Cognitive functioning in obsessive-compulsive disorder: a meta-analysis

N. Y. Shin¹, T. Y. Lee², E. Kim² and J. S. Kwon^{1,2,3}*

¹Interdisciplinary Cognitive Science Program, Seoul National University, Seoul, South Korea

² Department of Psychiatry, Seoul National University College of Medicine, Seoul, South Korea

³Department of Brain & Cognitive Sciences, Seoul National University College of Natural Science, Seoul, South Korea

Background. Substantial empirical evidence has indicated impairment in the cognitive functioning of patients with obsessive-compulsive disorder (OCD) despite inconsistencies. Although several confounding factors have been investigated to explain the conflicting results, the findings remain mixed. This study aimed to investigate cognitive dysfunction in patients with OCD using a meta-analytic approach.

Method. The PubMed database was searched between 1980 and October 2012, and reference lists of review papers were examined. A total of 221 studies were identified, of which 88 studies met inclusion criteria. Neuropsychological performance and demographic and clinical variables were extracted from each study.

Results. Patients with OCD were significantly impaired in tasks that measured visuospatial memory, executive function, verbal memory and verbal fluency, whereas auditory attention was preserved in these individuals. The largest effect size was found in the ability to recall complex visual stimuli. Overall effect estimates were in the small to medium ranges for executive function, verbal memory and verbal fluency. The effects of potentially confounding factors including educational level, symptom severity, medication status and co-morbid disorders were not significant.

Conclusions. Patients with OCD appear to have wide-ranging cognitive deficits, although their impairment is not so large in general. The different test forms and methods of testing may have influenced the performance of patients with OCD, indicating the need to select carefully the test forms and methods of testing used in future research. The effects of various confounding variables on cognitive functioning need to be investigated further and to be controlled before a definite conclusion can be made.

Received 5 April 2013; Revised 20 June 2013; Accepted 22 June 2013; First published online 19 July 2013

Key words: Cognitive function, meta-analysis, obsessive-compulsive disorder.

Introduction

Obsessive-compulsive disorder (OCD) is a neuropsychiatric disorder involving distressing intrusive thoughts and related compulsive behaviours. OCD has been reported to be associated with neurobiological abnormalities that are distinct from those associated with other anxiety disorders (Radua *et al.* 2010). Considerable evidence has indicated neurocognitive impairment in patients with OCD (Kuelz *et al.* 2004; Chamberlain *et al.* 2005; Muller & Roberts, 2005; Cavedini *et al.* 2006; Olley *et al.* 2007; Menzies *et al.* 2008; Goncalves *et al.* 2010; Melloni *et al.* 2012). The cognitive domain with the most consistently reported deficits is visual memory, especially for complex visual stimuli (Savage *et al.* 1999; Muller & Roberts, 2005; Olley et al. 2007), although it has also been suggested that such memory deficits arise from impairments in executive function, such as organizational strategy, rather than from memory difficulties per se (Savage et al. 1999). Other cognitive functions frequently investigated with regard to this disorder include executive function, verbal fluency and verbal memory. Individuals with OCD have been observed to experience difficulties in inhibiting ongoing cognitive and motor responses (Aycicegi et al. 2003; Penades et al. 2007; Abramovitch et al. 2011; Rajender et al. 2011; Tukel et al. 2012), shifting attention from one aspect of stimuli to others (Abbruzzese et al. 1995, 1997; Aycicegi et al. 2003; Fenger et al. 2005), engaging in executive planning (Cavedini et al. 2001, 2010; Nielen & Den Boer, 2003; Chamberlain et al. 2007; Wobrock et al. 2010; Rajender et al. 2011; Tukel et al. 2012) and decision making (Cavallaro et al. 2003, Cavedini et al. 2010; Starcke et al. 2010), generating words within a limited time (Schmidtke et al. 1998; Murphy et al. 2004; Rampacher et al. 2010;

^{*} Address for correspondence: J. S. Kwon, M.D., Ph.D., Department of Psychiatry, Seoul National University College of Medicine, 28 Yeongon-dong, Chongno-gu, Seoul, South Korea 110-744.

⁽Email: kwonjs@snu.ac.kr)

Borges et al. 2011; Tukel et al. 2012) and recalling verbal information (Savage et al. 2000; Ceschi et al. 2003; Deckersbach et al. 2004; Segalas et al. 2010; Rajender et al. 2011). However, findings of cognitive dysfunction in OCD have not been consistent across studies (Kuelz et al. 2004; Chamberlain et al. 2005). Discrepant findings may be attributable to confounding factors including sex (Savage et al. 2000; Deckersbach et al. 2004), duration of illness (Nakao et al. 2009), medication status (Nakao et al. 2009; Segalas et al. 2010), comorbidity (Aycicegi et al. 2003), age at onset of illness (Henin et al. 2001; Roth et al. 2005; Hwang et al. 2007), insight (Tumkaya et al. 2009), family history (Boone et al. 1991) and symptom-based subtype (Ceschi et al. 2003; Cha et al. 2008; Nedeljkovic et al. 2009). However, data on the impact of these confounding factors on cognitive functioning have been conflicting.

Although several reviews systematically reviewed the neuropsychological impairments (Kuelz et al. 2004; Chamberlain et al. 2005; Olley et al. 2007; Menzies et al. 2008; Melloni et al. 2012) and/or confounding variables (Kuelz et al. 2004) in OCD, they adopted a qualitative approach to the data and did not provide a systematic overview of the magnitude of the effects. Thus, to assess comprehensively the existence and magnitude of the cognitive impairments experienced by patients with OCD, this study used a meta-analytic approach to synthesize the available data. Additionally, we examined the effects of demographic and clinical variables on cognitive impairment in OCD patients to test our hypothesis that cognitive dysfunction in patients with OCD may be associated with these variables.

