

Exploring the relationship between education, living environment, and anxiety/depression among stable patients: insights from the COPD-AD China Registry study

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Research Article

Keywords: Anxiety, China, COPD, depression, education, living environment.

Posted Date: May 22nd, 2023

DOI: <https://doi.org/10.21203/rs.3.rs-2877766/v1>

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Additional Declarations: No competing interests reported.

Abstract

Background

The relationship between education and living environment is complex, and both factors are related to mental health. However, few studies have examined the independent and combined effects of education and living environment on mental health among patients with chronic obstructive pulmonary disease (COPD).

Methods

We analyzed data for 1064 patients with COPD enrolled in the COPD-AD China Registry study from 2017 to 2021. Binary logistic regression was used to analyze the independent and combined effects of education and living environment on patients' mental health. We also evaluated additive interactions with the relative excess risk ratio (RERI), attribution percentage (AP), and synergy index (SI).

Results

Our primary analysis revealed the incidence of anxiety and depression was associated with educational level and living environment. After adjustment for potential confounding factors, binary logistic regression analyses showed that low education level and urban living environment were independently associated with higher risks for anxiety (odds ratio [OR]: 1.56, 95% confidence interval [CI] 1.06–2.29 and OR:2.15, 95% CI 1.51–2.05) and depression (OR:1.62, 95% CI 1.17–2.27 and OR: 2.01, 95% CI 1.46–2.75) among patients with COPD. Our secondary analysis showed that the combination of low education level and living in urban areas was associated with the highest and significantly increased risk for anxiety (OR: 7.90, 95% CI 3.83–16.29, $P < .001$) or depression (OR: 11.79, 95% CI 5.77–24.10, $P < .001$) among these patients. Furthermore, we observed strong synergistic additive interactions between low education level and living in urban areas for anxiety (SI: 11.57, 95% CI 1.41–95.27; RERI: 6.31, 95% CI 1.60–11.01; AP: 0.8, 95% CI 0.66–0.94) and depression (SI: 31.31, 95% CI 1.59–617.04; RERI: 10.44, 95% CI 2.66–18.23; AP: 0.89, 95% CI 0.8–0.97).

Conclusions

Our study suggests that mental health among patients with COPD is influenced by their education and living environment. Patients who have low education levels and live in urban areas may experience more anxiety and depression.

Trial registration

The clinical trial registration code was obtained from the ClinicalTrials.gov (date of registration: 14/06/2017, registration number: NCT03187236).

Introduction

Chronic obstructive pulmonary disease (COPD) is a prevalent chronic airway disease that often co-occurs with anxiety and depression [1]. Our previous research showed that 13.93% of patients with stable COPD reported symptoms of anxiety, and 23.37% experienced depression [2]. Moreover, patients with COPD that had a history of anxiety or depression exhibited poor physical function, impaired health status, and frequent acute exacerbations [3, 4]. Therefore, identifying potentially modifiable risk factors that may improve mental health among patients with COPD is important.

Education and urban living environment are important factors that affect an individual's mental health [5, 6]. Education can positively impact mental health by enabling individuals to succeed and pursue fundamental goals [6], and a low education level can contribute to poor mental health. In addition, an urban living environment directly and indirectly affects residents' mental health through factors such as congestion, crime, high-rise housing, noise pollution, air pollution, traffic congestion, poor housing, and heightened competition [7]. Therefore, it is necessary to consider education and living environment when addressing mental health issues in different populations. Importantly, education has complex interactions with a person's living environment [8]. For example, a previous study found that migrant residents with a tertiary education had a higher incidence of depressive symptoms compared with Italian-born residents [9]. This suggested the protective effect of education against anxiety and depression may be influenced by other social-contextual variables. Although researches have suggested that systemic inflammation, gender, smoking, acute exacerbation of COPD, hypoxemia, and disease knowledge level were risk factors for anxiety and depression symptoms among patients with COPD [1, 10, 11]. However, few studies have examined the independent and combined effects of education and living environment on mental health among patients with COPD.

When we collated the Chronic Obstructive Pulmonary Disease with Anxiety and Depression in China (COPD-AD China Registry) study database, we found that patients participating in that study originated from diverse urban and rural regions, had varied educational backgrounds, and some suffered from symptoms of anxiety and depression. Therefore, this paper aimed to evaluate the independent effects of education and living environment on mental health among patients with COPD and determine whether the impact of education on mental health differed between rural and urban living environments. This information will support a better understanding of factors that contribute to mental health issues in patients with COPD and facilitate development of targeted interventions to improve their mental well-being.

METHODS

Data Source

The COPD-AD China Registry study comprises a comprehensive database. This database was a key national research and development project focused on COPD in China, and involved a national clinical registration study that was initiated in June 2017 and lasted for 3.5 years (Clinical Trials ID: NCT03187236).

