"REVOLVING DOOR" AND BIPOLAR DISORDERS: A RETROSPECTIVE STUDY IN AN ACUTE INPATIENT UNIT

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SUMMARY

Introduction: The present retrospective study investigated clinical correlates of the revolving door (RD) phenomenon in a population of subjects affected by Bipolar Disorders (BDs).

Subjects and methods: Medical records of subjects with BDs admitted to a psychiatric inpatient unit over a 5-year period of time were retrospectively reviewed and clinical data were extracted into an electronic dataset. "Revolving Door Subjects" (RDS) were defined as those who presented three or more "Revolving Door Hospitalizations" (RDH) during twelve months. Features of RDH were compared with non-RDH in order to identify characteristics associated with RD phenomenon and possible risk factors for readmission. To explore predictors of RDH, a stepwise backword logistic regression model was built, including the variables that were significantly associated with RDH in the bivariate analyses.

Results: In our sample of 176 subjects affected by BDs, 53 (19.9%) RDH were identified. In the RDH group, a higher prevalence of mixed episodes (p=0.029) and medical co-morbidities (p=0.004) was detected. Subjects with repeated hospitalizations were more often committed to psychiatric residential facilities at discharge (p=0.002). Treatment features related to RDH were represented by a higher prescription rate of atypical antipsychotics (p=0.030), benzodiazepines (p=0.001) and antidepressants (p=0.048).

Conclusions: Findings from the present study suggest that the early identification and treatment of medical comorbidities and specific clinical features of BDs may help reducing the RD phenomenon in this population of subjects.

Key words: revolving door - bipolar disorders - affective disorders - re-hospitalizations - serious mental illnesses

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INTRODUCTION

The term "revolving door" (RD) defines subjects who undergo repeated hospitalizations in psychiatric units (Morlino et al. 2011, Garrido & Saraiva 2012). The re-organization of mental health services, which took place during the last decades of the 20th century as a consequence of deinstituzionalization policies, led to the establishment of community-based services, with a gradual reduction of the number of acute and sub-acute inpatient psychiatric beds, thus contributing to the growth of the RD phenomenon (Botha et al. 2010, Di Lorenzo et al. 2016). During the last years, the interest on the topic increased, due to its relevant implications under both a clinical and financial point of view (Morlino et al. 2011). The RD phenomenon was described across different countries, with no significant variations in terms of prevalence, despite disparate organizations of healthcare systems (Botha et al. 2010, Di Lorenzo et al. 2016). Previous studies analyzing possible socio-demographic, clinical and treatment correlates of the RD phenomenon underlined the high burden that repeated hospitalizatons determine in the context of serious mental illnesses (Webb et al. 2007, Peritogiannis et al. 2008, Callaly et al. 2011, Frick et al.

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2013, Moss et al. 2014, Di Lorenzo et al. 2016). Bipolar disorders (BDs) are complex clinical conditions, characterized by high relapse rates and often complicated by frequent hospital readmissions (Um et al. 2020). Male gender, unsatisfactory socio-economic conditions, and global functioning impairment were identified as risk factors for rehospitalizations in BDs within the same year (Hamilton et al. 2016). As for psychopathological characteristics, Bipolar Disorder type II (BD-II) appears to be associated to a lesser extent with RD, whilst impulsivity and aggressive behaviors were linked to higher hospitalization rates (Ballester et al. 2012, Bega et al. 2012, Hamilton et al. 2016). Despite this, predictive factors for psychiatric readmissions in BDs are not yet well understood since most studies analyzed the RD phenomenon in mix-diagnosis samples and existing research on the topic mainly focuses on specific features, e.g. psychopharmacological treatment (Woo et al. 2014, Vigod et al. 2015, Um et al. 2020). Given the relevance of the problem under a clinical and economic point of view, the identification of specific risk factors for RD in subjects affected by BDs claims further attention. As a consequence, the present retrospective study was aimed at investigating clinical correlates of repeated hospitalizations in individuals diagnosed with

BDs, as well as features of psychiatric care that may be related to RD in this population. Such information could help a better identification of at-risk patients, shaping possible treatment strategies and allowing a more efficient distribution of health care resources.

