Dissociative states in borderline personality disorder and their relationships to psychotropic medication

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Background: According to recent data, dissociation may play an important role in borderline personality disorder (BPD), nevertheless specific influences of psychotropic medication on dissociative symptoms in BPD and their therapeutic indications are largely unknown. The purpose of this study was to assess relationships of dissociative symptoms in BPD patients with levels of psychotropic medication and compare these results with a subgroup of patients with schizophrenia.

Materials and methods: In this study, we investigated 52 BPD patients and compared the results with a control group of 36 schizophrenia patients. In all participants, we assessed actual day doses of antipsychotic medication in chlorpromazine equivalents and antidepressant medication in fluoxetine equivalents. Dissociative symptoms were measured by Dissociative Experiences Scale (DES), and other psychopathological symptoms were measured using Health of the Nation Outcome Scales.

Results: Results indicate that dissociative symptoms measured by DES were significantly correlated with antipsychotic medication (Spearman R = 0.45, P < 0.01) in chlorpromazine equivalents and antidepressant medication in fluoxetine equivalents (0.36, P < 0.01). These relationships between medication and dissociative symptoms were not found in the control group of schizophrenia patients.

Conclusion: The results suggest that levels of antipsychotic medication and antidepressant medication are significantly associated with dissociative symptoms in BPD but not in schizophrenia.

Keywords: dissociation, stress, antipsychotics, antidepressants

Introduction

Dissociation describes mental disintegration related to stress influences,1–4 which leads to alterations of neural activity that may dissociate certain external and internal information out of awareness leading to states of divided consciousness.1,5–8 In this context, dissociation also reflects a cognitive conflict related to implicitly consolidated traumatic memories.2,6 The process of mental disintegration and conflicting information processing according to recent findings is likely closely associated with activity of anterior cingulate cortex (ACC) that is related to detecting a cognitive conflict and selection among competing stimuli.1,9,10 Dissociative processes were also reported to have a significant role in borderline personality disorder (BPD) and other mental diseases, for example, in schizophrenia.11–14 Because of high prevalence of dissociative symptoms in schizophrenia and psychotic-like symptoms in BPD, these disorders...
have significant similarities that may help to understand manifestations of dissociative symptoms and their relationship to medication. 12–14

Recent findings indicate that treatment influences of antipsychotic and antidepressant medication are closely linked to ACC activity, 15–20 which significantly influences mechanisms of conscious awareness and processing conflicting information. 21–23 With the aim to study relationships between dissociative symptoms and antipsychotic and antidepressant medication, we compared occurrence of dissociation in BPD and the control group of schizophrenia patients, and its association with chlorpromazine equivalents of antipsychotic and fluoxetine equivalents of antidepressant medication.

Materials and methods
Participants
The participants were recruited from regular daily treatment programs at the Psychotherapeutic and Psychosomatic Clinic ESET in Prague. The participants had diagnosis of BPD and the control group of patients had diagnosis of schizophrenia. Exclusion criteria were organic illnesses involving the central nervous system, substance, and/or alcohol abuse and mental retardation (IQ Raven lower than 90). 24 Clinical diagnoses were based on DSM-IV criteria. Diagnosis of BPD patients was confirmed using semi-structured interview for BPD SCID-II 25 and diagnosis of schizophrenia patients was reassessed using the Mini-International Neuropsychiatric Interview. 26 For all participants, we calculated actual day doses of antipsychotic medication in equivalents of chlorpromazine (EC) and antidepressant medication in equivalents of fluoxetine (EF). 27–29 The clinical study was approved by Charles University Hospital ethical committee, and all participants provided written informed consent in compliance with the Declaration of Helsinki.

The sample included 52 patients with BPD (14 men and 38 women), mean age 31.5 years; SD=8.6 years; mean period of psychiatric treatment 7.23 years (SD=5.36 years) with average of 2.4 hospitalizations. The control group of schizophrenia patients included 36 participants (18 men and 18 women), mean age 37.7 years; SD=10.32 years, mean period of psychiatric treatment 13.78 years; SD=8.63 years with average of 4.81 hospitalizations.

Psychometric measures
Dissociative symptoms were assessed using Dissociative Experiences Scale (DES). 30,31 DES is a 28-item self-reported questionnaire that evaluates frequencies of various experiences of dissociative phenomena in everyday life. Each item ranges from 0 to 100, and the mean of all item scores is calculated as the DES score. In the present study, we used the Czech version of the DES that similarly as original English version displays high reliability and internal consistency (Cronbach’s alpha 0.92, test–retest reliability after week 0.91). 32,33

Psychotic manifestations in both groups of patients were measured using Health of the Nation Outcome Scales (HoNOS). 34 The scale includes 12 items (overactive, aggressive, disruptive or agitated behavior; non-accidental self-injury; problem drinking or drug-taking; cognitive problems; physical illness or disability problems; problems with hallucinations or delusions; problems with depressed mood; other mental and behavioral problems; problems with relationships; problems with activities of daily living; problems with living conditions; problems with occupation and activities). The assessment includes external evaluation by a mental health professional and the self-reported version for patients. The scale was translated into the Czech language (Cronbach’s alpha 0.797, test–retest reliability after one week 0.85). 35

Data analysis
Statistical evaluation of psychometric measures and doses of psychotropic medication included descriptive statistics and Spearman correlation coefficients. The non-parametric analyses were preferred because DES data did not have normal distribution. All the methods of statistical evaluation were performed using the software package Statistica version 6.

