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ABOUT COVER

Peer Reviewer of *World Journal of Psychiatry*, Qing-Zhong Wang, PhD, Associate Professor, Institute of Chinese Materia Medica, Shanghai University of Traditional Chinese Medicine, Shanghai 201203, China.
wangqingzhong3@gmail.com

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Retrospective Study

Meteorological factors, ambient air pollution, and daily hospital admissions for depressive disorder in Harbin: A time-series study

Ting Hu, Zhao-Yuan Xu, Jian Wang, Yao Su, Bing-Bing Guo

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Abstract

BACKGROUND

The literature has discussed the relationship between environmental factors and depressive disorders; however, the results are inconsistent in different studies and regions, as are the interaction effects between environmental factors. We hypothesized that meteorological factors and ambient air pollution individually affect and interact to affect depressive disorder morbidity.

AIM

To investigate the effects of meteorological factors and air pollution on depressive disorders, including their lagged effects and interactions.

METHODS

The samples were obtained from a class 3 hospital in Harbin, China. Daily hospital admission data for depressive disorders from January 1, 2015 to December 31, 2022 were obtained. Meteorological and air pollution data were also collected during the same period. Generalized additive models with quasi-Poisson regression were used for time-series modeling to measure the non-linear and delayed effects of environmental factors. We further incorporated each pair of environ-

mental factors into a bivariate response surface model to examine the interaction effects on hospital admissions for depressive disorders.

RESULTS

Data for 2922 d were included in the study, with no missing values. The total number of depressive admissions was 83905. Medium to high correlations existed between environmental factors. Air temperature (AT) and wind speed (WS) significantly affected the number of admissions for depression. An extremely low temperature (-29.0 °C) at lag 0 caused a 53% [relative risk (RR)= 1.53, 95% confidence interval (CI): 1.23-1.89] increase in daily hospital admissions relative to the median temperature. Extremely low WSs (0.4 m/s) at lag 7 increased the number of admissions by 58% (RR = 1.58, 95%CI: 1.07-2.31). In contrast, atmospheric pressure and relative humidity had smaller effects. Among the six air pollutants considered in the time-series model, nitrogen dioxide (NO₂) was the only pollutant that showed significant effects over non-cumulative, cumulative, immediate, and lagged conditions. The cumulative effect of NO₂ at lag 7 was 0.47% (RR = 1.0047, 95%CI: 1.0024-1.0071). Interaction effects were found between AT and the five air pollutants, atmospheric temperature and the four air pollutants, WS and sulfur dioxide.

CONCLUSION

Meteorological factors and the air pollutant NO₂ affect daily hospital admissions for depressive disorders, and interactions exist between meteorological factors and ambient air pollution.

Key Words: Mental health; Depressive disorder; Hospital admissions; Meteorological factors; Air pollution; Time-series

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Core Tip: This retrospective study assessed the influence of environmental factors on depressive disorders. Daily hospital admissions for depressive disorders at a hospital in Harbin, China from 2015 to 2022 were obtained. Four meteorological factors and six air pollutants were considered predictors in the time-series models. Air temperature, atmospheric pressure, wind speed, relative humidity, and nitrogen dioxide have effects on daily hospital admissions for depressive disorders, and interactions exist between meteorological factors and ambient air pollution.

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INTRODUCTION

Depressive disorder (also known as depression) is a common mental disorder that involves depressed mood and loss of pleasure or interest in activities for long periods of time[1]. Approximately 280 million people (3.8%) worldwide experience depression[2]. Depressive disorder is a complex phenomenon that represents the interplay between social, psychological, and biological mechanisms[1]. The crucial biochemical mechanism underlying depression lies in the disruption of synaptic chemical transmission, which leads to a functional deficiency in the brain's monoaminergic transmitters including norepinephrine, serotonin (5-hydroxytryptamine), and/or dopamine[3]. These mechanisms have become targets of antidepressant action; for example, selective serotonin reuptake inhibitors such as fluoxetine are typical antidepressant medications[1] used for regulating serotonergic transmission.

In the last couple of years, accumulated evidence suggests that depressive disorder is also affected by environmental factors[4], including meteorological factors[5-8], ambient air pollution[9-13], light exposure[14-16], and access to green space[17,18]. An increase in temperature has been found to exacerbate psychiatric disease hospitalizations[19] in Greece, deteriorate mental well-being, and increase depressive language and suicide rates[20] in North America. Exposure to fine particles (PM_{2.5}), inhalable particles (PM₁₀), nitrogen dioxide (NO₂), sulfur dioxide (SO₂), and carbon monoxide (CO) was positively associated with depression, and the risk of depression increased with exposure to high levels of these pollutants[21]. Sunlight intensity may also be associated with depressive symptoms[22]. People reporting inadequate natural light in their dwellings were more likely to report depression[15]. This strongly suggests that environmental factors play a role in altering serotonin levels, leading to depression. Corresponding therapies, such as bright-light therapy, have been developed and used, and have evolved into effective treatments that act on serotonin and other monoaminergic pathways[16].

While the literature has discussed the relationship between environmental factors and depressive disorders, the interactions between different environmental factors have rarely been considered. Moreover, the results of different studies have been inconsistent. This study aimed to investigate the effects of environmental factors on depressive disorders. We obtained environmental data containing four meteorological factors and six air pollutants and modeled the

interactions between these environmental factors simultaneously to gather more precise evidence and obtain a regional supplement for this research field.

MATERIALS AND METHODS

Study design

To measure the precise effects of and the interaction between meteorological factors and air pollution on depressive disorders, we obtained as our sample daily hospital admission data for depressive disorders at a class 3 hospital in Harbin, China from January 1, 2015 to December 31, 2022. Data on the meteorological factors and air pollution for each day of admission were also collected to construct a predictive model.

