European Psychiatry 37 (2016) 1-7

Contents lists available at ScienceDirect

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European Psychiatry

journal homepage: http://www.europsy-journal.com

Original article

# Anxiety symptoms in a major mood and schizophrenia spectrum disorders

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## ARTICLE INFO

Article history: Received 26 January 2016 Received in revised form 7 April 2016 Accepted 12 April 2016 Available online 21 July 2016

Keywords: Anxiety Schizophrenia Bipolar disorder Depression Comorbidity

## ABSTRACT

*Background:* Comorbid anxiety symptoms and disorders are present in many psychiatric disorders, but methodological variations render comparisons of their frequency and intensity difficult. Furthermore, whether risk factors for comorbid anxiety symptoms are similar in patients with mood disorders and schizophrenia spectrum disorders remains unclear.

*Methods:* The Overall Anxiety Severity and Impairment Scale (OASIS) was used to measure anxiety symptoms in psychiatric care patients with schizophrenia or schizoaffective disorder (SSA, n = 113), bipolar disorder (BD, n = 99), or depressive disorder (DD, n = 188) in the Helsinki University Psychiatric Consortium Study. Bivariate correlations and multivariate linear regression models were used to examine associations of depressive symptoms, neuroticism, early psychological trauma and distress, self-efficacy, symptoms of borderline personality disorder, and attachment style with anxiety symptoms in the three diagnostic groups.

*Results*: Frequent or constant anxiety was reported by 40.2% of SSA, 51.5% of BD, and 55.6% of DD patients; it was described as severe or extreme by 43.8%, 41.4%, and 41.2% of these patients, respectively. SSA patients were significantly less anxious (P = 0.010) and less often avoided anxiety-provoking situations (P = 0.009) than the other patients. In regression analyses, OASIS was associated with high neuroticism, symptoms of depression and borderline personality disorder and low self-efficacy in all patients, and with early trauma in patients with mood disorders.

*Conclusions:* Comorbid anxiety symptoms are ubiquitous among psychiatric patients with mood or schizophrenia spectrum disorders, and in almost half of them, reportedly severe. Anxiety symptoms appear to be strongly related to both concurrent depressive symptoms and personality characteristics, regardless of principal diagnosis.

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# 1. Introduction

Anxiety symptoms are conceptualized as anxiety disorders (ADs) when they constitute specified syndromes and are intensive, recurrent, and impede an individual's psychosocial functioning

http://dx.doi.org/10.1016/j.eurpsy.2016.04.007 0924-9338/© 2016 Elsevier Masson SAS. All rights reserved. [1]. ADs are the most common psychiatric conditions in the general population, with typical estimates for lifetime prevalence of 16–28% [2–5]. ADs also commonly co-occur with other psychiatric conditions. For instance, up to 38% of patients with schizophrenia [6], 45% of patients with bipolar disorder [7], and 73% of patients with depression [8] reportedly suffer from a lifetime comorbid AD(s). ADs impair quality of life and are associated with poorer prognosis and outcome of psychotic and affective disorders [9–13]. This is true also for comorbid subthreshold anxiety [14–16]. Thus, careful recognition and proper treatment of comorbid anxiety, either as diagnosable disorders or as subthreshold states, are important in clinical practice.

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Abundant literature on anxiety disorder comorbidity among patients with major mental disorders exists [6,17,18]. The majority of these studies have focused on the presence of specific comorbid disorders [19], rarely reporting on subthreshold anxiety symptoms, even if clinically relevant. Few studies on comorbid anxiety disorders or symptoms have included both uni- and bipolar mood as well as non-affective psychotic disorders, and methodological variations have rendered comparisons of the results difficult. Hence, it remains unclear whether prevalence of anxiety symptoms and their putative risk factors are similar in patients with schizophrenia or schizoaffective disorder (SSA), bipolar disorder (BD), and depressive disorder (DD).

