

APP Antibody (N-term)

Purified Rabbit Polyclonal Antibody (Pab) Catalog # AP6306A

Product Information

Application Primary Accession Other Accession	WB, IHC-P, FC, E <u>P05067</u> P53601
Reactivity	Human, Rat, Mouse
Predicted	Monkey
Host	Rabbit
Clonality	Polyclonal
Isotype	Rabbit IgG
Calculated MW	86943
Antigen Region	29-61

Additional Information

Gene ID	351
Other Names	Amyloid beta A4 protein, ABPP, APPI, APP, Alzheimer disease amyloid protein, Cerebral vascular amyloid peptide, CVAP, PreA4, Protease nexin-II, PN-II, N-APP, Soluble APP-alpha, S-APP-alpha, Soluble APP-beta, S-APP-beta, C99, Beta-amyloid protein 42, Beta-APP42, Beta-amyloid protein 40, Beta-APP40, C83, P3(42), P3(40), C80, Gamma-secretase C-terminal fragment 59, Amyloid intracellular domain 59, AICD-59, AID(59), Gamma-CTF(59), Gamma-secretase C-terminal fragment 57, Amyloid intracellular domain 57, AICD-57, AID(57), Gamma-CTF(57), Gamma-secretase C-terminal fragment 50, Amyloid intracellular domain 50, AICD-50, AID(50), Gamma-CTF(50), C31, APP, A4, AD1
Target/Specificity	This APP antibody is generated from rabbits immunized with a KLH conjugated synthetic peptide between 29-61 amino acids from the N-terminal region of human APP.
Dilution	WB~~1:2000 IHC-P~~1:100~500 FC~~1:10~50 E~~Use at an assay dependent concentration.
Format	Purified polyclonal antibody supplied in PBS with 0.09% (W/V) sodium azide. This antibody is prepared by Saturated Ammonium Sulfate (SAS) precipitation followed by dialysis against PBS.
Storage	Maintain refrigerated at 2-8°C for up to 2 weeks. For long term storage store at -20°C in small aliquots to prevent freeze-thaw cycles.
Precautions	APP Antibody (N-term) is for research use only and not for use in diagnostic or therapeutic procedures.

Name

Function

APP (<u>HGNC:620</u>)

Functions as a cell surface receptor and performs physiological functions on the surface of neurons relevant to neurite growth, neuronal adhesion and axonogenesis. Interaction between APP molecules on neighboring cells promotes synaptogenesis (PubMed:25122912). Involved in cell mobility and transcription regulation through protein-protein interactions. Can promote transcription activation through binding to APBB1-KAT5 and inhibits Notch signaling through interaction with Numb. Couples to apoptosis- inducing pathways such as those mediated by G(o) and JIP. Inhibits G(o) alpha ATPase activity (By similarity). Acts as a kinesin I membrane receptor, mediating the axonal transport of beta-secretase and presenilin 1 (By similarity). By acting as a kinesin I membrane receptor, plays a role in axonal anterograde transport of cargo towards synapses in axons (PubMed:<u>17062754</u>, PubMed:23011729). Involved in copper homeostasis/oxidative stress through copper ion reduction. In vitro, copper-metallated APP induces neuronal death directly or is potentiated through Cu(2+)-mediated low-density lipoprotein oxidation. Can regulate neurite outgrowth through binding to components of the extracellular matrix such as heparin and collagen I and IV. The splice isoforms that contain the BPTI domain possess protease inhibitor activity. Induces a AGER-dependent pathway that involves activation of p38 MAPK, resulting in internalization of amyloid-beta peptide and leading to mitochondrial dysfunction in cultured cortical neurons. Provides Cu(2+) ions for GPC1 which are required for release of nitric oxide (NO) and subsequent degradation of the heparan sulfate chains on GPC1. [Amyloid-beta protein 42]: More effective reductant than amyloid-beta protein 40. May activate mononuclear phagocytes in the brain and elicit inflammatory responses. The gamma-CTF peptides as well as the caspase-cleaved peptides, including C31, are potent enhancers of neuronal apoptosis.