Method

Search strategies and study selection

Potential articles were identified through a Pubmed literature search conducted by two researchers (N.Y.S. and T.Y.L.) for articles published between 1980 and October 2012. The following keywords were used for searching: ('obsessive-compulsive disorder' OR 'obsessive-compulsive') AND ('cognitive' OR 'cognition' OR 'neuropsychological' OR 'neuropsychology' OR 'neurocognitive' OR 'neurocognition' OR 'executive'). The reference lists of review articles were also checked. The following inclusion criteria were used to select studies for full-paper review: (1) published in English in a peer-reviewed journal; (2) restricted to adult patients (aged 18 years or older) diagnosed with OCD according to Diagnostic and Statistical Manual of Mental Disorders, third edition (DSM-III), fourth edition (DSM-IV), Ninth Revision of the International Classification of Diseases (ICD-9), or Tenth Revision (ICD-10); (3) used reliable neuropsychological tests to measure cognitive functioning and reported data for each individual test, not providing only composite scores; and (4) reported statistics for both patient and healthy control groups that were convertible to effect sizes. When several relevant articles from the same centre were identified, the study with the largest sample was selected.

Data extraction

The variables extracted for the meta-analysis were year of publication, mean and standard deviation for age, sex (ratio of males), mean and standard deviation of years of education, and cognitive performance (mean and standard deviation or t, F and p statistics). Additionally, we coded clinical variables including symptom severity, as evaluated by the Yale–Brown Obsessive-Compulsive Scale (YBOCS), percentage of patients receiving psychotropic medication, percentage of patients with co-morbid psychiatric disorders, and mean age at onset and duration of illness. The variables recorded were cross-checked by the two authors (N.Y.S. and T.Y.L.)

Statistical analyses

Analysis was conducted using Comprehensive Meta-Analysis version 2 (Biostat, Inc., USA) and Stata version 12 (StataCorp LP, USA) software. Effect sizes for outcome variables and cognitive domains were estimated by calculating Hedges' g, which is the difference between patient and control groups divided by the pooled standard deviation and weighted for sample size to control for small-sample bias (Hedges & Holkin, 1985). Negative values for effect sizes indicate poor performance by patients with OCD compared with healthy controls. The heterogeneity of effects across studies was tested with Cochran's Q statistic and the l^2 index. For homogeneous data (p>0.1 for Q statistic), a fixed-effect model was calculated, and in the presence of heterogeneity, a random-effect model was adopted for the effect size of each variable. In cases with significant heterogeneity across studies, potential sources of heterogeneity were investigated with a Galbraith plot. This scatter plot graphically displays studies that contribute heterogeneity by plotting each study's z score (the mean difference divided by the standard error of the difference) against the reciprocal of the standard error of the mean difference. Studies with high heterogeneity fell outside 2 standard deviations above and below the 95% confidence interval (CI) limit. We evaluated changes in Q statistics after removing outlier studies. Subgroup analysis and meta-regression were conducted to evaluate the effects of study characteristics on cognitive functioning. Subgroup analysis was performed to investigate the influence of categorical factors (test forms or methods of testing used). Between-group heterogeneity (Q_{bet}) was computed to test the significance of differences in the magnitudes of effect sizes between subgroups. Meta-regression analysis was used to examine the effects of continuous moderators on the effect sizes across all variables related to cognitive tasks (publication year, age, sex, years of education, symptom severity, percentage of medicated patients, and percentage of patients with co-morbid psychiatric disorders). We were not able to include the effects of age at onset and duration of illness in the analysis because too few studies reported relevant data. To control for type I errors due to multiple comparisons, we applied the adjusted *p* value for the meta-regression by dividing α by the number of moderators. Publication bias was examined by visually inspecting funnel plots and using the regression intercept developed by Egger et al. (1997). The trim-and-fill method (Duval & Tweedie, 2000) was applied to adjust for publication bias when indicated.

We analysed data from at least five studies for each neuropsychological variable. Cognitive tasks that are very similar in their set-up [e.g. the Auditory Verbal Learning Test and the California Verbal Learning Test; the Tower of London (ToL) and the Tower of Hanoi (ToH) tasks; the Object Alternation Test and the Delayed Alternation Test] were combined for analysis. To examine the cognitive domain specificity, we grouped individual test variables into six domains (visuospatial memory, verbal memory, verbal fluency, executive function, processing speed, and attention).

Results

Study characteristics

More than 2300 articles were identified through a two-step search strategy. After reviewing titles and abstracts, we selected 221 relevant articles for full-text analysis. Of these, 133 were excluded based on the inclusion criteria: 26 did not include a healthy control group, 10 included patients younger than 18 years of age, 12 did not use statistics appropriate for conversion into effect sizes, 43 did not use reliable standardized neuropsychological tests, and 15 used samples that overlapped with those used by studies that had been already included. Additionally, 25 studies were not included in the analysis because fewer than five studies reported results for each cognitive variable they addressed. Moreover, two studies were excluded because the statistics reported in tables did not match statements in the text. Therefore, 88 studies (Fig. 1) including a total of 3070 patients with OCD (mean

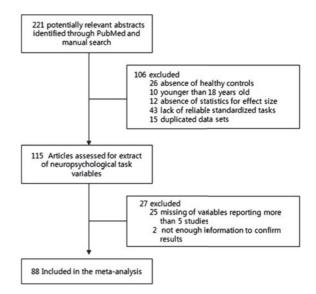


Fig. 1. Search strategy used for selection of studies included in the meta-analysis.

age 33.5 years, s.D.=4.7, 49.0% male) and 3024 control subjects (mean age 32.6 years, s.D.=4.7, 49.5% male) met the inclusion criteria (online Supplementary Table S1). The mean years of education in 63 studies was 13.5 (s.D.=2.0) in patients and 14.2 (s.D.=2.1) in controls. Of the patients, 56% with OCD included in 76 studies were drug-naive or drug-free, and 58% in 65 studies excluded patients with co-morbid psychiatric disorders. Symptom severity, as evaluated by the YBOCS, was 24.2 (s.D.=3.1) in 62 studies. The average duration of illness was 11.9 years in 42 studies, and the mean age at onset of OCD was 19 years in 28 studies.