Anxiety and depressive disorders constitute the main internalizing mental disorders [20,21], with a high level of temporal covariation [22]. Recent studies have found that bipolar disorder shares some etiological and pathogenetic connections with the internalizing domain as well [23,24]. The internalizing disorders are likely to share most of their genetic basis [25-27]. The personality trait of high neuroticism is the most significant risk factor for internalizing pathology [28,29] and a likely mediator of the underlying genetic diathesis for these disorders [30]. However, many other putative risk factors also contribute to the anxiety and depressive disorders. These factors include childhood and adolescence psychological trauma [31], low self-efficacy [32,33], borderline personality disorder [34], and negative experiences in close relationships [35]. Some findings indicate that the same factors could also affect the onset of schizophrenia and worsen its outcome [36-39]. However, whether similar covariation of depressive and anxiety symptoms exists and whether the same putative risk factors underlie anxiety in schizophrenia spectrum disorders and internalizing disorders remain unclear.

This study had both clinical and theoretical aims. The clinical aim was to compare the point prevalence of comorbid anxiety symptoms among psychiatric patients with depression, bipolar disorder, and schizophrenia or schizoaffective disorders. We hypothesized that the level of anxiety symptoms in patients with schizophrenia or schizoaffective disorder would be lower since, in contrast to mood disorders, these psychotic disorders are not diagnostically defined by the presence of negative affect as a central pathognomonic feature. The theoretical aim was to investigate the relationships of anxiety symptoms with neuroticism, depressive symptoms, and other putative risk factors. We expected that anxiety symptoms would show a clear association with these factors in patients with mood disorders, and explored whether the same relationships would apply to patients with schizophrenia spectrum disorders, in other words beyond the internalizing domain.

#### 2. Methods

#### 2.1. Setting

The current study was a part of the Helsinki University Psychiatric Consortium (HUPC) study performed in collaboration between the Faculty of Medicine, University of Helsinki; the Department of Psychiatry, Helsinki University Central Hospital; the Department of Health and the Mental Health Unit of the National Institute of Health and Welfare, Helsinki; the Department of Social Services and Health Care, Psychiatric Services, Helsinki; and the Department of Psychiatry, Helsinki City Health Department. The catchment area with 1,139,222 inhabitants in 2012 covered the metropolitan area of Helsinki, including the municipalities of Helsinki, Espoo, Vantaa, Kauniainen, Kerava, and Kirkkonummi. Specialized secondary mental health service is provided to these residents. The study was carried out in 10 community mental health centers, in 24 psychiatric inpatient units, in one day-care hospital, and in two residential communities. The HUPC study was approved by the Ethics Committee of Helsinki University Hospital and the pertinent institutional authorities.

# 2.2. Sampling

Stratified patient sampling was performed from 12 January 2011 to 20 December 2012. Patients were randomly drawn either by identifying all eligible patients on a certain day or week in a unit or from patient lists. Inclusion criteria were age from 18 to 64 years and provision of written informed consent. Patients with mental retardation, neurodegenerative disorders, and insufficient Finnish language skills were excluded. Of the 1361 eligible patients, 610 declined to participate and 304 were lost for other reasons. The final number of participants was 447, yielding a response rate of 33%. For the current study, patients with a principal diagnosis of anxiety disorder, eating disorder, neuropsychiatric disorder, or substance use disorder (n = 47) were excluded from the final analyses due to the low number of patients in each group. The total number of patients, thus, was 400.

#### 2.3. Diagnostic assessment

Diagnostic assessments were made according to the International Statistical Classification of Diseases and Related Health Problems, 10th Revision [40] following the principle of lifetime main diagnosis. The authors (K.A, I.B., M.K., and B.K.) verified the clinical diagnoses given by attending psychiatrists by re-examining information obtained from all available medical records. In cases of any diagnostic uncertainty, the senior research psychiatrists (G.J. and E.I.) were consulted. Altogether, 69 cases were consulted. According to the principal diagnosis, patients were divided into three diagnostic groups: schizophrenia or schizoaffective disorder (SSA, n = 113), bipolar disorder (BD, n = 99), and depressive disorder (DD, n = 188).

