

# KEAP1 Antibody

Catalog # ASC11552

## Specification

# **KEAP1** Antibody - Product Information

Application Primary Accession Other Accession Reactivity Host Clonality Isotype Calculated MW Application Notes WB, IF <u>O14145</u> <u>NP\_987096</u>, <u>45269145</u> Human, Mouse, Rat Rabbit Polyclonal IgG 69 kDa KDa KEAP1 antibody can be used for detection of KEAP1 by Western blot at 1 - 2 μg/mL. For immunofluorescence start at 20 μg/mL.

## **KEAP1** Antibody - Additional Information

Gene ID 9817 Target/Specificity KEAP1; At least two isoforms of KEAP1 are known to exist.

#### **Reconstitution & Storage**

KEAP1 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.

#### Precautions

KEAP1 Antibody is for research use only and not for use in diagnostic or therapeutic procedures.

## **KEAP1 Antibody - Protein Information**

Name KEAP1 {ECO:0000303|PubMed:14585973, ECO:0000312|HGNC:HGNC:23177}

#### Function

Substrate-specific adapter of a BCR (BTB-CUL3-RBX1) E3 ubiquitin ligase complex that regulates the response to oxidative stress by targeting NFE2L2/NRF2 for ubiquitination (PubMed:<a href="http://www.uniprot.org/citations/14585973" target="\_blank">14585973</a>, PubMed:<a href="http://www.uniprot.org/citations/15379550" target="\_blank">14585973</a>, PubMed:<a href="http://www.uniprot.org/citations/15572695" target="\_blank">15572695</a>, PubMed:<a href="http://www.uniprot.org/citations/15572695" target="\_blank">15572695</a>, PubMed:<a href="http://www.uniprot.org/citations/15601839" target="\_blank">15601839</a>, PubMed:<a href="http://www.uniprot.org/citations/15983046" target="\_blank">15983046</a>, PubMed:<a href="http://www.uniprot.org/citations/15983046" target="\_blank">15983046</a>, PubMed:<a href="http://www.uniprot.org/citations/15983046" target="\_blank">15983046</a>, PubMed:<a href="http://www.uniprot.org/citations/15983046" target="\_blank">15983046</a>, PubMed:<a href="http://www.uniprot.org/citations/37339955" target="\_blank">37339955</a>, A key sensor of oxidative and electrophilic stress: in normal conditions, the BCR(KEAP1) complex mediates ubiquitination and degradation of NFE2L2/NRF2, a transcription factor regulating expression of many cytoprotective genes (PubMed:<a href="http://www.uniprot.org/citations/15601839" target="\_blank">15601839</a>, PubMed:<a href="http://www.uniprot.org/citations/15601839" target="\_blank">15601839</a>, PubMed:<a href="http://www.uniprot.org/citations/37339955" target="\_blank">15601839</a>, PubMed:<a href="http://www.uniprot.org/citations/15601839" target="\_blank">15601839</a>, PubMed:<a href="http://www.uniprot.org/cita



href="http://www.uniprot.org/citations/16006525" target=" blank">16006525</a>). In response to oxidative stress, different electrophile metabolites trigger non-enzymatic covalent modifications of highly reactive cysteine residues in KEAP1, leading to inactivate the ubiguitin ligase activity of the BCR(KEAP1) complex, promoting NFE2L2/NRF2 nuclear accumulation and expression of phase II detoxifying enzymes (PubMed:<a href="http://www.uniprot.org/citations/16006525" target=" blank">16006525</a>, PubMed:<a href="http://www.uniprot.org/citations/17127771" target=" blank">17127771</a>, PubMed:<a href="http://www.uniprot.org/citations/18251510" target=" blank">18251510</a>, PubMed:<a href="http://www.uniprot.org/citations/19489739" target=" blank">19489739</a>, PubMed:<a href="http://www.uniprot.org/citations/29590092" target=" blank">29590092</a>). In response to selective autophagy, KEAP1 is sequestered in inclusion bodies following its interaction with SQSTM1/p62, leading to inactivation of the BCR(KEAP1) complex and activation of NFE2L2/NRF2 (PubMed: <a href="http://www.uniprot.org/citations/20452972" target=" blank">20452972</a>). The BCR(KEAP1) complex also mediates ubiquitination of SQSTM1/p62, increasing SQSTM1/p62 sequestering activity and degradation (PubMed:<a href="http://www.uniprot.org/citations/28380357" target=" blank">28380357</a>). The BCR(KEAP1) complex also targets BPTF and PGAM5 for ubiguitination and degradation by the

