlin phosphorylation is required for complex formation as well for its interaction with several isoforms of the regulatory 14-3-3 protein.¹¹ Tuberlin and hamartin are involved in the phosphoinositide 3-kinase/ Akt signal transduction pathway.⁴ Phosphorylation by Akt and mitogenic factors abrogates hamartin-tuberlin suppressor activity by inducing proteasome-mediated degradation of both proteins.^{12,13}

Reagent

Anti-Tuberin (VV-18) is supplied as a solution in 0.01 M phosphate buffered saline, pH 7.4, containing 15 mM sodium azide.

Antibody Concentration: Approx. 1.0 mg/ml

Precautions and Disclaimer

Due to the sodium azide content, a material safety data 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 also not recommended. If slight turbidity occurs upon prolonged storage, clarify the solution by centrifugation before use. Working dilutions should be discarded if not used within 12 hours.

Product Profile

By immunoblotting, a working antibody concentration of 0.2-0.4 µg/ml is recommended using whole extract of rat brain.

By immunoblotting, a working antibody concentration of 0.1-0.2 µg/ml is recommended using whole extract of mouse NIH-3T3 cells.

5-10 µg of the antibody immunoprecipitates tuberlin from 0.5 mg of RIPA extract of human HeLa cells.

Note: In order to obtain the best results using various techniques and preparations, we recommend determining the optimal working dilutions by titration.

References

1. Wienecke, R., et al., J. Biol. Chem., **270**, 16409-16414 (1995).
2. The European Chromosome 16 Tuberous Sclerosis Consortium, Cell, **75**, 1305-1315 (1993).
3. van Slechtenhorst, M., et al., Hum. Mol. Genet., **7**, 1053-1057 (1998).
4. Dan, H.C., et al., J. Biol. Chem., **277**, 35364-35370 (2002).
5. Gomez, M.R., (ed.) in: Tuberous Sclerosis, Raven Press, New York (1988).
6. Johnson, M.W., et al., Modern Pathol., **14**, 202-210 (2001).
7. Yamamoto, Y., et al., Arch. Biochem. Biophys., **404**, 210-217 (2002).
8. Wei, J., et al., Modern Pathol., **15**, 862-869 (2001).
9. Tee, A.R., et al., Curr. Biol., **13**, 1259-1268 (2003).
10. Krymskaya, V.P., Cell Signal., **15**, 729-739 (2003).
11. Nellist, M., et al., J. Biol. Chem., **277**, 39417-39424 (2002).
12. Aicher, L.D., et al., J. Biol. Chem., **276**, 21017-21021 (2001).
13. Plas, D.R., and Thompson, C.B., J. Biol. Chem., **278**, 12361-12366 (2003).

KAA/ST 09/04

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.