# 2.4. Measurement of symptoms and traits

Overall Anxiety Severity and Impairment Scale (OASIS) [41] is a brief, 5-item self-report questionnaire to assess severity and impairment associated with any anxiety disorder, multiple anxiety disorders, or subthreshold anxiety. The authors of the current article translated the OASIS into Finnish, which was then back translated into English and the translation revised in collaboration with the creator of OASIS, Dr. Sonya Norman. The questionnaire includes five questions regarding the frequency and severity of anxiety symptoms as well as anxiety-related avoidance behavior and decreased functioning at home/work/school and in social life. Responses range from zero (no anxiety or anxiety-related issues) to four (extreme anxiety and massive anxiety-related issues). A recommended cut-off score for screening of anxiety disorder is eight points [42]. Cronbach's alpha for OASIS in the total sample was 0.84, and specifically, 0.88 for SSA, 0.86 for BD, and 0.78 for DD patients, showing good internal consistency overall and in the subgroups.

Beck Depression Inventory (BDI) [43] is a 21-item self-report questionnaire for measuring the severity of depression symptoms. The "Short Five" (S5) [44] is a 60-item questionnaire constructed for measuring 30 facets of the Five-Factor Model identified by the NEO (Neuroticism-Extraversion-Openness) Personality Inventory. The current study used six items describing neuroticism (S5N). The S5N scale as well as the other four scales (Extraversion, Openness, Agreeableness, and Conscientiousness) showed good internal consistency (Cronbach's alpha for S5N see below, other values not shown). The Experiences in Close Relationships-Revised questionnaire (ECR-R) [45] is a self-report 36-item measure of adult attachment style on anxiety and avoidance subscales. The General Self-Efficacy scale (GSE) [46] is a self-report 10-item instrument to assess perceived self-efficacy regarding stressful life events. The McLean Screening Instrument for Borderline Personality Disorder (MSI-BPD, hereafter MSI) [47] is a self-report 10-item questionnaire to detect the possibility of borderline personality disorder (BPD). The Trauma and Distress Scale (TADS) [48,49] is a self-report 43-item scale for the assessment of early (childhood and early adulthood) traumatic experiences and distress. All of the scales had at least good internal consistency (Chronbach's alpha for BDI – 0.91; for S5N – 0.85; for ECR anxiety scale – 0.95 and avoidance scale – 0.97; for GSE – 0.93; for MSI – 0.92; and for TADS – 0.80).

# 2.5. Statistical analyses

The differences between nominal sociodemographic variables across diagnostic groups were explored with Chi-square test, and between continuous variables with the Kruskal-Wallis test. Nominal dichotomous variables, such as sex, presence or absence of children, education (primary or secondary and higher), smoking status, and care unit (in- or outpatients) were compared with mean OASIS scores using *t*-tests or Mann–Whitney *U*-tests; for marital status the Kruskal-Wallis test was used. The relationships between the OASIS and continuous variables (age, age of onset of illness, and duration of illness) were tested with bivariate correlation analysis. Age of onset and duration of illness were determined based on time of occurrence of the first symptoms reported by the patients. For investigation of the clinical hypothesis of the study, the differences between both the mean total scores and separate item scores of OASIS across the diagnostic groups the Kruskal-Wallis test was used. Bivariate correlation analysis (BCA; Spearman's coefficient) was used to estimate correlation of OASIS with BDI, S5N, MSI, GSE, TADS, and ECR anxiety and avoidance; analysis was performed for each group of patients separately. In order to test the theoretical hypothesis of the study, linear regression model was built to estimate the association between the OASIS (dependent variable) and measures correlated with it in BCA (independent variables) across all diagnostic groups. These measures were all of the abovementioned variables, with the exception of ECR avoidance. In addition, sex and age were included in the analysis. Separate regression models were constructed for each diagnostic group. As additional analysis and partly to avoid the problem of multicollinearity, regression analysis was performed for all independent variables and then with BDI and S5N excluded one at a time and simultaneously. Statistical significance was set at P < 0.05. Statistical analysis was performed using the Statistical Package for the Social Sciences [50].

