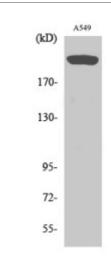


Anti-mTOR antibody



Description

Rabbit polyclonal to mTOR.

Model STJ94278

Host Rabbit

Reactivity Human, Mouse, Rat

Applications ELISA, IHC, WB

Immunogen Synthesized peptide derived from human mTOR around the non-

phosphorylation site of S2481.

Immunogen Region 2420-2500 aa

Gene ID <u>2475</u>

Gene Symbol MTOR

Dilution range WB 1:500-1:2000IHC 1:100-1:300ELISA 1:10000

Specificity mTOR Polyclonal Antibody detects endogenous levels of mTOR protein.

Tissue Specificity Expressed in numerous tissues, with highest levels in testis.

Purification The antibody was affinity-purified from rabbit antiserum by affinity-

chromatography using epitope-specific immunogen.

Note For Research Use Only (RUO).

Protein Name Serine/threonine-protein kinase mTOR FK506-binding protein 12-rapamycin

complex-associated protein 1 FKBP12-rapamycin complex-associated protein Mammalian target of rapamycin mTOR Mechanistic target of rapamycin

Rapamyc

Molecular Weight 289 kDa

Clonality Polyclonal

Conjugation Unconjugated

Isotype IgG

Formulation Liquid in PBS containing 50% glycerol, 0.5% BSA and 0.02% sodium azide.

Concentration 1 mg/ml

Storage Instruction Store at -20°C, and avoid repeat freeze-thaw cycles.

Database Links HGNC:3942OMIM:601231

Alternative Names Serine/threonine-protein kinase mTOR FK506-binding protein 12-rapamycin

complex-associated protein 1 FKBP12-rapamycin complex-associated protein Mammalian target of rapamycin mTOR Mechanistic target of rapamycin

Rapamyc

Function Serine/threonine protein kinase which is a central regulator of cellular

metabolism, growth and survival in response to hormones, growth factors, nutrients, energy and stress signals. MTOR directly or indirectly regulates the phosphorylation of at least 800 proteins. Functions as part of 2 structurally and functionally distinct signaling complexes mTORC1 and mTORC2 (mTOR complex 1 and 2). Activated mTORC1 up-regulates protein synthesis by phosphorylating key regulators of mRNA translation and ribosome synthesis. This includes phosphorylation of EIF4EBP1 and release of its inhibition toward the elongation initiation factor 4E (eiF4E). Moreover, phosphorylates and activates RPS6KB1 and RPS6KB2 that promote protein synthesis by modulating the activity of their downstream targets including ribosomal protein S6, eukaryotic translation initiation factor EIF4B, and the inhibitor of translation initiation PDCD4. Stimulates the pyrimidine biosynthesis pathway, both by acute regulation through RPS6KB1-mediated phosphorylation of the biosynthetic enzyme CAD, and delayed regulation, through transcriptional enhancement of the pentose phosphate pathway which produces 5phosphoribosyl-1-pyrophosphate (PRPP), an allosteric activator of CAD at a later step in synthesis, this function is dependent on the mTORC1 complex. Regulates ribosome synthesis by activating RNA polymerase III-dependent transcription through phosphorylation and inhibition of MAF1 an RNA polymerase III-repressor. In parallel to protein synthesis, also regulates lipid synthesis through SREBF1/SREBP1 and LPIN1. To maintain energy homeostasis mTORC1 may also regulate mitochondrial biogenesis through regulation of PPARGC1A. mTORC1 also negatively regulates autophagy through phosphorylation of ULK1. Under nutrient sufficiency, phosphorylates ULK1 at 'Ser-758', disrupting the interaction with AMPK and preventing activation of ULK1. Also prevents autophagy through phosphorylation of the autophagy inhibitor DAP. mTORC1 exerts a feedback control on upstream growth factor signaling that includes phosphorylation and activation of GRB10 a INSR-dependent signaling suppressor. Among other potential targets mTORC1 may phosphorylate CLIP1 and regulate microtubules. As

part of the mTORC2 complex MTOR may regulate other cellular processes including survival and organization of the cytoskeleton. Plays a critical role in

phosphoinositide 3-kinase, facilitating its activation by PDK1. mTORC2 may regulate the actin cytoskeleton, through phosphorylation of PRKCA, PXN and

the phosphorylation at 'Ser-473' of AKT1, a pro-survival effector of

activation of the Rho-type guanine nucleotide exchange factors RHOA and RAC1A or RAC1B. mTORC2 also regulates the phosphorylation of SGK1 at 'Ser-422'. Regulates osteoclastogensis by adjusting the expression of CEBPB isoforms .

Sequence and Domain Family

The kinase domain (PI3K/PI4K) is intrinsically active but has a highly restricted catalytic center. The FAT domain forms three discontinuous subdomains of alpha-helical TPR repeats plus a single subdomain of HEAT repeats. The four domains pack sequentially to form a C-shaped a-solenoid that clamps onto the kinase domain .

Cellular Localization

Endoplasmic reticulum membrane Golgi apparatus membrane Mitochondrion outer membrane Lysosome Cytoplasm Nucleus, PML body Microsome membrane. Shuttles between cytoplasm and nucleus. Accumulates in the nucleus in response to hypoxia . Targeting to lysosomes depends on amino acid availability and RRAGA and RRAGB .

Post-translational Modifications

Autophosphorylates when part of mTORC1 or mTORC2. Phosphorylation at Ser-1261, Ser-2159 and Thr-2164 promotes autophosphorylation.

Phosphorylation in the kinese domain modulates the interactions of MTOP.

Phosphorylation in the kinase domain modulates the interactions of MTOR with RPTOR and PRAS40 and leads to increased intrinsic mTORC1 kinase activity. Phosphorylation at Thr-2173 in the ATP-binding region by AKT1 strongly reduces kinase activity.

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