RESEARCH HIGHLIGHTS

Nature Reviews Neuroscience | AOP, published online 16 July 2014;

IN BRIEF

NEUROGENESIS

Adult neurogenesis gets a boost

The orphan nuclear receptor TLX promotes neural stem cell self-renewal, but its role in hippocampal neurogenesis is not known. Murai *et al.* generated transgenic (Tg) mice in which TLX expression was placed under the control of the promoter for nestin, a marker of neural precursors. Adult Tg mice exhibited increased numbers of proliferating neural progenitors and 5-bromodeoxyuridine-labelled newborn neurons in the dentate gyrus compared with wild-type mice, and performed better in the Morris water maze, a test of spatial learning and memory. Thus, TLX expression in neural progenitors cells is important for adult hippocampal neurogenesis and memory.

ORIGINAL RESEARCH PAPER Murai, K. *et al.* Nuclear receptor TLX stimulates hippocampal neurogenesis and enhances learning and memory in a transgenic mouse model. *Proc. Natl Acad. Sci. USA* **111**, 9115–9120 (2014)

NEUROTRANSMITTERS

Dopamine tone depends on DAT

The extracellular concentration of dopamine $([DA]_e)$ exhibits circadian oscillations, although the mechanisms underlying this phenomenon are unclear. Through the use of voltammetry in rat brain slices, Ferris *et al.* showed that, over a 24-hour cycle, the rate of DA reuptake by the DA transporter (DAT) was inversely related to $[DA]_e$. Slices from *Dat*-knockout mice showed no oscillation in $[DA]_e$, indicating that diurnal variation in $[DA]_e$ is governed by the DAT.

ORIGINAL RESEARCH PAPER Ferris, M. J. et al. Dopamine transporters govern diurnal variation in extracellular dopamine tone. Proc. Natl Acad. Sci. USA 111, E2751–E2759 (2014)

NEURODEVELOPMENTAL DISORDERS

Mind the SYNGAP

Haploinsufficiency of SYNGAP1 (synaptic RAS GTPase activating protein 1) in humans can lead to intellectual disability and epilepsy, and $Syngap1^{+/-}$ mice exhibit impaired cognition. Here, in mice, heterozygous knockout of Syngap1 in developing forebrain pyramidal neurons, but not in GABAergic neurons, was sufficient to replicate the $Syngap1^{+/-}$ mouse phenotype. Reduction of SYNGAP1 levels in adult mice had no effect on cognition or pyramidal neuron excitability, indicating that the cognitive impairment in $Syngap1^{+/-}$ mice is due to altered development of forebrain excitatory neurons.

ORIGINAL RESEARCH PAPER Ozkan, E. D. *et al.* Reduced cognition in *Syngap1* mutants is caused by isolated damage within developing forebrain excitatory neurons. *Neuron* **82**, 1317–1333 (2014)

PSYCHIATRIC DISORDERS

miR-135: a marker of mood

Serotonergic activity is dysregulated in depression and anxiety disorders, but less is known about the role of microRNAs (miRNAs) in mood disorders. miR-135, which inhibits expression of the serotonin transporter and inhibitory serotonin autoreceptor, was upregulated in serotonin neurons of animals that were treated with antidepressants. Mice that overexpressed miR-135 showed less anxiety- and depression-like behaviour after social defeat than did controls. Moreover, patients with depression exhibited lower circulating levels of miR-135. This study indicates that circulating miR-135 could be a marker of serotonin dysregulation or of response to antidepressant therapy.

ORIGINAL RESEARCH PAPER lssler, O. et al. MicroRNA 135 is essential for chronic stress resiliency, antidepressant efficacy, and intact serotonergic activity. Neuron <u>http://dx.doi.org/10.1016/j.neuron.2014.05.042</u> (2014)