Check for updates

OPEN ACCESS

EDITED BY Huixuan Zhou, Beijing Sport University, China

REVIEWED BY

Bartlomiej Stanczykiewicz, Wrocław Medical University, Poland Wei Zhang, National Institutes for Food and Drug Control, China

*CORRESPONDENCE Rongchun Yang ⊠ fyssyesk@126.com Huanzhong Liu ⊠ huanzhongliu@ahmu.edu.cn

[†]These authors have contributed equally to this work

SPECIALTY SECTION

This article was submitted to Public Mental Health, a section of the journal Frontiers in Psychiatry

RECEIVED 15 September 2022 ACCEPTED 16 December 2022 PUBLISHED 06 January 2023

CITATION

Liu Z, Zhang Y, Sun L, Wang J, Xia L, Yang Y, Sun F, Li W, Yao X, Yang R and Liu H (2023) Physical activity levels associated with insomnia and depressive symptoms in middle-aged and elderly patients with chronic schizophrenia. *Front. Psychiatry* 13:1045398. doi: 10.3389/fpsyt.2022.1045398

COPYRIGHT

© 2023 Liu, Zhang, Sun, Wang, Xia, Yang, Sun, Li, Yao, Yang and Liu. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.

Physical activity levels associated with insomnia and depressive symptoms in middle-aged and elderly patients with chronic schizophrenia

Zhiwei Liu^{1†}, Yulong Zhang^{2,3†}, Liang Sun^{1†}, Juan Wang⁴, Lei Xia^{2,3}, Yating Yang^{2,3}, Feng Sun¹, Wenzheng Li⁵, Xianhu Yao⁶, Rongchun Yang^{1*} and Huanzhong Liu^{2,3*}

¹Department of Psychiatry, The Third People's Hospital of Fuyang, Fuyang, China, ²Department of Psychiatry, Chaohu Hospital of Anhui Medical University, Hefei, China, ³Anhui Psychiatric Center, Anhui Medical University, Hefei, China, ⁴Department of Psychiatry, Chengdu Fourth People's Hospital, Chengdu, China, ⁵Department of Psychiatry, Hefei Fourth People's Hospital, Hefei, China, ⁶Department of Psychiatry, Ma'anshan Fourth People's Hospital, Ma'anshan, China

Background: Previous evidence suggested that physical activity had beneficial effects on psychopathological symptoms, insomnia, or depressive symptoms in people with schizophrenia. This study investigated the association between physical activity levels and insomnia and depressive symptoms in middle-aged and elderly hospitalized patients with chronic schizophrenia (CS).

Methods: 179 participants were enrolled. We used the 30-item Positive and Negative Syndrome Scale (PANSS $_{-30}$) to assess the psychopathological symptoms. We used the Insomnia Severity Index scale (ISI) and 17-item Hamilton Depression Scale (HAMD-17) to evaluate insomnia and depressive symptoms. Daily physical activity time less than 30 min, within 30-60 min, and more than 60 min were defined as physical inactivity, moderate physical activity, and vigorous physical activity, respectively. The Chisquare test, analysis of variance (ANOVA), and Mann-Whitney U-test were applied for categorical, continuous, and non-normal distribution variables, respectively. The Pearson or Spearman's correlation analyses were utilized to examine the association between physical activity levels, ISI total scores, HAMD total scores, and socio-demographic and clinical variables. Finally, socio-demographic variables with a P-value < 0.05 in the comparison between insomnia/depressive group and non-insomnia/depressive group were considered for inclusion in binary logistic regression analysis to determine the relationship between physical activity levels and insomnia or depressive symptoms.

Results: The ISI total scores (r = -0.247, P = 0.001) and HAMD total scores (r = -0.312, P < 0.001) were negatively correlated with physical activity

levels. Logistic regression analysis revealed that older age, higher depressive factor scores, and lower physical activity level were influential factors of insomnia symptoms in CS patients (P < 0.05). In addition, vigorous physical activity (compared with physical inactivity) and higher negative and depressive factor scores were independently associated with depressive symptoms in CS patients (P < 0.05).

Conclusion: Physical activity levels were influential factors in comorbid insomnia and depressive symptoms in CS patients. Given the benefits of physical activity, it should be strengthened as a routine adjunct to clinical treatment or psychiatric care so as to improve the physical and mental health of patients with psychiatric symptoms.

KEYWORDS

physical activity, insomnia, depressive, middle-aged, elderly, schizophrenia

Introduction

World Health Organization (WHO) statistics reported that approximately 15% of older adults aged 60 years and older suffer from mental disorders, such as depression, anxiety, and dementia, making it a category of public health challenges that seriously affects the elderly population (1). The results of the Global Burden of Disease Study (GBD) showed that mental disorders accounted for a significantly increasing proportion of global disability-adjusted life years (DALYs) and 14.6% of global disability life lost years (YLDs), making it one of the top 10 global burdens of disease (2). The latest, the most authoritative the national epidemiological survey of mental disorders in China indicated that the lifetime prevalence of mental illness in adults was 16.57% (3), with alone accounting for approximately 17% of the global burden of mental disorders (4), creating a heavy long-term burden at the individual, societal, and national levels.

Schizophrenia is a group of severe mental disorders of unknown etiology. Patients suffer from serious abnormalities in cognition, thinking, and behavior (5), with a lifetime prevalence of approximately 1% worldwide (6). Previous literature stated that the population with mental disorders suffered poorer sleep quality than the general population, and about 20-40% of patients with chronic schizophrenia (CS) were comorbid insomnia symptoms (7-9). Furthermore, research suggested that worse sleep quality was strongly related to more severe psychotic symptoms (10) and that insomnia symptoms could even exacerbate psychiatric symptoms and lead to increased somatic comorbidity. In contrast, healthy sleep hygiene significantly improved the severity of psychopathological symptoms (11). Strong evidence suggested that regular physical activity enhanced mental health (12) and could predict a better global and social quality of life in schizophrenic patients (13). Furthermore, higher physical activity levels were strongly related to fewer insomnia symptoms, better cognitive functioning, social functioning, and life satisfaction compared to lower levels of physical activity (14).