Cellular Location Cell membrane; Single-pass type I membrane protein. Membrane; Single-pass type I membrane protein. Perikaryon Cell projection, growth cone. Membrane, clathrin-coated pit. Early endosome. Cytoplasmic vesicle. Note=Cell surface protein that rapidly becomes internalized via clathrin-coated pits. Only a minor proportion is present at the cell membrane; most of the protein is present in intracellular vesicles (PubMed:20580937) During maturation, the immature APP (N-glycosylated in the endoplasmic reticulum) moves to the Golgi complex where complete maturation occurs (O-glycosylated and sulfated). After alpha-secretase cleavage, soluble APP is released into the extracellular space and the C-terminal is internalized to endosomes and lysosomes. Some APP accumulates in secretory transport vesicles leaving the late Golgi compartment and returns to the cell surface. APP sorts to the basolateral surface in epithelial cells. During neuronal differentiation, the Thr-743 phosphorylated form is located mainly in growth cones, moderately in neurites and sparingly in the cell body (PubMed:10341243). Casein kinase phosphorylation can occur either at the cell surface or within a post-Golgi compartment. Associates with GPC1 in perinuclear compartments. Colocalizes with SORL1 in a vesicular pattern in cytoplasm and perinuclear regions. [C99]: Early endosome [Amyloid-beta protein 40]: Cell surface [Gamma-secretase C-terminal fragment 59]: Nucleus. Cytoplasm Note=Located to both the cytoplasm and nuclei of neurons. It can be translocated to the nucleus through association with APBB1 (Fe65) (PubMed:11544248). In dopaminergic neurons, the phosphorylated Thr-743 form is localized to the nucleus (By similarity) {ECO:0000250|UniProtKB:P12023, ECO:0000269|PubMed:11544248} Expressed in the brain and in cerebrospinal fluid (at protein level) **Tissue Location** (PubMed:2649245). Expressed in all fetal tissues examined with highest levels

in brain, kidney, heart and spleen. Weak expression in liver. In adult brain, highest expression found in the frontal lobe of the cortex and in the anterior perisylvian cortex- opercular gyri. Moderate expression in the cerebellar cortex, the posterior perisylvian cortex-opercular gyri and the temporal associated cortex. Weak expression found in the striate, extra-striate and motor cortices. Expressed in cerebrospinal fluid, and plasma. Isoform APP695 is the predominant form in neuronal tissue, isoform APP751 and isoform APP770 are widely expressed in non-neuronal cells. Isoform APP751 is the most abundant form in T-lymphocytes. Appican is expressed in astrocytes.

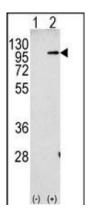
Background

APP is a cell surface receptor and transmembrane precursor protein that is cleaved by secretases to form a number of peptides. Some of these peptides are secreted and can bind to the acetyltransferase complex APBB1/TIP60 to promote transcriptional activation, while others form the protein basis of the amyloid plaques found in the brains of patients with Alzheimer disease. Mutations in this gene have been implicated in autosomal dominant Alzheimer disease and cerebroarterial amyloidosis (cerebral amyloid angiopathy).

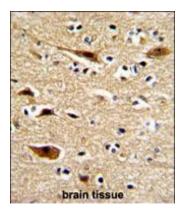
References

Borroni, B., et al., Arch. Neurol. 60(12):1740-1744 (2003). Lleo, A., et al., J. Biol. Chem. 278(48):47370-47375 (2003). Zhou, Y., et al., Science 302(5648):1215-1217 (2003). Murakami, K., et al., J. Biol. Chem. 278(46):46179-46187 (2003). Kim, J.Y., et al., Biochem. Biophys. Res. Commun. 310(3):904-909 (2003).

Images

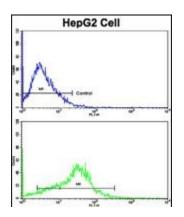


Western blot analysis of APP (arrow) using rabbit polyclonal APP Antibody (N-term) (RB05012). 293 cell lysates (2 ug/lane) either nontransfected (Lane 1) or transiently transfected with the APP gene (Lane 2) (Origene Technologies).



Formalin-fixed and paraffin-embedded human brain with APP Antibody (N-term), which was peroxidase-conjugated to the secondary antibody, followed by DAB staining. This data demonstrates the use of this antibody for immunohistochemistry; clinical relevance has not been evaluated.

Flow cytometric analysis of HepG2 cells using APP Antibody (N-term)(bottom histogram) compared to a negative control cell (top histogram). FITC-conjugated goat-anti-rabbit secondary antibodies were used for the analysis.



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