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Parkin Antibody

Rabbit mAb Catalog # AP91072

Product Information

Application WB, IHC, IF, FC, ICC, IP, IHF

Primary Accession <u>060260</u>

Reactivity Rat, Human, Mouse

Clonality Monoclonal

Other Names AR-JP; LPRS2; PARK2; parkin; parkin 2; PDJ; PRKN; PRKN2;

IsotypeRabbit IgGHostRabbitCalculated MW51641

Additional Information

Dilution WB 1:500~1:2000 IHC 1:50~1:200 ICC/IF 1:50~1:200 IP 1:50 FC 1:50

Purification Affinity-chromatography

Immunogen A synthesized peptide derived from human Parkin

Description Parkin is a zinc-finger protein that is related to ubiquitin at the amino

terminus. May protect neurons against alpha synuclein toxicity, proteasomal dysfunction, GPR37 accumulation, and kainate-induced excitotoxicity. May play a role in controlling neurotransmitter trafficking at the presynaptic terminal and in calcium-dependent exocytosis. Regulates cyclin-E during

neuronal apoptosis.

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 PRKN (HGNC:8607)

Synonyms PARK2

Function Functions within a multiprotein E3 ubiquitin ligase complex, catalyzing the

covalent attachment of ubiquitin moieties onto substrate proteins

(PubMed: 10888878, PubMed: 10973942, PubMed: 11431533, PubMed: 12150907, PubMed: 12628165, PubMed: 15105460, PubMed: 16135753, PubMed: 21376232, PubMed: 21532592, PubMed: 22396657, PubMed: 23620051, PubMed: 23754282, PubMed: 24660806, PubMed: 24751536, PubMed: 29311685,

PubMed:32047033). Substrates include SYT11 and VDAC1 (PubMed:29311685,

PubMed:32047033). Other substrates are BCL2, CCNE1, GPR37,

RHOT1/MIRO1, MFN1, MFN2, STUB1, SNCAIP, SEPTIN5, TOMM20, USP30, ZNF746, MIRO1 and AIMP2 (PubMed:10888878, PubMed:10973942,