Comorbid depressive symptoms were more frequent in patients with schizophrenia, with an estimated prevalence of 18.8–80% (15–17). One study noted that depressive symptoms were negatively associated with the severity of psychotic symptoms (16). Compared to patients with mildly depressed schizophrenia, patients with major depressive symptoms had more serious psychiatric symptoms and poorer quality of life (15). Furthermore, a cross-sectional survey showed that more severe depressive symptoms in patients with schizophrenia were strongly related to less daily physical activity (18). Strong evidence suggests that regular physical activity may reduce depressive symptoms and improve social functioning in schizophrenic patients significantly (19, 20).

Physical activity, or physical exercise, sports, etc., is considered to be an important factor related to mental health. The latest meta-analysis shows that participation in sports can significantly reduce the body fat content and improve the physical function and mental health of elderly people over 60 years old (21). Another meta-analysis suggests that aerobic exercise may have a significant effect on mood and anxiety symptoms (22). In addition, clinical study indicates that a combination of high sedentary behavior and low moderate-to-vigorous intensity physical activity (MVPA) is strongly associated with higher levels of depression and anxiety symptoms compared with populations with less sedentary behavior and with sufficient MVPA (23). According to a national survey, higher PA frequency was associated with lower levels of depression, anxiety in Chinese physicians (24). A systematic review of 31 studies found that reducing sedentary behavior and increasing MVPA were significantly associated with better mental health and quality of life (25). Meanwhile, physical activity has also proven to be a promising adjunctive intervention for mood disorders.

Although growing literature evidence reported the correlation between physical activity levels, insomnia symptoms, depressive symptoms and psychopathology, studies on middle-aged and elderly Chinese hospitalized schizophrenic patients are still lacking. This research investigated the association between physical activity levels, insomnia and depressive symptoms in middle-aged and elderly hospitalized patients with schizophrenia. Further, we analyzed the influencing factors of comorbidities of insomnia or depressive symptoms in schizophrenia.

Materials and methods

Subjects

The subjects were psychiatric inpatients from May to December 2018 from three tertiary hospitals (Chaohu Hospital of Anhui Medical University, Hefei Fourth People's Hospital, and Ma'anshan Fourth People's Hospital) in Anhui Province, China. This cross-sectional study was a secondary analysis of the physiological and psychological conditions of inpatients with CS. Please refer to our previous manuscripts for sample size calculation (26). Inclusion criteria: (1) age \geq 45 years; (2) it met the diagnostic criteria of schizophrenia according to the 5th edition of the Diagnostic and Statistical Manual of Mental Disorders (DSM-V); (3) disease duration \geq 5 years; and (4) able to complete clinical scale assessment. Exclusion criteria: (1) the presence of other serious mental retardation or neurological diseases; (2) unable to complete the assessment of clinical symptoms scales; (3) women who are pregnant or breastfeeding; (4) complicated with serious physical diseases (such as cardiovascular diseases, digestive system diseases, respiratory diseases, etc.).

The enrolled patients and their guardians agreed to participate in the project after knowing the study process and the related advantages and disadvantages and signed the paper informed consent. This research was approved by the Ethics Committee of Chaohu Hospital of Anhui Medical University (No. 201805-kyxm-03) and obtained registration number (No. ChiCTR1800017044) from the China Clinical Trials Registry.

Socio-demographic and clinical data

Demographic variables (age, gender, education, etc.) and clinical information of each patient were collected by questionnaire and the electronic medical record. Antipsychotic

dose in chlorpromazine equivalents was calculated using the defined daily dose method (27).

Blood indicators

Ten milliliters of fasting venous blood was collected on the second morning after enrollment. The fasting blood glucose (FBG), total cholesterol (TC), triglyceride (TG), high-density lipoprotein (HDL-C), low-density lipoprotein (LDL-C), and other blood lipid indexes were detected by a specialized laboratory technician.

Physical activity levels

According to the dietary guidelines for Chinese residents, it is recommended that middle-aged and older people exercise outdoors one to two times a day for 30-60 min each time (28). In addition, WHO suggested that adults complete 150-300 min of moderate physical activity per week (29). In this study, the daily physical activities included walking, rhythmic exercises, etc., and the activity time was comprehensively judged by patients' self-reported and medical care records. All subjects were measured by three simple questions and answered the questions "yes" or "no": (1) Did you participate in any form of physical activity during hospitalization, such as rhythmic exercises, jogging, walking, etc. (2) The amount of physical activity you did each day: less than 30 min, 30-60 min, or more than 60 min. (3) Did you have any physical discomfort when you participate in physical activity? Less than 30 min per day was considered physical inactivity, 30-60 min per day was considered moderate physical activity, and more than 60 min per day was considered vigorous physical activity.

Assessment of psychiatric symptoms

30-item Positive and Negative Syndrome Scale (PANSS₋₃₀) was utilized to assess the psychiatric symptoms of each inpatient. The five-factor model consists of positive, negative, cognitive, depressive, and excited factor scores were used for the statistical scoring, respectively (30).

Insomnia Severity Index scale (ISI) was used to assess insomnia symptoms with good reliability and validity (31, 32). The ISI total scores range from 0 to 28. We defined ISI total scores \geq 8 as comorbid insomnia symptoms (31).

17-Item Hamilton Depression Scale (HAMD₋₁₇) was used to assess depressive symptoms. Higher scores of HAMD indicate the different degrees of depression. We defined HAMD total scores ≥ 8 as comorbid depressive symptoms (33). The three clinical assessment scales' intraclass correlation coefficient (ICC) exceeded 0.8.

Statistical analysis

The SPSS 23.0 software package (*IBM*, *Chicago*, *IL*, *USA*) was used for data analysis. First, Chi-square test, analysis of variance (ANOVA), and Mann–Whitney *U*-test were applied for categorical, continuous, and non-normal distribution variables, respectively. Second, Pearson or Spearman's correlation analyses were utilized to examine the association between physical activity levels, ISI total scores, HAMD total scores, and socio-demographic and clinical variables. Finally, socio-demographic variables with a *P*-value < 0.05 in the comparison between insomnia/depressive group and non-insomnia/depressive group were considered for inclusion in binary logistic regression analysis to determine the relationship between physical activity levels and insomnia or depressive symptoms. The significance level was set as $\alpha = 0.05$ (2-tailed).