Cognitive functioning

The main results of the meta-analysis are presented in Table 1 and Fig. 2. Patients with OCD showed significantly poorer performances in all task variables except digit span and the extra-dimension trial of the Intra/Extra-Dimensional Set (IED ed) compared with that of healthy controls. The largest effect sizes were found for immediate recall in the Rey-Osterrieth Complex Figure test (RCFT ir) (g=-0.74), which measures visuospatial memory, and for number of moves in excess on the ToL/ToH (ToL/ToH em) (g=-0.73); this was followed by organizational strategies in the RCFT (RCFT organ) (g = -0.63). The effect sizes for the Stroop test, Wisconsin Card Sorting Test (WCST) and Corsi block-tapping tests (CBT) reflected a medium degree of impairment, whereas those for tests measuring other variables reflected a small to medium degree of impairment (g < 0.5). We found no significant differences between groups in the digit span test (p>0.2) and IED ed (p>0.06).

	Studies,			Effect						
Task	п	OCD, n	НС, п	size	95% CI	z	р	Q	I^2	$p(\mathbf{Q})$
Visuospatial										
memory										
CBT	6	241	170	-0.529	-0.731 to -0.327	-5.130	< 0.001	8.02	37.67	0.16
RCFT ir	21	956	873	-0.743	-0.928 to -0.558	14.53	< 0.001	66.39	69.88	< 0.01
SWM b/w se	5	158	168	-0.452	-0.836 to -0.067	-2.300	0.021	10.36	61.40	0.04
Verbal memory										
LM II	6	200	183	-0.418	-0.626 to -0.210	-3.941	< 0.001	5.33	6.23	0.38
VLT dr	21	629	595	-0.474	-0.721 to -0.227	-3.760	< 0.001	89.21	77.58	< 0.01
Executive function										
Block design	6	206	149	-0.449	-0.667 to -0.232	-4.047	< 0.001	5.00	0.00	0.42
Design fluency	6	142	185	-0.459	-0.890 to -0.027	-2.082	0.037	18.06	72.32	0.003
IED ed	6	173	185	-0.308	-0.639 to 0.022	-1.828	0.068	11.73	57.36	0.04
IED id	5	114	126	-0.373	-0.626 to -0.120	-2.893	0.004	3.93	0.00	0.42
OAT/DAT pe	5	152	135	-0.477	-0.715 to -0.239	-3.925	< 0.001	2.90	0.00	0.57
RCFT organ	15	609	582	-0.627	-0.827 to -0.427	-6.153	< 0.001	36.32	61.46	0.001
Stroop C-W	12	311	311	-0.547	-0.808 to -0.286	-4.106	< 0.001	26.75	58.88	0.01
TMT B	22	704	792	-0.486	-0.594 to -0.377	-8.766	< 0.001	26.68	21.3	0.18
ToL/ToH em	6	255	221	-0.732	-1.005 to -0.460	-5.275	< 0.001	10.12	50.6	0.07
VLT sc	9	278	248	-0.418	-0.591 to -0.245	-4.737	< 0.001	12.14	34.11	0.15
WCST pe	21	739	622	-0.511	-0.688 to -0.335	-5.682	0.001	49.93	59.95	< 0.01
Verbal fluency										
Category fluency	9	305	352	-0.385	-0.542 to -0.229	-4.82	< 0.001	12.4	35.24	0.14
Letter fluency	28	841	935	-0.415	-0.535 to -0.296	-6.796	< 0.001	37.71	28.40	0.08
Processing speed										
TMT A	23	796	838	-0.444	-0.548 to -0.340	-8.392	< 0.001	29.57	25.60	0.13
Attention										
CPT hits	6	172	172	-0.452	-0.807 to -0.096	-2.488	0.013	13.67	63.40	0.018
Digit span	11	336	332	-0.114	-0.270 to 0.042	-1.282	0.200	15.97	37.36	0.101

Table 1. Cognitive function in patients with OCD compared with controls subjects in individual cognitive test variables

OCD, Obsessive compulsive disorder; HC, healthy controls; CI, confidence interval; CBT, Corsi block-tapping test; RCFT, Rey–Osterrieth Complex Figure Test; ir, immediate recall; SWM b/w se, Spatial Working Memory between search errors; LM, logical memory; VLT, Verbal Learning Test; dr, delayed recall; IED, Intra/Extra Dimension; ed, extra-dimensional trial score; id, intra-dimensional trial score; OAT, Object Alternation Test; DAT, Delayed Alternation Test; pe, perseverative errors; organ, organizational strategies; Stroop C-W, Stroop Color–Word inference condition; TMT B, Trail Making Test part B; ToL, Tower of London; ToH, Tower of Hanoi; em, number of moves in excess; sc, semantic clustering; WCST, Wisconsin Card Sorting Test; TMT A, Trail Making Test part A; CPT, Continuous Performance Test.

Fig. 3 presents the results by cognitive domain. Patients with OCD showed significant impairment in the domains of visuospatial memory (g=-0.624, 95% CI -0.752 to -0.495, z=-9.508, p<0.001), executive function (g=-0.493, 95% CI -0.553 to -0.432, z=-15.937, p<0.001), verbal memory (g=-0.441, 95% CI -0.600 to -0.282, z=-5.436, p<0.001), processing speed (g=-0.444, 95% CI -0.548 to -0.340, z=-8.392, p<0.001) and verbal fluency (g=-0.404, 95% CI -0.499 to -0.309, z=-8.336, p<0.001), whereas no significant differences in the domain of attention (g=-0.244, 95% CI -0.566 to -0.078, z=-1.483, p=0.14) were observed. The grand mean effect size for all cognitive domains was g=-0.478 (p<0.001).

