

Rabbit Anti-phospho-Nrf2 (Ser40) antibody

Purified Rabbit Polyclonal Antibody (Pab) Catalog # AP52270

Product Information

Application WB, IHC-P, IHC-F, IF, ICC, E

Primary Accession Q16236

Reactivity Human, Mouse, Rat

Host Rabbit Clonality Polyclonal Calculated MW 67827 **Physical State** Liquid

KLH conjugated Synthesised phosphopeptide derived from human Nrf2 **Immunogen**

around the phosphorylation site of Ser40

Epitope Specificity DF(p-S)QR Isotype IgG

Purity affinity purified by Protein A

0.01M TBS (pH7.4) with 1% BSA, 0.02% Proclin300 and 50% Glycerol. **Buffer**

SUBCELLULAR LOCATION Cytoplasm, cytosol. Nucleus. Note=Cytosolic under unstressed conditions,

translocates into the nucleus upon induction by electrophilic agents.

SIMILARITY Belongs to the bZIP family. CNC subfamily. Contains 1 bZIP domain. Heterodimer. Forms a ternary complex with PGAM5 and KEAP1. May bind **SUBUNIT**

DNA with an unknown protein. Interacts via its leucine-zipper domain with

the coiled-coil domain of PMF1.

Phosphorylation of Ser-40 by PKC in response to oxidative stress dissociates Post-translational modifications

NFE2L2 from its cytoplasmic inhibitor KEAP1, promoting its translocation into

the nucleus.

Important Note This product as supplied is intended for research use only, not for use in

human, therapeutic or diagnostic applications.

Background Descriptions Nuclear factor erythroid 2-related factor 2 (Nrf2) is a transcription factor

which regulates the expression of many detoxification and antioxidant enzymes. Nrf2 can potentially play a significant role in adaptive responses to oxidative stress. Nrf2 belongs to the Cap N Collar (CNC-bZIP) subfamily of

basic /leucine zipper (bZIP) transcription factors.

Additional Information

Gene ID 4780

Other Names NRF2; Nuclear factor erythroid 2-related factor 2; NF-E2-related factor 2;

NFE2-related factor 2; HEBP1; Nuclear factor, erythroid derived 2, like 2;

Target/Specificity Widely expressed. Highest expression in adult muscle, kidney, lung, liver and

in fetal muscle.

Dilution WB=1:500-2000,IHC-P=1:100-500,IHC-F=1:100-500,ICC=1:100,IF=1:100-500,Flo

w-Cyt=1 Ig /test,ELISA=1:5000-10000

Format 0.01M TBS(pH7.4) with 1% BSA, 0.09% (W/V) sodium azide and 50% Glyce

Storage Store at -20 °C for one year. Avoid repeated freeze/thaw cycles. When

reconstituted in sterile pH 7.4 0.01M PBS or diluent of antibody the antibody

is stable for at least two weeks at 2-4 °C.

Protein Information

Name

NFE2L2 {ECO:0000303 | PubMed:29018201, ECO:0000312 | HGNC:HGNC:7782}

Function

Transcription factor that plays a key role in the response to oxidative stress: binds to antioxidant response (ARE) elements present in the promoter region of many cytoprotective genes, such as phase 2 detoxifying enzymes, and promotes their expression, thereby neutralizing reactive electrophiles (PubMed: 11035812, PubMed: 19489739, PubMed: 29018201, PubMed: 31398338). In normal conditions, ubiquitinated and degraded in the cytoplasm by the BCR(KEAP1) complex (PubMed: 11035812, PubMed: 15601839, PubMed: 29018201). In response to oxidative stress, electrophile metabolites inhibit activity of the BCR(KEAP1) complex, promoting nuclear accumulation of NFE2L2/NRF2, heterodimerization with one of the small Maf proteins and binding to ARE elements of cytoprotective target genes (PubMed: <u>19489739</u>, PubMed: <u>29590092</u>). The NFE2L2/NRF2 pathway is also activated in response to selective autophagy: autophagy promotes interaction between KEAP1 and SQSTM1/p62 and subsequent inactivation of the BCR(KEAP1) complex, leading to NFE2L2/NRF2 nuclear accumulation and expression of cytoprotective genes (PubMed: 20452972). The NFE2L2/NRF2 pathway is also activated during the unfolded protein response (UPR), contributing to redox homeostasis and cell survival following endoplasmic reticulum stress (By similarity). May also be involved in the transcriptional activation of genes of the beta-globin cluster by mediating enhancer activity of hypersensitive site 2 of the beta-globin locus control region (PubMed:7937919). Also plays an important role in the regulation of the innate immune response and antiviral cytosolic DNA sensing. It is a critical regulator of the innate immune response and survival during sepsis by maintaining redox homeostasis and restraint of the dysregulation of pro-inflammatory signaling pathways like MyD88- dependent and -independent and TNF-alpha signaling (By similarity). Suppresses macrophage inflammatory response by blocking pro- inflammatory cytokine transcription and the induction of IL6 (By similarity). Binds to the proximity of pro-inflammatory genes in macrophages and inhibits RNA Pol II recruitment. The inhibition is independent of the NRF2-binding motif and reactive oxygen species level (By similarity). Represses antiviral cytosolic DNA sensing by suppressing the expression of the adapter protein STING1 and decreasing responsiveness to STING1 agonists while increasing susceptibility to infection with DNA viruses (PubMed:30158636). Once activated, limits the release of pro-inflammatory cytokines in response to human coronavirus SARS-CoV-2 infection and to virus-derived ligands through a mechanism that involves inhibition of IRF3 dimerization. Also inhibits both SARS-CoV-2 replication, as well as the replication of several other pathogenic viruses including Herpes Simplex Virus-1 and-2, Vaccinia virus, and Zika virus through a type I interferon (IFN)- independent mechanism (PubMed:33009401).