proteasome (PubMed:<a href="http://www.uniprot.org/citations/15379550" target="\_blank">15379550</a>, PubMed:<a href="http://www.uniprot.org/citations/17046835" target="\_blank">17046835</a>).

#### **Cellular Location**

Cytoplasm. Nucleus. Note=Mainly cytoplasmic (PubMed:15601839). In response to selective autophagy, relocalizes to inclusion bodies following interaction with SQSTM1/p62 (PubMed:20452972).

**Tissue Location** Broadly expressed, with highest levels in skeletal muscle.

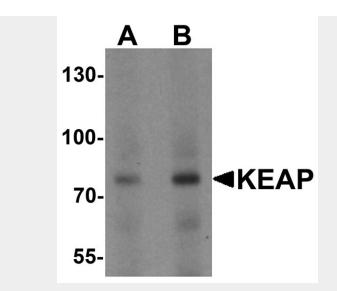
#### **KEAP1 Antibody - Protocols**

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

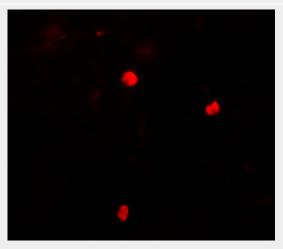
- <u>Western Blot</u>
- Blocking Peptides
- Dot Blot
- Immunohistochemistry
- Immunofluorescence
- Immunoprecipitation
- Flow Cytomety
- <u>Cell Culture</u>

**KEAP1 Antibody - Images** 





Western blot analysis of KEAP1 in human lung tissue lysate with KEAP1 antibody at (A) 1 and (B) 2  $\mu$ g/mL.



Immunofluorescence of KEAP1 in human lung tissue with KEAP1 antibody at 20 µg/mL.

# KEAP1 Antibody - Background

KEAP1 Antibody: KEAP1 (kelch-like ECH-associated protein 1) is a stress sensing adaptor for the Cullin3 (Cul3)-dependent E3 ubiquitin ligase complex that negatively regulates NRF2 (NF-E2-related factor 2) and plays a role in the oxidative stress response. It targets NFE2L2/NRF2 for ubiquitination and degradation by the proteasome. KEAP1 contains an amino terminal BTB/POZ domain and a carboxyl terminal KELCH domain which are required for interaction with NRF2, and in binding Cul3-E3 ubiquitin ligase. Altered expression of NRF2 is associated with chronic obstructive pulmonary disease (COPD). KEAP1 also targets the down regulation of NF-κB activity by targeting IKKβ degradation. Mutation of the KEAP1 gene is found in lung cancer.

## **KEAP1** Antibody - References

Zhang DD, Lo SC, Cross JV, et al. Keap1 is a redox-regulated substrate adaptor protein for a Cul3-dependent ubiquitin ligase complex. Mol. Cell. Biol. 2004;24:10941-53.

Kobayashi A, Kang MI, Okawa H, et al. Oxidative stress sensor Keap1 functions as an adaptor for Cul3-based E3 ligase to regulate proteasomal degradation of Nrf2. Mol. Cell. Biol. 2004; 24:7130-9. Jiang J, Mo ZC, Yin K, et al. Epigallocatechin-3-gallate prevents TNF- $\alpha$ -induced NF-kappaB activation thereby upregulating ABCA1 via the Nrf2/Keap1 pathway in macrophage foam cells. Int. J. Mol. Med. 2012; 29:946-56.



Devling TW, Lindsay CD, McLellan LI, et al. Utility of siRNA against Keap1 as a strategy to stimulate a cancer chemopreventive phenotype. Proc. Natl. Acad. Sci. USA 2005; 102: 7280-5A.