Data collection

Data from 2922 d were obtained from the information management system developed and used by the hospital. Patient information included the visit time and diagnosis. Individual identifiers were excluded. Patients with permanent residence in Harbin were selected, and those with other mental disorders were excluded. Daily depressive disorder admission data equaled the sum of depressive outpatient and hospitalization data. The diagnosis of depressive disorder (coded as F32-F33 ICD-10 diagnoses) refers to GB/T 14396-2016", classification and Code of Diseases"[23].

Meteorological data for the same period from 2015 to 2022 for Harbin, China, were collected from the China meteorological data service centre[24], including daily mean air temperature (AT), daily mean atmospheric pressure (AP), wind speed (WS), and relative humidity (RH).

Ambient air pollution data included six international standard air pollutants that have been extensively investigated previously[12]: PM_{2.5}, PM₁₀, SO₂, CO, NO₂, and ozone (O₃, usually measured at maximum 8-h average concentrations). The pollutant data were collected from the China Air Quality Online Monitoring and Analysis platform[25]. For all ambient air pollutants, we averaged all valid monitoring measurements each day to obtain their daily mean concentrations, in line with previous studies[12,26].

To conduct a time-series analysis, information on time-related variables was collected for modeling, such as the day of admission and public holidays. A sequence of calendar times was generated to indicate the time trends.

Statistical analysis

All data were inspected and preprocessed using Microsoft excel and IBM SPSS 25.0. No missing values or outliers were identified. Descriptive analysis, correlation calculations, time-series modeling, and plotting were conducted using R version 4.2.3. The mean value; SD; and minimum, maximum, median, and quartile (P₂₅ and P₇₅) were used to describe the data. The correlation coefficients between variables were calculated using Spearman's correlation analysis with the R package psych. We conducted a time-series analysis using the R packages dlnm[27], mgcv[28], rsm[29], and splines, modeling the effects of and interactions between various environmental factors. The R package ggplot2[30] was used to visualize the results. All statistical tests were two-tailed probability tests, $\alpha = 0.05$. Statistical results with $P < 0.05$, such as associations and differences, were considered statistically significant.

We used generalized additive models with quasi-Poisson regression to estimate the meteorological effects of each factor. The covariates in the models included a natural cubic spline of calendar time with 14 degrees of freedom, which modeled time trends in hospital admissions for depressive disorders, and two indicator variables for the day of the week and public holidays, accounting for potential variations in hospital admissions over weekdays, weekends, and holidays. Natural cubic splines of the daily AT, AP, WS, and RH with three degrees of freedom were included as predictors. To assess the potential lagged effects of meteorological factors, we included lags of up to seven days in the model for non-cumulative effects, similar to the methodology of a previously published study[31]. The choice of calendar time df was determined by calculating the minimum sum of the absolute values of the partial autocorrelation function based on the residuals of the underlying model, consistent with the residual independence principle[31]. Choices of meteorological factors dfs have been recommended by papers on time-series analyses[32] and have been adopted by most studies. The time-series analysis model for a single meteorological factor is as follows:

$$Y_t \sim \text{quasiPoisson}(\mu_t)$$

$$\log E([Y_t]) = \alpha + ns_t(\text{calendar time}, df = 14) + DOW_t + \text{holiday}_t + \beta_{it} \\ \times ns_{it}(\text{meteorological factor}, df = 3)$$

Where $E(Y_t)$ denotes the expected number of depressive disorder admissions on day t ; α is the intercept; ns represents the natural cubic spline function; and β_{it} is the exposure coefficients of meteorological factors and their lagged effects. We included the immediate and lagged effects of each air pollutant in the model and used meteorological factors as covariates. The time-series analysis model for a single air pollutant is as follows:

$$Y_t \sim \text{quasiPoisson}(\mu_t)$$

$$\log E([Y_t]) = \alpha + ns_t(\text{calendar time}, df = 14) + DOW_t + \text{holiday}_t + \beta_{it} \\ \times ns_{it}(\text{meteorological factor}, df = 3) + \beta_{jt} \times (\text{air pollutant})_{jt}$$

Where (air pollutant) represents lags of 0 to 7 for cumulative and non-cumulative exposures. Finally, we incorporated each meteorological factor and air pollutant into a bivariate response surface model to inspect the interaction effects of each pair of meteorological factors and air pollutants on hospital admissions for depressive disorders. The model for interaction effects is as follows:

