

# Phospho-GSK3 (Tyr216) polyconal antibody

Rabbit Polyclonal Antibody Catalog # ABV11747

#### **Product Information**

Application WB, E
Primary Accession P49841
Reactivity Human
Host Rabbit
Clonality Polyclonal
Isotype Rabbit IgG
Calculated MW 46744

### **Additional Information**

**Gene ID** 2932

Application & Usage Western blot, Immunoblot: 0.5-2 [g/ml, ELISA

Alias Symbol GSK3B

Other Names GSK-3 beta, GSK3B

**Appearance** Colorless liquid

Formulation 100 ug (1mg/ml) of antibody in 0.01M Tris-HCl, pH 8.0, 0.15M NaCl, and 0.02%

sodium azide.

Reconstitution & Storage -20 °C

**Background Descriptions** 

**Precautions** Phospho-GSK3 (Tyr216) polyconal antibody is for research use only and not

for use in diagnostic or therapeutic procedures.

#### **Protein Information**

Name GSK3B ( HGNC:4617)

**Function** Constitutively active protein kinase that acts as a negative regulator in the

hormonal control of glucose homeostasis, Wnt signaling and regulation of transcription factors and microtubules, by phosphorylating and inactivating glycogen synthase (GYS1 or GYS2), EIF2B, CTNNB1/beta-catenin, APC, AXIN1,

DPYSL2/CRMP2, JUN, NFATC1/NFATC, MAPT/TAU and MACF1 (PubMed: 11430833, PubMed: 12554650, PubMed: 14690523,

PubMed: 16484495, PubMed: 1846781, PubMed: 20937854, PubMed: 9072970).

Requires primed phosphorylation of the majority of its substrates

(PubMed: 11430833, PubMed: 16484495). In skeletal muscle, contributes to insulin regulation of glycogen synthesis by phosphorylating and inhibiting

