



## This week in therapeutics

Indication	Target/marker/pathway	Summary	Licensing status	Publication and contact information
Neurology				
Alzheimer's disease (AD)	Glycogen synthase kinase 3β (GSK3B)	Cell culture studies identified pyrazine-based inhibitors of GSK3B that could help treat neurodegenerative diseases such as AD. <i>In vitro</i> , a lead compound of the series inhibited GSK3B-mediated microtubule-associated protein-τ (MAPT; TAU; FTDP-17) phosphorylation with nanomolar IC <sub>50</sub> values, showed properties associated with good brain penetration and had good target selectivity across a panel of 26 kinases. Researchers did not disclose next steps, which could include evaluating the lead inhibitor in animal models of AD.  AstraZeneca plc's lead pyrazine analog GSK3B inhibitor is in preclinical development.  Tideglusib, a GSK3 inhibitor from Noscira S.A., is in Phase II testing to treat AD.  Neu-120, a selective uncompetitive NMDAR modulator and monoamine oxidase B (MAO-B) and GSK3B inhibitor from Neurim Pharmaceuticals Ltd., is in Phase II testing for Parkinson's disease (PD).  DiaMedica Inc.'s DM-99, an undisclosed naturally occurring protein that inhibits GSK3B, is in preclinical development for AD.		Berg, S. et al. J. Med. Chem.; published online April 10, 2012; doi:10.1021/jm201724m Contact: Stefan Berg, AstraZeneca R&D, Soedertaelje, Sweden e-mail: stefan.berg@astrazeneca.com
		SciBX 5(18); doi:10.1038/scibx.2012.472 Published online May 3, 2012		