Results

Comparisons between groups with and without insomnia or depressive symptoms

179 inpatients with CS were included, and the prevalence of insomnia or depressive symptoms was 22.9% (41/179) and 71.5% (128/179). Compared to patients without insomnia symptoms, the comorbid insomnia symptoms patients had older age, lower physical activity level, higher proportion of antipsychotic polypharmacy, higher FBG levels, and higher psychopathology symptoms such as positive/depressive factor scores, PANSS, ISI, and HAMD total scores. Compared to patients without depressive symptoms, the patients with depressive symptoms had lower physical activity levels, fewer hospitalizations, higher positive, negative, cognitive, and depressive factor scores, PANSS, ISI and HAMD total scores (see Table 1).

Correlation analysis between ISI total scores, HAMD total scores, and demographic and clinical variables

ISI total scores were positively related to age (r = 0.206), FBG (r = 0.169), positive (r = 0.234), negative (r = 0.181), depressive factor scores (r = 0.261), PANSS total scores (r = 0.167), and were negatively correlated with physical activity levels (r = -0.247), significantly. After controlling for other variables related to ISI total scores, ISI total scores were still negatively correlated with physical activity levels (r = -0.168, P = 0.027) (see Table 2).

HAMD total scores were positively correlated with negative (r = 0.316), cognitive (r = 0.251), depressive (r = 0.506), PANSS

total scores (r = 0.314), and were negatively correlated with number of admissions (r = -0.177) and physical activity levels (r = -0.312), significantly. After controlling for other variables related to HAMD total scores, HAMD total scores were still negatively correlated with physical activity levels (r = -0.188, P = 0.013) (see Table 2).

Influencing factors associated with insomnia symptoms in CS patients

The results showed that older age (OR = 1.07, 95% CI: 1.02–1.14, P = 0.013),vigorous physical activity (compared to physical inactivity) (OR = 0.22, 95% CI: 0.06–0.88, P = 0.032), and higher depressive factor scores (OR = 1.22, 95% CI: 1.02–1.46, P = 0.030) were independently correlated with insomnia symptoms in CS patients (see Table 3).

Influencing factors associated with depressive symptoms in CS patients

The results found that vigorous physical activity (compared to physical inactivity) (OR = 0.19, 95% CI: 0.07–0.48, P = 0.001), higher negative factor scores (OR = 1.10, 95% CI: 1.02–1.19, P = 0.019) and depressive factor scores (OR = 1.52, 95% CI: 1.21–1.91, P < 0.001) were related to depressive symptoms in CS patients (see Table 4).

Discussion

This study revealed that (1) insomnia and depressive symptoms in middle-aged and elderly CS inpatients were 22.9 and 71.5%. Patients with insomnia symptoms tend to be older, have lower levels of physical activity, higher rates of antipsychotic medication combination, higher FBG levels, and higher psychopathological symptoms such as positive, depressive factor scores, PANSS, ISI, and HAMD total scores. Patients with depressive symptoms tended to have fewer hospitalizations, lower levels of physical activity, higher positive, negative, cognitive, and depressive factor scores, PANSS, ISI, and HAMD total scores. (2) After controlling for potential confounders, the ISI total scores and HAMD total scores were still negatively correlated with physical activity levels. (3) Logistic regression analysis revealed that older age, higher depressive factor scores, and lower physical activity level were influential factors of insomnia symptoms in CS patients. In addition, vigorous physical activity (compared with physical inactivity) and higher negative and depressive factor scores were independently related to depressive symptoms in CS patients.

Our research noted that 22.9% of middle-aged and elderly CS patients had insomnia symptoms, which was more consistent with previous studies (7, 8). In addition, older age, higher

TABLE 1 Comparison of clinical and biological indicators between insomnia group and non-insomnia group, depressive group, and non-depressive group.

