INTRODUCTION

The goal of conventional treatment of schizophrenia has been to improve the psychotic symptoms (1). However, people gradually started to know that it would be sometimes difficult to achieve the goal (2). For example, increasing dose of antipsychotic drugs or using them for a long time to improve psychotic symptoms may cause various side effects such as acute dystonia, akathisia, parkinsonism, tardive dyskinesia, and hyperprolactinemia, which may adversely affect functional status of patients (2, 3). Therefore, therapists need to consider patient’s subjective well-being or quality of function in daily life as well as symptom reduction in the treatment of schizophrenia.

Since the 1980s, research and development of QOL assessment tools have been active (4, 5) and new assessment tools have been made for patients with schizophrenia (4-6). The usefulness of the concept of QOL as a framework for integrating information related to evaluation of services has been investigated (7, 8), and the concept of QOL has gradually expanded from an indicator focused on specific symptoms or diseases to a concept related to treatment outcome. However, because of a variety of scales for QOL evaluation, research results have been inconsistent (4). There are investigations reporting that there is a slight correlation between objective QOL assessed by medical professionals and subjective QOL assessed by patients themselves (8), but there are also reports showing that there is no correlation between them (6, 9, 10). These findings show that it is difficult to assess QOL appropriately (4). Objective QOL reflects function and lifestyle, and it can be used to address service programs more directly, but it is pointed out that it does not consider how patients feel about their lives and patients’ opinions (7). Therefore, recently, subjective QOL has been considered more important because it reflects the patients' satisfaction with treatment, perceived symptoms, and perceived health conditions (6).

Currently, treatment of patients with schizophrenia is shifting from a hospital-based approach to a community-based approach (11), and the goal of treatment is not only to improve patients’ symptoms but also to improve their QOL in social life from the perspective of enabling them to lead a psychologically and socially healthy daily life even if psychotic symptoms do not disappear completely (11, 12). Therefore, it is crucial to take patients' subjective QOL into consideration when they are receiving inpatient treatment. The purpose of this study is to clarify the relationship between QOL and clinical factors in inpatients with schizophrenia.

METHODS

Participants

Fifty inpatients with schizophrenia diagnosis according to the DSM-V (13) who were receiving inpatient treatment in department of psychiatry of Hospital A participated in this study. All participants gave written consent. Patients with organic brain diseases, epilepsy, and serious physical diseases were excluded when recruiting subjects for this study. Data were collected from November 1, 2018 to January 31, 2019.

Instruments

The following reliable and validated rating scales were used in this study.

Japanese version of the Schizophrenia Quality of Life Scale (JSQLS) (14-16)

The JSQLS is a self-report questionnaire that assesses cognition and interest of patients with schizophrenia. It consists of 90 questions and has three scales of Psychosocial (PS),

Received for publication November 17, 2021; accepted December 10, 2021.

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Motivation/Energy (ME), and Symptoms/Side effects (SS). It calculates score for each domain by scoring each item on a scale of 0 to 4 (0: never, 1: rarely, 2: sometimes, 3: often, 4: always), and lower score indicates better QOL. It is reported that the JSQLS has sufficient reliability and validity (16).

Subjective Well-being under Neuroleptic drug treatment Short form, Japanese version (SWNS-J) (17-19)

The SWNS-J is a self-report questionnaire consisting of 20 question items. It has five subscales: Mental Functioning (MF), Self-Control (SC), Emotional Regulation (ER), Physical Functioning (PF), and Social Integration (SI). Each item is rated on a 5-point scale. Higher score indicates better subjective well-being. It is reported that the SWNS-J has sufficient reliability and validity (19).

Mini Mental State Examination-Japanese (MMSE-J) (20-24)

The MMSE-J evaluates the following abilities: time awareness, place awareness, memorization, attention and calculation, replay, calling, recitation, comprehension, reading, writing, and drawing. When all responses are correct, the total score is 30 points (22,23). Higher score shows better cognitive function. It is reported that the MMSE-J has sufficient reliability and validity (22,23).

Japanese version of the Calgary Depression Scale for Schizophrenia (JCDSS) (25-27)

The JCDSS is a semi-structured interview instrument designed to assess severity of depression in patients with schizophrenia. It consists of eight question items and one observation item. Each item is scored on a scale of 0 to 3, and higher score indicates severer depression level (25,26). The JCDSS has sufficient reliability and validity (27).