Heterogeneity and publication bias

The *Q* tests showed significant heterogeneity for 11 of 21 cognitive variables. A Galbraith plot showed that several studies made significant contributions to the heterogeneity for the five variables. The study conducted by Rajender *et al.* (2011) reported the highest effect size between patients and controls in the Verbal Learning Test (VLT dr), and after removal of this study, significant heterogeneity disappeared (*Q*=18.95, l^2 <0.01, p=0.46; g=-0.344, 95% CI -0.450 to -0.218, z=-5.655, p<0.001). The study conducted by Starcke *et al.* (2010) was a significant source of heterogeneity on the letter fluency. After exclusion

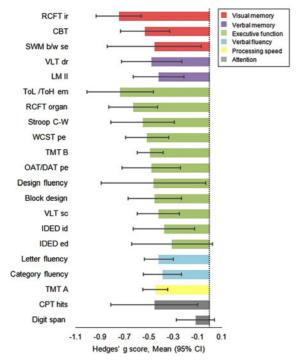


Fig. 2. Effect sizes of individual cognitive tasks in obsessive-compulsive disorder compared with controls. Negative values of Hedges' g mean worse performance in the patients compared with the controls. Values are means, with 95% confidence intervals (CIs) represented by vertical bars. RCFT, Rey-Osterrieth Complex Figure Test; ir, immediate recall; CBT, Corsi block-tapping test; SWM, spatial working memory; b/w se, between search errors; VLT, Verbal Learning Test; dr, delayed recall; LM, logical memory; ToL, Tower of London; ToH, Tower of Hanoi; em, number of moves in excess; organ, organizational strategies; Stroop C-W, Stroop Color-Word inference condition; WCST, Wisconsin Card Sorting Test; TMT B, Trail Making Test part B; OAT, Object Alternation Test; DAT, Delayed Alternation Test; pe, perseverative errors; VLT, Verbal Learning Test; sc, semantic clustering; IED, Intra/Extra Dimension; id, intra-dimensional trial score; ed, extra-dimensional trial score; TMT A, Trail Making Test part A; CPT, Continuous Performance Test.

of those data from the present study, the results of the *Q* test were non-significant (*Q*=30.70, l^2 =8.80, p=0.33; g=-0.449, 95% CI -0.550 to -0.348, z=-8.724, p<0.001). The study conducted by Cohen *et al.* (2003) reported the largest effect size for the Stroop test, and the *Q* test showed no significant heterogeneity after the data from that study were excluded (*Q*=17.05, l^2 =35.47, p=0.11; g=-0.429, 95% CI -0.594 to -0.264, z=-5.097, p<0.001). The study conducted by Trivedi *et al.* (2008) showed the largest effect size for the Continuous Performance Test (CPT); heterogeneity was no longer significant after the exclusion of the data from that study (*Q*=3.28, l^2 <0.01, p=0.51; g=-0.298,

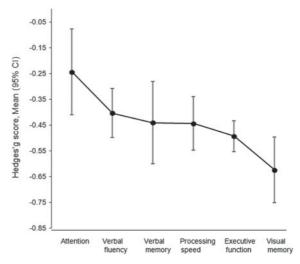


Fig. 3. Cognitive profile of neuropsychological domains in obsessive-compulsive disorder compared with controls. Negative values of Hedges' *g* mean worse performance in the patients compared with the controls. Values are means, with 95% confidence intervals (CIs) represented by vertical bars.

95% CI -0.530 to -0.067, z=-2.526, p=0.012). In terms of the IED ed, the effect reported by Nielen & Den Boer (2003) fell outside the margin in the Galbraith plot. When this study was removed, the heterogeneity of the IED ed was not significant (Q=1.30, $l^2<0.01$, p=0.86), and the effect size became significant (g=-0.439, 95% CI -0.660 to -0.218, z=-3.891, p<0.001). Visual inspection of funnel plots and Egger's test revealed significant publication bias for only the VLT dr (p=0.04). After adjusting for publication bias using the trim-and-fill method, the estimate of the effect size remained significant for this variable (g=-0.514, 95% CI -0.748 to -0.281).

Effects of moderators

Subgroup analysis revealed that the use of different forms of test was a significant contributor to heterogeneity for the WCST and the ToL/ToH. For the WCST, two forms were used: the classical manual (n=15) and the computerized (n=6, containing either 48 or 64 items) forms. Studies using the classical manual approach were identified homogenous (Q=18.66, I^2 =24.91, p=0.18), but those relying on the computerized version remained heterogeneous (Q=20.93, I^2 =76.12, p=0.001). The magnitudes of the effect sizes of the two methods differed significantly (Q_{bet} =10.34, p=0.001, g=-0.380, 95% CI -0.508 to -0.253 for the manual version; g=-0.794, 95% CI -1.012 to -0.577 for the computerized version) (Fig. 4a). When data from the ToL (n=3) were separated from data from

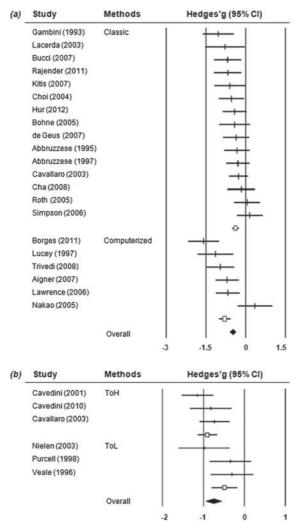


Fig. 4. Effect size of each study for the Wisconsin Card Sorting Test (*a*) and the Tower of Hanoi (ToH) and the Tower of London (ToL) tasks (*b*). The square indicates the overall estimate for each method and the diamond indicates the combined effect size for the two kinds of methods. CI, Confidence interval.

the ToH (*n*=3), each of the two groups of studies was homogeneous (Q=3.17, l^2 =36.88, p=0.21 for the ToL; Q=2.54, l^2 =21.34, p=0.28 for the ToH), and the two groups differed significantly from each other in their effect sizes (Q_{bet} =4.41, p=0.04; g=-0.904, 95% CI -1.137 to -0.671 for the ToH; g=-0.486, 95% CI -0.799 to -0.174 for the ToL) (Fig. 4b).

