# Patients With Borderline Personality Disorder and Major Depressive Disorder Are Not Distinguishable by Their Neuropsychological Performance: A Case-Control Study

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**Objective:** Patients with borderline personality disorder (BPD) and patients with major depressive disorder (MDD) exhibit a broad range of neuropsychological deficits. Studies in both groups of patients point to differences but also similarities. However, studies that compare both patient groups are missing from the literature. The present study aimed to compare neuropsychological functioning in BPD and MDD patients.

**Method:** Eighteen patients with BPD, 27 patients with MDD, 17 patients with BPD and MDD, and 76 healthy control subjects were included in the case-control study. Patients were treated for their disorders as inpatients of the Clinic of Psychiatry and Psychotherapy Bethel, Ev. Hospital Bielefeld (Bielefeld, Germany). All patients met *DSM-IV* diagnoses as assessed by trained psychotherapists within the first week of their admission. In addition to a comprehensive neuropsychological test battery, the inhibitory control of emotional stimuli was assessed. Data were collected between June 2004 and June 2007.

**Results:** Patients showed only a few impairments and no increased distractibility toward emotionally negative stimuli. Patients with BPD and patients with MDD were not distinguishable by the neuropsychological test results.

*Conclusions:* These data did not support the notion of specific neuropsychological profiles in BPD and MDD. Future research needs to clarify the overlap of symptoms between both disorders. *Prim Care Companion CNS Disord 2011;13(1):e1–e9* 

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Ognitive deficits of patients with borderline personality disorder (BPD) have been systematically investigated for 2 decades. Patients have been shown to exhibit deficits in the domains of executive functions, memory, attention, and visuospatial abilities.<sup>1</sup> Previously, we administered a comprehensive neuropsychological test battery and found that BPD patients exhibited impairments primarily in visual functions.<sup>2</sup> In agreement with these findings, it was concluded that nonverbal functions might be more affected than verbal functions,<sup>1,3</sup> especially in the domain of memory.<sup>4</sup> Some experimental studies in BPD revealed impairments of inhibitory functions,<sup>3,5</sup> especially the inhibition of stimuli with negative valence.<sup>6</sup> Domes et al<sup>7</sup> found BPD patients to show reduced inhibition of negative material in a directed forgetting task and in a negative priming task. However, the performance in the emotional stroop task was not affected. BPD patients in a study by Hurlemann et al<sup>8</sup> displayed enhanced retrograde and anterograde amnesia in response to negative stimuli. The authors concluded that there is a negative emotional response bias with a reduced ability to inhibit negatively valenced stimuli.

Patients with MDD exhibit deficits in the domains of executive functions, memory, and attention as well.9 In an early meta-analysis, Veiel10 concluded that patients with major depressive disorder (MDD) show a dominant deficit in cognitive flexibility/fluency and diffuse impairments in other cognitive domains. Data from our previous studies supported this conclusion.<sup>11,12</sup> Clark et al<sup>13</sup> found impairments in cognitive flexibility but no further neuropsychological deficits in relatives of depressed patients. With regard to fluency, semantic fluency seems to be more impaired than phonological fluency.<sup>14</sup> However, the profile and severity of impairment seem to depend on many factors such as comorbidity, subtype of the disorder, age, medication, and the experience of failure.<sup>15</sup> Recent studies also underline the influence of rumination,<sup>16</sup> motivation,<sup>17</sup> and sleep.<sup>18</sup> Further, experimental investigations indicate that cognitive changes in depression are more obvious with certain features of the neuropsychological tasks such as the consideration of affectively meaningful stimuli.<sup>19-21</sup> As with BPD patients, inhibitory dysfunction in MDD patients is most likely valence specific. In the study of Lau et al,<sup>21</sup> patients with MDD showed deficits in cognitive inhibition. These deficits were most pronounced for negatively valenced stimuli. An inhibition problem with negative distraction corresponds with an attentional bias toward negatively valenced information in MDD

#### **CLINICAL POINTS**

- On average, patients with major depressive disorder and patients with borderline personality disorder show only a few neuropsychological impairments as assessed by standardized neuropsychological tests. However, clinically significant deficits may occur in single cases, and, in particular, in more severely affected patients who were not included in the present study. In addition, clinically significant deficits in everyday life cannot be ruled out.
- Clinicians cannot differentiate patients with major depressive disorder from patients with borderline personality disorder by their neuropsychological performance.

