

## THE DISTILLERY

## This week in therapeutics

Indication	Target/marker/ pathway	Summary	Licensing status	Publication and contact information
Neurology				
Alzheimer's disease (AD)	Phospholipase A <sub>2</sub> , group IVA (cytosolic, calcium- dependent) (PLA <sub>2</sub> G4A)	Mouse and cultured neuron studies suggest that PLA <sub>2</sub> G4A could be targeted to treat AD. PLA <sub>2</sub> G4A cleaves phospholipids to produce arachidonic acid and other bioactive lipids. Mass spectroscopic analysis of hippocampus tissue from mice overexpressing human $\beta$ -amyloid (A4) precursor protein (APP) revealed higher levels of arachidonic acid than those seen in wild-type controls. Genetic disruption of PLA <sub>2</sub> G4A in APP-overexpressing mice improved learning and memory compared with what was seen in wild- type controls. Pretreatment of cultured neurons with a phospholipase A <sub>2</sub> (PLA <sub>2</sub> ) inhibitor lowered cell death caused by amyloid- $\beta$ peptide compared with what was seen in cultured neurons that did not receive a PLA <sub>2</sub> inhibitor. Amarin Corp. plc's AMR101 PLA <sub>2</sub> inhibitor is in Phase I, II and III trials to treat a range of metabolic, cardiovascular and neurological indications.	Unpatented; licensing status not applicable	Sanchez-Mejia, R.O. <i>et al. Nat.</i> <i>Neurosci.</i> ; published online Oct. 19, 2008; doi:10.1038/nn.2213 <b>Contact:</b> Lennart Mucke, Gladstone Institute of Neurological Disease, San Francisco, Calif. e-mail: Imucke@gladstone.ucsf.edu <b>Contact:</b> Rene O. Sanchez-Mejia, Massachusetts General Hospital, Boston, Mass. e-mail: rene_sanchez@post.harvard.edu

Morria Biopharmaceuticals plc's MRX-4  ${\rm PLA}_2$  inhibitor is in Phase II trials to treat allergy.

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