# 3. Results

## 3.1. Sociodemographic and background data

Table 1 shows the main sociodemographic characteristics of the sample. The patients were middle-aged and there was no significant difference in mean age between diagnostic groups (P = 0.112). The sex distribution differed markedly, with a preponderance of females in the DD and BD groups, but nearly equal distribution in the SSA group (P < 0.001). SSA patients had a family and children less often than BD and DD patients (P < 0.001). The proportion of childless patients in the DD group was higher than in the BD group. No significant differences in educational level or proportion of smokers were found. Of all the diagnostic groups, the SSA group had a highest proportion of inpatients.

# Table 1

Sociodemographic and background characteristics of the sample.

	SSA		BD	BD		DD			P-value
	n	%	n	%	n	%	n	%	
Number	113	28.2	99	24.8	188	47.0	400	100.0	
Female	54	47.8	63	63.6	146	77.7	263	65.7	$< 0.001^{a}$
Marital status									$< 0.001^{b}$
Married	2	1.8	20	20.2	39	21.0	61	15.4	
Cohabitation	8	7.3	17	17.2	29	15.6	54	13.7	
Divorced	16	14.5	29	29.3	36	19.4	81	20.5	
Widowed	3	2.7	1	1.0	3	1.6	7	1.8	
Unmarried	81	73.6	32	32.3	79	42.5	192	48.6	
Childless patients	97	89.0	58	59.8	130	70.7	285	73.1	$< 0.001^{a}$
Secondary/ higher education	68	61.8	71	71.7	121	65.1	260	65.8	0.307 <sup>a</sup>
Smokers	57	51.8	50	50.5	78	42.2	185	47.0	0.197 <sup>a</sup>
Inpatients	36	31.9	20	20.2	34	18.1	90	22.5	0.028 <sup>a</sup>
	Mean (	(SD)	Mear	n (SD)	Mea	n (SD)	Me	an (SD)	P-value
Age	44.3 (1	2.4)	43.4	(12.3)	41.2	(13.3)	42.0	5 (12.9)	0.112 <sup>b</sup>
Age of onset	30.4 (1	3.1)	34.7	(14.2)	35.2	(14.3)	33.0	0 (14.2)	0.009 <sup>b</sup>
Duration of illness	14.6 (1	3.8)	9.1 (	8.6)	6.3	(4.8)	9.8	(8.7)	0.001 <sup>b</sup>

SSA: schizophrenia or schizoaffective disorder; BD: bipolar disorder; DD: depressive disorder

<sup>a</sup> Chi-square test.

<sup>b</sup> Kruskal–Wallis test.

#### 3.2. Overall Anxiety Severity and Impairment Scale (OASIS)

The mean scores of OASIS (Table 2) from 9.4 to 11.0 were seemingly close to each other, but nevertheless differed significantly (P = 0.040). Of specific subgroups, childless SSA and DD patients had higher OASIS scores (P = 0.001 and P = 0.026, respectively), as did smokers with BD (P = 0.006). Analyses demonstrated no significant relations between OASIS scores and other sociodemographic and background variables (data not shown). Overall, from 40.2% to 55.6% of the patients of all groups reported experiencing anxiety frequently or constantly; from 41.2% to 43.8% felt anxiety as severe or extreme (Table 3). SSA patients felt frequent or constant anxiety less often than BD and DD patients (P = 0.010) and did not avoid anxiety-provoking situations as often as BD and DD patients (P = 0.009). Severe or extreme anxiety interfered with functioning at home, school, and work in 33.9% of SSA, 40.4% of BD, and 40.1% of DD patients (OASIS item 4). The corresponding figures for anxiety-induced impairment in social life and relationships were 35.7%, 33.4%, and 44.3% (OASIS item 5). However, the differences between diagnostic groups in these two last items were not statistically significant.