Brief Psychiatric Rating Scale (BPRS) (28-31)

The BPRS is widely used in research and clinical scene as an assessment scale of psychiatric symptoms. In this study, we used positive symptom subscale to assess suspiciousness, hallucinatory behavior, conceptual disorganization, and unusual thought content, and negative symptom subscale to assess emotional withdrawal, motor retardation, blunted affect, and disorientation (32,33). Higher score means severer psychotic symptoms (28-31). The BPRS has sufficient reliability (31).

Drug-induced Extrapyramidal Symptoms Scale (DIEPSS) (34,35)

The DIEPSS is a scale for evaluation of drug-induced extrapyramidal symptoms and consists of eight individual items and one summary item. It is rated on a 5-point scale from 0 to 4. In this study, we used total score of the eight individual items (34,35). Higher score indicates severer drug-induced extrapyramidal symptoms (34,35). It is reported that the DIEPSS has sufficient reliability and validity (34,35).

Data analysis

SPSS for Windows Ver.24 was used for statistical analysis. Shapiro-Wilk test was used to test for normality of the data. Since normality was not found for the scores of SWNS-J SI, MMSE-J, JCDSS, BPRS positive symptom subscale, and DIEPSS, Spearman’s rank correlation coefficient was used for correlation analysis. False discovery rate correction was done to correct significance probability. Stepwise regression analyses were performed using the scores of JSQLS and SWNS-J as dependent variables and the variables showed significant correlations with JSQLS or SWNS-J as independent variables.

Ethical considerations

This study was conducted with the approval of the Clinical Research Ethics Review Committee of Tokushima University Hospital (No.3280) and the Ethical Review Committee of Hospital A.

RESULTS

Table 1 shows demographic characteristics of subjects and results of Shapiro-Wilk test. Correlations between JSQLS and other clinical variables are shown in Table 2. Significant correlations were found between JSQLS PS and MMSE-J ($\rho = -0.382$, $p < 0.01$), between JSQLS PS and JCDSS ($\rho = 0.571$, $p < 0.01$), between JSQLS ME and JCDSS ($\rho = 0.470$, $p < 0.01$), and between JSQLS ME and BPRS negative symptom subscale ($\rho = 0.332$, $p < 0.05$).

Table 3 shows correlations between SWNS-J and other clinical variables. There were significant correlations between SWNS-J Total and JCDSS ($\rho = -0.418$, $p < 0.01$), between SWNS-J ME and MMSE-J ($\rho = 0.330$, $p < 0.05$), between SWNS-J MF and JCDSS ($\rho = -0.458$, $p < 0.01$), and between SWNS-J PF and JCDSS ($\rho = -0.405$, $p < 0.01$).

Results of stepwise regression analyses on JSQLS and SWNS-J are shown in Table 4 and Table 5, respectively. JCDSS significantly predicted JSQLS PS ($\beta = 0.249$, $p = 0.514$), JSQLS ME ($\beta = 0.276$, $p = 0.539$), SWNS-J Total ($\beta = 0.149$, $p = 0.407$), SWNS-J MF ($\beta = 0.212$, $p = 0.478$), and SWNS-J PF ($\beta = 0.174$, $p = 0.437$).

DISCUSSION

In this study, we found that depressive symptom affects schizophrenia inpatients' QOL, but no other symptoms such as positive symptom, negative symptom, cognitive dysfunction, and drug-induced extrapyramidal symptom have any significant effect on their QOL. To the best of our knowledge, there has been no study examining clinical factors affecting subjective QOL of patients with schizophrenia. So, we discussed our results in comparison with previous studies that examined clinical factors affecting subjective QOL of outpatients with schizophrenia.

