

WNK1 Antibody

Rabbit mAb Catalog # AP91839

Product Information

Application WB, IHC Primary Accession Q9H4A3

Reactivity Rat, Human, Mouse

Clonality Monoclonal

Other Names KDP; PSK; p65; HSN2; HSAN2; PRKWNK1;

IsotypeRabbit IgGHostRabbitCalculated MW250794

Additional Information

Dilution WB 1:500~1:2000 IHC 1:50~1:200

Purification Affinity-chromatography

Immunogen A synthesized peptide derived from human WNK1

Description Serine/threonine kinase which plays an important role in the regulation of

electrolyte homeostasis, cell signaling, survival, and proliferation. Acts as an activator and inhibitor of sodium-coupled chloride cotransporters and potassium-coupled chloride cotransporters respectively. Activates SCNN1A,

SCNN1B, SCNN1D and SGK1.

Storage Condition and Buffer Rabbit IgG in phosphate buffered saline, pH 7.4, 150mM NaCl, 0.02% sodium

azide and 50% glycerol. Store at +4°C short term. Store at -20°C long term.

Avoid freeze / thaw cycle.

Protein Information

Name WNK1 {ECO:0000303|PubMed:11571656, ECO:0000312|HGNC:HGNC:14540}

Function Serine/threonine-protein kinase component of the WNK1- SPAK/OSR1 kinase

cascade, which acts as a key regulator of blood pressure and regulatory

volume increase by promoting ion influx (PubMed: 15883153, PubMed: 17190791, PubMed: 31656913, PubMed: 34289367,

PubMed:<u>36318922</u>). WNK1 mediates regulatory volume increase in response to hyperosmotic stress by acting as a molecular crowding sensor, which senses cell shrinkage and mediates formation of a membraneless

compartment by undergoing liquid-liquid phase separation

(PubMed:36318922). The membraneless compartment concentrates WNK1

with its substrates, OXSR1/OSR1 and STK39/SPAK, promoting

WNK1-dependent phosphorylation and activation of downstream kinases OXSR1/OSR1 and STK39/SPAK (PubMed:15883153, PubMed:16263722,

PubMed:<u>17190791</u>, PubMed:<u>19739668</u>, PubMed:<u>21321328</u>, PubMed:<u>22989884</u>, PubMed:<u>25477473</u>, PubMed:<u>34289367</u>,

PubMed:36318922). Following activation, OXSR1/OSR1 and STK39/SPAK catalyze phosphorylation of ion cotransporters SLC12A1/NKCC2, SLC12A2/NKCC1, SLC12A5/KCC2 and SLC12A6/KCC3, regulating their activity (PubMed:16263722, PubMed:21321328). Phosphorylation of Na-K-Cl cotransporters SLC12A2/NKCC1 and SLC12A2/NKCC1 promote their activation and ion influx; simultaneously, phosphorylation of K-Cl cotransporters SLC12A5/KCC2 and SLC12A6/KCC3 inhibit their activity, blocking ion efflux (PubMed: <u>19665974</u>, PubMed: <u>21321328</u>). Also acts as a regulator of angiogenesis in endothelial cells via activation of OXSR1/OSR1 and STK39/SPAK: activation of OXSR1/OSR1 regulates chemotaxis and invasion, while STK39/SPAK regulates endothelial cell proliferation (PubMed: 25362046). Also acts independently of the WNK1- SPAK/OSR1 kinase cascade by catalyzing phosphorylation of other substrates, such as SYT2, PCF11 and NEDD4L (PubMed: 29196535). Mediates phosphorylation of SYT2, regulating SYT2 association with phospholipids and membrane-binding (By similarity). Regulates mRNA export in the nucleus by mediating phosphorylation of PCF11, thereby decreasing the association between PCF11 and POLR2A/RNA polymerase II and promoting mRNA export to the cytoplasm (PubMed:29196535). Acts as a negative regulator of autophagy (PubMed: 27911840). Required for the abscission step during mitosis, independently of the WNK1-SPAK/OSR1 kinase cascade (PubMed:21220314). May also play a role in actin cytoskeletal reorganization (PubMed:10660600). Also acts as a scaffold protein independently of its protein kinase activity: negatively regulates cell membrane localization of various transporters and channels, such as SLC4A4, SLC26A6, SLC26A9, TRPV4 and CFTR (By similarity). Involved in the regulation of epithelial Na(+) channel (ENaC) by promoting activation of SGK1 in a kinase-independent manner: probably acts as a scaffold protein that promotes the recruitment of SGK1 to the mTORC2 complex in response to chloride, leading to mTORC2-dependent phosphorylation and activation of SGK1 (PubMed:36373794). Acts as an assembly factor for the ER membrane protein complex independently of its protein kinase activity: associates with EMC2 in the cytoplasm via its amphipathic alpha-helix, and prevents EMC2 ubiquitination and subsequent degradation, thereby promoting EMC2 stabilization (PubMed:33964204).

Cellular Location

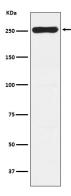
Cytoplasm. Nucleus. Cytoplasm, cytoskeleton, spindle. Note=Mediates formation and localizes to cytoplasmic membraneless compartment in response to hyperosmotic stress (PubMed:36318922). Also localizes to the nucleus (PubMed:29196535) Localizes to the mitotic spindle during mitosis (PubMed:21220314)

Tissue Location

Widely expressed, with highest levels observed in the testis, heart, kidney and skeletal muscle [Isoform 3]: This isoform is kidney-specific and specifically expressed in the distal convoluted tubule (DCT) and connecting tubule (CNT) of the nephron.

Images

Western blot analysis of WNK1 expression in Saos-2 cell lysate.



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