Study Cohort

Patients that were selected for this analysis were from the respiratory medicine outpatient departments in nine Chinese hospitals (Liangxiang Hospital of Fangshan District, Beijing; Affiliated Hospital of Binzhou Medical College; Ordos Central Hospital; Guangdong Provincial People's Hospital; Affiliated Hospital of Guangdong Medical College; the First Affiliated Hospital of Nanchang University; Qilu Hospital of Shandong University; Beijing Chaoyang Hospital of Capital Medical University; and the People's Hospital of Wuhan University). All patients were aged over 40 years and diagnosed with COPD via a lung function test following the 2017 Global Initiative Chronic Obstructive Lung Disease (GOLD): Global Strategy for the Diagnosis, Management, and Prevention of COPD [12]. None of the included patients were pregnant. Patients were informed about the purpose of the study and signed a consent form. Between 2017 and 2021, 1366 participants were enrolled in the COPD-AD China Registry Study. Patients with incomplete data (n = 233) were excluded from this analysis. Further exclusion criteria were: severe cognitive disorder; suicide attempt or psychiatric hospitalization in the past year; current suicidal ideation with plan or intent; mental illness, such as dementia, autism, schizophrenia (n = 3), or bipolar disorder; alcohol or drug abuse (n = 10); bronchial asthma (n = 24); bronchiectasis (n = 12); pulmonary fibrosis (n = 2); tuberculosis (n = 8); neuromuscular disease (n = 2); and cancer (n = 8). After applying the exclusion criteria, 1064 participants were included in the final baseline study sample.

Measures

Clinical measurements and spirometry

Demographic characteristics (age, sex, height, weight, body mass index [BMI], smoking history, education, living environment) and clinical characteristics, including oxygen therapy status, Charlson Comorbidity Index (CCI), COPD Assessment Test (CAT) score, and modified Medical Research Council (mMRC) score, were collected from the data management network (mzf.fwncpc.com). We performed spirometry according to the guidelines for lung function tests formulated by the Chinese Thoracic Society. We measured the percentage predicted forced expiratory volume in 1 second (FEV₁% predicted) for all participating patients.

Urban-rural classification

The Chinese administrative system has six levels: national, provincial, prefectural, county, township, and village levels. Each county-level unit (county or district) usually includes two types of township level units: townships and subdistricts. The former comprises a town center and dozens of surrounding villages. The

latter comprises urban communities or suburbs. We defined the rural population as people living in townships, and the urban population as people living in subdistricts [13]. There were no rural townships in 17 of the studied urban districts; therefore, only an urban population was defined in these districts. Data for subdistrict populations were unavailable for five counties with predominantly rural populations, and we treated the population of these entire counties as rural populations. Therefore, we defined 655 sub-county level units for assessment overall: 97 urban units and 558 rural units. There were 32 units with small populations, including seven urban units and 25 rural units.

Education classification

Education level was classified by years of formal education received: illiteracy (0 years), primary school (1–6 years), junior secondary school (7–9 years), senior secondary school (10–12 years), and university (≥ 13 years). We used 9 years as the cut-off value to categorize participants as having a low (≤ 9 years) or a high (> 9 years) education level because China has free compulsory education for 9 years.

Determination of anxiety and depression

Anxiety and depression were measured with the Hospital Anxiety and Depression Scale (HADS). The HADS comprises 14 items that detect anxiety and depression in patients with chronic diseases. There are seven anxiety items and seven depression items; each item is rated on a Likert scale from 0 to 3. The total scores for the anxiety and depression items are calculated separately, with scores of 0–7 indicating no anxiety/depression, 8–10 indicating mild anxiety/depression, 11–14 indicating moderate anxiety/depression, and 15–21 indicating severe anxiety/depression [14].

Statistical Analysis

All statistical analyses were performed with SPSS version 25.0 (IBM Corp., Armonk, NY, USA), with the level of statistical significance set at $P < .05$. Categorical data were reported as number (percentage), and continuous variables were presented as mean \pm standard deviation (SD) for normally distributed data. Chi-square tests were used for comparisons of categorical variables, and independent t -tests were used for comparisons of normally distributed continuous variables. Binary logistic regression was used to analyze the effects of education and living environment (individually or in combination) on the risk for anxiety and depression among patients with COPD. To investigate the additive interaction of education and living environment on the risk for anxiety and depression among patients with COPD, the Microsoft Excel table compiled by Andersson et al. was used to calculate the relative excess risk (RERI), attributive ratio (AP), and synergy index (SI) for the additive interactions [15]. Two models were used in this regression analysis. Model 1 was unadjusted, and Model 2 was adjusted for confounder variables, including age, sex, smoking history, BMI, CAT, FEV₁% predicted, oxygen therapy, mMRC score, and CCI score. We adjusted for these variables because we considered they were related to anxiety and depression.