The meta-regression analysis indicated that two tests (the IED ed and CPT) were significantly associated with demographic variables. Younger patients showed more pronounced impairment than did their age-matched controls on the IED ed (z=3.122, p=0.002). Additionally, deficits on the CPT were more pronounced with inclusion of fewer male patients (z=3.278, p=0.001). In terms of clinical variables, symptom severity, as measured by YBOCS total scores, was

significantly related to performance on the category fluency test (z=-2.928, p=0.003), with patients with severe symptoms showing more impairment on this test than those with less severe symptoms. The percentages of medicated patients and patients with co-morbid disorders were not significantly associated with performance on any of the tasks.

Discussion

The purpose of this meta-analysis was to provide a comprehensive overview of the magnitude of cognitive deficits in OCD. To our knowledge, this is the first meta-analysis examining the results of various neuropsychological tests addressing different cognitive domains with respect to OCD. Consistent with earlier reviews, we found impairments on the tasks measuring visuospatial memory, verbal memory, executive function, verbal fluency and processing speed among patients with OCD, with effect sizes ranging from -0.7 to -0.3 compared with healthy control subjects. No deficits were found in auditory attention, as measured by the digit span test. According to the conventional interpretation of effect size (Cohen, 1988), visuospatial memory, visual organizational skill and planning ability showed medium-to-large effects, whereas set-shifting ability, design fluency, cognitive inhibition, verbal memory, verbal fluency and processing speed showed medium or small-to-medium effect size. When the tasks were grouped according to cognitive domain, visuospatial memory showed medium-to-large effects, whereas executive function, verbal fluency, processing speed and verbal memory showed small-to-medium effects. We found no significant impairment in the domain of attention among patients with OCD. Patients with OCD appear to have broad, albeit not severe, cognitive dysfunction but preserved attentional ability.

The results of the current meta-analysis suggest that impairment in visuospatial memory is more pronounced than are deficits in executive function such as set shifting and inhibition in patients with OCD. The small number of studies using certain tests and the small sample sizes included in some studies require that additional research be conducted before definite conclusions are reached. However, this meta-analysis underscores the significance of findings reflecting OCD patients' difficulty in accurately recalling information about visual stimuli. This impairment was more profound when the configurations of complex figures were recalled than when sequences of spatial locations were retrieved. It has been assumed that memory deficits in patients with OCD may be secondary to defective organizational strategies (Savage et al.

1999; Melloni et al. 2012). Several studies found that organizational skill, which is required for efficient information processing, was impaired in patients with OCD and completely or partially mediated problems with non-verbal (Savage et al. 1999, 2000; Shin et al. 2004) and verbal memory (Savage et al. 2000). The current meta-analysis provides evidence about the relatively strong magnitude of effects related to visual organizational skill. Impaired performance seemed to be more pronounced in tasks involving the organization of complex visual configurations than in those involving the assembly of simple diagonal patterns. Additionally, this disability in organizational skill seems to be more evident in tasks that require processing of visuospatial than of verbal information, given the smaller effect size of semantic clustering on the verbal learning test. However, it should be noted that the deficit in visual memory was slightly larger in magnitude than was the deficit in visual organizational skill. This may imply that poor performance with regard to visuospatial memory is not explained by a single problem involving organizational skill.

Regarding executive function, the magnitude of the deficit differed among tasks, but in no case was it very large. A relatively large effect size was found for planning ability. However, as we included studies measuring excessive numbers of moves, many studies that used different outcome variables (n=17) were excluded from the analysis. Thus, our interpretation of these results should be considered with caution. It is somewhat surprising that executive function, in particular set shifting and inhibition, which have been considered core deficits in OCD (Chamberlain et al. 2005), had relatively small effect sizes. A considerable body of research using neuroimaging techniques with individuals with OCD has investigated executive function in OCD patients by employing neuropsychological tests known to be sensitive to abnormalities in certain brain regions, including the orbitofrontal cortex, anterior cingulate cortex and basal ganglia (Melloni et al. 2012). However, the neuropsychological findings are inconsistent, and our meta-analysis revealed that impairment in tasks involving set shifting and inhibition (i.e. alternation tests and the IED, Stroop, TMT, WCST) was moderate (medium or small to medium).

The subgroup analysis revealed that the use of different forms of tests explained a significant proportion of the heterogeneity in the effect estimates for the WCST and the ToL/ToH. The computerized version of the WCST appears to be more sensitive than the classic method for identifying deficits in patients with OCD. Similarly, the ToH seems to be more sensitive for detecting planning problems in these patients. Although additional studies are needed to draw conclusions, these results imply that the selection of the method of testing and the form of the test may be important considerations in efforts to detect neurocognitive dysfunction in OCD.

Our examination of confounding moderator variables revealed that age and sex significantly contributed to the variability in only two tests (the IED ed and CPT) and that educational level had no effect. These results suggest that cognitive deficits are not typically moderated by demographic variables in OCD. In terms of clinical variables, severe symptoms were associated with deficits in category fluency, whereas the prevalence of medicated patients or patients with co-morbid psychiatric disorders did not influence any outcome variables. However, these results should be interpreted with caution, as many studies included in this main meta-analysis did not report the relevant clinical information. Moreover, some important potential clinical moderators that have been proposed to influence cognitive functioning in OCD could not be analysed due to the small number of studies that reported the relevant information. Level of depressive symptoms, symptom subtypes, age at onset and duration of illness have been considered as possible moderators affecting cognitive functioning in OCD (Kuelz et al. 2004). These important moderators are needed to be investigated further and to be controlled before a definite conclusion can be made.