Variables	Insomnia group (n = 41)	Non- insomnia group (n = 138)	X ² /F	Р	Depressive group (n = 128)	Non- depressive group (n = 51)	X ² /F	Р
Age (years, $\bar{x} \pm s$)	55.71 ± 7.49	52.94 ± 5.97	-2.449	0.015 ^a	52 (49.25, 57)	52.45 ± 5.92	-1.385	0.166 ^b
Gender								
Male (%)	21 (51.2)	78 (56.5)	0.360	0.549 ^c	69 (53.9)	30 (58.8)	0.357	0.550 ^c
Female (%)	20 (48.8)	60 (43.5)			59 (46.1)	21 (41.2)		
Education (years)	7.68 ± 3.73	8 (5, 11)	-0.454	0.650 ^b	8 (5, 11)	8 (8, 11)	-1.154	0.249 ^b
Onset age (years)	29.12 ± 8.67	29.32 ± 8.62	0.128	0.898 ^a	29.41 ± 8.56	28.92 ± 8.78	-0.345	0.731 ^a
Illness duration (years)	26.39 ± 11.64	23.75 ± 9.78	-1.449	0.149 ^a	24.67 ± 10.31	23.57 ± 10.20	-0.648	0.518 ^a
Number of admissions	5 (3, 8.5)	5 (3, 9)	-0.017	0.986 ^b	5 (3, 8)	8.25 ± 5.82	-2.274	0.023 ^b
BMI	24.03 ± 3.48	23.96 ± 3.90	-0.107	0.915 ^a	24.05 ± 3.60	23.77 ± 4.27	-0.442	0.659 ^a
Antipsychotic categ	jory (%)							
Monotherapy	9 (22.0)	67 (48.6)	9.154	0.002 ^c	50 (39.1)	26 (51.0)	2.120	0.145 ^c
Polypharmacy	32 (78.0)	71 (51.4)			78 (60.9)	25 (49.0)		
Hematological inde	x					1		1
FBG (mmol/L)	5.30 ± 1.09	4.80 (4.40, 5.40)	-2.008	0.045 ^b	4.90 (4.50, 5.40)	5.20 ± 1.24	-0.005	0.996 ^b
TC (mmol/L)	5.08 ± 2.03	4.69 (4.03, 5.50)	-0.757	0.449 ^b	4.74 (4.09, 5.49)	4.86 ± 1.65	-0.413	0.680 ^b
TG (mmol/L)	2.30 ± 1.75	1.82 (1.35, 2.59)	-0.326	0.744 ^b	1.82 (1.36, 2.52)	1.96 (1.28, 2.93)	-0.272	0.786 ^b
HDL-C (mmol/L)	1.06 ± 0.31	0.98 (0.87, 1.19)	-0.888	0.375 ^b	1.04 ± 0.27	1.06 ± 0.26	0.422	0.674 ^a
LDL-C (mmol/L)	2.54 ± 0.64	2.45 ± 0.65	-0.823	0.412 ^a	2.48 ± 0.62	2.45 ± 0.73	-0.267	0.790 ^a
Physical activity leve	els		1	1		1		1
Physical inactivity	18 (43.9)	38 (27.6)	9.756	0.008 ^c	48 (37.5)	8 (15.7)	12.846	0.002 ^c
Moderate physical activity	20 (48.8)	58 (42.0)			56 (43.8)	22 (43.1)		
Vigorous physical activity	3 (7.3)	42 (30.4)			24 (18.7)	21 (41.2)		
PANSS								
Positive factor scores	11.8 ± 4.94	9 (6.75, 14)	-2.019	0.044 ^b	10.5 (7, 15)	9.47 ± 5.40	-2.509	0.012 ^b
Negative factor scores	19.66 ± 7.35	17.64 ± 6.61	-1.668	0.097 ^a	19.33 ± 6.35	15.04 ± 7.05	-3.951	<0.001 ^a
Cognitive factor scores	9.46 ± 3.23	9.12 ± 2.84	-0.666	0.506 ^a	9.57 ± 2.77	8.25 ± 3.14	-2.761	0.006 ^a
Depressive factor scores	7.90 ± 2.91	6 (4, 8)	-2.732	0.006 ^b	7.48 ± 2.84	5.16 ± 1.90	-5.365	<0.001 ^a
Excited factors scores	8.10 ± 3.27	6 (5, 9)	-1.538	0.124 ^b	7 (5, 9)	6 (4, 10)	-1.500	0.134 ^b
PANSS total scores	84.95 ± 24.45	76.73 ± 22.78	-1.995	0.048 ^a	82.97 ± 21.57	67.69 ± 24.30	-4.125	<0.001 ^a
ISI total scores	11.61 ± 3.29	1.82 ± 1.54	-26.596	< 0.001 ^a	3 (1, 8)	1 (0, 2)	-6.012	<0.001 ^b
HAMD total scores	14.49 ± 4.31	8.91 ± 4.45	-7.100	<0.001 ^a	12 (9, 15)	5 (3, 7)	-10.458	<0.001 ^b
Chlorpromazine equivalent (mg/d)	468.23 ± 221.92	360 (227.5, 600)	-1.682	0.093 ^b	428.02 ± 223.05	300 (240, 540)	-0.848	0.397 ^b
Hypnotics prescription (yes,%)	9 (22.0)	30 (21.7)	0.001	0.977 ^c	29 (22.7)	10 (19.6)	0.199	0.656 ^c
Antidepressants prescription (yes,%)	2 (4.9)	10 (7.2)	0.283	0.594 ^c	10 (7.8)	2 (3.9)	0.883	0.347 ^c

^aIndependent samples *t*-test. ^bMann-Whitney *U*-test. ^cPearson chi-square. Bold value means *P* < 0.05.

TABLE 2 Correlation analysis of clinical and biological indicators with
ISI total scores and HAMD total scores.

Variables	ISI tota	l scores	HAMD total scores		
	r	Р	r	Р	
Age (years)	0.206	0.006	0.145	0.052	
Gender	0.084	0.262	0.050	0.504	
Education (years)	-0.076	0.311	-0.147	0.050	
Illness duration (years)	0.062	0.413	0.063	0.402	
Number of admissions	-0.082	0.273	-0.177	0.018	
BMI	0.060	0.427	-0.045	0.552	
Monotherapy/ Polypharmacy	0.111	0.138	0.118	0.117	
FBG (mmol/L)	0.169	0.024	0.048	0.527	
TC(mmol/L)	0.098	0.191	0.013	0.862	
TG (mmol/L)	-0.011	0.884	-0.049	0.514	
HDL-C (mmol/L)	0.129	0.087	0.000	0.996	
LDL-C (mmol/L)	0.109	0.149	-0.020	0.791	
Physical activity levels	-0.247	0.001	-0.312	<0.001	
Positive factor scores	0.234	0.002	0.203	0.006	
Negative factor scores	0.181	0.016	0.316	<0.001	
Cognitive factor scores	0.077	0.305	0.251	0.001	
Excited factor scores	0.085	0.259	0.116	0.122	
Depressive factor scores	0.261	<0.001	0.506	<0.001	
PANSS total scores	0.167	0.026	0.314	<0.001	
Chlorpromazine equivalent (mg/d)	0.117	0.118	0.112	0.134	
Hypnotics prescription	0.000	0.996	0.058	0.440	
Antidepressants prescription	-0.014	0.852	0.025	0.741	

Bold value means P < 0.05.

depressive factor scores, and lower physical activity were related factors of insomnia symptoms. Previous studies pointed out that compared to the general population, schizophrenic patients had less sleep, and the prevalence of sleep-related problems was significantly higher. In contrast, patients with sleep problems had significantly lower life satisfaction and happiness (34). In a study of outpatients with schizophrenia, a high prevalence of sleep disorders was strongly associated with older age, physical inactivity, and severe psychopathological symptoms (9).

Historical literature suggested that physical inactivity was strongly correlated with poorer psychosomatic health in patients with schizophrenia (18). In contrast, regular physical activity significantly improved patients' insomnia symptoms and their subjective sleep quality to a consistent degree (14, 35). A randomized controlled study in Germany noted that patients with regular physical activity had significantly better sleep quality than psychiatric patients with physical inactivity (36). Second, patients with adequate physical activity had lower TABLE 3 Demographic and clinical variables independently associated with insomnia symptoms by binary logistic regression analysis.

Variables	В	Р	OR	95% CI
Age	0.072	0.013	1.07	1.02-1.14
FBG (mmol/L)	0.095	0.535	1.10	0.82-1.48
Physical activity levels (ref. physical inactivity)		0.098		
Moderate physical activity	-0.264	0.547	0.77	0.33-1.81
Vigorous physical activity	-1.517	0.032	0.22	0.06-0.88
Positive factor scores	0.071	0.297	1.07	0.94-1.23
Negative factor scores	0.038	0.421	1.04	0.95-1.14
Depressive factor scores	0.200	0.030	1.22	1.02-1.46
PANSS total scores	-0.025	0.241	0.98	0.94-1.02

Bold value means P < 0.05.

