

Phospho-p38 MAPK (Y182) antibody

Purified Rabbit Polyclonal Antibody (Pab) Catalog # AP3905a

Product Information

Application	WB, E
Primary Accession	<u>Q15759</u>
Other Accession	<u>Q9WUI1</u>
Reactivity	Human, Mouse
Predicted	Mouse
Host	Rabbit
Clonality	polyclonal
Isotype	Rabbit IgG
Clone Names	RB43190
Calculated MW	41357

Additional Information

Gene ID	5600
Other Names	Mitogen-activated protein kinase 11, MAP kinase 11, MAPK 11, 2.7.11.24, Mitogen-activated protein kinase p38 beta, MAP kinase p38 beta, p38b, Stress-activated protein kinase 2b, SAPK2b, p38-2, MAPK11, PRKM11, SAPK2, SAPK2B
Target/Specificity	This antibody is generated from a rabbit immunized with a KLH conjugated synthetic peptide between 155-186 amino acids from human.
Dilution	WB~~1:500 E~~Use at an assay dependent concentration.
Format	Purified polyclonal antibody supplied in PBS with 0.09% (W/V) sodium azide. This antibody is purified through a protein A column, followed by peptide affinity purification.
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	Phospho-p38 MAPK (Y182) antibody is for research use only and not for use in diagnostic or therapeutic procedures.

Protein Information

Name	MAPK11
Synonyms	PRKM11, SAPK2, SAPK2B

Serine/threonine kinase which acts as an essential component of the MAP kinase signal transduction pathway (PubMed:<u>12452429</u>, PubMed:<u>20626350</u>, PubMed:<u>35857590</u>). MAPK11 is one of the four p38 MAPKs which play an important role in the cascades of cellular responses evoked by extracellular stimuli such as pro-inflammatory cytokines or physical stress leading to direct activation of transcription factors (PubMed:12452429, PubMed:20626350, PubMed:<u>35857590</u>). Accordingly, p38 MAPKs phosphorylate a broad range of proteins and it has been estimated that they may have approximately 200 to 300 substrates each (PubMed:12452429, PubMed:20626350, PubMed:<u>35857590</u>). MAPK11 functions are mostly redundant with those of MAPK14 (PubMed:<u>12452429</u>, PubMed:<u>20626350</u>, PubMed:<u>35857590</u>). Some of the targets are downstream kinases which are activated through phosphorylation and further phosphorylate additional targets (PubMed:12452429, PubMed:20626350). RPS6KA5/MSK1 and RPS6KA4/MSK2 can directly phosphorylate and activate transcription factors such as CREB1, ATF1, the NF-kappa-B isoform RELA/NFKB3, STAT1 and STAT3, but can also phosphorylate histone H3 and the nucleosomal protein HMGN1 (PubMed: 9687510). RPS6KA5/MSK1 and RPS6KA4/MSK2 play important roles in the rapid induction of immediate- early genes in response to stress or mitogenic stimuli, either by inducing chromatin remodeling or by recruiting the transcription machinery. On the other hand, two other kinase targets, MAPKAPK2/MK2 and MAPKAPK3/MK3, participate in the control of gene expression mostly at the post-transcriptional level, by phosphorylating ZFP36 (tristetraprolin) and ELAVL1, and by regulating EEF2K, which is important for the elongation of mRNA during translation. MKNK1/MNK1 and MKNK2/MNK2, two other kinases activated by p38 MAPKs, regulate protein synthesis by phosphorylating the initiation factor EIF4E2 (PubMed:<u>11154262</u>). In the cytoplasm, the p38 MAPK pathway is an important regulator of protein turnover. For example, CFLAR is an inhibitor of TNF-induced apoptosis whose proteasome-mediated degradation is regulated by p38 MAPK phosphorylation. Ectodomain shedding of transmembrane proteins is regulated by p38 MAPKs as well. In response to inflammatory stimuli, p38 MAPKs phosphorylate the membrane-associated metalloprotease ADAM17. Such phosphorylation is required for ADAM17-mediated ectodomain shedding of TGF-alpha family ligands, which results in the activation of EGFR signaling and cell proliferation. Additional examples of p38 MAPK substrates are the FGFR1. FGFR1 can be translocated from the extracellular space into the cytosol and nucleus of target cells, and regulates processes such as rRNA synthesis and cell growth. FGFR1 translocation requires p38 MAPK activation. In the nucleus, many transcription factors are phosphorylated and activated by p38 MAPKs in response to different stimuli. Classical examples include ATF1, ATF2, ATF6, ELK1, PTPRH, DDIT3, TP53/p53 and MEF2C and MEF2A (PubMed:<u>10330143</u>, PubMed:<u>15356147</u>, PubMed:<u>9430721</u>). The p38 MAPKs are emerging as important modulators of gene expression by regulating chromatin modifiers and remodelers (PubMed: 10330143, PubMed: 15356147, PubMed: 9430721). The promoters of several genes involved in the inflammatory response, such as IL6, IL8 and IL12B, display a p38 MAPK-dependent enrichment of histone H3 phosphorylation on 'Ser-10' (H3S10ph) in LPS-stimulated myeloid cells. This phosphorylation enhances the accessibility of the cryptic NF- kappa-B-binding sites marking promoters for increased NF-kappa-B recruitment. Phosphorylates NLRP1 downstream of MAP3K20/ZAK in response to UV-B irradiation and ribosome collisions, promoting activation of the NLRP1 inflammasome and pyroptosis (PubMed:<u>35857590</u>). Phosphorylates methyltransferase DOT1L on 'Ser-834', 'Thr-900', 'Ser-902', 'Thr-984', 'Ser-1001', 'Ser-1009' and 'Ser-1104' (PubMed:38270553).