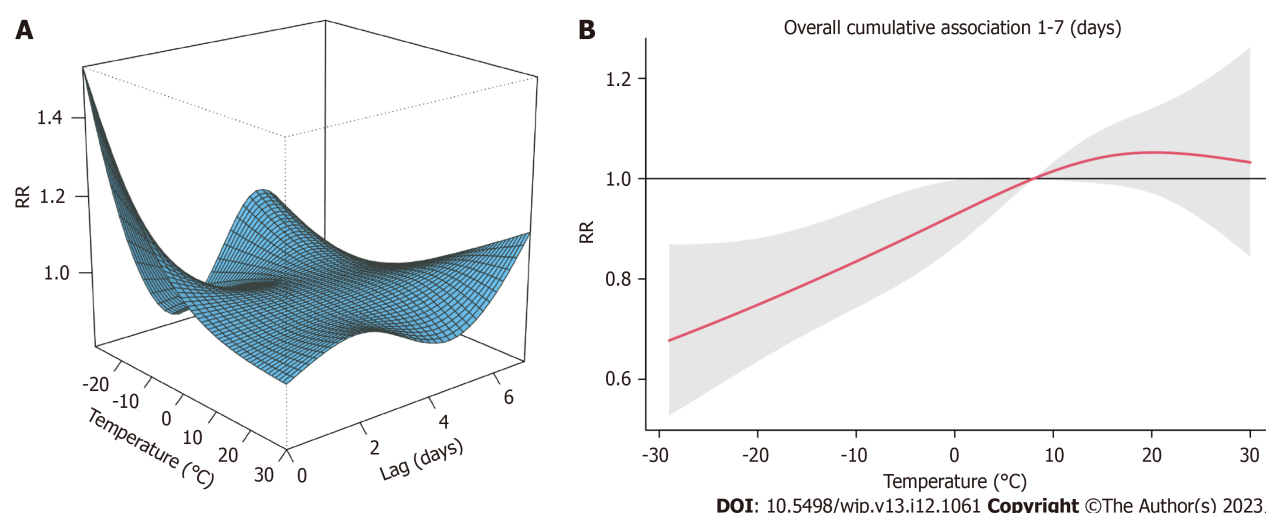


Figure 1 Effects of air temperature on hospital admissions for depressive disorder. A: 3D plot of relative risk (RR) along air temperature and lags, with reference to the median temperature of 7.8 °C; B: Overall RR of lags 1-7 by air temperature, with reference at 7.8 °C. RR: Relative risk.

$$Y_t \sim \text{quasiPoisson}(\mu_t)$$

$$\log E([Y_t]) = \alpha + ns_t(\text{calendar time, } df = 14) + DOW_t + \text{holiday}_t + t_e(\text{meteorological factor}_t, \text{air pollutant}_{t-i})$$

Where t_e is the interaction effect of meteorological factors and air pollutants (on the day with the strongest lag effect) on daily hospital admissions for depressive disorders. Results are expressed as relative percentage change (%) and estimated 95% confidence intervals (CIs) at hospital admissions for depressive disorders of certain meteorological factor levels or per unit increment of air pollutants.

RESULTS

Descriptive statistics and correlation

The general features of all the variables are summarized in Table 1. The total number of hospital admissions for depressive disorders from January 1, 2015 to December 31, 2022 included in the study was 83905, with a daily average admissions of 28.7. During the studied period, the daily average AT in Harbin was 5.4 °C, and the daily average AP was 749.3 mmHg. The average WS was 2.9 m/s. The mean RH was 66.2%. The climate in Harbin is cold but mild, which means that it is not excessively windy or dry most times of the year, and the AP remains slightly lower than the standard. The daily average concentrations of PM_{2.5}, PM₁₀, SO₂, CO, NO₂, and O₃-8h (the maximum 8-h average concentration of ozone) were 47.2 µg/m³, 71.3 µg/m³, 22.0 µg/m³, 0.9 mg/m³, 36.7 µg/m³, and 75.8 µg/m³, respectively. There is relatively high variability in air pollution, although the concentration levels are usually low according to the quartiles.

The correlation coefficients between the meteorological factors and air pollutants are shown in Table 2. All Spearman's correlation coefficients were statistically significant, except for the correlation between RH and CO. There was a moderate correlation among AT, AP, and air pollutants. Relatively high correlations among the air pollutants (except for O₃) were found.

Time-series analysis modeling

In the single meteorological factor model for AT, AP, WS, and RH, we reported relative risks (RRs) with 95%CIs of the depressive disorder admissions for each level of the meteorological factor. The overall relationships between each meteorological factor and hospital admissions for depressive disorders are summarized in Figures 1-4. As shown in Figure 1A, there was a sharp effect of extremely low temperature, although, in Figure 1B, the overall low temperature had no positive effect on RR. Figure 2 shows the mild effect of AP on RR, which is close to zero. Figure 3 shows a larger effect of low WS and a smaller effect of high WS. As shown in Figure 4, both moderate and high RH appeared to have relatively small effects.

To provide more certainty and precision in the estimates, we further investigated the effect of each meteorological factor by calculating and plotting the RRs by these factors at lags 0-7, as shown in Figures 5-8. The results for AT showed significant increases at lags 0-2 and 5-6. Figure 5 shows that an extremely low temperature (-29.0 °C) at lag 0 increased daily hospital admissions for depressive disorders by 1.53 times (95%CI: 1.23-1.89) relative to median temperature (7.8 °C), which was the largest effect of AT. As shown in Figure 6, the effect of AP was significant only at lags 5 and 6. A moderately high AP (762 mmHg) at lag 5 increased the number of admissions by 1.13 times (95%CI: 1.00-1.27) relative to the median AP (748.9 mmHg). Figure 7 shows that WS had a significant effect at lags 2-7. The largest effect was at lag 7, where an extremely low WS (0.4 m/s) increased the number of admissions by 1.58 times (95%CI: 1.07-2.31) relative to the median WS (2.6 m/s). Relatively high RHs at lags 4-7 had statistically significant effects, and the largest effect of RH