TABLE 4 Demographic and clinical variables independently associated with depressive symptoms by binary logistic regression analysis.

Variables	В	Р	OR	95% CI
Number of admissions	-0.038	0.216	0.96	0.91-1.02
Physical activity levels(ref. physical inactivity)		0.002		
Moderate physical activity	-0.873	0.057	0.42	0.17-1.03
Vigorous physical activity	-1.679	0.001	0.19	0.07-0.48
Negative factor scores	0.096	0.019	1.10	1.02-1.19
Cognitive factor scores	0.002	0.983	1.00	0.82-1.22
Depressive factor scores	0.418	<0.001	1.52	1.21-1.91
PANSS	-0.017	0.344	0.98	0.95-1.02

Bold value means P < 0.05.

rates of insomnia symptoms than those with physical inactivity (37, 38). In addition, schizophrenic patients who underwent weekly physical activity had less overall symptom severity and a higher quality of life than conventional treatment modalities without physical activity (39). A systematic review also indicated that higher levels of physical activity predicted a moderate increase in patients' motor activity (40) and that increased motor activity improved patients' insomnia symptoms and cognitive-related impairments (41) and was strongly associated with fewer negative symptoms and depressive symptoms (42). Furthermore, analysis of studies has shown that the beneficial effects of physical activity on sleep quality in psychiatric patients were attributed to its improvement of the overall health status of patients (43). The adjunctive use of physical activity interventions in schizophrenic populations has clear efficacy and no associated side effects (44).

In this study, we found that 71.5% of middle-aged and elderly CS patients had depressive symptoms, and

depressive symptoms were independently correlated with severe psychopathological symptoms such as (negative, cognitive, depressive factor scores, PANSS total scores), fewer hospital admissions, and lower physical activity levels. In addition, vigorous physical activity (compared to physical inactivity) and higher negative or depressive factor scores were related to depressive symptoms in CS patients. A study noted that the prevalence of depressive symptoms as a usual comorbidity of schizophrenia in the elderly ranged from 44 to 75% (45). Two studies of older Chinese patients with schizophrenia noted the incidence of depressive symptoms of 48.5% (patients aged: above 65 years old) (46) and 32.8% (patients aged: above 60 years old) (47), both using the Geriatric Depression Scale (GDS) to evaluate depressive symptoms. In addition, a Tunisian study assessed by Depression Anxiety and Stress scales (DASS-21) noted that older patients with schizophrenia (age: 66.9 ± 3.8 years old) had a prevalence of comorbid moderate-to-severe depressive symptoms of only about 25% (48). A French study that used the Center of Epidemiologic Studies Depression (CESD) scale to assess depressive symptoms in patients found that 78.1% presented with subsyndromes or syndromes depressive symptoms which were positively related to psychotic symptoms and were not associated with psychotropic medication use, or with the use of antidepressants (49). The results of our research were consistent with those of previous studies by using the HAMD $_{-17}$ scale to assess depressive symptoms. Previous studies have shown that weekly physical activity levels were negatively related to depressive symptoms in schizophrenic patients (50). Furthermore, evidence indicated that regular physical activity significantly reduced depressive symptoms in older adults (51), mainly because patients who engaged in regular physical activity had significantly reduced psychiatric symptoms and fewer physical complaints (52). Physical activity improved the prognosis of quality of schizophrenic patients, resulting in fewer accompanying depressive symptoms (53). According to a national survey in China, compared to population on physical activity in the last 1 year, people who sometimes or often participated in physical activity had significantly lower symptoms of depression and anxiety (24). Moreover, a systematic review indicated that compared to adults engaged in lower level of physical activity, MVPA was associated with better mental health and higher quality of life (25). The previous solid conclusions suggested that physical activity could be routinely incorporated into the regular treatment and daily care of patients with mental disorders because of its clear benefits (54).

Although current evidence pointed to a beneficial role of physical activity on psychopathological symptoms, insomnia, or depressive symptoms in CS patients, schizophrenic patients spend significantly less time physically active than the general population (10, 55), with only about one-fifth of patients meeting the minimum required level of physical activity (56). In contrast, physical inactivity associated with sedentary behavior in schizophrenic patients was closely related to a higher prevalence of depressive symptoms (57). Furthermore, due to their negative and depressive symptoms and lack of internal drive, patients with severe mental disorders obtained a lower level of physical activity (58). The relationship between depressive symptoms and physical activity levels is a two-way process (59). Moreover, the high rate of sleep disturbance in schizophrenic patients, resulting in reduced energy and deficits, was also associated with physical inactivity (60).

The Study on Global Ageing and Adult Health (SAGE) demonstrated that meeting physical activity guidelines (\geq 150 min of MVPA/week) was significantly associated with more happiness (61). A systematic review of 31 studies found that reducing sedentary behavior and increasing MVPA were significantly associated with better mental health and quality of life (25). Our study indicated that the prevalence of insomnia and depressive symptoms was high in CS patients. Furthermore, higher level of physical activity correlated to fewer insomnia and depressive symptoms. In clinical treatment, in addition to conventional pharmacotherapy or psychotherapy, physical activity can also be used to relieve the emotional symptoms of patients with mental disorders.

This study has the following limitations. First of all, this study was a cross-sectional study, which only explored the correlation between physical activity levels and insomnia and depressive symptoms of the enrolled patients, and could not be determined the causality. Second, the enrolled subjects were middle-aged and older adults, and the prevalence of depressive symptoms was slightly higher than that of the elderly reported in previous studies, which might be due to the differences in the age of the subjects or the assessment tools, which was also the innovation of this study. Third, although combined with electronic medical records as an objective reference, the physical activity level of the subjects in this study was assessed using subjective questions. Using objective methods like accelerometers will give stronger corroborating evidence than subjective description of the physical activity by patients. Moreover, the small sample size included in this study could lead to the bias of the study results, and further large-sample controlled studies are needed.

