

Centre de Neurosciences Psychiatriques

CNP SEMINAR

ANNOUNCEMENT

Wednesday, June 27, 2012, 13:30

"EORS, the epigenetic oxidative redox theory of aging: How redox is more important than ROS in the aging brain"

Prof Gregory J. Brewer, PhD

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Prof Gregory J. Brewer's RO1 project on Mitoenergetic Failure in Brain Aging (RO1 AG032431) indicates a focus on redox bioenergetics and epigenetic control of metabolism. He began studies of bioenergetics in 1999 in Zurich at the ETH. In 2008, he invented a neuron culture medium called NbActiv4 that results in a doubling of synapse density and a >4-fold increase in spike rates associated with a lower resting oxygen consumption and a greater O₂ max capacity. Other publications relating to metabolism include studies of glucose uptake, improvements in brain mitochondrial function with anti-oxidants and detailed analyses of whole neuron metabolism with age-related loss of function due to substrate limitations at complex I and cytochrome C as well as enzymatic deficits at complex IV. Most significantly, these age-related deficits could be reversed by simple treatment with estrogen for a few days or nicotinamide overnight and involve epigenetic changes.

Recent publications

- 1. Boehler, M.D., Leondopulos, S.S., Wheeler, B.C., Brewer, G.J. 2012. Hippocampal networks on reliable patterned substrates. J. Neurosci. Meth. 203:344-353.
- 2. Walker, MW, LaFerla, F; Oddo, S, and Brewer, GJ. 2012. Reversible early age epigenetic histone modifications in a mouse model of Alzheimer's disease: effects on BDNF expression, AGE, DOI: 10.1007/s11357-011-9375-5 Springer doi. (Major epigenetic changes in histone acetylation and methylation occur early in aging and in 3xTg-AD mice)
- 3. Leondopulos, S.S., Boehler, M.D., Wheeler, B.C., Brewer, G.J. 2012. Chronic stimulation of cultured neural networks boosts oscillatory activity in the beta and theta frequencies, J. Neural Eng. 9 026015 doi:10.1088/1741-2560/9/2/026015. PM:22361724. (EEG-like frequencies emerge spontaneously in networks that self-assemble on electrode arrays opening access to establishing I/O dependence and the mechanistic basis of each frequency.)
- 4. Ghosh, D, LeVault, K, Barnett, A, Brewer, GJ 2012. Oxidized redox state in cultured hippocampal neurons precedes cognitive losses and macromolecular ROS damage in 3xTg-AD mice, J. Neurosci. 32:5821-5832. PM:22539844 (The redox state is shifted in the oxidizing direction with age and in an AD mouse model, but the shift is rapidly reversible and can be decoupled from ROS damage)