The limitations of this meta-analysis should be addressed. First, studies using non-standardized cognitive tests were excluded because the psychometric properties of such tests have not been established. Second, the classification of individual tasks into cognitive domains was not based on reliable criteria, although it was based on existent psychometric evidence.

In conclusion, this meta-analysis indicated that patients with OCD experience significant impairments in visuospatial memory, executive function, verbal memory, verbal fluency and processing speed, whereas the attentional ability of these individuals is relatively preserved. Although the magnitude of the deficits is, in general, not large, visuospatial memory, visual organizational skill and planning ability appear to be the most impaired areas in patients with OCD. Different test forms and methods of testing probably influence the performance of patients with OCD, indicating the need to carefully select the form of each test and methods of testing used. Further exploration of the effects of various clinical variables on cognitive functioning in patients with OCD and additional investigation of whether the cognitive dysfunction associated with this disorder differs from or overlap with that in other anxiety disorders are needed.

Supplementary material

For supplementary material accompanying this paper visit http://dx.doi.org/10.1017/S0033291713001803.

Acknowledgements

This research was supported by the Mid-carrier Research Program through National Research Foundation grants funded by the Ministry of Education, Science and Technology, Republic of Korea (20110015639).

Declaration of Interest

None.

References

Abbruzzese M, Bellodi L, Ferri S, Scarone S (1995). Frontal lobe dysfunction in schizophrenia and obsessive– compulsive disorder: a neuropsychological study. *Brain and Cognition* **27**, 202–212.

- Abbruzzese M, Ferri S, Scarone S (1997). The selective breakdown of frontal functions in patients with obsessive– compulsive disorder and in patients with schizophrenia: a double dissociation experimental finding. *Neuropsychologia* 35, 907–912.
- Abramovitch A, Dar R, Schweiger A, Hermesh H (2011). Neuropsychological impairments and their association with obsessive–compulsive symptom severity in obsessive– compulsive disorder. *Archives of Clinical Neuropsychology* 26, 364–376.

Aigner M, Sachs G, Bruckmuller E, Winklbaur B, Zitterl W, Kryspin-Exner I, Gur R, Katschnig H (2007). Cognitive and emotion recognition deficits in obsessive-compulsive disorder. *Psychiatry Research* 149, 121–128.

Aycicegi A, Dinn WM, Harris CL, Erkmen H (2003). Neuropsychological function in obsessive–compulsive disorder: effects of comorbid conditions on task performance. *European Psychiatry* 18, 241–248.

Bohne A, Savage CR, Deckersbach T, Keuthen NJ, Jenike MA, Tuschen-Caffier B, Wilhelm S (2005). Visuospatial abilities, memory, and executive functioning in trichotillomania and obsessive-compulsive disorder. *Journal of Clinical and Experimental Neuropsychology* **27**, 385–399.

Boone KB, Ananth J, Philpott L, Kaur A (1991). Neuropsychological characteristics of nondepressed adults with obsessive–compulsive disorder. *Neuropsychiatry, Neuropsychology and Behavioral Neurology* **4**, 96–109.

Borges MC, Braga DT, Iego S, D'Alcante CC, Sidrim I, Machado MC, Pinto PS, Cordioli AV, do Rosario MC, Petribu K, Mendlowicz MV, Mari JJ, Miguel EC, Fontenelle LF (2011). Cognitive dysfunction in post-traumatic obsessive–compulsive disorder. Australian and New Zealand Journal of Psychiatry 45, 76–85. Bucci P, Galderisi S, Catapano F, Di Benedetto R, Piegari G, Mucci A, Maj M (2007). Neurocognitive indices of executive hypercontrol in obsessive-compulsive disorder. *Acta Psychiatry Scandinavica* 115, 380–387.

Cavallaro R, Cavedini P, Mistretta P, Bassi T, Angelone SM, Ubbiali A, Bellodi L (2003). Basal-corticofrontal circuits in schizophrenia and obsessive–compulsive disorder: a controlled, double dissociation study. *Biological Psychiatry* 54, 437–443.

Cavedini P, Cisima M, Riboldi G, D'Annucci A, Bellodi L (2001). A neuropsychological study of dissociation in cortical and subcortical functioning in obsessive– compulsive disorder by Tower of Hanoi task. *Brain and Cognition* **46**, 357–363.

Cavedini P, Gorini A, Bellodi L (2006). Understanding obsessive–compulsive disorder: focus on decision making. *Neuropsychology Review* 16, 3–15.

Cavedini P, Zorzi C, Piccinni M, Cavallini MC, Bellodi L (2010). Executive dysfunctions in obsessive–compulsive patients and unaffected relatives: searching for a new intermediate phenotype. *Biological Psychiatry* **67**, 1178–1184.

- Ceschi G, Van der Linden M, Dunker D, Perroud A, Bredart S (2003). Further exploration memory bias in compulsive washers. *Behaviour Research and Therapy* **41**, 737–748.
- Cha KR, Koo MS, Kim CH, Kim JW, Oh WJ, Suh HS, Lee HS (2008). Nonverbal memory dysfunction in obsessive–compulsive disorder patients with checking compulsions. *Depression and Anxiety* 25, E115–E120.

Chamberlain SR, Blackwell AD, Fineberg NA, Robbins TW, Sahakian BJ (2005). The neuropsychology of obsessive compulsive disorder: the importance of failures in cognitive and behavioural inhibition as candidate endophenotypic markers. *Neuroscience and Biobehavioral Reviews* **29**, 399–419.

Chamberlain SR, Fineberg NA, Blackwell AD, Clark L, Robbins TW, Sahakian BJ (2007). A neuropsychological comparison of obsessive–compulsive disorder and trichotillomania. *Neuropsychologia* **45**, 654–662.

