

# Ambral Antibody

Catalog # ASC10675

# Specification

# Ambra1 Antibody - Product Information

Application Primary Accession Other Accession Reactivity Host Clonality Isotype Application Notes

WB, IHC, IF <u>O9C0C7</u> <u>O9C0C7</u>, <u>166215833</u> Human, Mouse, Rat Rabbit Polyclonal IgG Ambra1 antibody can be used for the detection of Ambra1 by Western blot at 1 μg/mL. Antibody can also be used for immunohistochemistry starting at 5 μg/mL. For immunofluorescence start at 20 μg/mL.

# Ambral Antibody - Additional Information

Gene ID Target/Specificity AMBRA1;

### **Reconstitution & Storage**

Ambra1 antibody can be stored at 4°C for three months and -20°C, stable for up to one year. As with all antibodies care should be taken to avoid repeated freeze thaw cycles. Antibodies should not be exposed to prolonged high temperatures.

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**Precautions** Ambra1 Antibody is for research use only and not for use in diagnostic or therapeutic procedures.

# Ambra1 Antibody - Protein Information

Name AMBRA1 {ECO:0000303|PubMed:17589504, ECO:0000312|HGNC:HGNC:25990}

Function

Substrate-recognition component of a DCX (DDB1-CUL4-X-box) E3 ubiquitin-protein ligase complex involved in cell cycle control and autophagy (PubMed:<a

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href="http://www.uniprot.org/citations/20921139" target="_blank">20921139</a>, PubMed:<a
href="http://www.uniprot.org/citations/23524951" target="_blank">23524951</a>, PubMed:<a
href="http://www.uniprot.org/citations/24587252" target="_blank">24587252</a>, PubMed:<a
href="http://www.uniprot.org/citations/32333458" target="_blank">32333458</a>, PubMed:<a
href="http://www.uniprot.org/citations/32333458" target="_blank">3333458</a>, PubMed:<a
href="http://www.uniprot.org/citations/33854232" target="_blank">33854232</a>, PubMed:<a
href="http://www.uniprot.org/citations/33854235" target="_blank">33854232</a>, PubMed:<a
href="http://www.uniprot.org/citations/33854239" target="_blank">33854235</a>, PubMed:<a
href="http://www.uniprot.org/citations/33854239" target="_blank">33854239</a>). The
DCX(AMBRA1) complex specifically mediates the polyubiquitination of target proteins such as
BECN1, CCND1, CCND2, CCND3, ELOC and ULK1 (PubMed:<a
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href="http://www.uniprot.org/citations/23524951" target=" blank">23524951</a>, PubMed:<a href="http://www.uniprot.org/citations/33854232" target=" blank">33854232</a>, PubMed:<a href="http://www.uniprot.org/citations/33854235" target=" blank">33854235</a>, PubMed:<a href="http://www.uniprot.org/citations/33854239" target=" blank">33854239</a>). Acts as an upstream master regulator of the transition from G1 to S cell phase: AMBRA1 specifically recognizes and binds phosphorylated cyclin-D (CCND1, CCND2 and CCND3), leading to cyclin-D ubiguitination by the DCX(AMBRA1) complex and subsequent degradation (PubMed:<a href="http://www.uniprot.org/citations/33854232" target=" blank">33854232</a>, PubMed:<a href="http://www.uniprot.org/citations/33854235" target="\_blank">33854235</a>, PubMed:<a href="http://www.uniprot.org/citations/33854239" target="\_blank">33854239</a>). By controlling the transition from G1 to S phase and cyclin-D degradation, AMBRA1 acts as a tumor suppressor that promotes genomic integrity during DNA replication and counteracts developmental abnormalities and tumor growth (PubMed: <a href="http://www.uniprot.org/citations/33854232" target=" blank">33854232</a>, PubMed:<a href="http://www.uniprot.org/citations/33854235" target="\_blank">33854235</a>, PubMed:<a href="http://www.uniprot.org/citations/33854239" target=" blank">33854239</a>). AMBRA1 also regulates the cell cycle by promoting MYC dephosphorylation and degradation independently of the DCX(AMBRA1) complex: acts via interaction with the catalytic subunit of protein phosphatase 2A (PPP2CA), which enhances interaction between PPP2CA and MYC, leading to MYC dephosphorylation and degradation (PubMed: <a href="http://www.uniprot.org/citations/25438055" target=" blank">25438055</a>, PubMed:<a href="http://www.uniprot.org/citations/25803737" target=" blank">25803737</a>). Acts as a regulator of Cul5-RING (CRL5) E3 ubiguitin- protein ligase complexes by mediating ubiguitination and degradation of Elongin-C (ELOC) component of CRL5 complexes (PubMed:<a href="http://www.uniprot.org/citations/25499913" target=" blank">25499913</a>, PubMed:<a href="http://www.uniprot.org/citations/30166453" target=" blank">30166453</a>). Acts as a key regulator of autophagy by modulating the BECN1-PIK3C3 complex: controls protein turnover during neuronal development, and regulates normal cell survival and proliferation (PubMed:<a href="http://www.uniprot.org/citations/21358617" target=" blank">21358617</a>). In normal conditions, AMBRA1 is tethered to the cytoskeleton via interaction with dyneins DYNLL1 and DYNLL2 (PubMed:<a href="http://www.uniprot.org/citations/20921139" target=" blank">20921139</a>). Upon autophagy induction, AMBRA1 is released from the cytoskeletal docking site to induce autophagosome nucleation by mediating ubiguitination of proteins involved in autophagy (PubMed:<a href="http://www.uniprot.org/citations/20921139" target=" blank">20921139</a>). The DCX(AMBRA1) complex mediates 'Lys-63'-linked ubiguitination of BECN1, increasing the association between BECN1 and PIK3C3 to promote PIK3C3 activity (By similarity). In collaboration with TRAF6, AMBRA1 mediates 'Lys-63'-linked ubiquitination of ULK1 following autophagy induction, promoting ULK1 stability and kinase activity (PubMed:<a href="http://www.uniprot.org/citations/23524951" target="\_blank">23524951</a>). Also activates ULK1 via interaction with TRIM32: TRIM32 stimulates ULK1 through unanchored 'Lys-63'-linked polyubiquitin chains (PubMed: <a href="http://www.uniprot.org/citations/31123703" target=" blank">31123703</a>). Also acts as an activator of mitophagy via interaction with PRKN and LC3 proteins (MAP1LC3A, MAP1LC3B or MAP1LC3C); possibly by bringing damaged mitochondria onto autophagosomes (PubMed:<a href="http://www.uniprot.org/citations/21753002" target=" blank">21753002</a>, PubMed:<a href="http://www.uniprot.org/citations/25215947" target=" blank">25215947</a>). Also activates mitophagy by acting as a cofactor for HUWE1; acts by promoting HUWE1- mediated ubiquitination of MFN2 (PubMed:<a href="http://www.uniprot.org/citations/30217973" target=" blank">30217973</a>). AMBRA1 is also involved in regulatory T-cells (Treg) differentiation by promoting FOXO3 dephosphorylation independently of the DCX(AMBRA1) complex: acts via interaction with PPP2CA, which enhances interaction between PPP2CA and FOXO3, leading to FOXO3 dephosphorylation and stabilization (PubMed: <a href="http://www.uniprot.org/citations/30513302" target=" blank">30513302</a>). May act as a regulator of intracellular trafficking, regulating the localization of active PTK2/FAK and SRC (By similarity). Also involved in transcription regulation by acting as a scaffold for protein complexes at chromatin (By similarity).