SUBJECTS NAD METHODS

Subjects

The present retrospective cohort study was conducted in the Psychiatric Inpatient Unit of University/General Hospital of Perugia, Umbria, Italy, between June 20th 2011 and June 30th 2016. Subjects affected by BDs according to the latest editions of the Diagnostic and Statistical Manual of Mental Disorders (DSM-IV-TR, DSM-5; American Psychiatric Association 2000, 2013) were considered for inclusion in the study. "Revolving Door Subjects" (RDS) were defined as those who presented three or more "Revolving Door Hospitalizations" (RDH) during twelve months (Morlino et al. 2011, Di Lorenzo et al. 2016).

Methods

Data related to hospitalizations of subjects affected by BDs were extracted from medical charts and subsequently collected in an electronic datasheet for statistical analysis. The collected variables included sociodemographic characteristics of RDS and clinical features related to RDH, encompassing treatmentrelated factors and different modalities of discharge. The twelve-month period following the first hospitalization was analyzed for each subject in order to determine whether they belonged to the RDS group or not. Subsequently, hospitalizations were categorized as RDH and non-RDH. Good Epidemiologic Practice (GEP)-IEA Guidelines (http://ieaweb.org) were followed for proper conduct of epidemiologic research, as well as pertinent national, legal and regulatory requirements. All subjects gave informed consent for the use of personal and clinical data for research purposes during the hospitalizations.

Statistical analyses

A descriptive analysis was performed in order to assess the distributional properties of variables within the considered sample. Features of RDH were compared with non-RDH in order to identify characteristics associated with RD phenomenon and possible risk factors for readmission. Bivariate analyses were performed to compare the two groups, using Chisquare test for categorical variables, and Student's t-test for continuous variables, after assessing the normality of distribution of the variables evaluated. To explore predictors of RDH, a stepwise backword logistic regression model was built, including the variables that were significantly associated with RDH in the bivariate analyses. Odds ratios (OR) with 95% confidence intervals (CI) were assessed for observed associations. Statistical analyses were performed using the Statistical Package for Social Sciences (Statistical Package for Social Science-SPSS, 21.0 version for Windows Inc., Chicago, IL, USA). All p values were two-tailed and statistical significance was set at p<0.05.

RESULTS

The population of the present study consisted of 176 patients, 100 (56.8%) females and 76 (43.2%) males, with a mean age of 45.1±12.61. Among these, 10 (5.7%) of the subjects were RDS for a total of 53 (19.9%) RDH. When comparing clinical features among the two subgroups, significant differences were detected as for hospitalizations reasons. In the non-RDH subgroup a higher prevalence of hypo/manic episodes was depicted (29.9% vs 7.5%, p=0.002), whilst episodes characterized by the presence of mixed features were more frequent among RDH (9.4% vs 2.3%, p=0.029). As for diagnostic features, no differences in psychiatric co-morbidities were shown between the two subgroups. Conversely, in the RDH group a higher prevalence of medical illnesses was detected (47.2% vs 25.8%, p=0.004). At discharge from the hospital, subjects belonging to the RDH population were more frequently commited to psychiatric residential facilities (22.6% vs 7%, p=0.002). When analyzing treatment features, higher rates of benzodiazepines (68.6% vs 41.7%, p=0.001), oral antipsychotics (70.6% vs 52.9%, p=0.034), especially second generation ones (58.8% vs 40.7%, p=0.030), and antidepressants (27.5%) vs 14.6%, p=0.048) were demonstrated to be prescribed in the RDH group (see Table 1).

After performing a stepwise backward multivariate modeling procedure ($\chi^2(5)=38.024$, p<0.001) with RDH as the dependent variable, the model explained between 14% (COX and Snell R Square) and 22% (Nagelkerke R Square) of the variance. Statistical significance persisted for treatment with benzodiazepines (p=0.007, OR=2.60, CI=1.292-5.220), commitment to psychiatric residential facilities at discharge (p=0.013, OR=3.07, CI=1.264-7.436), and the presence of a medical co-morbidity (p=0.049, OR=1.99, CI=1.001-3.957), that were positively associated with RDH, whilst hospitalization due to a hypomanic episode displayed a negative association with RDH (p=0.011, OR=0.24, CI=0.080-0.721).