patients. This bias may lead to an enhanced memory for negatively valenced emotional material.<sup>22</sup>

These findings point to neuropsychological differences as well as similarities between BPD and MDD patients. Unfortunately, to our knowledge, there are no studies that compare neuropsychological profiles of both patient groups. Völker et al<sup>23</sup> investigated executive functions in BPD patients as well as in subjects with a lifetime diagnosis of depression and healthy subjects. No differences between the 3 groups were found. However, conclusions are limited because other neuropsychological functions were not considered and acute MDD patients were not investigated. Fertuck et al<sup>24</sup> administered a comprehensive neuropsychological battery in MDD patients with and without BPD as well as in healthy control subjects. Neuropsychological performance did not differ between the patient groups, thus indicating no additional negative effect of BPD in patients with MDD with regard to cognitive functioning. However, since a group with BPD only was missing, no clear conclusions about BPD can be drawn.<sup>24</sup> Thus, the question of a possible neuropsychological overlap in BPD and MDD is still a matter of debate. This question implies important implications. Clinically, knowledge about neuropsychological profiles in BPD and MDD is helpful for diagnostic purposes. From a theoretical point of view, neuropsychological similarities between both disorders would bring up questions about common etiologic pathways.

The present study aimed at a comparison of neuropsychological functioning in BPD and MDD. We hypothesized (1) that patients with BPD would show specific deficits in visual functions, primarily visual memory; (2) that MDD patients predominantly would exhibit deficits in cognitive flexibility and semantic fluency; and (3) that BPD and MDD patients would show an increased distractibility toward emotionally negative stimuli (that is, reduced learning performance with the presentation of emotionally negative stimuli).

#### METHOD

#### Subjects

The case-control study included 27 patients with MDD but without any personality disorder, 18 patients with BPD but without acute MDD or any history of MDD, 17 patients with acute MDD and BPD, and 76 healthy control subjects. Patients were treated for their disorders as inpatients of the Clinic of Psychiatry and Psychotherapy Bethel, Ev. Hospital Bielefeld (Bielefeld, Germany). All patients met DSM-IV diagnoses as assessed by trained psychotherapists within the first week of their admission. Healthy subjects were recruited by advertisements in a local newspaper. Exclusion criteria for participation in the study were further comorbid Axis I disorders apart from anxiety disorders, somatization disorder, substance abuse of more than 6 months ago, and bulimia. In addition, patients with MDD with psychotic symptoms were not included. Healthy control subjects were free of any Axis I or II disorders. The subjects underwent careful clinical examination and were assessed to exclude the following medical conditions: pregnancy, endocrine system disorders, malignant diseases, liver cirrhosis, a history of neurologic disorders with central nervous system involvement, and mental retardation. After a complete explanation of the study, written informed consent was obtained from all subjects. The study was accepted by the institutional review board (University of Muenster Ethics Committee, Muenster, Germany). Data were collected between June 2004 and June 2007.

### Instruments

*Clinical examination.* Psychiatric diagnoses were made using the Structured Clinical Interview for *DSM-IV* (SCID): SCID-I for Axis I disorders and SCID-II for personality disorders.<sup>25</sup> These interviews were applied by trained psychotherapists. The clinical examination also included the assessment of depressive mood using the Beck Depression Inventory (BDI).<sup>26</sup>

#### Neuropsychological Assessment

*Memory.* Neuropsychological assessment was performed by a neuropsychologist (C.M.) and a master's student. They were trained and supervised by a trained neuropsychologist (T.B.).

<u>Visual learning with and without distraction.</u> Subjects learned 3 lists (A, B, C) of 15 simple designs. The A and B sets of learning stimuli were drawn from the Rey Visual Design Learning Test.<sup>27</sup> List C was developed by the authors themselves using variations of the stimuli from lists A and B. Pretests of the sets used showed comparable results for all item lists. In contrast to the standard procedure, each list was presented only 3 times.