Table 2	
OASIS scores distributions: comparison between diagnostic groups.	

	SSA ( <i>n</i> = 113)	BD ( <i>n</i> = 99)	DD ( <i>n</i> =188)
Mean <sup>*</sup> (SD)	9.4 (5.5)	10.8 (4.4)	11.0 (4.8)
Percentiles			
10	4.0	4.0	4.0
25	5.0	7.0	8.0
50	10.0	12.0	12.0
75	14.0	14.0	15.0
90	16.0	16.0	17.0

SSA: schizophrenia or schizoaffective disorder; BD: bipolar disorder; DD: depressive disorder.

\* P=0.040 (Kruskal-Wallis test).

Table	3

4

Results of th	e OASIS	questionnaire	items	bv	diagnostic	group

	SSA (n = 113)		BD (n	ı=99)	DD (n=188)	
	n	%	n	%	n	%
How often have you fe						
No anxiety	18	16.1	4	4.0	10	5.3
Infrequent anxiety	21	18.8	15	15.2	29	15.5
Occasional anxiety	28	25.0	29	29.3	44	23.5
Frequent anxiety	32	28.6	41	41.4	76	40.6
Constant anxiety	13	11.6	10	10.1	28	15.0
When you have felt and	xious, ho	w intense	or severe	e was your	anxiety	
Little or None	16	14.3	3	3.0	7	3.7
Mild	18	16.1	20	20.2	35	18.7
Moderate	29	25.9	35	35.4	68	36.4
Severe	42	37.5	31	31.3	68	36.4
Extreme	7	6.3	10	10.1	9	4.8
How often did you avo	id situati	ons, places	s, objects	, or activiti	ies becau	ise of
anxiety or fear						
None	23	20.4	14	14.1	20	10.6
Infrequent	23	20.4	14	14.1	34	18.1
Occasional	38	33.6	29	29.3	54	28.7
Frequent	20	17.7	32	32.3	65	34.6
All the time	9	8.0	10	10.1	15	8.0
How much did your an	xiety into	erfere with	n your ab	ility to do	the thing	gs you
needed to do at work	, at scho	ol, or at ho	ome	-		
None	27	24.1	11	11.1	18	9.6
Mild	17	15.2	21	21.2	36	19.3
Moderate	30	26.8	27	27.3	58	31.0
Severe	26	23.2	32	32.3	52	27.8
Extreme	12	10.7	8	8.1	23	12.3
How much has anxiety	interfere	d with yo	ur social	life and re	lationshi	ps
None	22	19.6	8	8.1	16	8.6
Mild	22	19.6	24	24.2	46	24.6
Moderate	28	25.0	34	34.3	44	23.5
Severe	29	25.9	26	26.3	51	27.3
Extreme	11	9.8	7	7.1	30	16.0

SSA: schizophrenia or schizoaffective disorder; BD: bipolar disorder; DD: depressive disorder.

\* P = 0.010.

\* P=0.009 (Kruskal-Wallis test).

# 3.3. OASIS correlation with other measures

Overall, OASIS correlated mainly with the same scales in all groups (Table 4). The strong correlation between anxiety and depression symptoms was found in each diagnostic group. Noteworthy is that all patients experienced fairly severe depressive symptoms (data not shown). High neuroticism and anxiety correlated strongly in the SSA group and moderately in the BD and DD groups. In all patients, anxiety symptoms had a moderate direct correlation with the symptoms of borderline personality disorder

#### Table 4

Bivariate correlation between OASIS and other rating scales by diagnostic group (Spearman's rank).