Results

Participants' Characteristics

In total, 1064 patients with COPD were included in our analyses: 211 (19.8%) were complicated with anxiety and 319 (30.0%) were complicated with depression. Patients mean age was 69.97 years (SD: 8.68 years, range: 60–88 years). We observed significant differences in CAT scores, CCI scores, oxygen intake, education, and living environment between participants with and without anxiety. There were significant differences in CAT scores, FEV₁% predicted, CCI scores, oxygen intake, education, and living environment between patients with and without depression (Table 1).

Table 1
Characteristics of study participants.

Variables	COPD (N = 1064)		P	COPD (N = 1064)		P
	No anxiety n = 853 (80.2)	Anxiety n = 211 (19.8)		No depression n = 745 (70.0)	Depression n = 319 (30.0)	
Age (years), mean ± SD	70.07 ± 8.50	69.58 ± 9.36	0.466	69.98 ± 8.61	69.96 ± 8.84	0.974
Male, n (%)	677 (80.0)	169 (20.0)	0.815	592 (70.0)	254 (30.0)	0.953
BMI (kg/m ²), mean ± SD	21.84 ± 4.08	22.10 ± 4.09	0.399	21.74 ± 3.97	22.24 ± 4.32	0.068
CAT, mean ± SD	14.17 ± 6.83	17.67 ± 7.66	< 0.01	13.97 ± 6.97	16.98 ± 7.19	< 0.01
FEV ₁ % predicted, mean ± SD	51.13 ± 21.30	53.95 ± 22.42	0.089	50.67 ± 21.09	54.07 ± 22.43	0.018
CCI, mean ± SD	3.14 ± 1.22	2.88 ± 1.13	0.005	3.14 ± 1.24	2.96 ± 1.14	0.027
mMRC, n (%)			0.252			0.998
0–1	359(78.6)	98(21.4)		320(70.0)	137(30.0)	
2–4	494(81.4)	113(18.6)		425(70.0)	182(30.0)	
Oxygen therapy, n (%)			< 0.01			0.001
No	758(82.5)	161(17.5)		661(71.9)	258(28.1)	
Yes	95(65.5)	50(34.5)		84(57.9)	61(42.1)	
Smoker, n (%)	619(79.3)	162(20.7)	0.215	541(69.3)	240(30.7)	0.376
Education level, n (%)			0.021			0.004
High education level	241(84.9)	43(15.1)		218 (76.8)	66(23.2)	
Low education level	612(78.5)	168(21.5)		527(67.6)	253(32.4)	
Living environment, n(%)			< 0.01			< 0.01
Rural living environment	673(83.5)	133(16.5)		596(73.9)	210(26.1)	
Urban living environment	180(69.8)	78(30.2)		149(57.8)	109 (42.2)	

BMI, body mass index; FEV₁% predicted, percentage predicted forced expiratory volume in 1 second; CAT, COPD Assessment Test; CCI, Charlson Comorbidity Index; mMRC, modified Medical Research Council; SD, standard deviation.

Primary Analysis

Relationship between education and risk for anxiety/depression among patients with COPD

Patients with COPD with a low education level had an increased risk for anxiety and depression compared with those with a high education level. This effect remained after adjustment for potential confounding factors (odds ratio [OR] 1.56, 95% confidence interval [CI] 1.06–2.29 and OR 1.62, 95%CI 1.17–2.27, respectively) (Table 2).

Relationship between living environment and risk for anxiety/depression among patients with COPD

Patients with COPD living in urban areas had an increased risk for anxiety and depression compared with those living in rural areas. This effect remained after adjustment for potential confounding factors (OR 2.15, 95%CI 1.51–3.05 and OR 2.01, 95%CI 1.46–2.75, respectively) (Table 2).

Table 2

Risk for anxiety or depression among patients with COPD by education level and living environment.

Education level and living environment	Anxiety		Depression	
	Model 1	Model 2	Model 1	Model 2
Education level				
High education level	1	1	1	1
Low education level	1.54 (1.07–2.22) *	1.56 (1.06–2.29) *	1.59 (1.16–2.17) **	1.62 (1.17–2.27) *
Living environment				
Rural living environment	1	1	1	1
Urban living environment	2.19 (1.59–3.03) **	2.15 (1.51–3.05) **	2.08 (1.55–2.78) **	2.01 (1.46–2.75) **

Model 1: non-adjusted. Model 2: adjusted for age, sex, smoking history, body mass index, COPD Assessment Test, percentage predicted forced expiratory volume in 1 second; oxygen therapy, modified Medical Research Council score, and Charlson Comorbidity Index.

* $P < .05$, ** $P < .01$.