All stimuli were presented on a video screen of a standard personal computer using the software Presentation 0.76.<sup>28</sup> After each learning trial, the subjects were asked to draw the figures they remembered on a sheet of paper. The dependent variable was the sum of correctly drawn figures in trials 1-3. Three experimental conditions were presented to all subjects. (1) In order to assess baseline learning performance, subjects learned a design list without distraction. (2) In the first distraction condition, designs of a list were presented alternating with pictures from the International Affective Picture Series (IAPS) of neutral valence.<sup>29</sup> For the 3 learning trials, 45 different IAPS pictures were used. (3) The second distraction condition corresponded to the first distraction condition, but IAPS pictures with negatively emotional valence were used. The ratings of emotional valence from the neutral and negative pictures differed as indicated by results of an analysis of variance (ANOVA) ( $F_{1.88}$  = 723.83, P < .0001). The assignment of the learning conditions (1-3) with the design lists (A-C) was randomized, and the lists were displayed in the same order (A-B-C).

In the distraction conditions, presentation started with a 350-ms interval presenting a black screen, followed by an interval of 1,000 ms in which a distractor was presented. Then, for a 350-ms interval, a black screen was shown, followed by a 1,000-ms presentation of a learning stimulus (design). The baseline condition was comparable, but instead of the distractor, a white screen was presented (1,000 ms). Thus, in all conditions, the interval between 2 learning stimuli was 2,700 ms, maintaining the duration of 1 learning trial by 40.5 seconds in total.

The Complex Figure Test (CFT)<sup>30</sup> was applied for the additional assessment of visual memory. Subjects had to recall and draw a complex figure that they had previously been shown and had copied 30 minutes before.

Logical memory. In the subtest logical memory of the Wechsler Memory Scale-Revised,<sup>31,32</sup> subjects had to recall 2 short stories as accurately as possible. Recall performance was assessed immediately after each story was heard (immediate recall) and after 20 minutes (delayed recall). Working memory with and without interfering stimuli. Immediate visual memory spans were assessed by the Corsi Block Tapping Test.<sup>33</sup> The examiner tapped a series of blocks and then asked the subjects to tap the blocks in the same order. These blocks were irregularly arranged on a board. Additionally, a modified version of the Corsi Block Tapping Test, the Block Suppression Test,<sup>34</sup> was administered. Subjects were asked to reproduce only every second block beginning with the first block from a series of blocks tapped by the examiner.

Immediate verbal memory spans were assessed by the digit span forward subtest from the Wechsler Memory Scale-Revised. Subjects had to repeat a series of digits in a given order. The number of correctly recalled digit spans was assessed. In addition, the Digit Suppression Test <sup>34</sup> was administered. In the Digit Suppression Test, only every second digit of a series of orally presented digits had to be reproduced, beginning with the first digit. The number of correctly recalled digit spans was assessed.

Attention. Reaction time was assessed by means of the subtest "alertness" of the computerized Test-Battery of Attentional Performance (Testbatterie zur Aufmerksamkeitspruefung).35 Subjects had to press a button as fast as possible after a cross appeared on the screen. The subtest go/no-go assessed response selection and response inhibition. Two different crosses-1 target and 1 distractor-were presented in random order. The subjects had to respond to the target as quickly as possible. For the assessment of divided attention, in the divided attention subtest, subjects had to respond to visually and auditorily presented targets. Visuomotor tracking was assessed by means of the Trail Making Test, part A.<sup>36</sup> In the Trail Making Test, part A, subjects had to connect 25 numbers as quickly as possible. The Frankfurt Attention Inventory (Frankfurter Aufmerksamkeits-Inventar<sup>37</sup>) was used to assess speed of visual scanning and selective attention. In this cancellation task, 2 critical stimuli consisting of a shape (square or circle) and dots (2 or 3) had to be crossed out among different distractors.

<u>Executive functions.</u> Lexical verbal fluency was assessed by requiring the subject to name as many words as possible with the initial letters F, A, and S. One minute is given for each letter. In the semantic verbal fluency task, subjects had to name as many animals as possible within 1 minute. Cognitive flexibility was assessed by means of the Trail Making Test, part B.<sup>36</sup> This subtest required subjects to connect a series of numbers and letters in an alternating manner (1 to A, A to 2, 2 to B, etc). The test Logical Thinking of the *Leistungspruefsystem*<sup>38</sup> was also administered. This test consists of 40 items with each item consisting of a series of digits and letters arranged with 1 exception according to a basic rule. The test required the subject to identify the wrong element and included a time limit of 8 minutes.