-								
		BDI	S5N	MSI	GSE	TADS	ECR anxiety	ECR avoidance
	SSA ( <i>n</i> = 113) BD ( <i>n</i> = 99)	.700 <sup>°°°</sup> .729	.712 .569	.588 <sup>***</sup> .447	448 <sup>****</sup> 398 <sup>****</sup>	.498 <sup>***</sup> .498 <sup>***</sup>	.350 <sup>**</sup> .365 <sup>***</sup>	017 .232
	DD $(n = 188)$	.700	.584	.457	440***	.413	.273	.203

SSA: schizophrenia or schizoaffective disorder; BD: bipolar disorder; DD: depressive disorder; OASIS: Overall Anxiety Severity and Impairment Scale; BDI: Beck Depression Inventory; SSN: "Short Five" Neuroticism Scale; MSI: McLean Screening Instrument for Borderline Personality Disorder; GSE: General Self-Efficacy scale; TADS: Trauma and Distress Scale; ECR: Experiences in Close Relationships; ECR anxiety: ECR questionnaire items 1–18; ECR avoidance: ECR questionnaire items 19–36.

 $P \le 0.05.$ 

 $P \le 0.01.$ 

 $P \le 0.001.$ 

(MSI) and early trauma (TADS), and a weak direct correlation with anxious attachment style (ECR anxiety). Across all the diagnostic groups, patients with more severe anxiety symptoms tended to have a lower self-efficacy level, as there was a moderate inverse correlation between OASIS and GSE. In addition, avoidant attachment style (ECR avoidance) showed a weak direct correlation with anxiety symptoms only in the BD and DD groups.

# 3.4. Regression analysis

Of all the variables, symptoms of depression (BDI) and high neuroticism (S5N) were the most strongly associated with OASIS in different regression models (Table 5). Surprisingly, in the main model with all the variables, neuroticism showed a significant weight in the SSA and DD groups, but not in the BD group. In the same model, depressive symptoms were significantly associated with OASIS in the BD and DD groups. When BDI and S5N were both excluded from the regression model, the symptoms of borderline personality disorder and level of self-efficacy acquired a regression weight in each diagnostic group and the early trauma and distress in the BD and DD groups.

# 4. Discussion

The current study investigated comorbid anxiety symptoms from both clinical and theoretical viewpoints. The clinical aim was to examine the point prevalence and level of comorbid anxiety symptoms across the major psychiatric disorders in specialized

#### Table 5

Clinical correlates for OASIS by diagnosis group (linear regression analysis). The main analysis showed in the first model (analysis with all variables).

	SSA (n = 113)		BD ( <i>n</i> =99	9)	DD ( <i>n</i> =188)	
	В	Sig.	В	Sig.	В	Sig.
Analysis with all	variables					
Sex	845	.396	647	.370	595	.415
Age	005	.900	.037	.219	.036	.127
BDI	.081	.213	.198	.000	.180	.000
S5N	.148	.007	.086	.053	.094	.007
MSI	.388	.084	.008	.966	.214	.152
GSE	007	.934	.072	.330	.007	.913
TADS	.011	.674	.029	.094	.021	.110
ECR anxiety	.024	.214	.002	.879	014	.271
Analysis with BD	I excluded					
Sex	815	.415	749	.368	499	.540
Age	.001	.981	.053	.126	.037	.159
S5N	.184	.000	.145	.004	.144	.000
MSI	.410	.069	.124	.565	.257	.122
GSE	017	.836	.002	.979	106	.123
TADS	.019	.456	.050	.011	.043	.003
ECR anxiety	.025	.208	.000	.981	020	.170
Analysis with S5	N excluded					
Sex	-1.008	.338	888	.222	368	.620
Age	014	.741	.022	.450	.038	.117
BDI	.181	.002	.218	.000	.204	.000
MSI	.585	.011	.169	.331	.382	.007
GSE	090	.257	001	.982	061	.320
TADS	.050	.998	.023	.181	.018	.177
ECR anxiety	.031	.129	.012	.450	007	.564
Analysis with BD	I and S5N e	excluded				
Sex	-1.042	.356	-1.202	.163	104	.903
Age	002	.969	.029	.402	.041	.145
MSI	.812	.001	.436	.030	.544	.001
GSE	189	.018	144	.045	242	.000
TADS	.017	.554	.043	.035	.043	.005
ECR anxiety	.038	.084	.016	.390	010	.511