Choi JS, Kang DH, Kim JJ, Ha TH, Lee JM, Youn T, Kim IY, Kim SI, Kwon JS (2004). Left anterior subregion of orbitofrontal cortex volume reduction and impaired organizational strategies in obsessive-compulsive disorder. *Journal of Psychiatry Research* **38**, 193–199.

Cohen J (1988). Statistical Power Analysis for the Behavioral Sciences, 2nd edn. Lawrence Earlbaum: Hillsdale, NJ.

Cohen Y, Lachenmeyer JR, Springer C (2003). Anxiety and selective attention in obsessive–compulsive disorder. *Behaviour Research and Therapy* **41**, 1311–1323.

Deckersbach T, Savage CR, Reilly-Harrington N, Clark L, Sachs G, Rauch SL (2004). Episodic memory impairment in bipolar disorder and obsessive–compulsive disorder: the role of memory strategies. *Bipolar Disorders* 6, 233–244.

de Geus F, Denys DA, Sitskoorn MM, Westenberg HG (2007). Attention and cognition in patients with obsessive-compulsive disorder. *Psychiatry and Clinical Neurosciences* **61**, 45–53.

Duval S, Tweedie R (2000). Trim and fill: a simple funnel-plot-based method of testing and adjusting for publication bias in meta-analysis. *Biometrics* **56**, 455–463.

Egger M, Smith GD, Schneider M, Minder C (1997). Bias in meta-analysis detected by a simple, graphical test. *BMJ* **315**, 629–634.

Fenger MM, Gade A, Adams KH, Hansen ES, Bolwig TG, Knudsen GM (2005). Cognitive deficits in obsessive– compulsive disorder on tests of frontal lobe functions. *Nordic Journal of Psychiatry* 59, 39–44.

Gambini O, Abbruzzese M, Scarone S (1993). Smooth pursuit and saccadic eye movements and Wisconsin Card Sorting Test performance in obsessive-compulsive disorder. *Psychiatry Research* **48**, 191–200.

Goncalves OF, Marques TR, Lori NF, Sampaio A, Branco MC (2010). Obsessive–compulsive disorder as a visual processing impairment. *Medical Hypotheses* **74**, 107–109.

Hedges L, Holkin I (1985). *Statistical Methods for Meta-Analysis*. Academic Press: New York, NY.

Henin A, Savage CR, Rauch SL, Deckersbach T, Wilhelm S, Baer L, Otto MW, Jenike MA (2001). Is age at symptom onset associated with severity of memory impairment in adults with obsessive–compulsive disorder? *American Journal of Psychiatry* **158**, 137–139.

Hur JW, Shin NY, Kim SN, Jang JH, Choi JS, Shin YC, Kwon JS (2012). Do pathological gambling and obsessive-compulsive disorder overlap? A neurocognitive perspective. CNS Spectrums 17, 207–213.

Hwang SH, Kwon JS, Shin YW, Lee KJ, Kim YY, Kim MS (2007). Neuropsychological profiles of patients with obsessive–compulsive disorder: early onset *versus* late onset. *Journal of the International Neuropsychological Society* **13**, 30–37.

Kitis A, Akdede BB, Alptekin K, Akvardar Y, Arkar H, Erol A, Kaya N (2007). Cognitive dysfunctions in patients with obsessive-compulsive disorder compared to the patients with schizophrenia patients: relation to overvalued ideas. *Progress in Neuro-Psychopharmacology and Biological Psychiatry* **31**, 254–261.

Kuelz AK, Hohagen F, Voderholzer U (2004). Neuropsychological performance in obsessive– compulsive disorder: a critical review. *Biological Psychology* 65, 185–236.

Lacerda AL, Dalgalarrondo P, Caetano D, Haas GL, Camargo EE, Keshavan MS (2003). Neuropsychological performance and regional cerebral blood flow in obsessive-compulsive disorder. *Progress in Neuro-Psychopharmacology and Biological Psychiatry* 27, 657–665.

Lawrence NS, Wooderson S, Mataix-Cols D, David R, Speckens A, Phillips ML (2006). Decision making and set shifting impairments are associated with distinct symptom dimensions in obsessive-compulsive disorder. *Neuropsychology* **20**, 409–419.

Lucey JV, Burness CE, Costa DC, Gacinovic S, Pilowsky LS, Ell PJ, Marks IM, Kerwin RW (1997). Wisconsin Card Sorting Task (WCST) errors and cerebral blood flow in obsessive-compulsive disorder (OCD). *British Journal of Medical Psychology* **70** 403–411.

Melloni M, Urbistondo C, Sedeño L, Gelormini C, Kichic R, Ibanez A (2012). The extended fronto-striatal model of obsessive compulsive disorder: convergence from event-related potentials, neuropsychology and neuroimaging. *Frontiers in Human Neuroscience* 6, 259.

Menzies L, Chamberlain SR, Laird AR, Thelen SM, Sahakian BJ, Bullmore ET (2008). Integrating evidence from neuroimaging and neuropsychological studies of obsessive–compulsive disorder: the orbitofronto-striatal model revisited. *Neuroscience and Biobehavioral Reviews* 32, 525–549.

Muller J, Roberts JE (2005). Memory and attention in obsessive–compulsive disorder: a review. *Journal of Anxiety Disorders* 19, 1–28.

Murphy R, Nutzinger DO, Paul T, Leplow B (2004). Conditional-associative learning in eating disorders: a comparison with OCD. *Journal of Clinical and Experimental Neuropsychology* **26**, 190–199.

Nakao T, Nakagawa A, Yoshiura T, Nakatani E, Nabeyama M, Yoshizato C, Kudoh A, Tada K, Yoshioka K, Kawamoto M (2005). A functional MRI comparison of patients with obsessive-compulsive disorder and normal controls during a Chinese character Stroop task. *Psychiatry Research* **139**, 101–114.