	Healthy	BPD	MDD	BPD/MDD
	Controls	Group	Group	Group
Variable	(n = 76)	(n = 18)	(n = 27)	(n = 17)
Comorbidity, n (%)				
Anxiety disorders		8 (44)	12 (44)	12 (71)
Somatization disorder		1 (6)	2(7)	
Alcohol abuse, remitted		1 (6)		1 (6)
Bulimia		2 (11)		3 (18)
Dysthymia		5 (28)		2 (12)
Medication, n (%)				
Selective serotonin reuptake inhibitors		2 (11)	11 (41)	7 (41)
Tricyclics		2 (11)	4 (15)	2 (12)
Lithium		1 (6)	1 (4)	1 (6)
Other antidepressants		1 (6)	2(7)	1 (6)
Neuroleptics		3 (17)	5 (19)	4 (24)
Beta-blocker		1 (6)		
Benzodiazepines		1 (6)		
Antiepileptic medication			3 (11)	
Beck Depression Inventory score, mean (SD)	2.8 (3.2) <sup>a</sup>	21.6 (14.1)	19.5 (9.4)	27.3 (10.3)

Table 1. Acute Comorbidity, Medication, and Severity of Depression in Healthy Controls, Patients With Borderline Personality Disorder (BPD), Patients With Major Depressive Disorder (MDD), and Patients With BPD and MDD

<u>Visuospatial abilities.</u> Construction was assessed by means of the Complex Figure Test.<sup>30</sup> A complex figure had to be copied as accurately as possible.

#### Data Analysis

All statistical analyses were performed using the Statistical Package for the Social Sciences 14.0 (SPSS 14.0, IBM Corporation, Somers, New York). The level of significance was set at P < .05 and 2-tailed for all analyses. The effect of distraction on visual learning was investigated by a repeated measure 3 (learning condition: no distraction, neutral distraction, negative distraction) × 4 (group: healthy subjects, MDD, BPD, MDD/BPD) analysis of covariance (ANCOVA) with age as covariate. Post hoc tests were calculated if indicated.

For the analysis of the neuropsychological profile, all other group differences were compared for explorative purposes with ANCOVAs and age as covariate. We used Pearson correlations to investigate associations between depression, as assessed by the BDI, and neuropsychological performance. Given the large number of tests and a relatively small sample size, a strict  $\alpha$  correction (eg, Bonferroni) would have resulted in a huge  $\beta$  error. We therefore decided to regard these outcomes strictly as exploratory. Post hoc tests were calculated if indicated. Demographic and clinical data were analyzed by ANOVAs or  $\chi^2$  tests.

# RESULTS

#### Sample Characteristics and Clinical Data

The mean age of the MDD patients was 38.0 years (SD = 13.9). All other patient groups were younger

( $F_{3,134}$  = 4.4, P = .005), and age did not differ between BPD patients (mean = 28.4, SD = 11.6), BPD/MDD patients (mean = 28.9, SD = 8.6), and healthy subjects (mean = 29.2, SD = 11.2). Most subjects had completed secondary school (*Sekundarstufe II*, 12 or 13 years of German basic school education). Groups did not differ with regard to their school education (healthy subjects: n = 6 [9 years, *Hauptschule*], n = 21 [10 years, *Realschule*], n = 49 [12/13 years, *Abitur*]; BPD: n = 1, n = 5, and n = 12, respectively; MDD: n = 7, n = 5, and n = 14; and BPD/ MDD: n = 4, n = 7, and n = 6). Distribution by sex was comparable in all 4 groups, with 61% women (n = 11) in the BPD group, 59% women (n = 16) in the MDD group, and 64% women (n = 49) in the healthy subject group.

Severe comorbidity was excluded, but as expected, patients in all groups suffered from some comorbid Axis I disorders, primarily in the form of anxiety disorders (Table 1). Medication is also shown in Table 1.

With regard to the applied questionnaires, groups differed on the BDI ( $F_{3,132} = 74.9$ , P < .001) due to the low scores of the healthy controls (Table 1). Significant differences were not revealed between the 3 clinical groups for the BDI. For the 3 clinical groups, the results of the BDI indicate a relevant burden with symptoms of depression.