Conclusion

This study investigated the association between physical activity levels, insomnia, and depressive symptoms in middleaged and elderly patients with CS. The results pointed out that lower physical activity levels were influential factors in comorbid insomnia and depressive symptoms in CS patients. Due to the advantages of physical activity on psychotic symptoms, we should enhance the use of physical activity as a regular adjunct in clinical treatment or psychiatric care to improve the psychosomatic health and quality of life of patients with psychiatric symptoms.

Data availability statement

The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation.

Ethics statement

This study was reviewed and approved by the Biomedical Ethics Committee of Chaohu Hospital of Anhui Medical University (Grant no. 201805-kyxm-03). The patients/participants provided their written informed consent to participate in this study. Written informed consent was obtained from the individual(s) for the publication of any potentially identifiable images or data included in this article.

Author contributions

ZL, YZ, LS, JW, and HL collected and statistically analyzed the data and wrote the first draft. All authors revised and approved the manuscript.

Funding

This study was funded by grants from the Anhui Institute of Translational Medicine Research Fund (No.

References

1. World Health Organization [WHO]. *Mental health of older adults.* Geneva: World Health Organization (2017).

2. Whiteford H, Ferrari A, Degenhardt L, Feigin V, Vos T. The global burden of mental, neurological and substance use disorders: an analysis from the global burden of disease study 2010. *PLoS One*. (2015) 10:e0116820. doi: 10.1371/journal. pone.0116820

3. Huang Y, Wang Y, Wang H, Liu Z, Yu X, Yan J, et al. Prevalence of mental disorders in China: a cross-sectional epidemiological study. *Lancet Psychiatry*. (2019) 6:211-24. doi: 10.1016/s2215-0366(18)30511-x

 Charlson F, Baxter A, Cheng H, Shidhaye R, Whiteford H. The burden of mental, neurological, and substance use disorders in China and India: a systematic analysis of community representative epidemiological studies. *Lancet.* (2016) 388:376–89. doi: 10.1016/s0140-6736(16)30590-6

5. Jauhar S, Johnstone M, McKenna P. Schizophrenia. *Lancet.* (2022) 399:473–86. doi: 10.1016/s0140-6736(21)01730-x

6. Jongsma H, Turner C, Kirkbride J, Jones P. International incidence of psychotic disorders, 2002-17: a systematic review and meta-analysis. *Lancet Public Health*. (2019) 4:e229–44. doi: 10.1016/s2468-2667(19)30056-8

7. Batalla-Martín D, Belzunegui-Eraso A, Miralles Garijo E, Martínez Martín E, Romaní Garcia R, Heras J, et al. Insomnia in schizophrenia patients: prevalence and quality of life. *Int J Environ Res Public Health*. (2020) 17:1350. doi: 10.3390/ ijerph17041350

8. Miller B, McCall W, Xia L, Zhang Y, Li W, Yao X, et al. Insomnia, suicidal ideation, and psychopathology in Chinese patients with chronic schizophrenia. *Prog Neuropsychopharmacol Biol Psychiatry.* (2021) 111:110202. doi: 10.1016/j. pnpbp.2020.110202

2017zhyx17), the 2019 National Clinical Key Discipline Ability Enhancement Project (Provincial Project), and the Anhui Provincial Key R&D Programme (No. 202004j07020030).

Acknowledgments

We thank all participants and the staff in the clinical department.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

9. Hombali A, Seow E, Yuan Q, Chang S, Satghare P, Kumar S, et al. Prevalence and correlates of sleep disorder symptoms in psychiatric disorders. *Psychiatry Res.* (2019) 279:116–22. doi: 10.1016/j.psychres.2018.07.009

10. Kiwan N, Mahfoud Z, Ghuloum S, Chamali R, Yehya A, Hammoudeh S, et al. Self-reported sleep and exercise patterns in patients with schizophrenia: a cross-sectional comparative study. *Int J Behav Med.* (2020) 27:366–77. doi: 10.1007/s12529-019-09830-2

11. Taliercio J, Bonasera B, Portillo C, Ramjas E, Serper M. Physical activity, sleep-related behaviors and severity of symptoms in schizophrenia. *Psychiatry Res.* (2020) 294:113489. doi: 10.1016/j.psychres.2020.113489

12. Imboden C, Claussen M, Seifritz E, Gerber M. The importance of physical activity for mental health. *Praxis.* (2022) 110:e186–90. doi: 10.1024/1661-8157/a003820

13. Costa R, Bastos T, Probst M, Seabra A, Abreu S, Vilhena E, et al. Association of lifestyle-related factors and psychological factors on quality of life in people with schizophrenia. *Psychiatry Res.* (2018) 267:382–93. doi: 10.1016/j.psychres.2018.06. 022

14. Pengpid S, Peltzer K. Physical activity, health and well-being among a nationally representative population-based sample of middle-aged and older adults in India, 2017-2018. *Heliyon*. (2021) 7:e08635. doi: 10.1016/j.heliyon.2021.e08635

15. Grover S BN. Depression in schizophrenia: prevalence and its impact on quality of life, disability, and functioning. *Asian J Psychiatr.* (2020) 54:102425. doi: 10.1016/j.ajp.2020.102425

16. Li W, Yang Y, An F, Zhang L, Ungvari G, Jackson T, et al. Prevalence of comorbid depression in schizophrenia: a meta-analysis of observational studies. *J Affect Disord.* (2020) 273:524–31. doi: 10.1016/j.jad.2020.04.056

17. Golubović B, Gajić Z, Ivetić O, Milatović J, Vuleković P, Đilvesi Đ, et al. Factors associated with depression in patients with schizophrenia. *Acta Clin Croat.* (2020) 59:605–14. doi: 10.20471/acc.2020.59.04.06

18. Leutwyler H, Hubbard E, Jeste D, Miller B, Vinogradov S. Associations of schizophrenia symptoms and neurocognition with physical activity in older adults with schizophrenia. *Biol Res Nurs.* (2014) 16:23–30. doi: 10.1177/1099800413500845

19. Rosenbaum S, Tiedemann A, Sherrington C, Curtis J, Ward P. Physical activity interventions for people with mental illness: a systematic review and meta-analysis. J Clin Psychiatry. (2014) 75:964–74. doi: 10.4088/JCP.13r08765