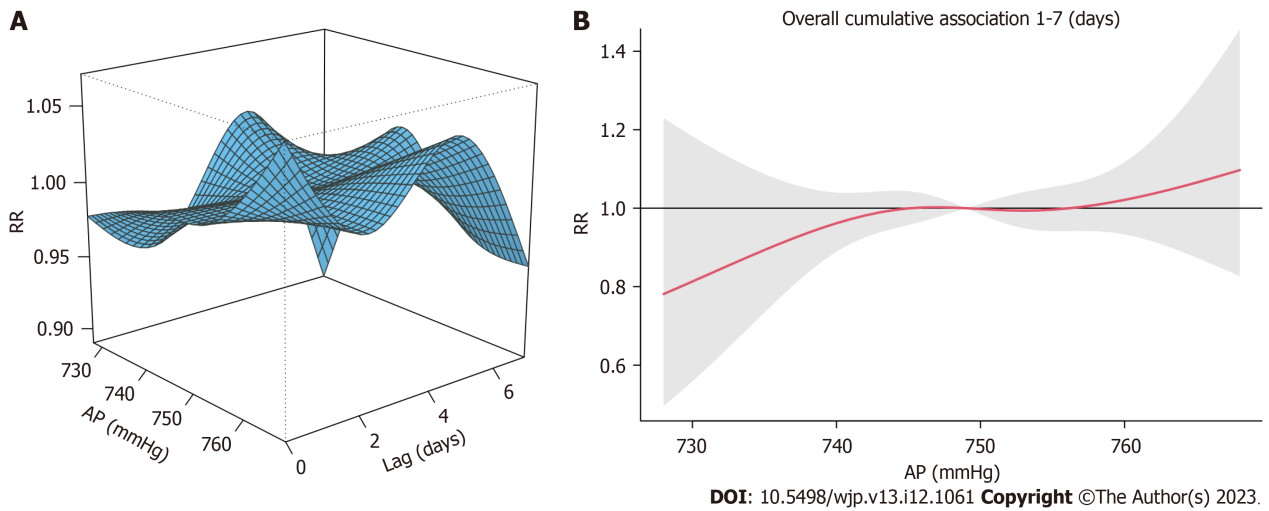


Figure 2 Effects of atmospheric pressure on hospital admissions for depressive disorder. A: 3D plot of relative risk (RR) along atmospheric pressure (AP) and lags, with reference to the median AP of 748.9 mmHg; B: Overall RR of lags 1-7 by atmospheric pressure, with reference at 748.9 mmHg. RR: Relative risk; AP: Atmospheric pressure.

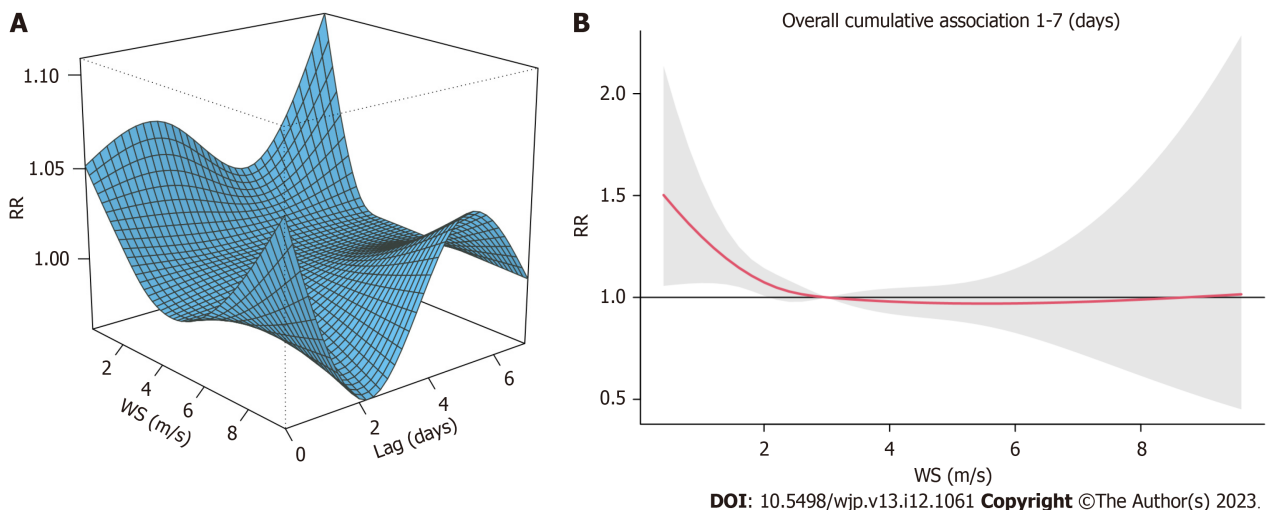


Figure 3 Effects of wind speed on hospital admissions for depressive disorder. A: 3D plot of relative risk (RR) along wind speed (WS) and lags, with reference to the median WS of 2.6 m/s; B: Overall RR of lags 1-7 by wind speed, with reference at 2.6 m/s. RR: Relative risk; WS: Wind speed.

resulted from a moderately high RH (86%) at lag 7, which increased the number of admissions by 1.09 times (95%CI: 1.01-1.17) relative to the median RH (67.9%), as shown in [Figure 8](#).

Next, the concentrations on the same day (lag 0) and 1-7 d ago (lag 1-7) of the six air pollutants were included in the time-series model. The results are shown in [Figure 9](#). Among these air pollution variables, only NO₂ showed significant effects, and the effects of concentration increases in NO₂ at lags 0-7 were all statistically significant, despite their weakness. The cumulative exposure to air pollution was also modeled and plotted, as shown in [Figure 10](#). Consistent with the non-cumulative results, only NO₂ exhibited a significant cumulative effect. As NO₂ accumulated, the exposure effect on daily hospital admissions for depressive disorders increased slightly and smoothly, reaching its highest value at lag 7, where RR = 1.0047 (95%CI: 1.0024-1.0071).