#### **Visual Learning With and Without Distraction**

ANCOVA revealed that the 4 groups differed in their visual learning performance ( $F_{3,125}$  = 4.1, P = .008) due to a superior performance by the healthy subjects. However, post hoc analysis revealed no group differences between the 3 clinical groups. Compared to healthy subjects,

Table 2. Visual Lear Healthy Controls, P Disorder (BPD), Pa (MDD), and Patient	atients Wit tients With	h Borderli Major Dej	ne Person pressive Di	ality
	Healthy	BPD	MDD	BPD/MI

	Healthy	BPD	MDD	BPD/MDD		
	Controls	Group	Group	Group		
Variable	(n = 76)	(n = 18)	(n = 27)	(n = 17)		
Without distraction	27.3 (7.0) <sup>b</sup>	23.2 (7.2)	23.6 (5.9)	22.9 (7.3)		
Neutral distraction	25.3 (7.2) <sup>b</sup>	21.4 (7.7)	21.6 (6.1)	21.1 (6.2)		
Negative distraction	<b>22.1 (6.9)</b> <sup>b</sup>	17.1 (6.2)	17.4 (7.4)	18.1 (8.6)		
<sup>a</sup> Data are presented as mean (SD). <sup>b</sup> Bolding indicates > the 3 clinical groups: $F_{0,eee} = 4.1$ P = 008						

BPD patients ( $F_{1,88}$  = 7.0, P < .010) and BPD/MDD patients ( $F_{1,87}$  = 5.9, P < .018) performed worse. MDD patients showed no deficits compared to healthy subjects. However, since the 3 patient groups did not differ, no specificity of deficits can be claimed for the BPD group; therefore, hypothesis 1 was not confirmed (Patients with BPD show specific deficits in visual functions.).

Subjects performed better when no distraction or a neutral distraction was presented; subjects performed worse when presented with negative distraction (main effect "condition":  $F_{2,124}$ =7.4, P<.001; without distraction vs neutral distraction: not significant; and neutral distraction vs negative distraction:  $F_{1,125}$ =12.3, P<.001). However, hypothesis 3 (patients are disproportionally more distracted by emotionally relevant distractors) was not confirmed (no group × condition interaction). All means and standard deviations are given in Table 2.

# **Further Neuropsychological Test Results**

Comparison of the groups regarding their performance on other neuropsychological tests did not show group differences except in construction (copy of Rey's Complex Figure:  $F_{3,132} = 6.0$ , P = .001) and verbal learning (logical memory, immediate recall:  $F_{3,131} = 3.0, P = .031$ ). For construction, post hoc analysis showed an inferior performance of the BPD/MDD group compared to healthy subjects ( $F_{1.89}$  = 14.8, P < .001) and compared to the MDD patients ( $F_{1,41} = 8.8$ , P = .005) but not compared to the BPD patients (Table 3). BPD patients did not differ from the other groups; therefore, hypothesis 1 (BPD patients show specific deficits in visual functions) was not supported by the data. For verbal learning, post hoc analysis revealed that MDD patients performed worse than healthy subjects ( $F_{1,99} = 7.9, P = .006$ ) but did not perform worse than the other patient groups; that is, poor performance on verbal learning was not specific to the MDD patient group. The BPD and BPD/ MDD patient groups' verbal learning performance was similar to the healthy subjects' performance.

# Associations Between Clinical Symptoms and Neuropsychological Test Results

We did not find any associations between the severity of depression (as assessed by the BDI) and neuropsychological performance (Table 4).

# Control of Comorbidity in the BPD and MDD Samples

In order to examine whether the reduced performances of the BPD and MDD group were due to comorbid disorders, comorbidity-defined subgroups (BPD patients with Axis I comorbidity vs BPD patients without, MDD patients with comorbidity vs MDD patients without) were compared with regard to the parameters with inferior performance of the patients (BPD: visual learning, MDD: verbal learning) via *t* tests. Subgroups did not differ in these exploratory analyses; that is, comorbid disorders were not a plausible explanation for reduced performances.

# DISCUSSION

To our knowledge, this is the first study comparing the neuropsychological profiles of subjects with BPD, MDD, and BPD/MDD and healthy subjects. The main finding of the study is that the patient groups did not show a specific profile of neuropsychological deficits. In general, patient groups showed only a few impairments. Even the distractibility toward emotionally negative stimuli did not differ between the clinical groups and healthy subjects.