Secondary Analysis

We used the binary educational level indicator (high education level vs. low education level) and living environment indicator (rural living environment vs. urban living environment) to examine the combined interaction. As expected, the combination group of low education level and urban living environment showed the highest and significantly elevated risk for anxiety or depression among patients with COPD (OR 9.14, 95%CI 4.56–18.32, $P < .001$ and OR 13.76, 95%CI 6.88–27.53, respectively; $P < .001$). The correlation was only slightly weakened after adjusting for potential confounding factors (OR 7.90, 95%CI 3.83–16.29, $P < .001$ and OR 11.79, 95%CI 5.77–24.10, respectively; $P < .001$) (Table 3).

Table 3

Combined effect of education level (high vs. low) and living environment (rural vs. urban) used to calculate interactions.

Combined effect (Educational level and living environment)		Anxiety		Depression	
		Model 1	Model 2	Model 1	Model 2
High educational level	Rural living environment	1	1	1	1
Low educational level	Rural living environment	1.31 (0.73–2.34)	1.34 (0.74–2.42)	1.25 (0.78–2.01)	1.28 (0.78–2.09)
High educational level	Urban living environment	1.24 (0.63–2.44)	1.26 (0.62–2.55)	1.07 (0.61–1.88)	1.07 (0.59–1.93)
Low educational level	Urban living environment	9.14 (4.56–18.32) **	7.90 (3.83–16.29) **	13.76 (6.88–27.53) **	11.79 (5.77–24.10) **

Model 1: non-adjusted. Model 2: adjusted for age, sex, smoking history, body mass index, COPD Assessment Test, percentage predicted forced expiratory volume in 1 second, oxygen therapy, modified Medical Research Council score, and Charlson Comorbidity Index.

* $P < .05$, ** $P < .01$.

Figure 1 shows the excess risks due to low education level, urban living environment, and their interactions in an analysis of the risk for anxiety (A) or depression (B) among patients with COPD after adjusting for all confounders. The STROBE Statement advises reporting interaction analyses using an additive and multiplicative scale when evaluating the joint effect of exposures [16]. This is because an additive interaction model is closer to the nature of biological interaction and has more relevant public health significance than a multiplication model [17]. Therefore, as recommended in a previous study, we used an excel sheet to calculate the additive interaction [15]. As shown in Table 4, the estimated RERI values were 6.31 (95%CI 1.60–11.01) for anxiety and 10.44 (95%CI 2.66–18.23) for depression,

indicating that of the additive interaction between low education level and urban living environment resulted in 6.31 relative excess risks for anxiety and 10.44 relative excess risks for depression. The AP revealed that 80% of the total odds of anxiety and 89% of the total odds of depression were attributable to the interaction between low education level and urban living environment. In addition, SI values were 11.57 (95% CI 1.41–95.27) and 31.31 (95%CI 1.59–617.04), suggesting that the risks for anxiety or depression in patients with both low education level and an urban living environment were 11.57 and 31.31 times as high as the sum of risks for patients presenting with a single factor.

Table 4

Measures for estimation of biological interaction between low education level and urban living environment for the risk for anxiety or depression among patients with COPD.

Measures of biological interaction	Anxiety		Depression	
	Model 1	Model 2	Model 1	Model 2
RERI (95% CI)	7.60 (1.91–13.29)	6.31 (1.60–11.01)	12.45 (3.25–21.64)	10.44 (2.66–18.23)
AP (95% CI)	0.83 (0.72–0.95)	0.80 (0.66–0.94)	0.90 (0.84–0.97)	0.89 (0.80–0.97)
SI (95% CI)	14.95 (2.21–101.16)	11.57 (1.41–95.27)	40.45 (1.57–1043.07)	31.31 (1.59–617.04)

Note: *If there was no biological interaction the 95%CI for RERI and AP would include 0 and that for SI would include 1.

RERI: relative excess risk due to interaction; AP: attribution ratio; SI: synergy index; CI: confidence interval.

Model 1: non-adjusted. Model 2: adjusted for age, sex, smoking history, body mass index, COPD Assessment Test, percentage predicted forced expiratory volume in 1 second, oxygen therapy, modified Medical Research Council score, and Charlson Comorbidity Index.

Discussion

The COPD-AD China Registry study is a nationwide registration study in China that has a comprehensive database that tracks patients with COPD who also suffer from anxiety or depression. Its goal is to screen for anxiety and depression among patients with COPD at an early stage, and track how these conditions impact the overall progression and prognosis of COPD. In this cross-sectional study, we investigated a sample of 1064 patients with COPD to evaluate the independent and combined effects of education level and living environment on their mental health.

