

¹H MRSI Study of Effects of Cognitive-Behavioral Therapy on Obsessive-Compulsive Disorder

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Introduction: Obsessive-compulsive disorder (OCD) is a common, debilitating psychiatric condition that responds to cognitive-behavioral therapy (CBT). CBT reduces hyperintense FDG-PET signals in anterior cingulate, caudate, thalamus and other brain regions¹⁻⁴ but its effects are little explored with MRS⁵. We examined ¹H MRSI metabolites in multiple brain regions implicated in adult OCD before and after 4 weeks of daily CBT with no medication changes.

Methods: 8 DSM-IV OCD patients (5 ♀, 37.1 ± 9.8 yr) were treated with the Gorbis integrated method of intensive CBT; 7 completed therapy. Clinical response was evaluated by the Yale-Brown Obsessive-Compulsive Scale (Y-BOCS). MRSI (1.5 T, Siemens PRESS, TR/TE = 1500/30 ms, 11 x 11 mm² in-plane, 9 mm thick) was acquired before and after CBT from three slabs: basal ganglia (Fig. 1), perigenual anterior cingulate, and anterior middle cingulate. MRS peaks were fit and normed to water by LCModel. Voxels were selected with MRSI Voxel Picker⁶ which rejected spectra with linewidth > 0.1 Hz or SNR < 3 and rejected peaks with SD > 20%. Pretreatment-results were compared to those of 18 healthy controls (14 ♀, 37.6 ± 10.4 yr). Statistics were non-parametric Mann-Whitney U and Wilcoxon tests.

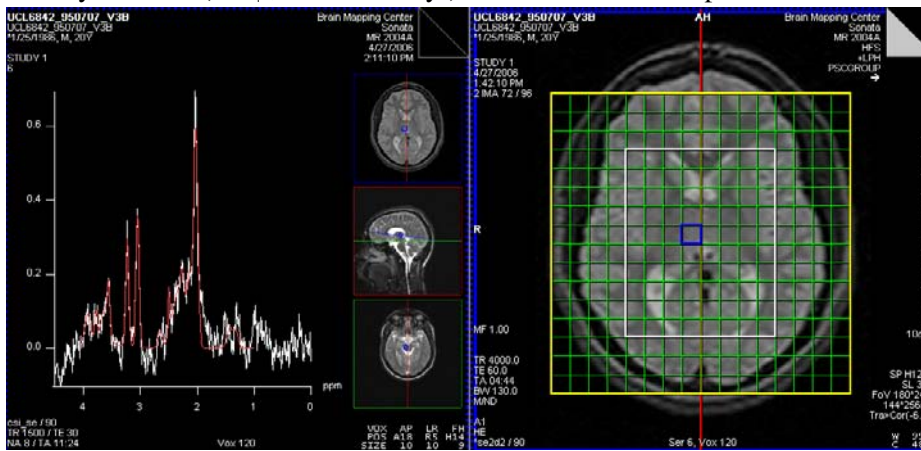


Fig. 1. (Right) Positioning of basal ganglia MRSI slab (yellow) and sample spectrum (left, from blue box). The PRESS volume (white) samples bilateral caudate, putamen, thalamus and other structures. Other slabs sampled bilateral perigenual anterior cingulate and anterior middle cingulate.

Results: Mean Y-BOCS in patients decreased from 26.2 ± 9.0 (severe OCD) to 10.3 ± 5.8 (subclinical). In bilateral thalamus, mean pretreatment Cr was 13-19 % below control ($p < 0.001-0.05$). Cr was below control mean for 7-8/8 patients. In 6-7/7 patients, Cr increased

post-treatment (mean 10-15%, $p < 0.05$). Other post-treatment increases were: NAA +NAAG in right thalamus (10%), Cr in left putamen (15%), Cho (13%) and mI (16%) in right anterior middle cingulate (all $p < 0.05$).

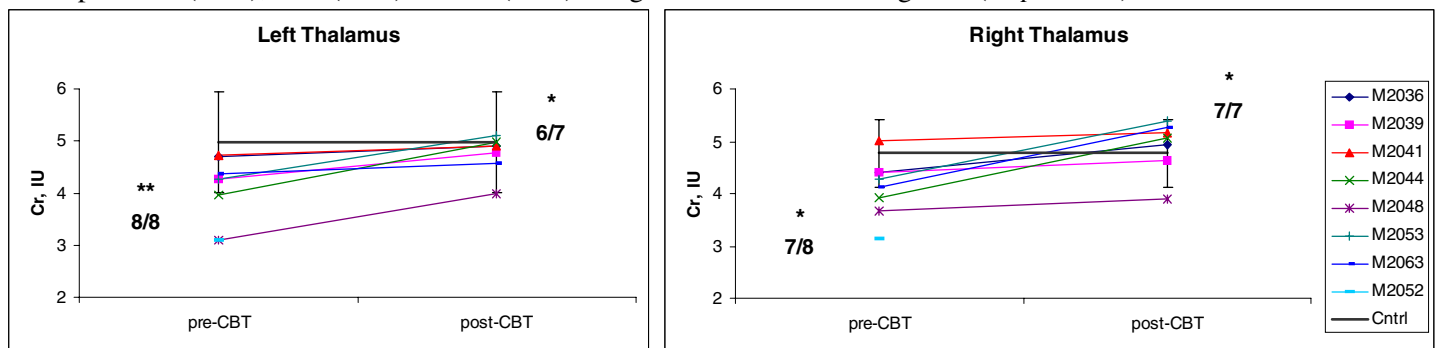


Fig. 2. Pre- and post-CBT Cr in thalamus in 8 OCD patients; mean ± SD of controls. $p < *0.05$, $**0.001$. “IU” = Institutional Units

Discussion: Abnormal Cr has been measured in bilateral thalamus in pediatric OCD⁷. In our patients (16 weeks stable on medication), metabolic response to CBT was observed in bilateral thalamus and right anterior middle cingulate, the same regions where we saw FDG-PET response to intensive integrated CBT in OCD³. MRSI response was also seen in left putamen. Anterior middle cingulate, putamen, and thalamus together form the “action-compulsive behaviors circuit” proposed to be disturbed in OCD in a recent model⁸. CBT response may be mediated by this circuit.

¹Baxter et al. *Arch Gen Psychiat* (1992), ²Schwartz et al. *Arch Gen Psychiat* (1996), ³Saxena, Gorbis, O’Neill et al. (submitted), ⁵Benazon et al. *JAACAP* (2003), ⁶O’Neill et al. ISMRM Workshop (2006), ⁷Mirza, O’Neill, Rosenberg et al. *J Child Neurol* (2006), ⁸Middleton in Vogt BA (ed.) *Cingulate Neurobiology & Disease* (in press). Supported by NIMH R01 MH069433; Center for Research Resources RR12169, RR13642, RR08655; and private donors