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SIRT2 Antibody

Purified Rabbit Polyclonal Antibody (Pab) Catalog # AP50816

Product Information

Application WB
Primary Accession Q8IX|6

Reactivity Human, Mouse, Rat

Host Rabbit
Clonality polyclonal
Calculated MW 43182

Additional Information

Gene ID 22933

Other Names NAD-dependent protein deacetylase sirtuin-2, 351-, Regulatory protein SIR2

homolog 2, SIR2-like protein 2, SIRT2, SIR2L, SIR2L2

Dilution WB~~ 1:1000

Format Rabbit IgG in phosphate buffered saline (without Mg2+ and Ca2+), pH 7.4,

150mM NaCl, 0.09% (W/V) sodium azide and 50% glycerol.

Storage Conditions -20°C

Protein Information

Name SIRT2

Synonyms SIR2L, SIR2L2

Function NAD-dependent protein deacetylase, which deacetylates internal lysines on

histone and alpha-tubulin as well as many other proteins such as key

transcription factors (PubMed:<u>12620231</u>, PubMed:<u>16648462</u>, PubMed:<u>18249187</u>, PubMed:<u>18332217</u>, PubMed:<u>18995842</u>, PubMed:<u>20543840</u>, PubMed:<u>20587414</u>, PubMed:<u>21081649</u>, PubMed:<u>21726808</u>, PubMed:<u>21949390</u>, PubMed:<u>22014574</u>, PubMed:<u>22771473</u>, PubMed:<u>23468428</u>, PubMed:<u>23908241</u>, PubMed:<u>24177535</u>, PubMed:<u>24681946</u>, PubMed:<u>24769394</u>,

PubMed:<u>24940000</u>). Participates in the modulation of multiple and diverse biological processes such as cell cycle control, genomic integrity, microtubule

dynamics, cell differentiation, metabolic networks, and autophagy

(PubMed: 12620231, PubMed: 16648462, PubMed: 18249187, PubMed: 18332217, PubMed: 18995842, PubMed: 20543840, PubMed: 20587414, PubMed: 21081649, PubMed: 21726808, PubMed: 21949390, PubMed: 22014574, PubMed: 22771473,