Dickerson, et al. (36) studied 72 outpatients with schizophrenia and reported that depressive symptom score of the Positive and Negative Syndromes Scale (PANSS) was related to the Global Subjective Quality of Life score, but scores of positive and negative symptoms of the PANSS were not. Fitzgerald, et al. (37) conducted a similar study with 174 schizophrenia outpatients and reported that life satisfaction scale was significantly correlated with the Montgomery and Asberg Depression Rating Scale, but there was no significant correlation between it and the PANSS positive and negative symptoms scores, and the Extrapyramidal Side Effects Rating Scale score. Reine, et al. (38) investigated the relationship between subjective QOL and depression in 67 outpatients with schizophrenia and found that depressive symptom was more significantly related to subjective QOL than extrapyramidal symptom. Noman, et al. (39) performed multivariate analyses using the General Well-Being Scale (GWB), the Scales for the Assessment of Positive Symptoms (SAPS), the Scale for the Assessment of Negative symptoms, and reported that the GWB correlated significantly with the reality distortion score, which is a subset of the SAPS, concluding that subjective well-being was strongly related to positive symptom. Awad, et al. (2) reported that severity of psychiatric symptoms, subjective re-
Table 1. Demographic characteristics and results of Shapiro-Wilk test

<table>
<thead>
<tr>
<th></th>
<th>Mean (SD)</th>
<th>Median (QD)</th>
<th>p - value</th>
</tr>
</thead>
<tbody>
<tr>
<td>N (men/women)</td>
<td>50 (22/28)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Age (years)</td>
<td>56.48 (11.93)</td>
<td>58.00 (9.00)</td>
<td>0.300</td>
</tr>
<tr>
<td>Age of onset (years)</td>
<td>24.60 (9.79)</td>
<td>22.00 (4.50)</td>
<td>0.000**</td>
</tr>
<tr>
<td>Duration of illness (years)</td>
<td>32.08 (14.16)</td>
<td>33.00 (11.50)</td>
<td>0.244</td>
</tr>
<tr>
<td>Number of hospitalization</td>
<td>4.44 (2.03)</td>
<td>4.00 (1.00)</td>
<td>0.005**</td>
</tr>
<tr>
<td>JSQLS</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>JSQLS PS</td>
<td>45.03 (16.26)</td>
<td>44.16 (9.58)</td>
<td>0.121</td>
</tr>
<tr>
<td>JSQLS ME</td>
<td>47.30 (14.37)</td>
<td>46.42 (11.16)</td>
<td>0.522</td>
</tr>
<tr>
<td>JSQLS SS</td>
<td>34.93 (14.73)</td>
<td>32.81 (6.64)</td>
<td>0.170</td>
</tr>
<tr>
<td>SWNS-J</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>SWNS-J Total</td>
<td>71.84 (11.92)</td>
<td>71.50 (7.50)</td>
<td>0.700</td>
</tr>
<tr>
<td>SWNS-J MF</td>
<td>13.48 (3.61)</td>
<td>13.50 (2.50)</td>
<td>0.431</td>
</tr>
<tr>
<td>SWNS-J SC</td>
<td>15.04 (3.04)</td>
<td>15.00 (2.50)</td>
<td>0.517</td>
</tr>
<tr>
<td>SWNS-J ER</td>
<td>13.74 (3.30)</td>
<td>14.00 (2.00)</td>
<td>0.771</td>
</tr>
<tr>
<td>SWNS-J PF</td>
<td>15.12 (3.25)</td>
<td>15.00 (2.00)</td>
<td>0.125</td>
</tr>
<tr>
<td>SWNS-J SI</td>
<td>14.46 (3.21)</td>
<td>14.00 (1.50)</td>
<td>0.023*</td>
</tr>
<tr>
<td>MMSE-J</td>
<td>24.20 (4.46)</td>
<td>26.00 (4.00)</td>
<td>0.003**</td>
</tr>
<tr>
<td>JCDSS</td>
<td>5.26 (4.75)</td>
<td>5.50 (3.50)</td>
<td>0.000**</td>
</tr>
<tr>
<td>BPRS</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>BPRS positive symptom</td>
<td>9.64 (3.98)</td>
<td>9.00 (3.50)</td>
<td>0.040*</td>
</tr>
<tr>
<td>BPRS negative symptom</td>
<td>9.88 (3.23)</td>
<td>10.00 (2.50)</td>
<td>0.188</td>
</tr>
<tr>
<td>DIEPSS</td>
<td>2.70 (2.50)</td>
<td>2.50 (2.00)</td>
<td>0.000**</td>
</tr>
</tbody>
</table>

Note: *p < 0.05, **p < 0.01, Shapiro-Wilk test.