DISCUSSION

In the present study, the prevalence of the RD phenomenon among subjects affected by BDs appears to be lower than what detected in previous literature (Hamilton et al. 2016). This could be related to different definitions of RD and distinct timeframes analyzed through various

able 1. Comparison between subjects with BDs with and without RDH				
	RDH (n=53, 80.1%)	non-RDH (n=214, 19.9%)	χ^2	р
Admission-related features				
Involuntary hospitalization	15 (28.3)	83 (38.8)	1.584	0.208
Committed from a Mental Health Service	32 (60.4)	114 (53.8)	0.504	0.478
Reasons of hospitalization				
Hypo/manic episode	4 (7.5)	64 (29.9)	10.041	0.002
Mixed episode	5 (9.4)	5 (2.3)	4.130	0.029
Depressive episode	4 (7.5)	13 (6.1)	0.006	0.937
Acute psychosis	6 (11.3)	15 (7)	0.576	0.478
Psychomotor agitation	12 (22.6)	43 (20.1)	0.049	0.825
Self-harm	5 (9.4)	19 (8.9)	0.000	1.000
Suicidal ideation	1 (1.9)	4 (1.9)	0.000	1.000
Social reasons	3 (5.7)	2 (0.9)	2.911	0.055
Diagnostic features				
Substance abuse	7 (13.2)	10 (4.7)	3.857	0.052
Personality disorders	5 (9.4)	33 (15.4)	0.805	0.307
Medical comorbidity	25 (47.2)	55 (25.8)	8.210	0.004
Treatment features				
Antidepressants	14 (27.5)	30 (14.6)	3.920	0.048
Benzodiazepines	35 (68.6)	86 (41.7)	10.801	0.001
Long acting injectable antipsychotics	2 (3.9)	11 (5.3)	0.003	1.000
Mood stabilizers	27 (52.9)	87 (42.2)	1.490	0.222
Oral antipsychotics	36 (70.6)	108 (52.9)	4.476	0.034
First-generation antipsychotics	13 (24.5)	43 (20.1)	0.272	0.602
Second-generation antipsychotics	30 (58.8)	83 (40.7)	4.720	0.030
Discharge-related features				
Community Residential Facility	12 (22.6)	15 (7)	9.765	0.002
Outpatient Mental Health Service	30 (56.6)	147 (68.7)	2.263	0.132
	Mean (SD)	Mean (SD)	t	р
Length of stay	16.64 (10.69)	13.33 (18.26)	-1.266	0.210

Notes: BDs = Bipolar Disorders; nRDH = Non-Revolving Door Hospitalizations; RDH = Revolving Door hospitalizations

studies. Several factors appeared to be associated with repeated hospitalizations in BDs. As for diagnostic correlates, mixed episodes were significantly more frequent in subjects belonging to the RDH group. The presence of mixed features, occurring in about 30-40% of affective episodes during the course of BDs, contributes to the overall complexity of such disorders (McIntyre et al. 2015, Verdolini et al. 2015). The presence of mixed episodes was associated with higher comorbidity rates, more frequent suicide attempts, longer duration and resistance to pharmacological treatment, determining a less favourable course of illness (Solè et al. 2017). The correct identification and treatment of mixed states represents a still debated issue in psychiatry, with an increasing amount of literature focusing of these clinical entities (Stahl et al. 2017, Verdolini et al. 2018). Results frrom the present study confirm that further research on mixed features may help reducing the global burden of disease in BDs. In the present study, admission to the psychiatric unit for manic/hypomanic was inversely associated with RDH.

