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IN BRIEF

ALZHEIMER DISEASE

Mutation in APP protects against Alzheimer disease

In a whole-genome sequencing study of 1,795 Icelandic people, researchers have identified a coding variant in amyloid- β (A β) precursor protein (APP) that protects against Alzheimer disease (AD) and cognitive decline. The A673T substitution occurs close to the target site of BACE1, the APP-cleaving enzyme that generates toxic A β fragments. The mutation leads to reduced production of A β by BACE1, and provides proof of principle for the hypothesis that blocking BACE1 cleavage of APP protects against AD.

Original article Jonsson, T. et al. A mutation in APP protects against Alzheimer's disease and age-related cognitive decline. Nature doi:10.1038/nature11283

EPILEPSY

A role for GABA receptor signalling in granule cell ectopia and pathogenesis of adult epilepsy

As temporal lobe epilepsy (TLE) is associated with abnormal localization of hippocampal granule cells, Koyama et al. investigated the mechanisms underlying granule cell ectopia. Using a rat model of complex febrile seizures, they found upregulation of γ -aminobutyric acid (GABA) receptors in granule cells, which caused a reversal of migration by these cells. This abnormality was avoided by inhibition of a cotransporter that regulates GABA activation, indicating a potential therapeutic approach.

Original article Koyama, R. et al. GABAergic excitation after febrile seizures induces ectopic granule cells and adult epilepsy. *Nat. Med.* doi:10.1038/nm.2850

SPINAL CORD INJURY

Fluoxetine shows promise as a neuroprotective therapy following spinal cord injury

Fluoxetine, an antidepressant drug, has shown promise as a therapy in spinal cord injury (SCI), but the mechanism underlying this neuroprotection was unknown. Using a mouse contusion-injury model, Lee *et al.* show that fluoxetine prevents breakdown of the blood–spinal cord barrier by inhibiting activation of matrix metalloproteinases and maintaining tight-junction integrity. The study highlights fluoxetine as a promising therapy for brain injury and SCI.

Original article Lee, J. Y. et al. Fluoxetine inhibits matrix metalloproteinase activation and prevents disruption of blood-spinal cord barrier after spinal cord injury. Brain doi:10.1093/brain/aws171

CEREBROVASCULAR DISEASE

Amyloid imaging enables prediction of haemorrhage sites in cerebral amyloid angiopathy

In a recent study, published in *Neurology*, Edip Gurol *et al.* sought to determine whether MRI combined with Pittsburgh compound B (PiB)-PET could be used to predict the locations of future haemorrhages in patients with cerebral amyloid angiopathy (CAA). In 11 patients who underwent baseline and follow-up MRI with PiB-PET, CAA-related haemorrhages tended to occur at sites with greatest amyloid deposition. The researchers suggest PiB-PET could be a useful tool for prediction of haemorrhages in CAA.

Original article Edip Gurol, E. et al. Predicting sites of new haemorrhage with amyloid imaging in cerebral amyloid angiopathy. *Neurology* doi:10.1212/ WNL.0b013e31826043a9