Our results showed that the incidences of anxiety and depression were associated with education and living environment. After adjustment for potential confounding factors, binary logistic regression analyses showed that a lower education level and an urban living environment were independently associated with higher risks for anxiety and depression among patients with COPD. This suggested that

lower education level and urban living environment were stable factors predicting the risk for anxiety and depression among patients with COPD. Education is a comprehensive reflection of a person's knowledge, skills, and values, and has a profound impact on that individual's cognition, behavior, and emotions [18, 19]. Patients with COPD with a lower education level may lack relevant health knowledge, lack understanding and confidence about their disease and treatment methods, and feel helpless and fearful. They may also be more dependent on the advice of doctors and lack the ability and willingness to self-manage than patients with a higher education level [20–23]. In addition, they may face social barriers and discrimination and feel lonely and lost. These factors may lead to them being more susceptible to symptoms of anxiety and depression. Living environment refers to the conditions in which a person lives, and includes social, economic, and cultural aspects. An urban living environment has previously been considered an important factor affecting mental health [24]. Urban living environments have serious air pollution that impacts patients with COPD. Exhaust from cars, industrial waste gases, and dust from construction sites and other sources of pollution are constantly emitted into the atmosphere, causing concentrations of harmful substances such as PM_{2.5} in the air [25, 26]. The respiratory system of patients with COPD has been damaged, and air pollution can therefore aggravate respiratory difficulties and symptoms [27]. Moreover, there is serious noise pollution in urban living environments [28]. Traffic, industrial, and community noise sources continuously interfere with the lives of patients with COPD. These noises may reduce the quality of patients' sleep and affect their emotional and mental state [29, 30]. Finally, there may also be social problems in urban living environments such as tense interpersonal relationships [31]. The pace of urban life is fast, and interpersonal relationships are complex and many patients with COPD may feel lonely. All of these factors may cause patients to feel lost and helpless, further exacerbating their anxiety and depression. Therefore, we believed that a low education level and an urban living environment may increase the risks for anxiety and depression in patients with COPD.

We also examined the combined effect of a low education level and an urban living environment on the risks for anxiety and depression in patients with COPD. Our findings indicated that the combination of these two factors was statistically significant and exhibited a positive interactive effect. Specifically, the risk for anxiety or depression in patients with COPD was higher when they had a low education level and lived in urban areas, with this interactive effect being greater than the sum of their individual effects (11.57 times for anxiety and 31.31 times for depression). There are several possible explanations for the synergistic interaction between a low education level and an urban living environment in terms of anxiety and depression. First, these risk factors are associated with increased stress. Living in an urban area can be stressful because of noise pollution, traffic congestion, and overcrowding. In addition, a low education level may be associated with lower socioeconomic status, which can lead to stress related to financial difficulties and inadequate access to healthcare [32–34]. Second, a low education level and an urban living environment may impact patient's social support network and social activities. A low education level may limit individuals' access to resources and opportunities, which in turn may impact their social support networks and social activities. In addition, the fast-paced, busy lifestyle of an urban living environment often leaves people without sufficient time and opportunities to connect with others and participate in social activities [35, 36]. Third, a low education level may lead to a lack of necessary health

knowledge and behaviors when facing health problems, resulting in a deterioration of health conditions. This situation can leave patients feeling frustrated and helpless, and in combination with the fast-paced, high-pressure, and competitive urban living environment, may also make individuals feel lonely, anxious, and even more helpless, meaning they feel unable to cope with the challenges of their illness and life [37, 38]. Therefore, it is crucial for healthcare professionals to consider multiple risk factors when assessing risks for anxiety and depression among patients with COPD, especially those with low education levels that live in urban areas. Health policymakers should develop targeted interventions to reduce the risk for anxiety and depression in patients with COPD by increasing the greening rate in urban areas, strengthening governance of noise and pollution, providing regular health services and cultural activities, and offering economic and psychological support.

There were certain limitations in our study that should be noted. First, this study was conducted among patients from nine hospitals in China, which may limit the generalizability of the findings to other settings. Second, this study used a cross-sectional design, which limited our ability to establish causal relationships between urban living environment, low education level, and anxiety and depression among patients with COPD. Longitudinal studies are needed to further explore these relationships. Finally, we did not assess other potential confounding variables, such as socioeconomic status and medication complexity, which may affect our results.

In conclusion, our study provides evidence that the mental health of patients with COPD is influenced by their education and living environment. Patients with low education levels that live in urban areas may experience more anxiety and depression than better educated, rural dwelling patients, which can be compounded by their interactive effects. It is crucial to consider educational and environmental factors for patients with COPD, offer them optimal medical care and social support, and help them manage psychological distress. These efforts can enhance their quality of life and treatment outcomes.

Abbreviations

COPD

Chronic obstructive pulmonary disease

COPD-AD China Registry

Chronic Obstructive Pulmonary Disease with Anxiety and Depression in China

GOLD

Global Initiative for Chronic Obstructive Lung Disease

BMI

Body mass index

CAT

COPD Assessment Test

mMRC

Modified Medical Research Council

FEV₁% predicted

Percentage predicted forced expiratory volume in 1 second

HADS

Hospital Anxiety and Depression Scale

SD

Standard deviation

RERI

Relative excess risk

AP

Attributive ratio

SI

Synergy index

OR

Odds ratio

CI

Confidence interval.