manic subjects in this subgroup, confirming the significantly low association of the diagnosis of BD type II with repeated hospitalizations (Bega et al. 2012). Future prospective studies should further clarify this result, stratifying subjects with BDs according to their diagnostic subgroup and other clinical characteristics, e.g. predominant polarity. Another finding was that the comorbidity with medical diseases represents a predictive factor for RD. Previous studies already highlighted the role of medical comorbidities as possible risk factors for psychiatric readmissions (Kim et al. 2011, Šprah et al. 2017). Medical burden in BDs is connected to a more chronic course, longer duration of illness, poorer global functioning and quality of life (Magalhães et al. 2012). Furthermore, pathways leading to both mental and organic disorders are considered to be inter-related and to interact (Šprah et al. 2017). Subsequently, the course of comorbid psychiatric and physical disorders could be mutually influenced, leading to a worsening of both conditions and a consequent

This finding may be related to the presence of hypo-

higher readmission rate due to psychiatric and/or nonpsychiatric reasons (Schoepf & Heun 2014, Šprah et al. 2017). Due to the high mortality of patients with serious mental illnesses and to their unhealthy lifestyle (Correll et al. 2017, Holt et al. 2019), integrated interventions that could better target physical comorbidities were developed (Sampogna et al. 2018). Hospitalization-related factors also turned out to be statistically significant in determining RD. Commitment to residential community facilities at discharge was a predictor of RD in BDs, which was consistent with previous studies in mix-samples (Botha et al. 2010, Di Lorenzo et al. 2016, Morlino et al. 2011). Indeed, poor aftercare attendance and lack of adherence to pharmacological treatment represent major risk factors for frequent rehospitalizations in subjects with serious mental illnesses, including BDs and schizophrenia spectrum disorders (Bodén et al. 2011, Hamilton et al. 2015). On one hand, the commitment to residential facilities may represent a possible solution to the above-stated risk factors, but on the other side it reflects a higher clinical severity and a lack of social and familiar support, that significantly contribute to the RD phenomenon in BDs (Schmutte et al. 2010, Hamilton et al. 2016). In addition, the discharge to psychiatric residential facilities could be related to an overall lower functioning, which represents a further risk factor for frequent readmission in bipolar subjects (Hamilton et al. 2016). As for treatment-related features, our study partially contrasts with previous research underlining the role of some mood stabilizers in preventing rehospitalizations in BDs (Woo et al. 2014). On the other side, the higher prescription rate of compounds belonging to other classes in the RDH group may reflect a more complex clinical condition, encompassing several symptomatological dimensions thus requiring specific targeted interventions. Benzodiazepines prescription could be explained by the higher rate of mixed episodes among RDH, since anxiety represents a core symptom in these clinical entities (McElroy & Keck 2016). Similarly, atypical antipsychotics were demonstrated to be firstline treatments for mixed episodes according to recent guidelines (Verdolini et al. 2018). Subjects in the RDH group were also more frequently prescribed antidepressants, which may result from the presence of a depressive predominant polarity, more often associated to a higher clinical complexity and unfavourable clinical course (Rosa et al. 2018, Garcia-Jimenez et al. 2019). This study presents some limitations. First, the small sample and the retrospective nature of the study limited the generalizability of findings. Since data were collected from clinical records, the number of variables that could be analyzed was finite and no specific psychopathological assessment instruments were used. Finally, since information about outpatient mental health service utilization was not available it was not possible to evaluate the adherence to outpatient programs.

CONCLUSIONS

Findings from the present study suggest that targeted strategies focused on early identification of specific symptomatological dimensions and on the management of physical health in subjects affected by mental disorders might help reducing the RD phenomenon. The improvement of treatment adherence and social support by means of integrated treatment strategies may also be useful to reduce readmission rate and the global burden of BDs.

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Contribution of individual authors:

- Giulia Menculin, Chiara Gobbicchi & Norma Verdolini conceived and designed the study.
- Giulia Menculini, Chiara Gobbicchi, Norma Verdolini & Federica Cirimbilli extracted data in the electronic database.
- Giulia Menculini, Chiara Gobbicchi & Norma Verdolini performed the statystical analysis.
- Giulia Menculini, Chiara Gobbicchi & Federica Cirimbilli wrote the first draft of the manuscript.
- Norma Verdolini corrected the first draft of the manuscript;
- Alfonso Tortorella, Patrizia Moretti & Norma Verdolini supervised all phases of the study design and writing of the manuscript.

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