With regard to BPD, we found that visual memory was in fact impaired. This finding is in line with many other studies (eg, Swirsky-Sacchetti et al<sup>39</sup>) and with the conclusion of LeGris and van Reekum<sup>4</sup> that visual memory is among the most reported deficits in BPD. A reduced volume of the hippocampus is a wellconfirmed finding in BPD and may be associated with memory deficits.<sup>40,41</sup> Interestingly, the study of Irle et al<sup>41</sup> documented a visual memory deficit that was correlated to a volume reduction of the right hippocampus. Apart from visual memory, we did not find further visual deficits in BPD patients; construction, visual working memory, and other nonverbal neuropsychological functions were not impaired. These findings contrast with some other studies (eg, O'Leary et al<sup>42</sup>). It might be speculated that the well-selected study patients without severe comorbidity do not present such deficits. In accordance with this interpretation, patients with BPD and comorbid MDD did show deficits in construction. In general, study samples without substantial comorbidity and medication (eg, Kunert et al43) rarely present neuropsychological deficits, whereas other studies (eg, Monarch et al<sup>44</sup>) that also included patients with substantial comorbidity and medication showed a broader range of impairments. In addition, the small sample size of our study might have prevented detection of subtle

	Healthy Controls	BPD Group	MDD Group	BPD/MDD Group (n=17)	
Variable	(n=76)	(n=18)	(n=27)		
Memory					
Logical memory (immediate)	33.0 (7.1)	30.1 (4.8)	28.1 (6.2) <sup>b</sup>	30.2 (8.6)	
Logical memory (delayed)	28.8 (7.8)	27.2 (5.9)	24.8 (7.7)	25.7 (9.2)	
Complex Figure Test (recall)	19.7 (7.2)	19.6 (5.5)	20.0 (7.4)	18.9 (4.2)	
Working memory, raw score					
Digit forward	7.6 (1.9)	8.4 (1.7)	7.1 (1.3)	7.6 (1.8)	
Digit suppression	11.1 (3.2)	12.3 (3.8)	10.6 (3.2)	9.8 (3.5)	
Block forward	8.6 (1.6)	8.8 (1.9)	8.3 (2.3)	8.2 (2.3)	
Block suppression	10.1 (3.4)	10.8 (5.6)	9.5 (2.9)	10.1 (3.6)	
Flexibility/fluency					
Trail Making Test, part B (sec)	56.8 (22.9)	65.1 (21.5)	68.7 (25.9)	69.7 (21.9)	
Animals (semantic fluency)	25.8 (5.5)	25.4 (4.9)	23.6 (5.7)	23.7 (6.0)	
FAS (lexical fluency)	37.3 (9.3)	36.5 (6.4)	36.9 (10.7)	37.5 (11.9)	
Construction					
Complex Figure Test (copy)	34.4 (1.6)	34.0 (2.3)	34.3 (1.7)	32.2 (4.1) <sup>c</sup>	
Attention					
Alertness (msec)	242 (27)	249 (29)	244 (23)	235 (22)	
Go/no-go (msec)	396 (74)	417 (48)	438 (74)	429 (79)	
Divided attention (msec)	672 (72)	718 (84)	702 (104)	711 (79)	
Frankfurt Attention Inventory	203 (47)	192 (44)			
Trail Making Test, part A (sec)	25.0 (7.8)	26.2 (9.1)			
Reasoning					
Leistungspruefsystem-4	28.5 (4.7)	28.9 (4.0)	26.9 (3.7)	26.4 (5.1)	

Table 3. Further Neuropsychological Test Outcomes in Healthy Controls, Patients With Borderline Personality Disorder (BPD), Patients With Major Depressive Disorder (MDD), and Patients With BPD and **MDD**<sup>a</sup>

<sup>a</sup>Data are presented as mean (SD). <sup>b</sup>Bolding indicates < healthy controls ( $F_{1,99}$  = 7.9, P = .006). <sup>c</sup>Bolding indicates < healthy controls ( $F_{1,89}$  = 14.8, P < .001) and the MDD group ( $F_{1,41}$  = 8.8, P = .005). Abbreviation: FAS = words with the letters F, A, and S (production 1 minute each).

