Revision 1

SB203580	Cell Signaling
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For Research Use Only. Not for Use in Diagnostic Procedures.

Background	p38 MAP kinase (MAPK), also called RK (1) or CSBP (2), is the mammalian orthologue of the yeast HOG kinase that participates in a signaling cascade controlling cellular responses to cytokines and stress (1-4). Four isoforms of p38 MAPK, p38α, β , γ (also known as Erk6 or SAPK3), and δ (also known as SAPK4) have been identified. Similar to the SAPK/JNK pathway, p38 MAPK is activated by a variety of cellular stresses, including osmotic shock, inflammatory cytokines, lipopolysaccharide (LPS), UV light, and growth factors (1-5). MKK3, MKK6, and SEK activate p38 MAPK by phosphorylation at Thr180 and Tyr182. Activated p38 MAPK has been shown to phosphorylate and activate MAPKAP kinase 2 (3) and to phosphorylate the transcription factors ATF-2 (5), Max (6), and MEF2 (5-8). SB203580 (4-(4-fluorophenyl)-2-(4-methylsulfinylphenyl)-5-(4-pyridyl)-imidazole) is a selective inhibitor of p38 MAPK. This compound inhibits the activation of MAPKAPK-2 by p38 MAPK and subsequent phosphorylation of HSP27 (9). SB203580 inhibits p38 MAPK catalytic activity by binding to the ATP-binding pocket, but does not inhibit phosphorylation of p38 MAPK by upstream kinases (10).
Molecular Formula	C ₂₁ H ₁₆ FN ₃ OS
Molecular Weight	377.4 g/mol
Purity	>98%
CAS	152121-47-6
Solubility	Soluble in DMSO at 100mg/ml and EtOH at 10mg/ml.
Storage	Store lyophilized or in solution at -20°C, desiccated. Protect from light. In lyophilized form, the chemical is stable for 24 months. Once in solution, use within 3 months to prevent loss of potency. Aliquot to avoid multiple freeze/thaw cycles.
Directions for Use	SB203580 is supplied as a lyophilized white powder. For 10 mM stock, resuspend 5 mg of the inhibitor in 1.32 ml DMSO. Aliquot and freeze at -20°C to avoid multiple freeze/thaw cycles which can degrade the inhibitor. For experiments with cultured cells, we recommend pre-treating the cells with SB203580 at 10 μ M for one to two hours prior to stimulation. This product is for <i>in vitro</i> research use only and is not intended for use in humans or animals.
Background References	 Rouse, J. et al. (1994) <i>Cell</i> 78, 1027-37. Han, J. et al. (1994) <i>Science</i> 265, 808-11. Lee, J.C. et al. (1994) <i>Nature</i> 372, 739-46. Freshney, N.W. et al. (1994) <i>Cell</i> 78, 1039-49. Raingeaud, J. et al. (1995) <i>J Biol Chem</i> 270, 7420-6. Zervos, A.S. et al. (1995) <i>Proc Natl Acad Sci U S A</i> 92, 10531-4. Zhao, M. et al. (1999) <i>Mol Cell Biol</i> 19, 21-30. Yang, S.H. et al. (1995) <i>FEBS Lett</i> 364, 229-33. Kumar, S. et al. (1999) <i>Biochem Biophys Res Commun</i> 263, 825-31.
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