Cellular Location

Cytoplasm. Nucleus.

Tissue Location

Highest levels in the brain and heart. Also expressed in the placenta, lung,

Background

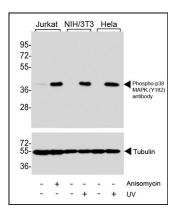
Serine/threonine kinase which acts as an essential component of the MAP kinase signal transduction pathway. MAPK11 is one of the four p38 MAPKs which play an important role in the cascades of cellular responses evoked by extracellular stimuli such as proinflammatory cytokines or physical stress leading to direct activation of transcription factors. Accordingly, p38 MAPKs phosphorylate a broad range of proteins and it has been estimated that they may have approximately 200 to 300 substrates each. MAPK11 functions are mostly redundant with those of MAPK14. Some of the targets are downstream kinases which are activated through phosphorylation and further phosphorylate additional targets. RPS6KA5/MSK1 and RPS6KA4/MSK2 can directly phosphorylate and activate transcription factors such as CREB1, ATF1, the NF-kappa-B isoform RELA/NFKB3, STAT1 and STAT3, but can also phosphorylate histone H3 and the nucleosomal protein HMGN1. RPS6KA5/MSK1 and RPS6KA4/MSK2 play important roles in the rapid induction of immediate-early genes in response to stress or mitogenic stimuli, either by inducing chromatin remodeling or by recruiting the transcription machinery. On the other hand, two other kinase targets, MAPKAPK2/MK2 and MAPKAPK3/MK3, participate in the control of gene expression mostly at the post-transcriptional level, by phosphorylating ZFP36 (tristetraprolin) and ELAVL1, and by regulating EEF2K, which is important for the elongation of mRNA during translation. MKNK1/MNK1 and MKNK2/MNK2, two other kinases activated by p38 MAPKs, regulate protein synthesis by phosphorylating the initiation factor EIF4E2. In the cytoplasm, the p38 MAPK pathway is an important regulator of protein turnover. For example, CFLAR is an inhibitor of TNF-induced apoptosis whose proteasome-mediated degradation is regulated by p38 MAPK phosphorylation. Ectodomain shedding of transmembrane proteins is regulated by p38 MAPKs as well. In response to inflammatory stimuli, p38 MAPKs phosphorylate the membrane- associated metalloprotease ADAM17. Such phosphorylation is required for ADAM17-mediated ectodomain shedding of TGF-alpha family ligands, which results in the activation of EGFR signaling and cell proliferation. Additional examples of p38 MAPK substrates are the FGFR1. FGFR1 can be translocated from the extracellular space into the cytosol and nucleus of target cells, and regulates processes such as rRNA synthesis and cell growth. FGFR1 translocation requires p38 MAPK activation. In the nucleus, many transcription factors are phosphorylated and activated by p38 MAPKs in response to different stimuli. Classical examples include ATF1, ATF2, ATF6, ELK1, PTPRH, DDIT3, TP53/p53 and MEF2C and MEF2A. The p38 MAPKs are emerging as important modulators of gene expression by regulating chromatin modifiers and remodelers. The promoters of several genes involved in the inflammatory response, such as IL6, IL8 and IL12B, display a p38 MAPK-dependent enrichment of histone H3 phosphorylation on 'Ser-10' (H3S10ph) in LPS-stimulated myeloid cells. This phosphorylation enhances the accessibility of the cryptic NF-kappa-B-binding sites marking promoters for increased NF-kappa-B recruitment.

References

Jiang Y.,et al.J. Biol. Chem. 271:17920-17926(1996). Jiang Y.,et al.Submitted (APR-1997) to the EMBL/GenBank/DDBJ databases. Kumar S.,et al.Biochem. Biophys. Res. Commun. 235:533-538(1997). Enslen H.,et al.J. Biol. Chem. 273:1741-1748(1998). Goedert M.,et al.EMBO J. 16:3563-3571(1997).

Images

Western blot analysis of extracts from Jurkat cells, untreated or treated with anisomycin (25 µg/ml), NIH/3T3 and Hela cells, untreated or treated with UV (30 minutes), using Phospho-p38 MAPK (Y182) antibody (upper) or Tubulin (lower).



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