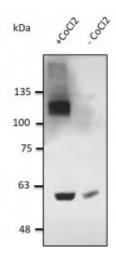


Anti-HIF1a antibody



Description

Goat polyclonal to alpha subunit of transcription factor hypoxia-inducible factor-1 (HIF-1), which is a heterodimer composed of an alpha and a beta subunit. HIF-1 functions as a master regulator of cellular and systemic homeostatic response to hypoxia by activating transcription of many genes, including those involved in apoptosis, angiogenesis, energy metabolism, and other genes whose protein products increase oxygen delivery or facilitate metabolic adaptation to hypoxia. HIF-1 thus plays an essential role in tumour angiogenesis, embryonic vascularization, and pathophysiology of ischemic disease.

Model STJ140124

Host Goat

Reactivity Avian, Bovine, Canine, Donkey, Feline, Goat, Guinea Pig, Hamster, Horse,

Human, Mouse, Other, Porcine, Rabbit, Rat, Sheep, Simian

Applications WB

Immunogen Recombinant peptide derived from within residues 470 to 590 aa of human

HIF1a produced in E. coli.

Immunogen Region 470-590 aa

Gene ID 3091

Gene Symbol HIF1A

Dilution range Western blot 1:500-1:2,000Immunofluorescence NDImmunohistochemistry

(paraffin) NDImmunohistochemistry (frozen) ND

Tissue Specificity Expressed in most tissues with highest levels in kidney and heart.

Overexpressed in the majority of common human cancers and their

metastases, due to the presence of intratumoral hypoxia and as a result of mutations in genes encoding oncoproteins and tumor suppressors. A higher level expression seen in pituitary tumors as compared to the pituitary gland.

Purification This antibody is epitope-affinity purified from goat antiserum.

Note For research use only (RUO).

Protein Name Hypoxia-inducible factor 1-alpha (HIF-1-alpha) (HIF1-alpha) (ARNT-

interacting protein) (Basic-helix-loop-helix-PAS protein MOP1) (Class E basic helix-loop-helix protein 78) (bHLHe78) (Member of PAS protein 1)

(PAS domain-containing protein 8)

Molecular Weight 93 kDa

Clonality Polyclonal

Conjugation Unconjugated

Isotype IgG

Formulation PBS, 20% glycerol and 0.05% sodium azide.

Concentration 3 mg/mL

Storage Instruction Store at -20°, and avoid repeated freeze-thaw cycles.

Database Links HGNC:49100MIM:603348

Alternative Names Hypoxia-inducible factor 1-alpha (HIF-1-alpha) (HIF1-alpha) (ARNT-

interacting protein) (Basic-helix-loop-helix-PAS protein MOP1) (Class E basic helix-loop-helix protein 78) (bHLHe78) (Member of PAS protein 1)

(PAS domain-containing protein 8)

Function Functions as a master transcriptional regulator of the adaptive response to

hypoxia. Under hypoxic conditions, activates the transcription of over 40 genes, including erythropoietin, glucose transporters, glycolytic enzymes, vascular endothelial growth factor, HILPDA, and other genes whose protein products increase oxygen delivery or facilitate metabolic adaptation to hypoxia. Plays an essential role in embryonic vascularization, tumor angiogenesis and pathophysiology of ischemic disease. Binds to core DNA

hypoxia. Plays an essential role in embryonic vascularization, tumor angiogenesis and pathophysiology of ischemic disease. Binds to core DNA sequence 5'-[AG]CGTG-3' within the hypoxia response element (HRE) of target gene promoters. Activation requires recruitment of transcriptional coactivators such as CREBBP and EP300. Activity is enhanced by interaction with both, NCOA1 or NCOA2. Interaction with redox regulatory protein APEX seems to activate CTAD and potentiates activation by NCOA1 and

CREBBP. Involved in the axonal distribution and transport of mitochondria in

neurons during hypoxia.

Sequence and Domain Family Contains two independent C-terminal transactivation domains, NTAD and

CTAD, which function synergistically. Their transcriptional activity is

repressed by an intervening inhibitory domain (ID).

Cellular Localization Cytoplasm Nucleus Speckle. Colocalizes with HIF3A in the nucleus

and speckles. Cytoplasmic in normoxia, nuclear translocation in response to

hypoxia.

Post-translational In normoxia, is hydroxylated on Pro-402 and Pro-564 in the oxygen-

Modifications dependent degradation domain (ODD) by EGLN1/PHD2 and EGLN2/PHD1.

EGLN3/PHD3 has also been shown to hydroxylate Pro-564. The hydroxylated

prolines promote interaction with VHL, initiating rapid ubiquitination and subsequent proteasomal degradation. Deubiquitinated by USP20. Under hypoxia, proline hydroxylation is impaired and ubiquitination is attenuated, resulting in stabilization. In normoxia, is hydroxylated on Asn-803 by HIF1AN, thus abrogating interaction with CREBBP and EP300 and preventing transcriptional activation. This hydroxylation is inhibited by the Cu/Zn-chelator, Clioquinol. S-nitrosylation of Cys-800 may be responsible for increased recruitment of p300 coactivator necessary for transcriptional activity of HIF-1 complex. Requires phosphorylation for DNA-binding. Phosphorylation at Ser-247 by CSNK1D/CK1 represses kinase activity and impairs ARNT binding. Phosphorylation by GSK3-beta and PLK3 promote degradation by the proteasome. Sumoylated; with SUMO1 under hypoxia. Sumovlation is enhanced through interaction with RWDD3. Both sumovlation and desumoylation seem to be involved in the regulation of its stability during hypoxia. Sumoylation can promote either its stabilization or its VHLdependent degradation by promoting hydroxyproline-independent HIF1A-VHL complex binding, thus leading to HIF1A ubiquitination and proteasomal degradation. Desumoylation by SENP1 increases its stability amd transcriptional activity. There is a disaccord between various publications on the effect of sumoylation and desumoylation on its stability and transcriptional activity. Acetylation of Lys-532 by ARD1 increases interaction with VHL and stimulates subsequent proteasomal degradation. Deacetylation of Lys-709 by SIRT2 increases its interaction with and hydroxylation by EGLN1 thereby inactivating HIF1A activity by inducing its proteasomal degradation. Polyubiquitinated; in normoxia, following hydroxylation and interaction with VHL. Lys-532 appears to be the principal site of ubiquitination. Clioquinol, the Cu/Zn-chelator, inhibits ubiquitination through preventing hydroxylation at Asn-803. Ubiquitinated by a CUL2-based E3 ligase. The iron and 2oxoglutarate dependent 3-hydroxylation of asparagine is (S) stereospecific within HIF CTAD domains.

St John's Laboratory Ltd

F +44 (0)207 681 2580

T +44 (0)208 223 3081

W http://www.stjohnslabs.com/ E info@stjohnslabs.com