UNIVERSITÀ DEGLI STUDI DI MILANO



THE DEPARTMENT OF PHARMACOLOGICAL AND BIOMOLECULAR SCIENCES

presents



Thursday, July 2nd 2015

Via Balzaretti 9, Milan

Registration is free, via email at nextstep6.register@gmail.com

Deadline for registration: June 12th 2015



UNIVERSITÀ DEGLI STUDI DI MILANO Department of Pharmacological and Biomolecular Sciences (DiSFeB) Via Balzaretti 9, Milano

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Thursday July 2nd 2015 - SCIENTIFIC PROGRAM

Registration

8:00 - 8:45 Opening Prof. Giorgio Racagni, DiSFeB Director Oral presentations, 1st session 8:45 - 9:00 9:00 - 10.45

Neuroscience Field	Metabolism Field
Chairmen: Davide Lecca, Tanmoy Samaddar	Chairmen: Alessandra Ferrari, Paola Corsetto
9.00 – 9.15 Veronica Begni BDNF modulation in response to stressful life events: role of DNA methylation and miRNAs 9.15 – 9.30 Paolo Tornese Time-dependent modulation of AMPA/NMDA receptor subunits and changes in working memory induced by acute stress 9.30 – 9.45 Luca Castelnovo GABA-A receptor subunits in Schwann cells are cross-regulated in GABA-B null mice 9.45 – 10.00 Daniele Riccio The P27 ₂ purinergic receptor: a new therapeutic target in trigeminal inflammatory pain 10.00 – 10.15 Maria Elena Cicardi The role of the Protein Quality Control system in motorneuronal and muscular models of motor neuron diseases 10.15 – 10.30 Maria Serena Paladini Genome-wide analysis of LPS-induced inflammatory response in the rat ventral hippocampus: modulatory activity of the antidepressant agomelatine 10.30 – 10.45 Giada Amodeo Human adipose-derived stromal cells and their conditioned medium induce a long lasting relief of painful symptoms in STZ-induced diabetic neuropathy	9.00 – 9.15 Matteo Audano A cytoplasmic RNA binding protein (RBP) controls skeletal muscle differentiation 9.15 – 9.30 Simone Romano Cholesterol and diabetic encephalopathy 9.30 – 9.45 Alessandra Fidone Epigenetics and metabolism: the role of histone modifications in metabolic diseases 9.45 – 10.00 Elena Signoretto Characterization of Dictyostelium discoideum Nramp1 and Nramp2 transporters expressed in Xenopusocytes 10.00 – 10.15 Federia Lolli Hepatic ERa coordinates metabolic and inflammatory response to nutritional changes in female liver 10.15 – 10.30 Chiara Macchi Glucose and lipid homeostasis during confinement in a 520-day simulated interplanetary mission to Mars 10.30 – 10.45 Chiara Ricci STAT3 inhibition induces PCSK9 in hepatic cell line: possible involvement in hypertriglyceridemia associated with insulin resistance

10:45 - 11.15 Coffee break 11.15 - 13.15 Round Table: «50 Shades of Research» <u>13.15 - 14.30</u> Lunch 14.30 - 16.00 16.00 - 16.15 16.15 - 17.30 Oral presentations, 2nd and 3rd sessions Short Break Oral presentations, 2nd and 3rd sessions