20. Gouveia R, Ferreira-Junior A, Schuch F, Zanetti G, da Silva A, Del-Ben C, et al. Physical activity, quality of life and global functioning an early stages of psychosis. *Psychiatr Danub*. (2020) 32:373–9. doi: 10.24869/psyd.2020.373

21. Oliveira JS, Gilbert S, Pinheiro MB, Tiedemann A, Macedo LB, Maia L, et al. Effect of sport on health in people aged 60 years and older: a systematic review with meta-analysis. *Br J Sports Med.* (2022). [Epub ahead of print]. doi: 10.1136/bjsports-2022-105820

22. Tang Z, Wang Y, Liu J, Liu Y. Effects of aquatic exercise on mood and anxiety symptoms: a systematic review and meta-analysis. *Front Psychiatry*. (2022) 13:1051551. doi: 10.3389/fpsyt.2022.1051551

23. Choi J, Cho Y, Kim Y, Lee S, Lee J, Yi Y, et al. The relationship of sitting time and physical activity on the quality of life in elderly people. *Int J Environ Res Public Health*. (2021) 18:1459. doi: 10.3390/ijerph18041459

24. Luo J, Liu H, Liu Y, Jiang F, Tang Y. Physical activity and mental health among physicians in tertiary psychiatric hospitals: a national crosssectional survey in China. *Front Psychol.* (2021) 12:731525. doi: 10.3389/fpsyg.2021.731525

25. Hakimi S, Kaur S, Ross-White A, Martin L, Rosenberg M. A systematic review examining associations between physical activity, sedentary behaviour, and sleep duration with quality of life in older adults aged 65 years and above. *Appl Physiol Nutr Metab.* (2022). [Epub ahead of print]. doi: 10.1139/apnm-2022-0298

26. Wang J, Zhang Y, Yang Y, Liu Z, Xia L, Li W, et al. The prevalence and independent influencing factors of obesity and underweight in patients with schizophrenia: a multicentre cross-sectional study. *Eat Weight Disord.* (2021) 26:1365–74. doi: 10.1007/s40519-020-00920-9

27. Leucht S, Samara M, Heres S, Davis J. Dose equivalents for antipsychotic drugs: the DDD method. *Schizophr Bull.* (2016) 42:S90–4. doi: 10.1093/schbul/sbv167

28. Wang S, Lay S, Yu H, Shen S. Dietary guidelines for Chinese residents (2016): comments and comparisons. *J Zhejiang Univ Sci B.* (2016) 17:649–56. doi: 10.1631/jzus.B1600341

29. Bull F, Al-Ansari S, Biddle S, Borodulin K, Buman M, Cardon G, et al. World health organization 2020 guidelines on physical activity and sedentary behaviour. *Br J Sports Med.* (2020) 54:1451–62. doi: 10.1136/bjsports-2020-102955

30. Wu B, Lan T, Hu T, Lee S, Liou J. Validation of a five-factor model of a Chinese Mandarin version of the positive and negative syndrome scale (CMV-PANSS) in a sample of 813 schizophrenia patients. *Schizophr Res.* (2015) 169:489–90. doi: 10.1016/j.schres.2015.09.011

31. Bastien C, Vallières A, Morin C. Validation of the insomnia severity index as an outcome measure for insomnia research. *Sleep Med.* (2001) 2:297–307. doi: 10.1016/s1389-9457(00)00065-4

32. Yu D. Insomnia severity index: psychometric properties with Chinese community-dwelling older people. J Adv Nurs. (2010) 66:2350–9. doi: 10.1111/j. 1365-2648.2010.05394.x

33. Han J, Feng Y, Li N, Feng L, Xiao L, Zhu X, et al. Correlation between word frequency and 17 items of hamilton scale in major depressive disorder. *Front Psychiatry.* (2022) 13:902873. doi: 10.3389/fpsyt.2022.902873

34. Parletta N, Aljeesh Y, Baune B. Health behaviors, knowledge, life satisfaction, and wellbeing in people with mental illness across four countries and comparisons with normative sample. *Front Psychiatry.* (2016) 7:145. doi: 10.3389/fpsyt.2016. 00145

35. Xie Y, Liu S, Chen X, Yu H, Yang Y, Wang W. Effects of exercise on sleep quality and insomnia in adults: a systematic review and meta-analysis of randomized controlled trials. *Front Psychiatry.* (2021) 12:664499. doi: 10.3389/fpsyt.2021.664499

36. Zeibig J, Seiffer B, Sudeck G, Rösel I, Hautzinger M, Wolf S. Transdiagnostic efficacy of a group exercise intervention for outpatients with heterogenous psychiatric disorders: a randomized controlled trial. *BMC Psychiatry*. (2021) 21:313. doi: 10.1186/s12888-021-03307-x

37. Duncan M, Holliday E, Burton N, Glozier N, Oftedal S. Prospective associations between joint categories of physical activity and insomnia symptoms with onset of poor mental health in a population-based cohort. *J Sport Health Sci.* (2022). [Epub ahead of print]. doi: 10.1016/j.jshs.2022.02.002

38. Stubbs B, Vancampfort D, Firth J, Hallgren M, Schuch F, Veronese N, et al. Physical activity correlates among people with psychosis: data from 47 low- and middle-income countries. *Schizophr Res.* (2018) 193:412–7. doi: 10.1016/j.schres. 2017.06.025

39. Dauwan M, Begemann M, Heringa S, Sommer I. Exercise improves clinical symptoms, quality of life, global functioning, and depression in schizophrenia: a systematic review and meta-analysis. *Schizophr Bull.* (2016) 42:588–99. doi: 10. 1093/schbul/sbv164

40. Pearsall R, Smith D, Pelosi A, Geddes J. Exercise therapy in adults with serious mental illness: a systematic review and meta-analysis. *BMC Psychiatry.* (2014) 14:117. doi: 10.1186/1471-244x-14-117

41. Schmitt A, Maurus I, Rossner M, Röh A, Lembeck M, von Wilmsdorff M, et al. Effects of aerobic exercise on metabolic syndrome, cardiorespiratory fitness, and symptoms in schizophrenia include decreased mortality. *Front Psychiatry.* (2018) 9:690. doi: 10.3389/fpsyt.2018.00690