GYS1 activity and hence glycogen synthesis (PubMed:<u>8397507</u>). May also mediate the development of insulin resistance by regulating activation of transcription factors (PubMed:<u>8397507</u>). Regulates protein synthesis by controlling the activity of initiation factor 2B (EIF2BE/EIF2B5) in the same manner as glycogen synthase (PubMed:8397507). In Wnt signaling, GSK3B forms a multimeric complex with APC, AXIN1 and CTNNB1/beta-catenin and phosphorylates the N-terminus of CTNNB1 leading to its degradation mediated by ubiquitin/proteasomes (PubMed:12554650). Phosphorylates JUN at sites proximal to its DNA-binding domain, thereby reducing its affinity for DNA (PubMed: 1846781). Phosphorylates NFATC1/NFATC on conserved serine residues promoting NFATC1/NFATC nuclear export, shutting off NFATC1/NFATC gene regulation, and thereby opposing the action of calcineurin (PubMed: 9072970). Phosphorylates MAPT/TAU on 'Thr-548', decreasing significantly MAPT/TAU ability to bind and stabilize microtubules (PubMed:14690523). MAPT/TAU is the principal component of neurofibrillary tangles in Alzheimer disease (PubMed: 14690523). Plays an important role in ERBB2-dependent stabilization of microtubules at the cell cortex (PubMed: 20937854). Phosphorylates MACF1, inhibiting its binding to microtubules which is critical for its role in bulge stem cell migration and skin wound repair (By similarity). Probably regulates NF-kappa-B (NFKB1) at the transcriptional level and is required for the NF-kappa-B-mediated antiapoptotic response to TNF-alpha (TNF/TNFA) (By similarity). Negatively regulates replication in pancreatic beta-cells, resulting in apoptosis, loss of beta-cells and diabetes (By similarity). Through phosphorylation of the anti-apoptotic protein MCL1, may control cell apoptosis in response to growth factors deprivation (By similarity). Phosphorylates MUC1 in breast cancer cells, decreasing the interaction of MUC1 with CTNNB1/beta-catenin (PubMed: 9819408). Is necessary for the establishment of neuronal polarity and axon outgrowth (PubMed: 20067585). Phosphorylates MARK2, leading to inhibition of its activity (By similarity). Phosphorylates SIK1 at 'Thr-182', leading to sustainment of its activity (PubMed: 18348280). Phosphorylates ZC3HAV1 which enhances its antiviral activity (PubMed: 22514281). Phosphorylates SNAI1, leading to its ubiquitination and proteasomal degradation (PubMed: 15448698, PubMed: 15647282, PubMed: 25827072, PubMed: 29059170). Phosphorylates SFPQ at 'Thr-687' upon T-cell activation (PubMed:20932480). Phosphorylates NR1D1 st 'Ser-55' and 'Ser-59' and stabilizes it by protecting it from proteasomal degradation. Regulates the circadian clock via phosphorylation of the major clock components including BMAL1, CLOCK and PER2 (PubMed: 19946213, PubMed: 28903391). Phosphorylates FBXL2 at 'Thr-404' and primes it for ubiquitination by the SCF(FBXO3) complex and proteasomal degradation (By similarity). Phosphorylates CLOCK AT 'Ser-427' and targets it for proteasomal degradation (PubMed: 19946213). Phosphorylates BMAL1 at 'Ser-17' and 'Ser-21' and primes it for ubiquitination and proteasomal degradation (PubMed: <u>28903391</u>). Phosphorylates OGT at 'Ser-3' or 'Ser-4' which positively regulates its activity. Phosphorylates MYCN in neuroblastoma cells which may promote its degradation (PubMed:24391509). Regulates the circadian rhythmicity of hippocampal long-term potentiation and BMAL1 and PER2 expression (By similarity). Acts as a regulator of autophagy by mediating phosphorylation of KAT5/TIP60 under starvation conditions, activating KAT5/TIP60 acetyltransferase activity and promoting acetylation of key autophagy regulators, such as ULK1 and RUBCNL/Pacer (PubMed:30704899). Negatively regulates extrinsic apoptotic signaling pathway via death domain receptors. Promotes the formation of an anti-apoptotic complex, made of DDX3X, BRIC2 and GSK3B, at death receptors, including TNFRSF10B. The anti-apoptotic function is most effective with weak apoptotic signals and can be overcome by stronger stimulation (PubMed:18846110). Phosphorylates E2F1, promoting the interaction between E2F1 and USP11, stabilizing E2F1 and promoting its activity (PubMed: 17050006, PubMed: 28992046). Phosphorylates mTORC2 complex component RICTOR at 'Ser-1235' in

response to endoplasmic stress, inhibiting mTORC2 (PubMed:21343617). Phosphorylates mTORC2 complex component RICTOR at 'Thr-1695' which facilitates FBXW7-mediated ubiquitination and subsequent degradation of RICTOR (PubMed:25897075). Phosphorylates FXR1, promoting FXR1 ubiquitination by the SCF(FBXO4) complex and FXR1 degradation by the proteasome (By similarity). Phosphorylates interleukin-22 receptor subunit IL22RA1, preventing its proteasomal degradation (By similarity).

**Cellular Location** Cytoplasm. Nucleus. Cell membrane. Note=The phosphorylated form shows

localization to cytoplasm and cell membrane (PubMed:20937854) The MEMO1-RHOA-DIAPH1 signaling pathway controls localization of the phosphorylated form to the cell membrane (PubMed:20937854)

**Tissue Location** Expressed in testis, thymus, prostate and ovary and weakly expressed in lung,

brain and kidney. Colocalizes with EIF2AK2/PKR and TAU in the Alzheimer

disease (AD) brain

## **Background**

Glycogen synthase kinase 3 (GSK-3) is a serine/threonine protein kinase that has been implicated in the regulation of cell fate and in the Wnt signaling pathway. GSK-3 plays an important role in the PI3 kinase and Akt mediated cell survival pathways, and its activity can be inhibited by Akt-mediated phosphorylation at Ser21 of GSK-3 $\alpha$  and Ser9 of GSK-3 $\beta$ . GSK-3 has also been implicated in alzheimer's disease. Six Tau protein isoforms have been identified, all of which are phosphorylated by GSK-3.

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