Declarations

Ethics approval and consent to participate

The research was approved by the Ethics Committee of the Affiliated Hospital of Guangdong Medical University with the committee's reference number 2017-879 and the registration number is NCT03187236, all participants have provided their written informed consent for study participation. All methods were performed in accordance with the Declaration of Helsinki in the study.

Consent for publication

Not applicable.

Availability of data and materials

The datasets used and/or analyzed during the current study available from the corresponding author on reasonable request.

Competing interests

The authors declare that they have no competing interests.

Funding

This research was supported by the National Key Research and Development Program of China (grant number: 2016YFC1304404), the Guangdong Medical Science and Technology Research Fund Project (grant number: B2021294), the Discipline Construction Project of Guangdong Medical University (grant

number: 4SG21231G), the Clinical Research Project of Affiliated Hospital of Guangdong Medical University (grant numbers: LCYT2017A003, LCYJ2020B008, LCYJ2021B007, LCYJ2022DL01), and the Project of Zhanjiang City (grant numbers: 2021A05052, 2021A05077).

Authors' contributions

BW, DL, MC and JW conceived the study design and are responsible for the overall content. WZ, QH, XS, XC and XZ analyzed and interpreted the data. XZ, BL and BH prepared the manuscript. DW, DH and DL edited the manuscript. All authors approved the submitted and final versions.

Acknowledgment

We thank Analisa Avila, MPH, ELS, of Liwen Bianji (Edanz) (www.liwenbianji.cn) for editing the language of a draft of this manuscript, all the patients participating in the study and all the staff involved at the hospital.

References

1. Martínez-Gestoso S, García-Sanz M-T, Carreira J-M, Salgado F-J, Calvo-Álvarez U, Doval-Oubiña L, et al. Impact of anxiety and depression on the prognosis of copd exacerbations. *BMC Pulmonary Medicine*. 2022;22(1):169.
2. Wu D, Zhao X, Huang D, Dai Z, Chen M, Li D, et al. Outcomes associated with comorbid anxiety and depression among patients with stable COPD: A patient registry study in China. *J Affect Disord*. 2022;313:77-83.
3. Ouellette DR, Lavoie KL. Recognition, diagnosis, and treatment of cognitive and psychiatric disorders in patients with COPD. *International Journal of Chronic Obstructive Pulmonary Disease*. 2017;12:639-50.
4. Wen-Tao D, Xue-Xiu C, Zun-Jiang C, Wei C, Cheng-Feng P, Xing-Ken F. The Relationship Between Hospitalization Frequency of Acute Exacerbation of Chronic Obstructive Pulmonary Disease and Anxiety and Depression. *Front Genet*. 2022;13:817727.
5. Bjelland I, Krokstad S, Mykletun A, Dahl AA, Tell GS, Tambs K. Does a higher educational level protect against anxiety and depression? The HUNT study. *Soc Sci Med*. 2008;66(6):1334-45.
6. McKenzie K, Murray A, Booth T. Do urban environments increase the risk of anxiety, depression and psychosis? An epidemiological study. *J Affect Disord*. 2013;150(3):1019-24.
7. Yang X. Rural-urban migration and mental and sexual health: a case study in Southwestern China. *Health Psychol Behav Med*. 2014;2(1).
8. Wang Y, Jiang Y, Wu W, Xu K, Zhao Q, Tan Z, et al. Education, neighborhood environment, and cognitive decline: Findings from two prospective cohort studies of older adults in China. *Alzheimers Dement*. 2023;19(2):560-8.