SSA: schizophrenia or schizoaffective disorder; BD: bipolar disorder; DD: depressive disorder; BDI: Beck Depression Inventory; S5N: "Short Five" Neuroticism Scale; MSI: McLean Screening Instrument for Borderline Personality Disorder; GSE: General Self-Efficacy scale; ECR anxiety: Experiences in Close Relationships questionnaire items 1–18; TADS: Trauma and Distress Scale.

psychiatric care. Overall, almost half of the patients of all diagnostic groups experienced frequently or constantly severe or extreme anxiety. However, anxiety was somewhat less frequent in schizophrenia spectrum disorders (SSA) patients than in their mood disorders counterparts. The theoretical aim was to explore the relationship of anxiety with likely covariates and putative risk factors, and determine whether these are similar across the disorders investigated, which indeed they mostly were.

Strengths of the study include investigation of the similarities and differences in comorbid anxiety symptoms using the same methodology in a relatively large sample (total n 400) of psychiatric patients with different principal diagnoses from the Helsinki metropolitan area psychiatric services. This enabled investigating the covariates and putative risk factors of anxiety symptoms across the major diagnostic groups simultaneously. Anxiety symptoms were measured using the Overall Anxiety Severity and Impairment Scale (OASIS), which have been found to be a valid and reliable brief scale [42]. In addition to frequency and intensity of anxiety symptoms and avoidance due to these symptoms, the OASIS also captures anxiety-related functional and behavioral impairment [51].

Our study had several limitations. First, it was a cross-sectional study, thus not enabling causal inferences regarding risk factors for anxiety symptoms, or any analyses of temporal variations. Second, we used only a self-report measure of anxiety symptoms and did not have interview-based measures of anxiety symptoms. Third, the response rate was only 33%, probably due to sampling conducted during busy routine clinical practice and the length of the survey. However, according to the analysis of representativeness, our sample did not differ from the total patient population regarding age or gender. In terms of other demographic characteristics, our sample corresponded to the large screeningbased Vantaa Depression Study and Jorvi Bipolar Study [18,52]. Fourth, the presence, intensity, and quality of current psychotic symptoms were not measured, and thus, their role in comorbidity of anxiety remains unclear. Fifth, retrospective bias may exist in relation to some measurement scales, as patients may not always recollect past events and symptoms. Sixth, the principal clinical diagnoses were not based on structured interviews, although they were validated by the authors based on patients' psychiatric records. Seventh, the study includes multiple statistical analyses, so problems of multiple testing need to be considered. However, there were two hypotheses and one statistical test for each. The remaining analyses are either presented for descriptive purposes, or to confirm coherence and robustness of the hypothesis-related findings irrespective of methodological details.

The clinical aim of the study was to investigate prevalence and patterns of comorbid anxiety symptoms across the disorders. The mean OASIS total scores in all three subgroups clearly exceeded the cut-off score of eight points, usually indicating presence of an AD [42]. Nearly half of our patients in all groups frequently or constantly experienced severe or extreme anxiety. The proportions of our patients with frequent and severe anxiety were similar to findings of lifetime comorbid AD in the same diagnostic groups in earlier reports [6-8]. However, direct comparison of our results with those of previous studies is difficult due to methodological differences and since the published reports rely mostly on categorically diagnosed AD rather than on anxiety symptoms. Of all three subgroups, the SSA patients reported frequent anxiety and anxiety-related avoidance behavior less often than their mood disorder counterparts. The lower rate of comorbid anxiety symptoms in the SSA group could be explained in several ways. First, more frequent anxiety symptoms in patients with mood disorders could be expected because of strong co-incidence of internalizing disorders [21,24–26] as well as temporal covariation of depressive and anxiety symptoms among them [22,53]. However, virtually all of the patients, irrespective of their principal diagnosis, suffered from clinically significant depressive symptoms, which strongly correlated with anxiety symptoms, albeit more in patients with mood disorders than in those with SSA. Second, the majority of SSA patients were outpatients, and thus, in relatively stable condition. For this reason, they probably less often had florid positive symptoms or primary disorder-induced anxiety symptoms to report [9]. Third, avoidance behavior may be less prominent in SSA patients due to their common withdrawal from social roles, and hence, less frequent exposure to common anxiety-provoking situations [54-56]. Furthermore, these patients often experience negative symptoms, rendering some of them emotionally numb and indifferent to situations that tend to cause anxiety in other populations [57]. Nevertheless, despite the observed subgroup differences, we found comorbid anxiety symptoms to be ubiquitous among psychiatric patients with major mood or schizophrenia spectrum disorders, and in almost half of them, reportedly severe. These findings highlight the importance of the recognition and treatment of comorbid anxiety symptoms.