A bivariate response surface model was generated to analyze the interactions between environmental factors. AT significantly interacted with all pollutants included in this study, except for NO₂. As shown in [Figure 11](#), extremely low ATs combined with high concentrations of PM_{2.5} and PM₁₀ increased the number of admissions for depressive disorders. Combination of low temperature, low SO₂, and of high temperature and high SO₂ affected hospital admissions. Extremely low temperatures at any CO concentration increased the admission rate. When hot, a low CO concentration had an increasing effect, whereas a high CO concentration did not. Low temperatures and high O₃ concentrations had decreasing effects. [Figure 12](#) shows the interactions between the AP and air pollutants. The interactions between the AP and PM_{2.5}, CO, NO₂, and O₃ 8h were statistically significant. Low APs combined with low CO, low NO₂, or high O₃ had increasing effects, whereas high APs combined with high PM_{2.5}, high CO, or low O₃ increased hospital admissions for depressive disorders. The interaction patterns of AP and PM_{2.5}/CO/NO₂ were similar; however, the pattern between AP and O₃ was the opposite. The interactions between WS and air pollutants are shown in [Figure 13](#). SO₂ had the only significant

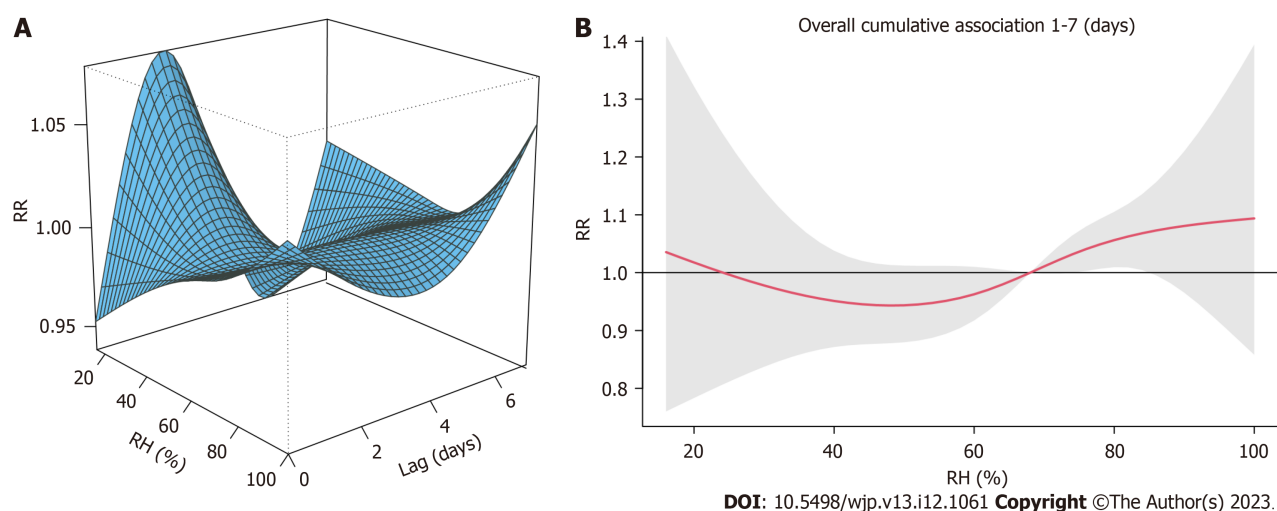


Figure 4 Effects of relative humidity on hospital admissions for depressive disorder. A: 3D plot of relative risk (RR) along relative humidity (RH) and lags, with reference to the median RH of 67.9%; B: Overall RR of lags 1-7 by RH, with reference at 67.9%. RR: Relative risk; RH: Relative humidity.