42. Sailer P, Wieber F, Pröpster K, Stoewer S, Nischk D, Volk F, et al. A brief intervention to improve exercising in patients with schizophrenia: a controlled pilot study with mental contrasting and implementation intentions (MCII). *BMC Psychiatry.* (2015) 15:211. doi: 10.1186/s12888-015-0513-y

43. Lederman O, Ward P, Firth J, Maloney C, Carney R, Vancampfort D, et al. Does exercise improve sleep quality in individuals with mental illness? A systematic review and meta-analysis. *J Psychiatr Res.* (2019) 109:96–106. doi: 10. 1016/j.jpsychires.2018.11.004

44. Firth J, Solmi M, Wootton R, Vancampfort D, Schuch F, Hoare E, et al. A meta-review of "lifestyle psychiatry": the role of exercise, smoking, diet and sleep in the prevention and treatment of mental disorders. *World Psychiatry.* (2020) 19:360–80. doi: 10.1002/wps.20773

45. Cohen C, Ryu HHA. Longitudinal study of the outcome and associated factors of subsyndromal and syndromal depression in community-dwelling older adults with schizophrenia spectrum disorder. *Am J Geriatr Psychiatry.* (2015) 23:925–33. doi: 10.1016/j.jagp.2014.06.011

46. Chen Y, Li W. Prevalence, influencing factors, and cognitive characteristics of depressive symptoms in elderly patients with schizophrenia. *Neuropsychiatr Dis Treat.* (2021) 17:3645–54. doi: 10.2147/ndt.s341297

47. Zou C, Chen S, Shen J, Zheng X, Wang L, Guan L, et al. Prevalence and associated factors of depressive symptoms among elderly inpatients of a Chinese tertiary hospital. *Clin Interv Aging*. (2018) 13:1755–62. doi: 10.2147/cia.s17 0346

48. Fekih-Romdhane F, Ben Ali S, Ghazouani N, Tira S, Cheour M. Burden in Tunisian family caregivers of older patients with schizophrenia spectrum and bipolar disorders; associations with depression, anxiety, stress, and quality of life. *Clin Gerontol.* (2020) 43:545–57. doi: 10.1080/07317115.2020.1728600

49. Hoertel N, Jaffré C, Pascal de Raykeer R, McMahon K, Barrière S, Blumenstock Y, et al. Subsyndromal and syndromal depressive symptoms among older adults with schizophrenia spectrum disorder: prevalence and associated factors in a multicenter study. *J Affect Disord*. (2019) 251:60–70. doi: 10.1016/j.jad. 2019.03.007

50. Aas M, Djurovic S, Ueland T, Mørch R, Fjæra Laskemoen J, Reponen E, et al. The relationship between physical activity, clinical and cognitive characteristics and BDNF mRNA levels in patients with severe mental disorders. *World J Biol Psychiatry.* (2019) 20:567–76. doi: 10.1080/15622975.2018.1557345

51. Ashdown-Franks G, Firth J, Carney R, Carvalho A, Hallgren M, Koyanagi A, et al. Exercise as medicine for mental and substance use disorders: a meta-review of the benefits for neuropsychiatric and cognitive outcomes. *Sports Med.* (2020) 50:151–70. doi: 10.1007/s40279-019-01187-6

52. Ng S, Leung T, Ng P, Ng R, Wong A. Activity participation and perceived health status in patients with severe mental illness: a prospective study. *East Asian Arch Psychiatry*. (2020) 30:95–100. doi: 10.12809/eaap1970

53. Marquez D, Aguiñaga S, Vásquez P, Conroy D, Erickson K, Hillman C, et al. A systematic review of physical activity and quality of life and well-being. *Transl Behav Med.* (2020) 10:1098–109. doi: 10.1093/tbm/ibz198

54. Schuch F, Vancampfort D. Physical activity, exercise, and mental disorders: it is time to move on. *Trends Psychiatry Psychother*. (2021) 43:177–84. doi: 10.47626/2237-6089-2021-0237

55. Scheewe T, Jörg F, Takken T, Deenik J, Vancampfort D, Backx F, et al. Low physical activity and cardiorespiratory fitness in people with schizophrenia: a comparison with matched healthy controls and associations with mental and physical health. *Front Psychiatry.* (2019) 10:87. doi: 10.3389/fpsyt.2019.00087

56. Sunhary De Verville P, Stubbs B, Etchecopar-Etchart D, Godin O, Andrieu-Haller C, Berna F, et al. Recommendations of the schizophrenia expert center network for adequate physical activity in real-world schizophrenia (FACE-SZ). *Eur Arch Psychiatry Clin Neurosci.* (2022)272:1273–82. doi: 10.1007/s00406-022-01384-x

57. Biviá-Roig G, Soldevila-Matías P, Haro G, González-Ayuso V, Arnau F, Peyró-Gregori L, et al. The impact of the COVID-19 pandemic on the lifestyles and levels of anxiety and depression of patients with schizophrenia: a retrospective observational study. *Healthcare*. (2022) 10:128. doi: 10.3390/healthcare10010128

58. Stubbs B, Firth J, Berry A, Schuch F, Rosenbaum S, Gaughran F, et al. How much physical activity do people with schizophrenia engage in? A systematic review, comparative meta-analysis and meta-regression. *Schizophr Res.* (2016) 176:431–40. doi: 10.1016/j.schres.2016.05.017

59. Falkai P, Schmitt A, Rosenbeiger C, Maurus I, Hattenkofer L, Hasan A, et al. Aerobic exercise in severe mental illness: requirements from the perspective of sports medicine. *Eur Arch Psychiatry Clin Neurosci.* (2022) 272:643–77. doi: 10. 1007/s00406-021-01360-x

60. Stubbs B, Koyanagi A, Schuch F, Firth J, Rosenbaum S, Gaughran F, et al. Physical activity levels and psychosis: a mediation analysis of factors influencing physical activity target achievement among 204 186 people across 46 low- and middle-income countries. *Schizophr Bull.* (2017) 43:536–45. doi: 10.1093/schbul/ sbw111

61. Felez-Nobrega M, Haro J, Stubbs B, Smith L, Koyanagi A. Moving more, ageing happy: findings from six low- and middle-income countries. *Age Ageing.* (2021) 50:488–97. doi: 10.1093/ageing/afaa137