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PubMed: 11431533, PubMed: 12150907, PubMed: 12628165,
PubMed: 15105460, PubMed: 16135753, PubMed: 21376232,
PubMed:21532592, PubMed:22396657, PubMed:23620051,
PubMed:23754282, PubMed:24660806, PubMed:24751536). Mediates
monoubiquitination as well as 'Lys-6', 'Lys-11', 'Lys-48'-linked and
'Lys-63'-linked polyubiquitination of substrates depending on the context
(PubMed: 19229105, PubMed: 20889974, PubMed: 25474007,
PubMed: <u>25621951</u>, PubMed: <u>32047033</u>). Participates in the removal and/or
detoxification of abnormally folded or damaged protein by mediating
'Lys-63'-linked polyubiquitination of misfolded proteins such as PARK7:
'Lys-63'-linked polyubiquitinated misfolded proteins are then recognized by
HDAC6, leading to their recruitment to aggresomes, followed by degradation
(PubMed: 17846173, PubMed: 19229105). Mediates 'Lys-63'-linked
polyubiquitination of a 22 kDa O-linked glycosylated isoform of SNCAIP,
possibly playing a role in Lewy-body formation (PubMed: 11431533,
PubMed: 11590439, PubMed: 15105460, PubMed: 15728840,
PubMed:19229105). Mediates monoubiquitination of BCL2, thereby acting as
a positive regulator of autophagy (PubMed: 20889974). Protects against
mitochondrial dysfunction during cellular stress, by acting downstream of
PINK1 to coordinate mitochondrial quality control mechanisms that remove
and replace dysfunctional mitochondrial components (PubMed: 11439185,
PubMed: 18957282, PubMed: 19029340, PubMed: 19966284,
PubMed: 21376232, PubMed: 22082830, PubMed: 22396657,
PubMed: 23620051, PubMed: 23933751, PubMed: 24660806,
PubMed: 24784582, PubMed: 24896179, PubMed: 25474007,
PubMed: 25527291, PubMed: 32047033). Depending on the severity of
mitochondrial damage and/or dysfunction, activity ranges from preventing
apoptosis and stimulating mitochondrial biogenesis to regulating
mitochondrial dynamics and eliminating severely damaged mitochondria via
mitophagy (PubMed: 11439185, PubMed: 19029340, PubMed: 19801972,
PubMed: 19966284, PubMed: 21376232, PubMed: 22082830,
PubMed: 22396657, PubMed: 23620051, PubMed: 23685073,
PubMed: 23933751, PubMed: 24896179, PubMed: 25527291,
PubMed:32047033, PubMed:33499712). Activation and recruitment onto the
outer membrane of damaged/dysfunctional mitochondria (OMM) requires
PINK1-mediated phosphorylation of both PRKN and ubiquitin
(PubMed:24660806, PubMed:24784582, PubMed:25474007,
PubMed: 25527291). After mitochondrial damage, functions with PINK1 to
mediate the decision between mitophagy or preventing apoptosis by inducing
either the poly- or monoubiquitination of VDAC1, respectively;
polyubiquitination of VDAC1 promotes mitophagy, while monoubiquitination
of VDAC1 decreases mitochondrial calcium influx which ultimately inhibits
apoptosis (PubMed: 27534820, PubMed: 32047033). When cellular stress
results in irreversible mitochondrial damage, promotes the autophagic
degradation of dysfunctional depolarized mitochondria (mitophagy) by
promoting the ubiquitination of mitochondrial proteins such as TOMM20,
RHOT1/MIRO1, MFN1 and USP30 (PubMed: 19029340, PubMed: 19966284,
PubMed:21753002, PubMed:22396657, PubMed:23620051,
PubMed:23685073, PubMed:23933751, PubMed:24896179,
PubMed: 25527291). Preferentially assembles 'Lys-6'-, 'Lys-11'- and
'Lys-63'-linked polyubiquitin chains, leading to mitophagy (PubMed: 25621951,
PubMed:32047033). The PINK1-PRKN pathway also promotes fission of
damaged mitochondria by PINK1-mediated phosphorylation which promotes
the PRKN-dependent degradation of mitochondrial proteins involved in
fission such as MFN2 (PubMed:23620051). This prevents the refusion of
unhealthy mitochondria with the mitochondrial network or initiates
mitochondrial fragmentation facilitating their later engulfment by
autophagosomes (PubMed: 23620051). Regulates motility of damaged
mitochondria via the ubiquitination and subsequent degradation of MIRO1
and MIRO2; in motor neurons, this likely inhibits mitochondrial intracellular
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anterograde transport along the axons which probably increases the chance of the mitochondria undergoing mitophagy in the soma (PubMed:22396657). Involved in mitochondrial biogenesis via the 'Lys-48'-linked polyubiquitination of transcriptional repressor ZNF746/PARIS which leads to its subsequent proteasomal degradation and allows activation of the transcription factor PPARGC1A (PubMed:21376232). Limits the production of reactive oxygen species (ROS) (PubMed: 18541373). Regulates cyclin-E during neuronal apoptosis (PubMed: 12628165). In collaboration with CHPF isoform 2, may enhance cell viability and protect cells from oxidative stress (PubMed: 22082830). Independently of its ubiquitin ligase activity, protects from apoptosis by the transcriptional repression of p53/TP53 (PubMed: 19801972). May protect neurons against alpha synuclein toxicity, proteasomal dysfunction, GPR37 accumulation, and kainate-induced excitotoxicity (PubMed: 11439185). May play a role in controlling neurotransmitter trafficking at the presynaptic terminal and in calcium-dependent exocytosis. May represent a tumor suppressor gene (PubMed:12719539).

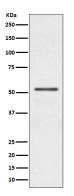
Cellular Location

Cytoplasm, cytosol. Nucleus. Endoplasmic reticulum. Mitochondrion. Mitochondrion outer membrane {ECO:0000250 | UniProtKB:Q9WVS6}. Cell projection, neuron projection. Postsynaptic density {ECO:0000250 | UniProtKB:Q9WVS6}. Presynapse {ECO:0000250 | UniProtKB:Q9WVS6}. Note=Mainly localizes in the cytosol (PubMed:19029340, PubMed:19229105). Co-localizes with SYT11 in neutrites (PubMed:12925569). Co-localizes with SNCAIP in brainstem Lewy bodies (PubMed:10319893, PubMed:11431533). Translocates to dysfunctional mitochondria that have lost the mitochondrial membrane potential; recruitment to mitochondria is PINK1-dependent (PubMed:18957282, PubMed:19966284, PubMed:23620051, PubMed:24898855) Mitochondrial localization also gradually increases with cellular growth (PubMed:22082830).

Tissue Location

Highly expressed in the brain including the substantia nigra (PubMed:19501131, PubMed:9560156). Expressed in heart, testis and skeletal muscle (PubMed:9560156). Expression is down-regulated or absent in tumor biopsies, and absent in the brain of PARK2 patients (PubMed:12719539, PubMed:14614460). Overexpression protects dopamine neurons from kainate-mediated apoptosis (PubMed:12628165) Found in serum (at protein level) (PubMed:19501131)

Images



Western blot analysis of Parkin expression in Jurkat cell lysate.

Please note: All products are 'FOR RESEARCH USE ONLY. NOT FOR USE IN DIAGNOSTIC OR THERAPEUTIC PROCEDURES'.