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IN VIVO MOLECULAR IMAGING REVEALS DISTINCT DISTRIBUTIONS OF THE SEROTONIN TRANSPORTER, THE MAJOR INHIBITORY AND EXCITATORY SEROTONIN RECEPTORS

M. Savli¹, A. Bauer², D. Häusler³, T. Kroll², A. Hahn¹, F. Rattay⁴, M. Mitterhauser³, W. Wadsak³, S. Kasper¹, R. Lanzenberger¹

¹Psychiatry and Psychotherapy, Medical Universitiy of Vienna, Vienna, Austria, ²Institute of Neuroscience and Medicine (INM-2), Research Centre Jülich, Jülich, Germany, ³Department of Nuclear Medicine, Medical University of Vienna, ⁴Institute for Analysis and Scientific Computing, Vienna University of Technology, Vienna, Austria

Introduction: Based on evidences in molecular neuroimaging, postmortem and genetic studies, impaired serotonergic neurotransmission has been implicated with affective disorders. Moreover, a growing number of evidences showed strong interrelations within the serotonergic system suggesting a common mechanism in the modulation of receptor and transporter densities.

Objective: Here we directly investigated the regional expression of the 5-HT_{1A}, 5-HT_{2A} and 5-HTT using PET and the three highly selective and specific radioligands [carbonyl-¹¹C]WAY-100635, [¹⁸F]Altanserin and [¹¹C]DASB in healthy subjects.

Methods: A total of 55 healthy subjects (5-HT_{1A}: 36 subjects, 18 males, age=26.0 \pm 4.9; 5-HT_{2A}: 19 subjects, 11 males, age=28.2 \pm 5.9; 5-HTT: 8 males, age=28.12 \pm 3.6) were included in this study. Binding potential (BP_{ND}) values were quantified according to the AAL parcellation scheme.

Results: BP_{ND} values averaged over both hemispheres ranged from 0.40-6.35 for the 5-HT_{1A} receptor; 0.01-2.01 for the 5-HT_{2A} receptor and 0.09-2.05 for the 5-HTT, respectively. There was a specific topological pattern according to the ratio between the 5-HT_{1A}, 5-HT_{2A} receptors and 5-HTT ("fingerprints").

Conclusions: Such information can be essential for detecting potential local alterations in the ratio between different binding proteins on a network level in pathological conditions. Moreover, these data might provide further insight in area-specific effects of frequently prescribed selective serotonin re-uptake inhibitors (SSRI):

- 1) due to the distinct local receptor and transporter availability:
- 2) SSRI application alters the postsynaptic receptor expression and thus;
- 3) leads to a modified interaction of inhibitory and exhibitory receptors.