The theoretical focus of our study was in investigating the clinical correlates of comorbid anxiety symptoms and their potential similarities across major psychiatric disorders. We found numerous quite similar associations; in addition to the strongest correlation of the OASIS score with symptoms of depression (BDI) and neuroticism (S5) in all patients, associations emerged also for low self-efficacy (GSE) and symptoms of borderline personality (MSI) across all diagnostic groups, and for early trauma and distress (TADS) in BD and DD patients. In multivariate regression analyses of all clinical variables, neuroticism in SSA patients was associated with comorbid anxiety symptoms as strongly as in DD patients. Therefore, the personality trait of neuroticism seems to be an underlying factor for comorbid anxiety beyond the internalizing domain, thus possibly also within schizophrenia spectrum disorders.

Presence of depressive symptoms and high neuroticism, thus, persisted as independent covariates for anxiety symptoms in multivariate regression models. There were also other correlates associated with anxiety, but not consistently after controlling for the above two factors. These other correlates were mostly the same across the diagnostic groups, with only TADS not being associated with OASIS in the SSA group. Numerous studies suggest an association between experienced childhood trauma and psychotic and mood disorders [31,37,58]. Early traumatic experiences may be connected to a higher level of neuroticism as well [59,60]. Hence, trauma could potentially contribute to comorbid anxiety as a distal cause as well as a neuroticism-mediated condition. In addition, in our patients self-reported symptoms of borderline personality disorder were associated with anxiety symptoms in all diagnostic groups. This finding is consistent with other studies showing that up to 90% of patients with borderline personality disorder experience comorbid anxiety [34,61]. Probably unsurprisingly, also self-efficacy was inversely associated with the level of anxiety and regardless of the primary diagnoses. Poor self-efficacy appears to be a significant factor in development, severity, and treatment of anxiety disorders [32,62]. Our finding suggests that the same logic applies to comorbid anxiety as a continuum. In short, the broad similarity of correlates across all diagnostic groups supports the view that comorbid anxiety symptoms have numerous common background factors, and thus, could be due to a non-aligned condition rather than a direct consequence of the primary psychiatric pathology. While these associations are interesting, it is important to bear in mind their inconsistent significance in multivariate analyses. Analyses of mediation or moderation were beyond the scope of this study. Overall, the most robust and consistent associations with symptoms of anxiety in all subgroups were those with current depressive symptoms and neuroticism.

# 5. Conclusion

Comorbid anxiety symptoms are highly prevalent among psychiatric patients with major mood or schizophrenia spectrum disorders, and in almost half of them, reportedly severe. The prevalence of symptoms is somewhat higher in the former group than in the latter. In addition, anxiety-related avoidance behavior is less frequent in patients with schizophrenia spectrum disorders. Anxiety symptoms appear strongly related to both concurrent presence of depressive symptoms and personality characteristics, particularly high neuroticism, regardless of the principal diagnosis.

# **Disclosure of interest**

The authors declare that they have no competing interest.

## Acknowledgements

The authors acknowledge the kind help of Dr. Sonya Norman in translation and backtranslation of the Finnish version of the Overall Anxiety Severity and Impairment Scale (OASIS).

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