9. Sesti F, Minardi V, Baglio G, Bell R, Goldblatt P, Marceca M, et al. Social determinants of mental health in Italy: the role of education in the comparison of migrant and Italian residents. *Int J Equity Health*. 2022;21(1):116.
10. Zhang Q, Liao J, Liao X, Wu X, Wan M, Wang C, et al. Disease knowledge level is a noteworthy risk factor of anxiety and depression in patients with chronic obstructive pulmonary disease: a cross-sectional study. *BMC Pulmonary Medicine*. 2014;14:92.
11. Pelgrim CE, van den Heuvel JM, Folkerts G, Garssen J, Maitland-van der Zee AH, Kraneveld AD. Higher prescription of antidepressants and/or anxiolytics among chronic obstructive pulmonary disease patients. *Therapeutic Advances In Respiratory Disease*. 2021;15:1753466620961696.
12. Vogelmeier CF, Criner GJ, Martinez FJ, Anzueto A, Barnes PJ, Bourbeau J, et al. Global Strategy for the Diagnosis, Management, and Prevention of Chronic Obstructive Lung Disease 2017 Report. GOLD Executive Summary. *American Journal of Respiratory and Critical Care Medicine*. 2017;195(5):557-82.
13. Chu J, Zhou C, Guo X, Sun J, Xue F, Zhang J, et al. Female Breast Cancer Mortality Clusters in Shandong Province, China: A Spatial Analysis. *Sci Rep*. 2017;7(1):105.
14. Turon H, Carey M, Boyes A, Hobden B, Dilworth S, Sanson-Fisher R. Agreement between a single-item measure of anxiety and depression and the Hospital Anxiety and Depression Scale: A cross-sectional study. *PloS One*. 2019;14(1):e0210111.
15. Andersson T, Alfredsson L, Källberg H, Zdravkovic S, Ahlbom A. Calculating measures of biological interaction. *European Journal of Epidemiology*. 2005;20(7):575-9.
16. Vandenberg JP, von Elm E, Altman DG, Gøtzsche PC, Mulrow CD, Pocock SJ, et al. Strengthening the Reporting of Observational Studies in Epidemiology (STROBE): explanation and elaboration. *Int J Surg*. 2014;12(12):1500-24.
17. Wang Z, Yang T, Fu H. Prevalence of diabetes and hypertension and their interaction effects on cardio-cerebrovascular diseases: a cross-sectional study. *BMC Public Health*. 2021;21(1):1224.
18. Galobardes B, Shaw M, Lawlor DA, Lynch JW, Davey Smith G. Indicators of socioeconomic position (part 1). *J Epidemiol Community Health*. 2006;60(1).
19. Paalanen L, Härkänen T, Kontto J, Tolonen H. Inequalities by education and marital status in the co-occurrence of cardiovascular risk factors in Finland persisted between 1997-2017. *Scientific Reports*. 2020;10(1):9123.
20. Fischer C, Jörres RA, Alter P, Trudzinski FC, Yildirim Ö, Bals R, et al. Basic Determinants of Disease Knowledge in COPD Patients: Results from COSYCONET. *Patient Prefer Adherence*. 2022;16:1759-70.
21. Dong H, Hao Y, Li D, Su Z, Li W, Shi B, et al. Risk Factors for Acute Exacerbation of Chronic Obstructive Pulmonary Disease in Industrial Regions of China: A Multicenter Cross-Sectional Study. *International Journal of Chronic Obstructive Pulmonary Disease*. 2020;15:2249-56.
22. Yang H, Wang H, Du L, Wang Y, Wang X, Zhang R. Disease knowledge and self-management behavior of COPD patients in China. *Medicine*. 2019;98(8):e14460.

23. Mao Y, Ning W, Zhang N, Xie T, Liu J, Lu Y, et al. The Therapeutic Relationship in China: A Systematic Review and Meta-Analysis. *Int J Environ Res Public Health*. 2021;18(7).
24. Breslau J, Marshall GN, Pincus HA, Brown RA. Are mental disorders more common in urban than rural areas of the United States? *J Psychiatr Res*. 2014;56:50-5.
25. Strosnider H, Kennedy C, Monti M, Yip F. Rural and Urban Differences in Air Quality, 2008-2012, and Community Drinking Water Quality, 2010-2015 - United States. *MMWR Surveill Summ*. 2017;66(13).
26. Shi Z, Wang Y, Zhao Q. Analysis of Spatiotemporal Changes of Ecological Environment Quality and Its Coupling Coordination with Urbanization in the Yangtze River Delta Urban Agglomeration, China. *Int J Environ Res Public Health*. 2023;20(2).
27. Yan P, Liu P, Lin R, Xiao K, Xie S, Wang K, et al. Effect of ambient air quality on exacerbation of COPD in patients and its potential mechanism. *International Journal of Chronic Obstructive Pulmonary Disease*. 2019;14:1517-26.
28. Liu Y, Lan B, Shirai J, Austin E, Yang C, Seto E. Exposures to Air Pollution and Noise from Multi-Modal Commuting in a Chinese City. *Int J Environ Res Public Health*. 2019;16(14).
29. Brown AL, van Kamp I. WHO Environmental Noise Guidelines for the European Region: A Systematic Review of Transport Noise Interventions and Their Impacts on Health. *Int J Environ Res Public Health*. 2017;14(8).
30. Smith MG, Cordoza M, Basner M. Environmental Noise and Effects on Sleep: An Update to the WHO Systematic Review and Meta-Analysis. *Environ Health Perspect*. 2022;130(7):76001.
31. Cyril S, Oldroyd JC, Renzaho A. Urbanisation, urbanicity, and health: a systematic review of the reliability and validity of urbanicity scales. *BMC Public Health*. 2013;13:513.
32. Wang R, Xue D, Liu Y, Chen H, Qiu Y. The relationship between urbanization and depression in China: the mediating role of neighborhood social capital. *International Journal For Equity In Health*. 2018;17(1):105.
33. Halme M, Rautava P, Sillanmäki L, Sumanen M, Suominen S, Vahtera J, et al. Educational level and the use of mental health services, psychotropic medication and psychotherapy among adults with a history of physician diagnosed mental disorders. *Int J Soc Psychiatry*. 2023;69(2):493-502.
34. Lim HK, Ghazali SM, Kee CC, Lim KK, Chan YY, Teh HC, et al. Epidemiology of smoking among Malaysian adult males: prevalence and associated factors. *BMC Public Health*. 2013;13:8.
35. Kong F, Xu L, Kong M, Li S, Zhou C, Li J, et al. The Relationship between Socioeconomic Status, Mental Health, and Need for Long-Term Services and Supports among the Chinese Elderly in Shandong Province-A Cross-Sectional Study. *Int J Environ Res Public Health*. 2019;16(4).
36. Fukuda Y, Nakamura K, Takano T. Municipal health expectancy in Japan: decreased healthy longevity of older people in socioeconomically disadvantaged areas. *BMC Public Health*. 2005;5:65.
37. Yang X, Li L, Wang J, Huang J, Lu S. Cardiovascular mortality associated with low and high temperatures: determinants of inter-region vulnerability in China. *Int J Environ Res Public Health*. 2015;12(6):5918-33.

