

Motor Inhibition and Cognitive Flexibility in Obsessive-Compulsive Disorder and Trichotillomania

Samuel R. Chamberlain, M.A.
Naomi A. Fineberg, M.D.
Andrew D. Blackwell, Ph.D.
Trevor W. Robbins, Ph.D.
Barbara J. Sahakian, Ph.D.

Objective: Problems with inhibiting certain pathological behaviors are integral to obsessive-compulsive disorder (OCD), trichotillomania, and other putative obsessive-compulsive spectrum disorders. The authors assessed and compared motor inhibition and cognitive flexibility in OCD and trichotillomania for the first time, to their knowledge.

Method: The Stop-Signal Task and the Intradimensional/Extradimensional Shift Task were administered to 20 patients with OCD, 17 patients with trichotillomania, and 20 healthy comparison subjects.

Results: Both OCD and trichotillomania showed impaired inhibition of motor responses. For trichotillomania, the deficit was worse than for OCD, and the degree of the deficit correlated significantly with symptom severity. Only patients with OCD showed deficits in cognitive flexibility.

Conclusions: Impaired inhibition of motor responses (impulsivity) was found in OCD and trichotillomania, whereas cognitive inflexibility (thought to contribute to compulsivity) was limited to OCD. This assessment will advance the characterization and classification of obsessive-compulsive spectrum disorders and aid the development of novel treatments.

(*Am J Psychiatry* 2006; 163:1–3)

Growing evidence suggests that obsessive-compulsive disorder (OCD), trichotillomania (compulsive hair pulling), and other conditions may represent an obsessive-compulsive spectrum or family of disorders (1). The symptoms of OCD and trichotillomania suggest problems inhibiting motor behavior (impulsivity). However, OCD is also characterized by repetitive mental acts or behaviors performed according to rigid rules (DSM-IV). It has been proposed that these compulsive symptoms may be mediated by problems with cognitive flexibility, such as the ability to shift attentional focus (2). We compared motor inhibition and cognitive flexibility in subjects with OCD and trichotillomania using two well-validated neurocognitive tasks. Motor inhibition was assessed with the Stop-Signal Task (3), which provides a sensitive estimate of the time taken to internally suppress motor responses. This task has been shown to be sensitive to motor impulsivity associated with attention deficit hyperactivity disorder (ADHD) and damage to the right inferior frontal gyrus (4, 5). Cognitive flexibility was assessed with the Intradimensional/Extradimensional Shift Task, developed from the Wisconsin Card Sorting Test of frontal lobe integrity (6). The Intradimensional/Extradimensional Shift Task examines different components of attentional flexibility, including reversal learning, set formation, and the ability to shift attention between stimulus dimensions (7). We predicted that subjects with OCD would show deficits in motor inhibition and cognitive flexibility but that subjects with trichotillomania would show deficits in motor inhibition only.

Method

Twenty OCD patients, 17 trichotillomania patients, and 20 healthy comparison subjects gave written informed consent and

were administered the tasks. The groups were matched for age, education, and IQ (8) (Table 1) (analysis of variance [ANOVA], all $p > 0.10$). DSM-IV diagnoses were made by a fully certified consultant psychiatrist using an extended clinical interview supplemented by the Mini-International Neuropsychiatric Inventory (9). For OCD, only archetypal illness involving contamination or checking was allowed (no hoarding). The participants were excluded if they had significant comorbidities, including major depression (Montgomery-Åsberg Depression Rating Scale score [MADRS] > 16) (10), Tourette's syndrome, psychosis, eating disorders, and other anxiety disorders. OCD severity was assessed with the Yale-Brown Obsessive Compulsive Scale (11). The patients with OCD scored > 16 on the scale; 16 of 20 were receiving selective serotonin reuptake inhibitors (SSRIs), and the remainder were unmedicated (free from psychotropic medications for at least 6 months before cognitive testing). The severity of trichotillomania was assessed with the Massachusetts General Hospital Hairpulling Scale (12), and patients with trichotillomania were unmedicated (free from psychotropic medications for at least 6 months before cognitive testing).

On the computerized Stop-Signal Task, subjects respond rapidly to left- or right-facing arrows on a computer screen with corresponding motor responses and attempt to inhibit responses when an auditory "stop signal" sounds. With a tracking algorithm, this task estimates the time taken to internally suppress prepotent motor responses (stop-signal reaction time) (3, 5).