Nakao T, Nakagawa A, Yoshiura T, Nakatani E, Nabeyama M, Sanematsu H, Togao O, Yoshioka K, Tomita M, Kuroki T, Kanba S (2009). Duration effect of obsessive–compulsive disorder on cognitive function: a functional MRI study. *Depression and Anxiety* **26**, 814–823.

Nedeljkovic M, Kyrios M, Moulding R, Doron G,
Wainwright K, Pantelis C, Purcell R, Maruff P (2009).
Differences in neuropsychological performance between subtypes of obsessive–compulsive disorder. *Australian New Zealand Journal of Psychiatry* 43, 216–226.

Nielen MM, Den Boer JA (2003). Neuropsychological performance of OCD patients before and after treatment with fluoxetine: evidence for persistent cognitive deficits. *Psychological Medicine* **33**, 917–925.

Olley A, Malhi G, Sachdev P (2007). Memory and executive functioning in obsessive–compulsive disorder: a selective review. *Journal of Affective Disorders* **104**, 15–23.

Penades R, Catalan R, Rubia K, Andres S, Salamero M, Gasto C (2007). Impaired response inhibition in obsessive compulsive disorder. *European Psychiatry* 22, 404–410.

Purcell R, Maruff P, Kyrios M, Pantelis C (1998). Neuropsychological deficits in obsessive-compulsive disorder: a comparison with unipolar depression, panic disorder, and normal controls. *Archives of General Psychiatry* 55, 415–423.

Radua J, van den Heuvel OA, Surguladze S, Mataix-Cols D (2010). Meta-analytical comparison of voxel-based morphometry studies in obsessive–compulsive disorder vs other anxiety disorders. *Archives of General Psychiatry* **67**, 701–711.

Rajender G, Bhatia MS, Kanwal K, Malhotra S, Singh TB, Chaudhary D (2011). Study of neurocognitive endophenotypes in drug-naive obsessive–compulsive disorder patients, their first-degree relatives and healthy controls. *Acta Psychiatrica Scandinavica* **124**, 152–161. Rampacher F, Lennertz L, Vogeley A,

Schulze-Rauschenbach S, Kathmann N, Falkai P, Wagner M (2010). Evidence for specific cognitive deficits in visual information processing in patients with OCD compared to patients with unipolar depression. *Progress in Neuro-psychopharmacology and Biological Psychiatry* 34, 984–991.

Roth RM, Milovan D, Baribeau J, O'Connor K (2005). Neuropsychological functioning in early- and late-onset obsessive–compulsive disorder. *Journal of Neuropsychiatry* and Clinical Neuroscience 17, 208–213.

Savage CR, Baer L, Keuthen NJ, Brown HD, Rauch SL, Jenike MA (1999). Organizational strategies mediate nonverbal memory impairment in obsessive–compulsive disorder. *Biological Psychiatry* **45**, 905–916.

Savage CR, Deckersbach T, Wilhelm S, Rauch SL, Baer L, Reid T, Jenike MA (2000). Strategic processing and episodic memory impairment in obsessive compulsive disorder. *Neuropsychology* **14**, 141–151.

Schmidtke K, Schorb A, Winkelmann G, Hohagen F (1998). Cognitive frontal lobe dysfunction in obsessive–compulsive disorder. *Biological Psychiatry* 43, 666–673.

Segalas C, Alonso P, Real E, Garcia A, Minambres A, Labad J, Pertusa A, Bueno B, Jimenez-Murcia S, Menchon JM (2010). Memory and strategic processing in first-degree relatives of obsessive compulsive patients. *Psychological Medicine* 40, 2001–2011.

Shin MS, Park SJ, Kim MS, Lee YH, Ha TH, Kwon JS (2004). Deficits of organizational strategy and visual memory in obsessive–compulsive disorder. *Neuropsychology* 18, 665–672. Simpson HB, Rosen W, Huppert JD, Lin SH, Foa EB, Liebowitz MR (2006). Are there reliable neuropsychological deficits in obsessive-compulsive disorder? *Journal of Psychiatric Research* 40, 247–257.

Starcke K, Tuschen-Caffier B, Markowitsch HJ, Brand M (2010). Dissociation of decisions in ambiguous and risky situations in obsessive–compulsive disorder. *Psychiatry Research* 175, 114–120.

Trivedi JK, Dhyani M, Goel D, Sharma S, Singh AP, Sinha PK, Tandon R (2008). Neurocognitive dysfunction in patients with obsessive compulsive disorder. *African Journal* of *Psychiatry* **11**, 204–209.

Tukel R, Gurvit H, Ertekin BA, Oflaz S, Ertekin E, Baran B, Kalem SA, Kandemir PE, Ozdemiroglu FA, Atalay F (2012). Neuropsychological function in obsessive– compulsive disorder. *Comprehensive Psychiatry* **53**, 167–175.

Tumkaya S, Karadag F, Oguzhanoglu NK, Tekkanat C, Varma G, Ozdel O, Atesci F (2009). Schizophrenia with obsessive–compulsive disorder and obsessive–compulsive disorder with poor insight: a neuropsychological comparison. *Psychiatry Research* **165**, 38–46.

Veale DM, Sahakian BJ, Owen AM, Marks IM (1996). Specific cognitive deficits in tests sensitive to frontal lobe dysfunction in obsessive-compulsive disorder. *Psychological Medicine* 26, 1261–1269.

Wobrock T, Gruber O, McIntosh AM, Kraft S, Klinghardt A, Scherk H, Reith W, Schneider-Axmann T, Lawrie SM, Falkai P, Moorhead TW (2010). Reduced prefrontal gyrification in obsessive–compulsive disorder. *European Archives of Psychiatry and Clinical Neurosciences* 260, 455–464.