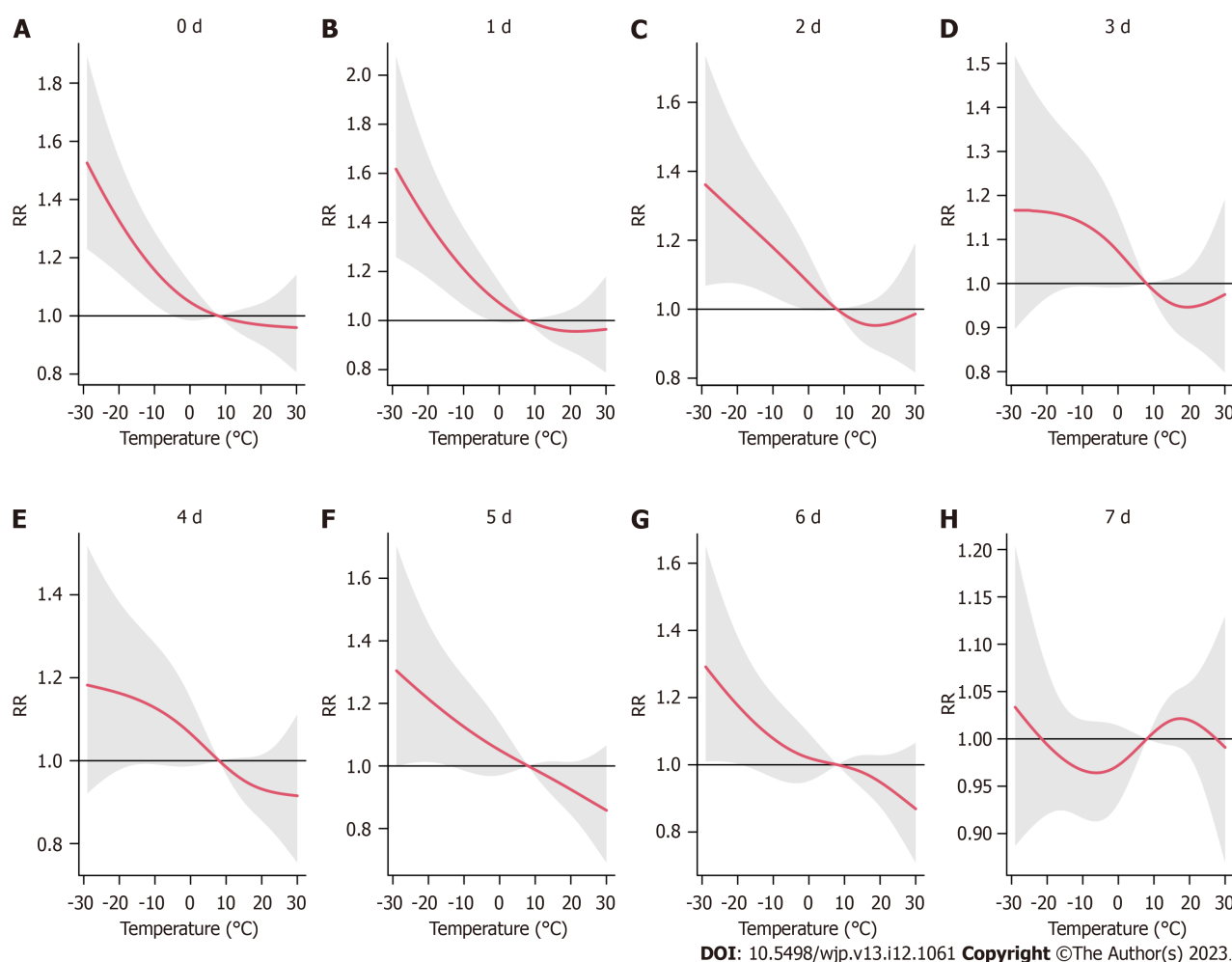
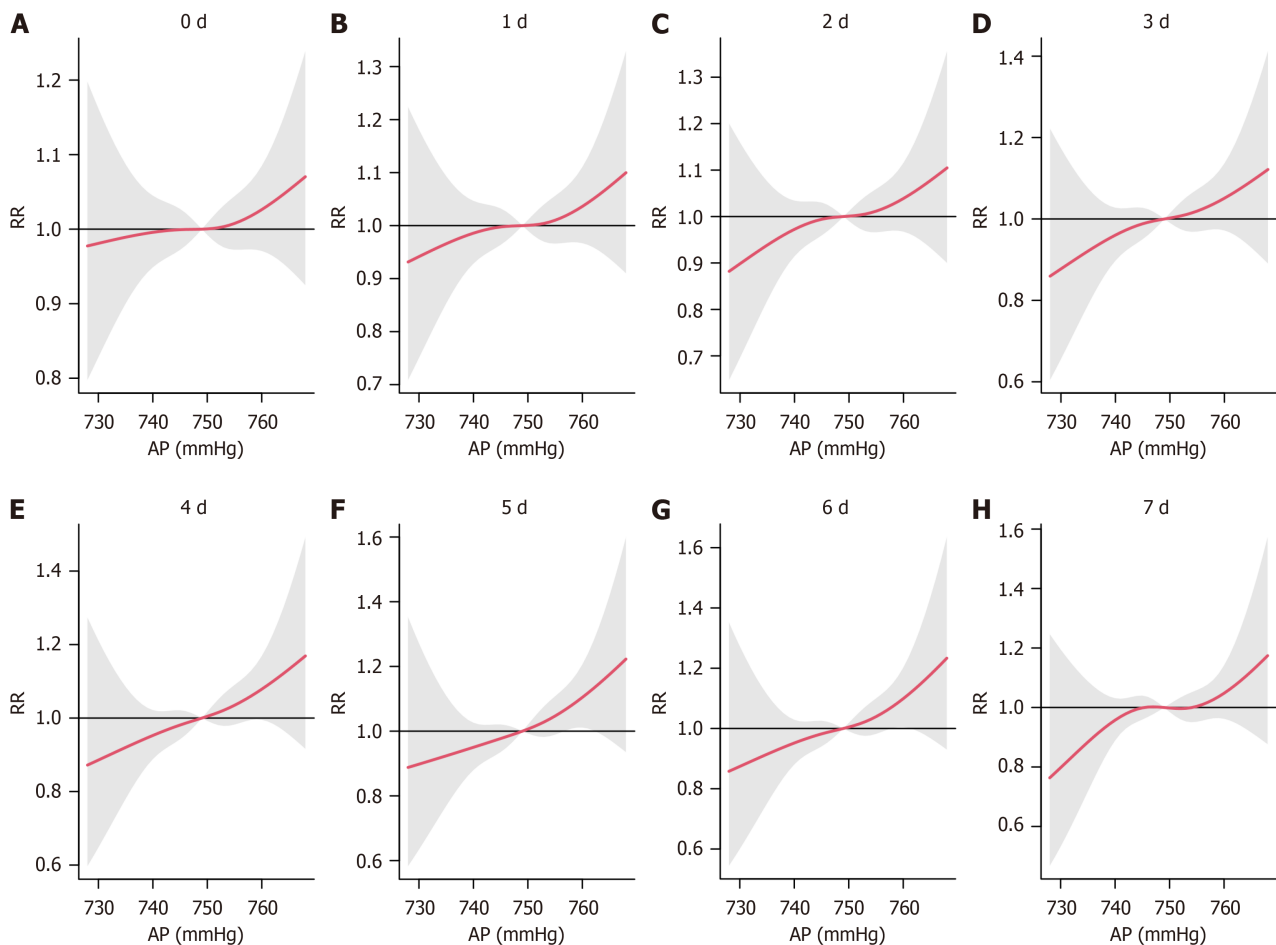


Figure 5 Plots of relative risk by air temperature at lags 0-7. A: Relative risk (RR) by air temperature (AT) at lag 0; B: RR by AT at lag 1; C: RR by AT at lag 2; D: RR by AT at lag 3; E: RR by AT at lag 4; F: RR by AT at lag 5; G: RR by AT at lag 6; H: RR by AT at lag 7. Reference at 7.8 °C. RR: Relative risk.