PubMed: 23468428, PubMed: 23908241, PubMed: 24177535, PubMed:24681946, PubMed:24769394, PubMed:24940000). Plays a major role in the control of cell cycle progression and genomic stability (PubMed:12697818, PubMed:16909107, PubMed:17488717, PubMed: 17726514, PubMed: 19282667, PubMed: 23468428). Functions in the antephase checkpoint preventing precocious mitotic entry in response to microtubule stress agents, and hence allowing proper inheritance of chromosomes (PubMed:<u>12697818</u>, PubMed:<u>16909107</u>, PubMed:<u>17488717</u>, PubMed: 17726514, PubMed: 19282667, PubMed: 23468428). Positively regulates the anaphase promoting complex/cyclosome (APC/C) ubiquitin ligase complex activity by deacetylating CDC20 and FZR1, then allowing progression through mitosis (PubMed:22014574). Associates both with chromatin at transcriptional start sites (TSSs) and enhancers of active genes (PubMed:<u>23468428</u>). Plays a role in cell cycle and chromatin compaction through epigenetic modulation of the regulation of histone H4 'Lys-20' methylation (H4K20me1) during early mitosis (PubMed: 23468428). Specifically deacetylates histone H4 at 'Lys-16' (H4K16ac) between the G2/M transition and metaphase enabling H4K20me1 deposition by KMT5A leading to ulterior levels of H4K20me2 and H4K20me3 deposition throughout cell cycle, and mitotic S-phase progression (PubMed: 23468428). Deacetylates KMT5A modulating KMT5A chromatin localization during the mitotic stress response (PubMed:23468428). Also deacetylates histone H3 at 'Lys-57' (H3K56ac) during the mitotic G2/M transition (PubMed: 20587414). Upon bacterium Listeria monocytogenes infection, deacetylates 'Lys-18' of histone H3 in a receptor tyrosine kinase MET- and PI3K/Akt-dependent manner, thereby inhibiting transcriptional activity and promoting late stages of listeria infection (PubMed: <u>23908241</u>). During oocyte meiosis progression, may deacetylate histone H4 at 'Lys-16' (H4K16ac) and alpha-tubulin, regulating spindle assembly and chromosome alignment by influencing microtubule dynamics and kinetochore function (PubMed: 24940000). Deacetylates histone H4 at 'Lys-16' (H4K16ac) at the VEGFA promoter and thereby contributes to regulate expression of VEGFA, a key regulator of angiogenesis (PubMed: 24940000). Deacetylates alpha-tubulin at 'Lys-40' and hence controls neuronal motility, oligodendroglial cell arbor projection processes and proliferation of non-neuronal cells (PubMed:18332217, PubMed:18995842). Phosphorylation at Ser-368 by a G1/S-specific cyclin E-CDK2 complex inactivates SIRT2-mediated alpha- tubulin deacetylation, negatively regulating cell adhesion, cell migration and neurite outgrowth during neuronal differentiation (PubMed: 17488717). Deacetylates PARD3 and participates in the regulation of Schwann cell peripheral myelination formation during early postnatal development and during postinjury remyelination (PubMed:<u>21949390</u>). Involved in several cellular metabolic pathways (PubMed:20543840, PubMed:21726808, PubMed:24769394). Plays a role in the regulation of blood glucose homeostasis by deacetylating and stabilizing phosphoenolpyruvate carboxykinase PCK1 activity in response to low nutrient availability (PubMed: 21726808). Acts as a key regulator in the pentose phosphate pathway (PPP) by deacetylating and activating the glucose-6-phosphate G6PD enzyme, and therefore, stimulates the production of cytosolic NADPH to counteract oxidative damage (PubMed: 24769394). Maintains energy homeostasis in response to nutrient deprivation as well as energy expenditure by inhibiting adipogenesis and promoting lipolysis (PubMed: 20543840). Attenuates adipocyte differentiation by deacetylating and promoting FOXO1 interaction to PPARG and subsequent repression of PPARG-dependent transcriptional activity (PubMed: 20543840). Plays a role in the regulation of lysosome- mediated degradation of protein aggregates by autophagy in neuronal cells (PubMed: 20543840). Deacetylates FOXO1 in response to oxidative stress or serum deprivation, thereby negatively regulating FOXO1- mediated autophagy (PubMed: 20543840). Deacetylates a broad range of transcription factors and co-regulators regulating target gene expression. Deacetylates transcriptional factor FOXO3 stimulating the

ubiquitin ligase SCF(SKP2)-mediated FOXO3 ubiquitination and degradation (By similarity). Deacetylates HIF1A and therefore promotes HIF1A degradation and inhibition of HIF1A transcriptional activity in tumor cells in response to hypoxia (PubMed:24681946). Deacetylates RELA in the cytoplasm inhibiting NF-kappaB-dependent transcription activation upon TNF-alpha stimulation (PubMed:21081649). Inhibits transcriptional activation by deacetylating p53/TP53 and EP300 (PubMed:18249187, PubMed:18995842). Also deacetylates EIF5A (PubMed:22771473). Functions as a negative regulator on oxidative stress-tolerance in response to anoxia-reoxygenation conditions (PubMed:24769394). Plays a role as tumor suppressor (PubMed:22014574). In addition to protein deacetylase activity, also has activity toward long-chain fatty acyl groups and mediates protein-lysine demyristoylation and depalmitoylation of target proteins, such as ARF6 and KRAS, thereby regulating their association with membranes (PubMed:25704306, PubMed:29239724, PubMed:32103017).