#### Table 4. Beck Depression Inventory Score Correlations Between Depressive Mood and Neuropsychological Performance in Healthy Controls, Patients With Borderline Personality Disorder (BPD), Patients With Major Depressive Disorder (MDD), and Patients With BPD and MDD

	Healthy Controls (n=76)		BPD G	BPD Group		MDD Group		BPD/MDD Group	
Variable			(n=18)		(n=27)		(n=17)		
	r	Р	r	Р	r	Р	r	Р	
Memory									
Visual learning (without distraction)	0.05	.68	-0.14	.61	-0.01	.97	-0.07	.81	
Visual learning (neutral distraction)	-0.08	.48	-0.29	.29	-0.04	.86	-0.01	.99	
Visual learning (negative distraction)	0.01	.91	-0.48	.08	-0.17	.48	0.12	.68	
Logical memory (immediate)	-0.05	.64	-0.06	.84	0.36	.10	-0.04	.88	
Logical memory (delayed)	-0.01	.95	-0.23	.41	0.27	.22	0.04	.90	
Complex Figure Test (recall)	0.04	.72	-0.22	.46	0.33	.13	-0.18	.49	
Working memory, raw score									
Digit forward	-0.08	.48	-0.25	.38	-0.21	.36	0.06	.81	
Digit suppression	-0.07	.54	-0.21	.44	-0.10	.67	-0.08	.75	
Block forward	-0.08	.48	-0.31	.27	-0.05	.82	-0.18	.48	
Block suppression	-0.19	.11	-0.31	.28	-0.15	.51	-0.28	.27	
Flexibility/fluency									
Trail Making Test, part B (sec)	0.06	.59	0.40	.16	0.16	.47	-0.05	.85	
Animals (semantic fluency)	0.00	.99	0.12	.68	0.00	.99	-0.05	.85	
FAS (lexical fluency)	0.08	.49	0.41	.14	0.02	.93	0.06	.82	
Construction									
Complex Figure Test (copy)	-0.02	.89	-0.31	.26	-0.01	.95	0.17	.52	
Attention									
Alertness (msec)	0.17	.17	0.23	.43	0.25	.29	0.17	.53	
Go/no-go (msec)	0.19	.11	-0.40	.16	0.37	.11	0.27	.31	
Divided attention (msec)	0.09	.47	0.04	.88	-0.24	.30	0.41	.12	
Frankfurt Attention Inventory	-0.17	.15	-0.26	.30	-0.08	.71	0.07	.79	
Trail Making Test, part A (sec)	0.04	.77	-0.10	.72	-0.08	.73	-0.02	.93	
Reasoning									
Leistungspruefsystem-4	-0.09	.47	-0.31	.26	0.08	.74	-0.09	.74	
Abbreviation: FAS = words with the letters	F. A. and S (	productio	n 1 minut	e each).					

deficits. However, Table 3 indicates almost identical performances between healthy subjects and BPD subjects regarding construction and further visual functions.

The hypothesis that BPD patients would show an increased distractibility toward emotionally negative stimuli (hypothesis 3) was not supported by the data. In a previous study,<sup>45</sup> we investigated the interference caused by emotional stimuli by using the emotional stroop paradigm. Only BPD patients with comorbid PTSD showed an increased interference to emotionally negative words. The majority of BPD patients in the present study did not suffer from PTSD. Korfine and Hooley<sup>6</sup> and Domes et al<sup>7</sup> demonstrated that BPD patients showed a decreased forgetting rate for negative words using the directed forgetting paradigm. These results might be interpreted as an impaired inhibition of emotionally negative stimuli. However, the paradigm of the present study requires the inhibition of distractors during task performance. In the directed forgetting paradigm, participants are initially asked to learn stimuli, and the instruction to forget them is placed later. Therefore, inhibition processes are relevant at a later stage of the process.

The MDD patients in the present study seem to show a reduced performance in cognitive flexibility and semantic fluency (Table 3). However, the results did not reach statistical significance, perhaps due to sample size and the associated lack of statistical power. However, as mentioned above, the profile of neuropsychological deficits in depression was proven to depend upon additional variables such as the patient's type of affective disorder.<sup>15</sup> Airaksinen et al<sup>46</sup> found that problems in mental flexibility primarily characterize patients with dysthymia. MDD patients and patients with mixed anxiety-depressive disorders instead show impairments in verbal memory. In agreement with these findings, MDD patients in the present study showed a reduced performance in verbal memory. Some authors also discuss the patient's age as an important confound for the presence of executive functions such as fluency. Porter et al stated that "... executive tasks were disproportionately impaired in the older depressed group."47(p119) By contrast, Castaneda et al concluded that "Executive dysfunction seems to be a key factor of young adulthood MDD ...," whereas "Results about verbal memory and learning functions are inconsistent."9(p17) However, as with executive dysfunction, memory impairment is consistent with neurobiological findings in depression, such as alterations of the prefrontal cortex or hippocampus.<sup>48</sup>