The Intradimensional/Extradimensional Shift Task (2) is a nine-stage visual discrimination task with multidimensional stimuli. Two stimuli are displayed at a time, and feedback is provided so that the subject can learn which stimulus is correct. To pass each stage, six consecutive correct responses are required within 50 trials; otherwise, the task ends. The rule for correct responding is modified at the start of each task stage in order to dissociate different aspects of cognitive flexibility (7). For example, the intradimensional shift stage examines rule generalization when novel stimuli are introduced, whereas the extradimensional shift stage examines the ability to inhibit or shift attention away from previously relevant stimulus dimensions (akin to a category shift on the

TABLE 1. Demographic and Clinical Characteristics of Patients With Obsessive-Compulsive Disorder (OCD) or Trichotillomania and Healthy Comparison Subjects

Characteristic	OCD (N=20) (four men and 16 women)		Trichotillomania (N=17) (two men and 15 women)		Comparison Subjects (N=20) (four men and 16 women)		p ^a
	Mean	SD	Mean	SD	Mean	SD	
Demographic characteristics							
Age (years)	35.3	14.0	36.1	12.6	34.0	8.1	
Education score (maximum=4)	2.8	1.0	2.8	0.8	3.0	0.8	
National Adult Reading Test premorbid verbal IQ estimate	115.7	5.9	117.7	7.5	118.0	5.5	
Stop-Signal Task score							
Stop-signal reaction time (motor inhibition) (msec)	211.6	57.9	264.9	72.2	167.8	48.6	<0.01
Median response time (msec)	428.6	85.0	397.2	36.1	421.2	63.7	
Directional errors	5.4	6.1	6.8	9.2	2.2	2.6	
Intradimensional/Extradimensional Shift Task score							
Intradimensional shift trials until criterion met	6.5	0.7	6.8	1.3	6.5	0.5	
Extradimensional shift trials until criterion met	25.9	18.1	17.1	8.4	16.3	13.1	<0.05

^a Significant differences between groups (ANOVA: $df=2, 54$).

Wisconsin Card Sorting Test). The subjects who failed to pass a stage were assigned a score of 50 trials to the criterion for that stage and were excluded from data analysis for subsequent stages not attempted.

To detect overall group differences for demographic and cognitive measures, ANOVA was employed. In the instances in which significant group differences were found with ANOVA, exploratory pairwise analyses were conducted with Fisher's least significant difference tests as appropriate.

Results

The mean Yale-Brown Obsessive Compulsive Scale score for the OCD group was 20.4 ($SD=4.1$). The mean Massachusetts General Hospital Hairpulling Scale score for the trichotillomania group was 16.4 ($SD=4.7$). MADRS scores for all groups were beneath the cutoff point for clinically significant depression (<10) (13)—OCD patients: mean=6.9 ($SD=4.4$), trichotillomania patients: mean=4.2 ($SD=3.6$), healthy comparison subjects: mean=3.1 ($SD=4.4$). There was a significant overall group difference on MADRS scores (ANOVA: $F=8.44, df=2, 54, p<0.01$). Post hoc least significant difference tests showed that the subjects with OCD had significantly higher MADRS scores than other groups (OCD group versus trichotillomania group: $p<0.05$; OCD group versus healthy comparison group: $p<0.01$), whereas the scores of the trichotillomania and healthy comparison groups did not differ on the MADRS ($p>0.10$).

On the Stop-Signal Task, the groups differed significantly on stop-signal reaction times ($F=12.19, df=2, 54, p<0.01$). Post hoc analysis revealed that the patients with trichotillomania had longer stop-signal reaction times than the OCD patients, who had longer reaction times than the healthy comparison subjects (trichotillomania group versus OCD group: $p<0.01$; trichotillomania group versus healthy comparison group: $p<0.001$; OCD group versus healthy comparison group: $p<0.05$). The groups did not differ significantly in the number of directional errors overall ($p>0.05$) or on median "go" response times ($p>0.10$).

In terms of the number of trials taken to reach the criteria for the intradimensional shift and extradimensional shift stages, there was a significant interaction of stage and group ($F=3.59, df=2, 54, p<0.05$). The groups differed from each other only at the extradimensional shift stage ($F=3.44, df=2, 54, p<0.05$), with post hoc analysis revealing that the OCD group required more trials at the extradimensional shift stage compared to both of the other groups (OCD group versus trichotillomania group: $p<0.05$; OCD group versus healthy comparison group: $p<0.05$). The trichotillomania and healthy comparison subjects did not differ at the extradimensional shift stage (all $p>0.10$). For other stages of the task (including those necessitating rule reversals), the performance of the OCD, trichotillomania, and healthy comparison groups did not differ significantly (ANOVAs: all $p>0.10$).

Analyses were undertaken with Pearson's correlations. Symptom severity in trichotillomania (Massachusetts General Hospital Hairpulling Scale scores) correlated significantly with stop-signal reaction times ($r=0.564, p<0.02$). Symptom severity in OCD (measured with the Yale-Brown Obsessive-Compulsive Scale) did not correlate with any task measures ($p>0.10$). MADRS scores did not correlate significantly with performance on either task within the clinical groups (all $p>0.10$).

Discussion

The clinical phenotypes of OCD and trichotillomania suggest an overlap in terms of impulsivity or problems inhibiting motor behavior. However, only OCD is associated with rigid mental acts or behaviors that imply problems with cognitive flexibility (2). Obtained with objective measures, our results provide direct support for these contentions: both clinical groups showed severely impaired motor inhibition (the Stop-Signal Task), but only the OCD group showed a deficit in cognitive flexibility (with the Intradimensional/Extradimensional Shift Task). Impairment of motor inhibition correlated significantly with symptom severity in trichotillomania. In OCD, the cognitive flexibil-

ity deficit was specific to the stage in which it was necessary to inhibit or shift attentional focus away from a previously relevant stimulus dimension. Individual scores for unmedicated OCD patients were comparable to medicated patients (within 0.5 SDs of the mean for the medicated subgroup on key task measures).