Cellular Location

Cytoplasm, cytosol. Nucleus {ECO:0000255 | PROSITE-ProRule:PRU00978, ECO:0000269 | PubMed:11035812, ECO:0000269 | PubMed:15601839, ECO:0000269 | PubMed:21196497, ECO:0000269 | PubMed:29983246 }.

Note=Cytosolic under unstressed conditions: ubiquitinated and degraded by the BCR(KEAP1) E3 ubiquitin ligase complex (PubMed:15601839, PubMed:21196497). Translocates into the nucleus upon induction by electrophilic agents that inactivate the BCR(KEAP1) E3 ubiquitin ligase complex (PubMed:21196497)

Tissue Location

Widely expressed. Highest expression in adult muscle, kidney, lung, liver and in fetal muscle

Background

Transcription activator that binds to antioxidant response (ARE) elements in the promoter regions of target genes. Important for the coordinated up-regulation of genes in response to oxidative stress. May be involved in the transcriptional activation of genes of the beta-globin cluster by mediating enhancer activity of hypersensitive site 2 of the beta-globin locus control region.

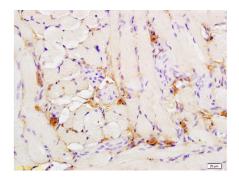
References

Ota T., et al. Nat. Genet. 36:40-45(2004). Totoki Y., et al. Submitted (MAR-2005) to the EMBL/GenBank/DDBJ databases. Hillier L.W., et al. Nature 434:724-731(2005). Mural R.J., et al. Submitted (SEP-2005) to the EMBL/GenBank/DDBJ databases. Bechtel S., et al. BMC Genomics 8:399-399(2007).

Images

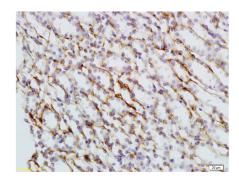


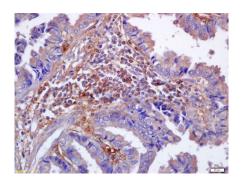
L1 mouse heart, L2 mouse muscle lysates probed (AP52270) at 1:200 in 4°C. Followed by conjugation to secondary antibody at 1:3000 90min in 37°C. Predicted and observed band size: 66kDa.



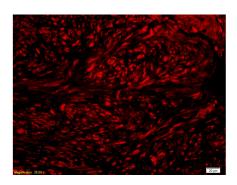
Formalin-fixed and paraffin embedded rat tongue tissue labeled with Anti-phospho-Nrf2 (Ser40) Polyclonal Antibody (AP52270), Unconjugated 1:200 followed by conjugation to the secondary antibody and DAB staining

Formalin-fixed and paraffin embedded rat kidney tissue labeled with Anti-phospho-Nrf2 (Ser40) Polyclonal Antibody (AP52270), Unconjugated 1:200 followed by conjugation to the secondary antibody and DAB staining





Formalin-fixed and paraffin embedded human lung carcinoma labeled with Anti-phospho-Nrf2 (Ser40) Polyclonal Antibody, Unconjugated (AP52270) at 1:200 followed by conjugation to the secondary antibody and DAB staining



Formalin-fixed and paraffin embedded human colon carcinoma tissue labeled with Anti-phospho-Nrf2 (Ser40) Polyclonal Antibody (AP52270), Unconjugated 1:200 followed by conjugation to the secondary Cy5 antibody

Citations

- Nrf2 Promotes Inflammation in Early Myocardial Ischemia-Reperfusion Recruitment and Activation of Macrophages
- <u>Scutellarin Prevents Nonalcoholic Fatty Liver Disease (NAFLD) and Hyperlipidemia via PI3K/AKT-Dependent Activation of Nuclear Factor (Erythroid-Derived 2)-Like 2 (Nrf2) in Rats.</u>

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