As with BPD patients, in MDD patients, we found no increased distractibility toward emotionally negative stimuli. This was surprising, as there is some evidence for an inhibitory dysfunction in depression for emotionally negative stimuli. In the study by Lau et al,<sup>21</sup> MDD patients needed more time to read stories that were presented with emotionally negative distractor words (prose distraction task). Goeleven et al<sup>20</sup> found that depressed patients showed a specific deficit when it came to inhibiting negative information during a priming task with pictures of sad and happy facial expressions. Investigations with the emotional stroop task revealed conflicting results, with some studies showing an increased reaction time specific to emotionally negative stimuli.49,50 Other studies did not show increased reaction times.<sup>51,52</sup> We speculate that the extent of similarities between targets and distractors may account for differences between study results. In the present study, targets and distractors were very easy to distinguish, whereas different stimuli were very similar in the other studies. The combined distraction effect of similarity and emotion might be necessary to create a greater interference in MDD patients.

The results of the present study do not support the view of distinguishable neuropsychological deficits in BPD and MDD patients. In addition and in accordance with Fertuck et al,<sup>24</sup> the results also fail to show that possible negative effects of BPD and MDD add to an inferior performance in the BPD/MDD group. BPD patients showed visual memory deficits, but their performance was not impaired when compared with MDD patients. Similarly, MDD patients showed verbal memory impairment, but the performance was not significantly worse than the performance of BPD patients. The BPD/MDD patients showed an inferior performance in construction when compared to healthy subjects and MDD patients but not when compared to BPD patients. In the majority of the tests applied, the 3 patient groups were not impaired. These results are in agreement with the outcomes of 2 studies<sup>23,24</sup> that also failed to show neuropsychological differences between the patient groups. However, Keilp et al<sup>53</sup> found that early visual information processing is impaired in MDD patients with BPD but not in MDD patients without comorbid BPD. Taken together, the results of the present study indicate that patients with BPD and MDD show a large overlap of neuropsychological performance, even though the study samples were well defined (BPD patients without MDD and MDD patients without BPD). Future research needs to clarify whether the overlap of symptoms point to common etiologic pathways in both disorders.

The main shortcoming of the present study is the small subsample sizes, especially in the BPD and BPD/ MDD groups. Related to this shortcoming, we were not able to perform multivariate statistics or a strict a correction. Therefore, the reported impairments in visual and verbal memory have to be interpreted with caution. Furthermore, the small subsample sizes of each group may have prevented us from detecting small effect sizes. Therefore, we can only conclude that we did not find evidence for large differences between the patient groups. The BPD patients without present and past MDD are hard to find. Therefore, we decided to focus on a rather small but very well-selected sample. Furthermore, most of the patients took psychotropic medication with possible cognitive side effects. Since we did not find any impairment in attentional performance, it is unlikely that our results can be explained by cognitive side effects due to medication. By contrast, it cannot to be ruled out that some antidepressant drugs such as selective serotonin reuptake inhibitors might have a slightly stimulating effect on the neuropsychological performance.<sup>54</sup> Although we controlled for some comorbidity, many patients showed comorbid disorders, primarily anxiety disorders. To control for this factor, we compared BPD patients with and without comorbid Axis I disorders as well as MDD patients with and without comorbid disorders. We found no differences in the performance of these patient subgroups. Therefore, it is unlikely that comorbid diagnoses explain the results of the present study.

In sum, the outcomes of the present study suggest only very few neuropsychological impairments in patients with BPD and MDD. Deficit profiles are not distinguishable between the clinical study samples. Our neuropsychological findings challenge the notion of disorder-specific neuropsychological findings in MDD and BPD patients. However, findings need to be replicated in larger samples in order to search for moderate group differences. The overlap of symptoms between BPD and MDD patients in the present study may point to common etiologic pathways in BPD and MDD. Since studies are missing from the literature that investigate patients with BPD and patients with MDD, the relation between the disorders remains unclear and requires further investigation.

Drug name: lithium (Lithobid and others).

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