This study is the first to our knowledge to demonstrate motor impulsivity across obsessive-compulsive symptom disorders, with deficits in cognitive flexibility limited to OCD. This implies distinct but also overlapping neurobiological underpinnings to disorders whose nosological status is currently under review. Limitations of this study include the group size and the possible confounding effect of SSRI medications in the OCD group. Future research should expand on the concepts of impulsivity and compulsivity, examine the effects of pharmacological and psychological treatments on behavioral inhibition, and delineate the contribution of frontal-striatal circuitry and neurochemical modulation of these processes. This will be relevant to the treatment of impulsive as well as compulsive aspects of the obsessive-compulsive symptom disorder profile.

Received Feb. 21, 2005; revision received April 5, 2005; accepted April 26, 2005. From the Department of Psychiatry, Addenbrooke's Hospital, University of Cambridge School of Clinical Medicine; the Department of Psychiatry, Queen Elizabeth II Hospital, Welwyn Garden City, Hertfordshire, U.K.; and the Behavioural and Clinical Neuroscience Institute and Department of Experimental Psychology, University of Cambridge, Cambridge, U.K. Address correspondence and reprint requests to Dr. Chamberlain, Department of Psychiatry, University of Cambridge School of Clinical Medicine, Addenbrooke's Hospital, Box 189, Cambridge, CB2 2QQ U.K.; src33@cam.ac.uk (e-mail).

Funded by the Wellcome Trust (program grant 019407) and the Medical Research Council (center grant G0001354). The Behavioural and Clinical Neuroscience Institute, University of Cambridge, is funded jointly by the Wellcome Trust and the Medical Research Council. Dr. Chamberlain works with the Cambridge M.B.-Ph.D. program and is funded by a Medical Research Council research studentship.

The authors thank the volunteers who took part and Dr. Ulrich Muller for comments on the manuscript.

Drs. Blackwell, Robbins, and Sahakian perform consulting for Cambridge Cognition.

References

- Stein DJ, Hollander E: Obsessive-compulsive spectrum disorders (letter). *J Clin Psychiatry* 1995; 56:265–266
- Chamberlain SR, Blackwell AD, Fineberg NA, Robbins TW, Sahakian BJ: The neuropsychology of obsessive compulsive disorder: the importance of failures in cognitive and behavioural inhibition as candidate endophenotypic markers. *Neurosci Biobehav Rev* 2005; 29:399–419
- Logan GD, Cowan WB, Davis KA: On the ability to inhibit simple and choice reaction time responses: a model and a method. *J Exp Psychol Hum Percept Perform* 1984; 10:276–291
- Aron AR, Dowson JH, Sahakian BJ, Robbins TW: Methylphenidate improves response inhibition in adults with attention-deficit/hyperactivity disorder. *Biol Psychiatry* 2003; 54:1465–1468
- Aron AR, Fletcher PC, Bullmore ET, Sahakian BJ, Robbins TW: Stop-signal inhibition disrupted by damage to right inferior frontal gyrus in humans. *Nat Neurosci* 2003; 6:115–156
- Lezak MD, Howieson DB, Loring DW: *Neuropsychological Assessment*. New York, Oxford University Press, 1995
- Lawrence AD, Sahakian BJ, Robbins TW: Cognitive functions and corticostriatal circuits: insights from Huntington's disease. *Trends Cogn Sci* 1998; 2:379–388
- Willshire D, Kinsella G, Prior M: Estimating WAIS-R IQ from the National Adult Reading Test: a cross-validation. *J Clin Exp Neuropsychol* 1991; 13:204–216
- Sheehan DV, Lecrubier Y, Sheehan KH, Amorim P, Janavs J, Weiller E, Hergueta T, Baker R, Dunbar GC: The Mini-International Neuropsychiatric Interview (M.I.N.I.): the development and validation of a structured diagnostic psychiatric interview for DSM-IV and ICD-10. *J Clin Psychiatry* 1998; 59(suppl 20):22–57
- Montgomery SA, Åsberg M: A new depression scale designed to be sensitive to change. *Br J Psychiatry* 1979; 134:382–389
- Goodman WK, Price LH, Rasmussen SA, Mazure C, Fleischmann RL, Hill CL, Heninger GR, Charney DS: The Yale-Brown Obsessive Compulsive Scale, I: Development, use, and reliability. *Arch Gen Psychiatry* 1989; 46:1006–1011
- Keuthen NJ, O'Sullivan RL, Ricciardi JN, Shera D, Savage CR, Borgmann AS, Jenike MA, Baer L: The Massachusetts General Hospital (MGH) HairPulling Scale: 1. development and factor analyses. *Psychother Psychosom* 1995; 64:141–145
- Hawley CJ, Gale TM, Sivakumaran T: Defining remission by cut-off score on the MADRS: selecting the optimal value. *J Affect Disord* 2002; 72:177–184