Abbreviations: SD, Standard Deviation; QD, Quartile Deviation; JSQLS, Japanese version of the Schizophrenia Quality of Life Scale; JSQLS PS, Psychosocial scale of JSQLS; JSQLS ME, Motivation/Energy scale of JSQLS; JSQLS SS, Symptoms/Side effects scale of JSQLS; SWNS-J, Subjective Well-being under Neuroleptic drug treatment short form, Japanese version; SWNS-J Total, Total score of SWNS-J; SWNS-J MF, Mental Functioning subscale of SWNS-J; SWNS-J SC, Self-Control subscale of SWNS-J; SWNS-J ER, Emotional Regulation subscale of SWNS-J; SWNS-J PF, Physical Functioning subscale of SWNS-J; SWNS-J SI, Social Integration subscale of SWNS-J; MMSE-J, Mini Mental State Examination-Japanese; JCDSS, Japanese version of the Calgary Depression Scale for Schizophrenia; BPRS, Brief Psychiatric Rating Scale; DIEPSS, Drug-induced Extrapyramidal Symptoms Scale.

Table 2. Correlation between JSQLS scores and other clinical variables

<table>
<thead>
<tr>
<th>JSQLS</th>
<th>JSQLS PS</th>
<th>JSQLS ME</th>
<th>JSQLS SS</th>
</tr>
</thead>
<tbody>
<tr>
<td>MMSE-J</td>
<td>-0.382**</td>
<td>-0.295</td>
<td>0.130</td>
</tr>
<tr>
<td>JCDSS</td>
<td>0.571**</td>
<td>0.470**</td>
<td>0.299</td>
</tr>
<tr>
<td>BPRS positive symptom</td>
<td>-0.014</td>
<td>-0.113</td>
<td>0.107</td>
</tr>
<tr>
<td>BPRS negative symptom</td>
<td>0.214</td>
<td>0.352*</td>
<td>0.157</td>
</tr>
<tr>
<td>DIEPSS</td>
<td>0.222</td>
<td>0.064</td>
<td>0.301</td>
</tr>
</tbody>
</table>

Notes: *p < 0.05, **p < 0.01, Spearman rank correlation. (False Discovery Rate correction)

Abbreviations: JSQLS, Japanese version of the Schizophrenia Quality of Life Scale; JSQLS PS, Psychosocial scale of JSQLS; JSQLS ME, Motivation/Energy scale of JSQLS; JSQLS SS, Symptoms/Side effects scale of JSQLS; MMSE-J, Mini Mental State Examination-Japanese; JCDSS, Japanese version of the Calgary Depression Scale for Schizophrenia; BPRS, Brief Psychiatric Rating Scale; DIEPSS, Drug-induced Extrapyramidal Symptoms Scale.
sponses to antipsychotic drugs, and side effects of antipsychotic drugs were considered as important predictors of QOL. However, Larsen, et al. (6) studied the relationship between psychiatric symptoms, medication side effects, and subjective QOL in chronic outpatients with schizophrenia visiting an outpatient clinic and reported that no significant correlation between subjective QOL and mental status, and side effects of drugs.

When summarizing the results of previous studies, we could say that depressive symptom is considered important as a clinical factor affecting the subjective QOL of outpatients with schizophrenia (40). In this study, we focused on inpatients with schizophrenia who had severer positive or negative symptom and higher level of drug-induced extrapyramidal symptom than outpatients. So, we expected that those symptoms would affect their subjective QOL significantly. Moreover, recently, cognitive dysfunction of patients with schizophrenia has been reported to be more important than psychiatric symptoms in terms of affecting daily functioning and interpersonal relationships (41-43). Therefore, we also expected that cognitive function would affect their subjective QOL. However, the results of this study clearly show that depressive symptom is only clinical factor independently affecting subjective QOL. This may be because subjects of this study had severer depression level than outpatients in the previous study (4). But the results suggest that improvement of depressive symptom may lead to better subjective QOL in the inpatient treatment of schizophrenia.

**CONCLUSION**

We found a clinical factor influencing subjective QOL of inpatients with schizophrenia. Although inpatients have severer psychiatric symptoms than outpatients, positive and negative symptoms do not affect their QOL significantly, but only depressive symptom significantly influences it. This is a new finding we found in this study, suggesting that focusing on treatment of...
depressive symptom may lead to better QOL of inpatients with schizophrenia.

CONFLICT OF INTEREST

The authors declare no conflict of interest.

ACKNOWLEDGEMENTS

We would like to express our deepest gratitude to the patients who participated in this study and the medical staff of Jouzan Hospital who supported this study.

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