Figures

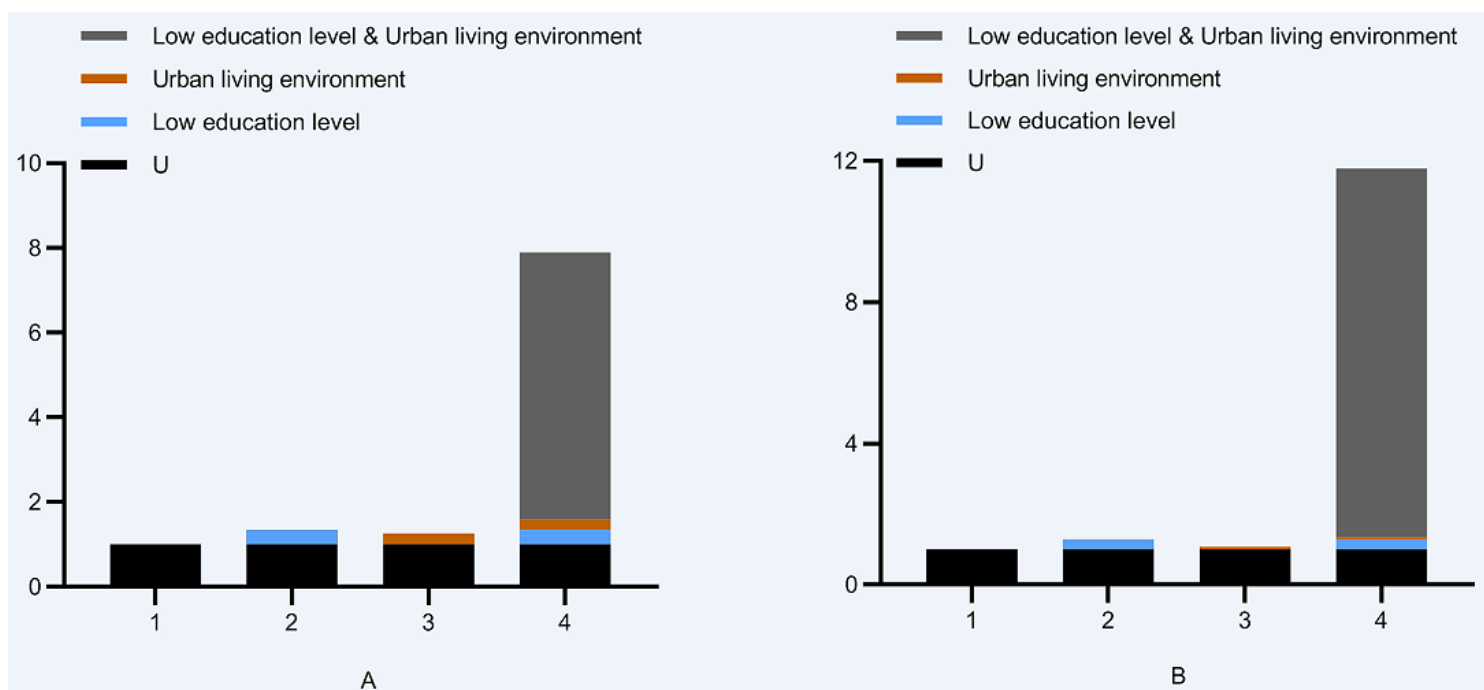


Figure 1

Relative risk with contributions from low education level, urban living environment, or a combination of both factors for anxiety (A) and depression (B). Multiple confounder adjustment included age, sex, smoking history, body mass index, COPD Assessment Test, percentage predicted forced expiratory volume in 1 second, oxygen therapy, modified Medical Research Council score, and Charlson Comorbidity Index.

Note: U means the odds ratio was equal to 1.