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This talk will review the work conducted within Moodinflamm around the role of the glucocorticoid receptor in both cellular models of neurogenesis and in clinical samples of depressed patients. In our established model of 'depression in a dish', using human neuronal stem cells, we have found that the glucocorticoid receptor is the target of both glucocorticoid (stress) hormones and of antidepressants, with the ability of both to inhibit and stimulate neurogenesis in the hippocampus. In addition, reduced glucocorticoid receptor function and expression in the peripheral blood of depressed patients is a consistent finding across clinical samples as diverse as young adults, older depressed patients with coronary heart disease, and patients with cytokine-induced depression. This body of research confirms the pivotal role of the glucocorticoid receptor, and the potential clinical relevance of targeting it for therapeutic antidepressant purposes.

References:

- Anacker C, Cattaneo A, Musaelyan K, Zunszain PA, Horowitz M, Molteni R, Luoni A, Calabrese F, Tansey K, Gennarelli M, Thuret S, Price J, Uher R, Riva MA, Pariante CM. Role for the kinase SGK1 in stress, depression, and glucocorticoid effects on hippocampal neurogenesis. *Proc Natl Acad Sci U S A*. 2013 May 21;110(21):8708-13.
- Anacker C, Cattaneo A, Luoni A, Musaelyan K, Zunszain PA, Milanese E, Rybka J, Berry A, Cirulli F, Thuret S, Price J, Riva MA, Gennarelli M, Pariante CM. Glucocorticoid-related molecular signaling pathways regulating hippocampal neurogenesis. *Neuropsychopharmacology*. 2013 Apr;38(5):872-83.
- Cattaneo A, Gennarelli M, Uher R, Breen G, Farmer A, Aitchison KJ, Craig IW, Anacker C, Zunszain PA, McGuffin P, Pariante CM. Candidate genes expression profile associated with antidepressants response in the GENDEP study: differentiating between baseline 'predictors' and longitudinal 'targets'. *Neuropsychopharmacology*. 2013 Feb;38(3):377-85.
- Zunszain PA, Anacker C, Cattaneo A, Choudhury S, Musaelyan K, Myint AM, Thuret S, Price J, Pariante CM. Interleukin-1 β : a new regulator of the kynurenine pathway affecting human hippocampal neurogenesis. *Neuropsychopharmacology*. 2012 Mar;37(4):939-49.
- Anacker C, Zunszain PA, Cattaneo A, Carvalho LA, Garabedian MJ, Thuret S, Price J, Pariante CM. Antidepressants increase human hippocampal neurogenesis by activating the glucocorticoid receptor. *Mol Psychiatry*. 2011 Jul;16(7):738-50.