

## Phospho-CAD (Ser1859) Antibody



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Applications: W	<b>Reactivity:</b> H M R Mk	<b>Sensitivity:</b> Endogenous	<b>MW (kDa):</b> 240	Source/Isotype: Rabbit	UniProt ID: #P27708	Entrez-Gene Id: 790	
Product Usage Information		<b>Application</b> Western Blotting			<b>Dilution</b> 1:1000		
Storage		Supplied in 10 mM sodium HEPES (pH 7.5), 150 mM NaCl, 100 $\mu$ g/ml BSA and 50% glycerol. Store at – 20°C. Do not aliquot the antibody.					
Specificity/Sensitivity		Phospho-CAD (Ser1859) Antibody recognizes endogenous levels of CAD protein only when phosphorylated at Ser1859.					
Source / Purification		Polyclonal antibodies are produced by immunizing animals with a synthetic phosphopeptide corresponding to residues surrounding Ser1859 of human CAD protein. Antibodies are purified by protein A and peptide affinity chromatography.					
Background		CAD is essential for the <i>de novo</i> synthesis of pyrimidine nucleotides and possesses the following enzymatic activities: glutamine amidotransferase, carbamoyl-phosphate synthetase, aspartate transcarbamoylase, and dihydroorotase. Thus, the enzyme converts glutamine to uridine monophosphate, a common precursor of all pyrimidine bases, and it is necessary for nucleic acid synthesis (1). In resting cells, CAD is localized mainly in the cytoplasm where it carries out pyrimidine synthesis. As proliferating cells enter S phase, MAP Kinase (Erk1/2) phosphorlyates CAD at Thr456, resulting in CAD translocation to the nucleus. As cells exit S phase, CAD is dephosphorylated at Thr456 and phosphorylated at Ser1406 by PKA, returning the pathway to basal activity (2). Various research studies have shown increased expression of CAD in several types of cancer, prompting the development of pharmacological inhibitors such as PALA. Further studies have identified CAD as a potential predictive early marker of prostate cancer relapse (3).  mTORC1 is a protein kinase that works to regulate the growth and proliferation of cells by sensing and integrating various growth signals. S6 kinase 1 (S6K1) is a downstream ribosomal protein target of mTORC1 and directly phosphorylates Ser1859 on CAD. This phosphorylation stimulates the first three steps of the <i>de novo</i> primidine synthesis and thus helps to advance the cells overall progression through S phase of the cell cycle (4,5).					
Background References		2. Sigoillot, F.D. et al. 3. Morin, A. et al. (201 4. Ben-Sahra, I. et al.	man, P.F. et al. (1977) <i>J Biol Chem</i> 252, 6379-85. illot, F.D. et al. (2005) <i>J Biol Chem</i> 280, 25611-20. in, A. et al. (2012) <i>FASEB J</i> 26, 460-7. ·Sahra, I. et al. (2013) <i>Science</i> 339, 1323-8. taille, A.M. et al. (2013) <i>Science</i> 339, 1320-3.				
Species Reacti	vity	Species reactivity is d	etermined by testin	g in at least one approve	ed application (e.g.,	western blot).	

**Western Blot Buffer** 

IMPORTANT: For western blots, incubate membrane with diluted primary antibody in 5% w/v nonfat dry milk, 1X TBS, 0.1% Tween® 20 at 4°C with gentle shaking, overnight.

**Applications Key** W: Western Blotting

**Cross-Reactivity Key** H: Human M: Mouse R: Rat Mk: Monkey

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