Table 1 Descriptive statistics of variables (sample size = 2922 d)

Variable (unit)	mean \pm SD	Percentile				
		Min	P ₂₅	Median	P ₇₅	Max
Admissions for depressive disorder (n)	28.7 \pm 21.2	0	9	28	42	130
AT ($^{\circ}$ C)	5.4 \pm 15.1	-29.2	-8.4	7.8	19.0	30.6
AP (mmHg)	749.3 \pm 7.0	727.9	743.7	748.9	754.8	768.0
WS (m/s)	2.9 \pm 1.2	0.4	2.0	2.6	3.5	9.8
RH (%)	66.2 \pm 15.6	14.9	56.1	67.9	77.8	100.0
PM _{2.5} (μ g/m ³)	47.2 \pm 54.6	0.0	16.0	29.0	58.0	906.0
PM ₁₀ (μ g/m ³)	71.3 \pm 59.3	0.0	35.0	53.0	87.0	644.0
SO ₂ (μ g/m ³)	22.0 \pm 23.6	3.0	9.0	13.0	26.0	222.0
CO (mg/m ³)	0.9 \pm 0.4	0.3	0.6	0.8	1.0	4.3
NO ₂ (μ g/m ³)	36.7 \pm 17.8	8.0	24.0	33.0	44.0	145.0
O _{3_8h} (μ g/m ³)	75.8 \pm 34.1	0.0	50.0	71.0	96.0	253.0

AT: Air temperature; AP: Atmospheric pressure; WS: Wind speed; RH: Relative humidity; PM_{2.5}: Fine particles; PM₁₀: Inhalable particles; SO₂: Sulfur dioxide; CO: Carbon monoxide; NO₂: Nitrogen dioxide; O_{3_8h}: Maximum 8-h average concentration of ozone.



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Figure 6 Plots of relative risk by atmospheric pressure at lags 0-7. A: Relative risk (RR) by atmospheric pressure (AP) at lag 0; B: RR by AP at lag 1; C: RR by AP at lag 2; D: RR by AP at lag 3; E: RR by AP at lag 4; F: RR by AP at lag 5; G: RR by AP at lag 6; H: RR by AP at lag 7. Reference at 748.9 mmHg. RR: Relative risk. AP: Atmospheric pressure.

Table 2 Spearman correlation coefficients between environmental factors

	AP	WS	RH	PM _{2.5}	PM ₁₀	SO ₂	CO	NO ₂	O ₃ _8h
AT	-0.75 ^c	0.07 ^c	0.18 ^c	-0.61 ^c	-0.46 ^c	-0.78 ^c	-0.49 ^c	-0.42 ^c	0.60 ^c
AP		-0.27 ^c	-0.17 ^c	0.50 ^c	0.40 ^c	0.62 ^c	0.41 ^c	0.44 ^c	-0.48 ^c
WS			-0.35 ^c	-0.16 ^c	-0.13 ^c	-0.11 ^c	-0.26 ^c	-0.41 ^c	0.17 ^c
RH				-0.17 ^c	-0.25 ^c	-0.19 ^c	-0.02	-0.06 ^c	-0.21 ^c
PM _{2.5}					0.92 ^c	0.77 ^c	0.78 ^c	0.76 ^c	-0.19 ^c
PM ₁₀						0.69 ^c	0.71 ^c	0.75 ^c	-0.09 ^c
SO ₂							0.67 ^c	0.64 ^c	-0.35 ^c
CO								0.76 ^c	-0.21 ^c
NO ₂									-0.24 ^c

^c*P* < 0.001.
AT: Air temperature; AP: Atmospheric pressure; WS: Wind speed; RH: Relative humidity; PM_{2.5}: Fine particles; PM₁₀: Inhalable particles; SO₂: Sulfur dioxide; CO: Carbon monoxide; NO₂: Nitrogen dioxide; O₃_8h: Maximum 8-h average concentration of ozone.