Cellular Location

Nucleus, Cytoplasm, perinuclear region {ECO:0000250|UniProtKB:Q8VDQ8}. Cytoplasm. Cytoplasm, cytoskeleton. Cytoplasm, cytoskeleton, microtubule organizing center, centrosome. Cytoplasm, cytoskeleton, microtubule organizing center, centrosome, centriole Cytoplasm, cytoskeleton, spindle. Midbody. Chromosome. Perikaryon {ECO:0000250 | UniProtKB:Q8VDQ8}. Cell projection {ECO:0000250 | UniProtKB:Q8VDQ8}. Cell projection, growth cone {ECO:0000250|UniProtKB:Q8VDQ8}. Myelin membrane {ECO:0000250|UniProtKB:Q8VDQ8}. Note=Localizes in the cytoplasm during most of the cell cycle except in the G2/M transition and during mitosis, where it is localized in association with chromatin and induces deacetylation of histone at 'Lys-16' (H4K16ac) (PubMed:17726514, PubMed:23468428). Colocalizes with KMT5A at mitotic foci (PubMed:23468428). Colocalizes with CDK1 at centrosome during prophase and splindle fibers during metaphase (PubMed:17488717) Colocalizes with Aurora kinase AURKA at centrosome during early prophase and in the centrioles and growing mitotic spindle throughout metaphase (PubMed:17488717). Colocalizes with Aurora kinase AURKB during cytokinesis with the midbody (PubMed:17488717). Colocalizes with microtubules (PubMed:12620231). Detected in perinuclear foci that may be aggresomes containing misfolded, ubiquitinated proteins (By similarity). Shuttles between the cytoplasm and the nucleus through the CRM1 export pathway (PubMed:17726514). Colocalizes with EP300 in the nucleus (PubMed:24177535). Translocates to the nucleus and chromatin upon bacterium Listeria monocytogenes infection in interphase cells (PubMed:23908241). Deacetylates FOXO3 in the cytoplasm (By similarity) Colocalizes with PLP1 in internodal regions, at paranodal axoglial junction and Schmidt-Lanterman incisures of myelin sheath (By similarity). Colocalizes with CDK5R1 in the perikaryon, neurites and growth cone of hippocampal neurons (By similarity). Colocalizes with alpha-tubulin in neuronal growth cone (By similarity). Localizes in the cytoplasm and nucleus of germinal vesicle (GV) stage oocytes (By similarity). Colocalizes with alpha-tubulin on the meiotic spindle as the oocytes enter into metaphase, and also during meiotic anaphase and telophase, especially with the midbody (By similarity). Colocalizes with PARD3 in internodal region of axons (By similarity). Colocalizes with acetylated alpha-tubulin in cell projection processes during primary oligodendrocyte precursor (OLP) differentiation (By similarity). {ECO:0000250|UniProtKB:Q8VDQ8, ECO:0000269|PubMed:12620231, ECO:0000269 | PubMed:17488717, ECO:0000269 | PubMed:17726514, ECO:0000269 | PubMed:23468428, ECO:0000269 | PubMed:23908241, ECO:0000269 | PubMed:24177535 | [Isoform 2]: Cytoplasm. Nucleus Note=Predominantly localized in the cytoplasmic

Tissue Location

Isoform 1 is expressed in heart, liver and skeletal muscle, weakly expressed in the cortex. Isoform 2 is strongly expressed in the cortex, weakly expressed in heart and liver. Weakly expressed in several malignancies including breast, liver, brain, kidney and prostate cancers compared to normal tissues. Weakly expressed in glioma cell lines compared to normal brain tissues (at protein level). Widely expressed. Highly expressed in heart, brain and skeletal muscle, while it is weakly expressed in placenta and lung. Down-regulated in many gliomas suggesting that it may act as a tumor suppressor gene in human gliomas possibly through the regulation of microtubule network

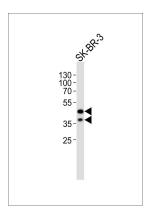
Background

NAD-dependent protein deacetylase, which deacetylates internal lysines on histone and non-histone proteins such as ACLY, PCK1 and RELA. Deacetylates 'Lys-40' of alpha-tubulin. Involved in the control of mitotic exit in the cell cycle, probably via its role in the regulation of cytoskeleton. Deacetylates PCK1, opposing proteasomal degradation. Deacetylates 'Lys-310' of RELA.

References

Frye R.A.,et al.Biochem. Biophys. Res. Commun. 260:273-279(1999). Afshar G.,et al.Gene 234:161-168(1999). De Smet C.,et al.J. Neurochem. 81:575-588(2002). Lennerz V.,et al.Submitted (AUG-2002) to the EMBL/GenBank/DDBJ databases. Hu R.-M.,et al.Proc. Natl. Acad. Sci. U.S.A. 97:9543-9548(2000).

Images



Western blot analysis of lysate from SK-BR-3 cell line, using SIRT2 Antibody. This antibody was diluted at 1:1000. A goat anti-rabbit IgG H&L(HRP) at 1:5000 dilution was used as the secondary antibody. Lysate at 35 ug.

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