Immunology Field Chairmen: Diletta Scaccabarozzi, Silvia Franchi	Nutraceutical Field Chairmen: Chiara Di Lorenzo, Enrico Sangiovanni
14.30 – 14.45 Giulia Dell'Omo The oscillatory expression of ERα as a new drug target for resensitizing hormone- refractory breast cancer 14.45 – 15.00 Angela Papale Molecular mechanisms involved in chemical allergen-induced IL-18 production in a human keratinocyte cell line 15.00 – 15.15 Giovanna Pepe Genome-wide expression analysis of estrogen action in macrophages	14.30 – 14.45 Monica Marzagalli Vitamin E-derived delta-tocotrienol against melanoma: molecular pathways leading to cell death 14.45 – 15.00 Francesca Colombo Development of novel rapid methods to assess the antioxidant activity associated with food compounds 15.00 – 15.15 Marco Fumagalli Anti-Inflammatory activity of different varieties of raisin (Vitis vinifera L.) in human gastric epithelial cells: a comparative study
Neuroscience Field Chairmen: Silvia Giatti, Manuela Mellone	Cardiovascular Field Chairmen: Alice Ossoli, Gianpaolo Tibolla
15.15 – 15.30 Giusy Coppolino Promoting re-myelination in Multiple Sclerosis via the GPR17 receptor, a new key actor in oligodendrogenesis 15.30 – 15.45 Nathalie Sala Brain area-specific changes in glucose metabolism in rats after acute stress 15.45 – 16.00 Riccardo Cristofani Misfolded proteins toxicity in motor neuron diseases 	15.15 – 15.30 Gioria Balzarotti PI3K-C2b plays a key role in the activation and the proliferation of T lymphocytes: impact on vascular diseases 15.30 – 15.45 Eva Tarantino ASA decreased venous thrombosis by reducing tissue factor in a thromboxane dependent-manner in mouse 15.45 – 16.00 Federica Dellera Effect of homoarginine on neointima formation in balloon-injured rat carotid arteries Short Break
16.15 – 16.30 Stefano Musardo Development of innovative tools for Alzheimer's disease therapy 16.30 – 16.45 Chiara Malipihi Repeated exposure to cocaine alters molecular mechanisms involved in the regulation of cognition in adolescent rats 16.45 – 17.00 Simona Melfi Decrease in NF2 expression induces changes in Schwann cell differentiation 17.00 – 17.15 Natalia Marchetti Interleukin-1beta induces long-term effects on the development of glutamatergic neurons	16.15 – 16.30 Giulia Ganzetti Effect of a single infusion of Tetranectin-ApoA-I on atherosclerosis in rabbits 16.30 – 16.45 Fabio Lucca Role of LpX in the development of renal disease in LCAT deficiency 16.45 – 17.00 Andrea Baragetti Genetic variants affecting Leukocytes Telomere Length and its shortening are associated with cardio-metabolic impairment and cardiovascular prognosis in the general population 17.00 – 17.15 Paola Canzano Impact of chronic kidney disease on the platelet phenotype and the plasma proteomic profile of CAD patients 17.15 – 17.30 Laura Rossetti Hypertension upregulates platelet tissue factor expression in stroke-prone rats

17:30 - 18.00

Closing remarks and Award Ceremony for the best oral presentations Happy Hour

18.00

Brain area-specific changes in glucose metabolism in rats after acute stress

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Neuroscience

Keywords: Acute stress, Glucose metabolism, Positron Emission Tomography, Hexokinase, Glucose uptake

ABSTRACT

Stress is known to be one of the main risk factors for neuropsychiatric disorders. The stress response is a physiological mechanism of adaptation essential for survival but, when the stressful event is prolonged or overwhelming, maladaptive mechanisms could occur, thus increasing the risk to develop a stress-related pathology. Several human and *ex vivo* animal studies tried to investigate how brain responds to stressors and which cerebral areas are involved, but only in the last years, with the recent progress in neuroimaging techniques such as Positron Emission Tomography (PET), it has been possible to study metabolic and neurobiological changes induced *in vivo* by stress. Compelling evidence shows that energy metabolism and mitochondrial activity are affected by stress: at a cellular level stress results to be linked to premature cellular ageing and shortened telomeres. Because mitochondria actively regulate synaptic transmission, brain seems to be vulnerable to bioenergetic fluctuations and mitochondrial defects induced by stress.

Main aim of the present study was to evaluate the effect of acute foot shock (FS) stress on brain glucose metabolism in rats. We took advance of [18F]FDG-PET studies on FS-stressed rats to highlight which brain areas were activated/inactivated by stress. Our results showed an increase of energy consumption in rostral regions, while in more caudal areas acute stress induced a decrease in glucose metabolism. In order to understand whether these changes in activation of selected brain areas were related with modifications in synaptic glucose metabolism, thus suggesting changes in synaptic function, we measured enzymatic activity of hexokinase, the rate-limiting enzyme for glycolysis in the brain. In line with previous results, hexokinase activity resulted to be increased by acute stress in synaptosomes from prefrontal cortex and dorsal hippocampus while showing an opposite trend in those from ventral hippocampus. Moreover, as we wanted to investigate *ex vivo* whether our previous results reflected real changes in synaptic glucose consumption, we analyzed synaptic glucose uptake by measuring the uptake of the glucose analogous 2-Deoxy [3H] glucose in perfused purified synaptosomes from stressed rats. Data from the uptake of 2-Deoxy [3H] glucose are also consistent with previous results.

Taken together, these results suggest that acute stress induces area-specific changes in glucose metabolism, immediately after the stress episode.