# **Cellular Location**

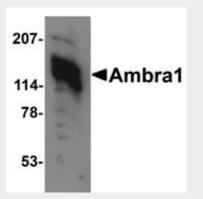
Endoplasmic reticulum. Cytoplasm, cytoskeleton. Cytoplasmic vesicle, autophagosome {ECO:000250|UniProtKB:A2AH22}. Mitochondrion. Cytoplasm, cytosol {ECO:000250|UniProtKB:A2AH22}. Nucleus. Cell junction, focal adhesion {ECO:0000250|UniProtKB:A2AH22}. Note=Localizes to the cytoskeleton in absence of autophagy induction (PubMed:20921139). Upon autophagy induction, AMBRA1 relocalizes to the endoplasmic reticulum to enable autophagosome nucleation (PubMed:20921139). Partially localizes at mitochondria in normal conditions (PubMed:21358617). Localizes also to discrete punctae along the ciliary axoneme (By similarity) {ECO:0000250|UniProtKB:A2AH22, ECO:0000269|PubMed:20921139, ECO:0000269|PubMed:21358617}

### Ambra1 Antibody - Protocols

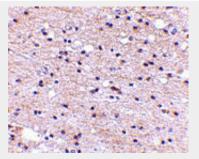
Provided below are standard protocols that you may find useful for product applications.

- <u>Western Blot</u>
- Blocking Peptides
- Dot Blot
- Immunohistochemistry
- Immunofluorescence
- Immunoprecipitation
- Flow Cytomety
- <u>Cell Culture</u>

### Ambra1 Antibody - Images

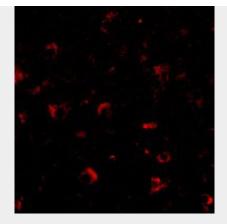


Western blot analysis of Ambra1 in 3T3 cell lysate with Ambra1 antibody at 1 µg/mL.



Immunohistochemistry of Ambra1 in human brain with Ambra1 antibody at 5 µg/mL.





Immunofluorescence of Ambra1 in Human Brain cells with Ambra1 antibody at 20 µg/mL.

# Ambra1 Antibody - Background

Ambra1 Antibody: Autophagy, the process of bulk degradation of cellular proteins through an autophagosomic-lysosomal pathway is important for normal growth control and may be defective in tumor cells. It is involved in the preservation of cellular nutrients under starvation conditions as well as the normal turnover of cytosolic components. Beclin-1, a principal regulator of autophagosome formation, is in turn regulated by Ambra1. Ambra1 associates with Beclin-1 through a region near its center as determined by yeast two-hybrid assay. Null mutations in this gene in mice resulted in embryonic lethality with severe neural tube defects associated with autophagy impairment, accumulation of ubiquitinated proteins, unbalanced cell proliferation and excessive apoptotic death. Furthermore, down-regulation of Ambra1 in cultured cells though RNA interference decreased the level of rapamycin- and nutrient starvation-induced autophagy. Multiple isoforms of Ambra1 are known to exist.

### Ambra1 Antibody - References

Gozuacik D and Kimchi A. Autophagy as a cell death and tumor suppressor mechanism. Oncogene2004; 23:2891-906.

Kisen GO, Tessitore L, Costelli P, et al. Reduced autophagic activity in primary rat hepatocellular carcinoma and ascites hepatoma cells. Carcinogenesis1993; 14:2501-5.

Liang XH, Jackson S, Seaman M, et al. Induction of autophagy and inhibition of tumorigenesis by beclin 1. Nature1999; 402:672-6.

Fimia GM, Stoykova A, Romagnoli A, et al. Ambra1 regulates autophagy and development of the nervous system. Nature2007; 447:1121-5.