Results
Results show statistically significant correlations of dissociative symptoms measured by DES with the levels of antipsychotic medication (EC) (Spearman R=0.45; P=0.0007 refined Fisher Z=0.51), and with the levels of antidepressant medication (EF) (Spearman R=0.36; P=0.008 refined Fisher Z=0.50) in patients with BPD (Figure 1) but not in schizophrenia patients (for EC Spearman R=0.19; P>0.05, for EF Spearman R=0.07; P>0.05). The results also did not show statistically significant correlations between psychotropic medication and psychotic symptoms measured by HoNOS (for BPD group – EC Spearman R=0.10; P>0.05, EF Spearman R=0.19; P>0.05; for the schizophrenia group – EC Spearman R=0.28; P>0.05, EF Spearman R=−0.28; P>0.05). For the BPD group, we also calculated correlations: between EC and the HoNOS item “problems with hallucinations or delusions” (Spearman R=0.007; P>0.05); between EF and the item of HoNOS “problems with depressed mood” (Spearman
This relationship between medication and dissociative symptoms may be linked to a specific and very important role of stress in BPD. These results suggest that dissociative states represent more significant factor in BPD etiology than in schizophrenia.4,14,36,37

Recent findings indicate that antipsychotic and antidepressant treatment decreases activation of ACC.15–20,38 On the other hand, increased ACC activity is also linked to a detection of cognitive conflict.39–41 These data suggest a hypothesis for further research that decreased ACC activity due to antipsychotic and antidepressant medication in BPD may also decrease conscious awareness of conflicting stressful experiences, which may produce dissociative symptoms measured by DES.

The results also show that dissociative symptoms correlate with the medication equivalents (EC, EF) just in the BPD group but not in the schizophrenia group. Because the patients’ medications were prescribed without a previous knowledge about the level of dissociative symptoms, these results also suggest important hypothesis for further research.

Nevertheless, these results simply may mean that prescribed higher doses of medication do not reflect clinical assessment of symptoms. Consequently, preliminary judgments about relationships between dissociative symptom and assessed medication equivalents can be made. These implications might be reasonable because these correlations between DES and medication were found only in BPD. On the other hand in schizophrenia patients, dissociative and other psychopathological symptoms did not correlate with medication. This finding suggests another hypothesis that the relationship between DES and medication likely does

### Table 1 Comparison of demographic data between BPD and schizophrenia group

|                      | Mean values | Student’s t-test | P-value |
|----------------------|-------------|------------------|---------|
|                      | BPD         | Schizophrenia    |         |
| Age                  | 31.5        | 37.72            | -3.074  | 0.002  |
| Number of hospitalizations | 2.4231       | 4.8056           | -3.0418 | 0.003  |
| Years of psychiatric treatment | 7.2308       | 13.7778          | -4.3866 | 0      |

**Abbreviation:** BPD, borderline personality disorder.
not reflect simple relationship “higher symptoms, higher medication” as usually may be expected. Certain limitations of this study need to be taken into account, it means mainly relatively small sample sizes that may cause a selection bias of patients regarding their comorbidities and polypharmacy effects. Nevertheless, it can be hypothesized that borderline and schizophrenic patients may differ with respect to side effects of psychotropic medication and that dissociative symptoms in BPD can be partially influenced by an iatrogenic effect of pharmacotherapy, especially polypharmacy in BPD and in BPD. Another possible hypothesis is that pharmacotherapy is more helpful for dissociative symptoms in schizophrenia than in BPD. In longitudinal research we plan to continue this research and we expect that certain issues regarding alternative hypotheses and research limitations might be resolved and will enable clinical applications of these findings.

## Conclusion

These results are in agreement with recent recommendations for BPD treatment focusing mainly on psychological therapies and state that no psychotropic drug has a specific marketing authorization for the treatment in BPD. The results also emphasize rigorous diagnostic distinctions of BPD, affective disorders, and schizophrenia. In this context, the results are in agreement with findings indicating an important role of dissociative processes in BPD, which might be hypothetically linked with antipsychotic and antidepressant medication.

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## Disclosure

The authors report no conflicts of interest in this work.

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