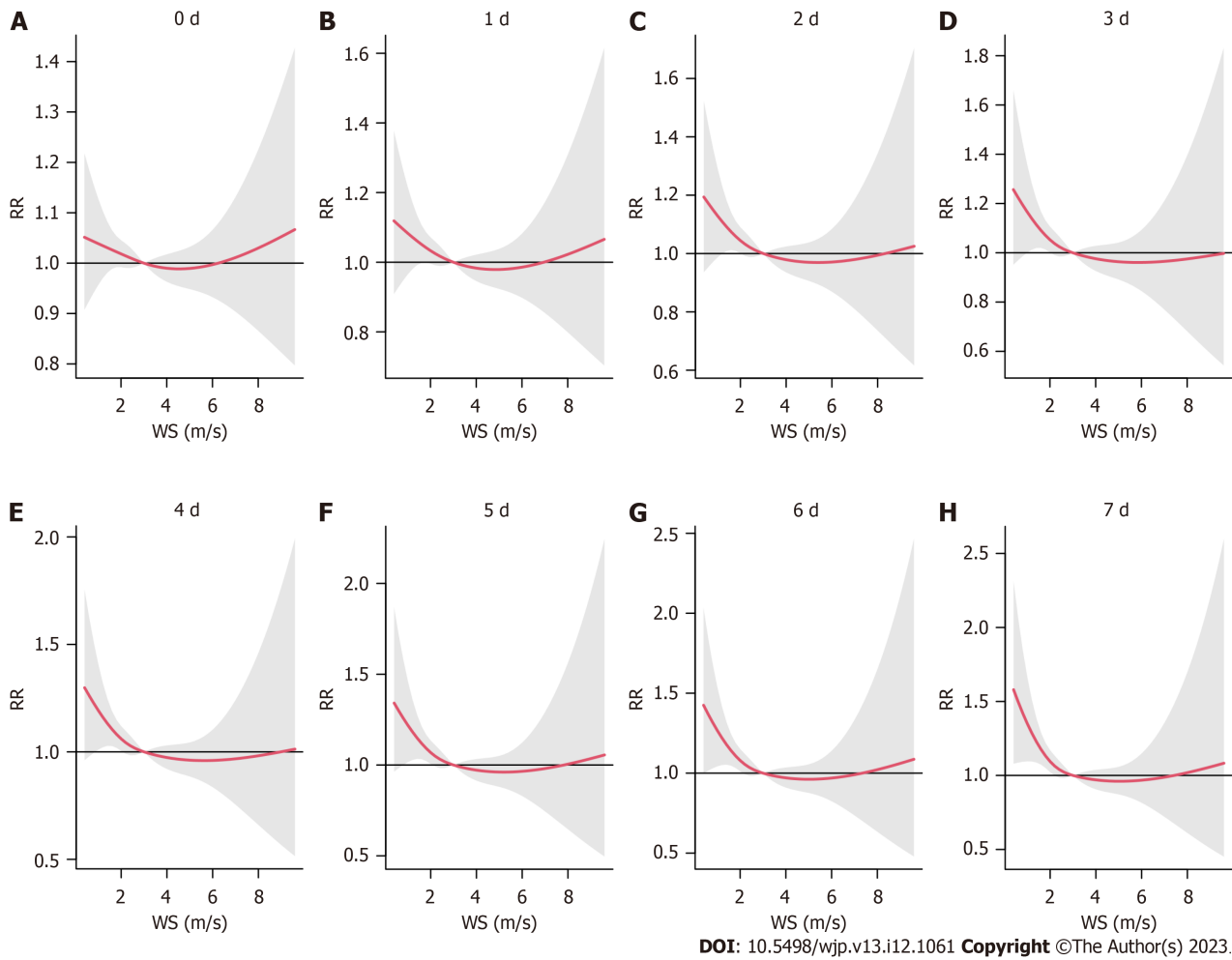


Figure 7 Plots of relative risk by wind speed at lags 0-7. A: Relative risk (RR) by wind speed (WS) at lag 0; B: RR by WS at lag 1; C: RR by WS at lag 2; D: RR by WS at lag 3; E: RR by WS at lag 4; F: RR by WS at lag 5; G: RR by WS at lag 6; H: RR by WS at lag 7. Reference at 2.6 m/s. RR: Relative risk; WS: Wind speed.

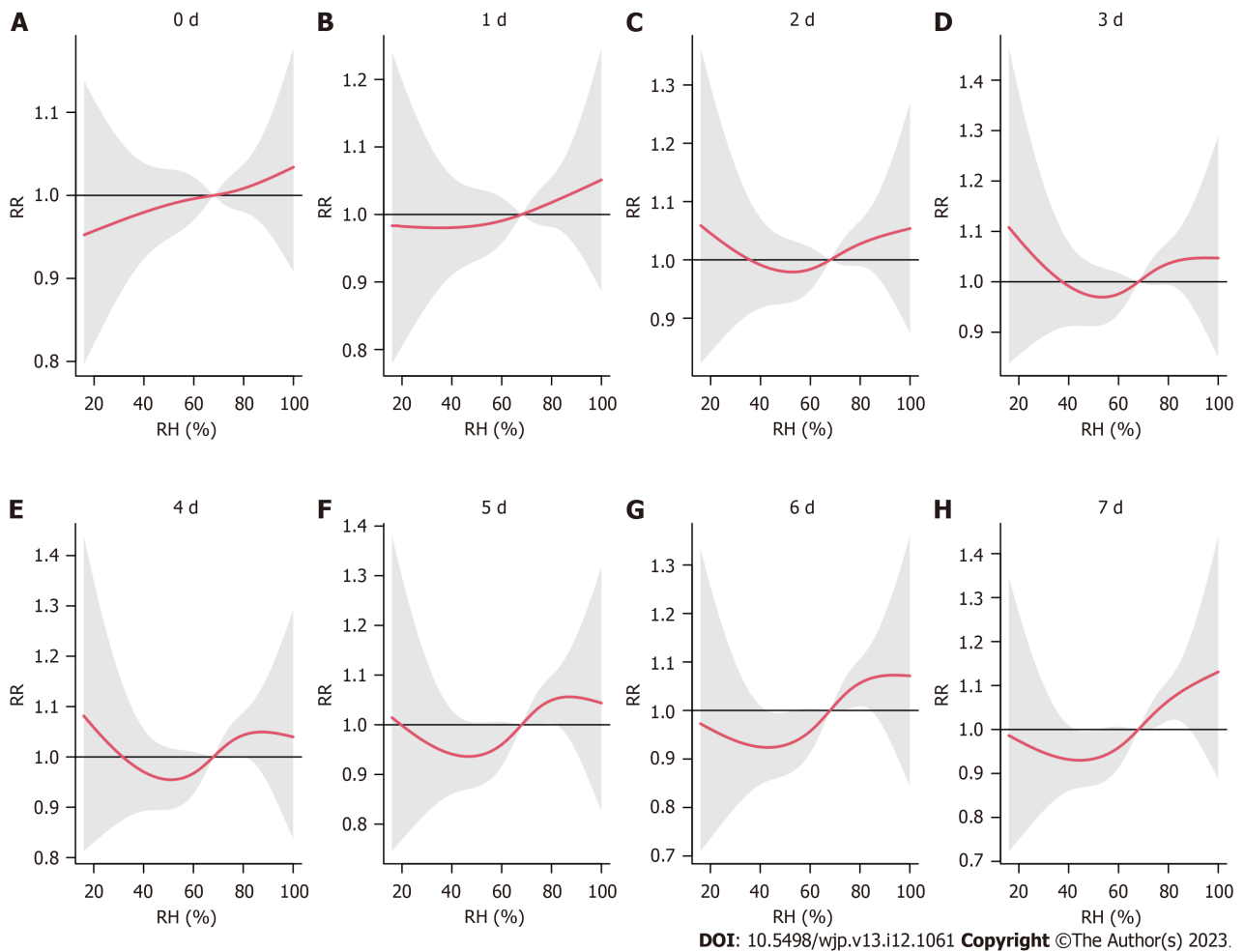


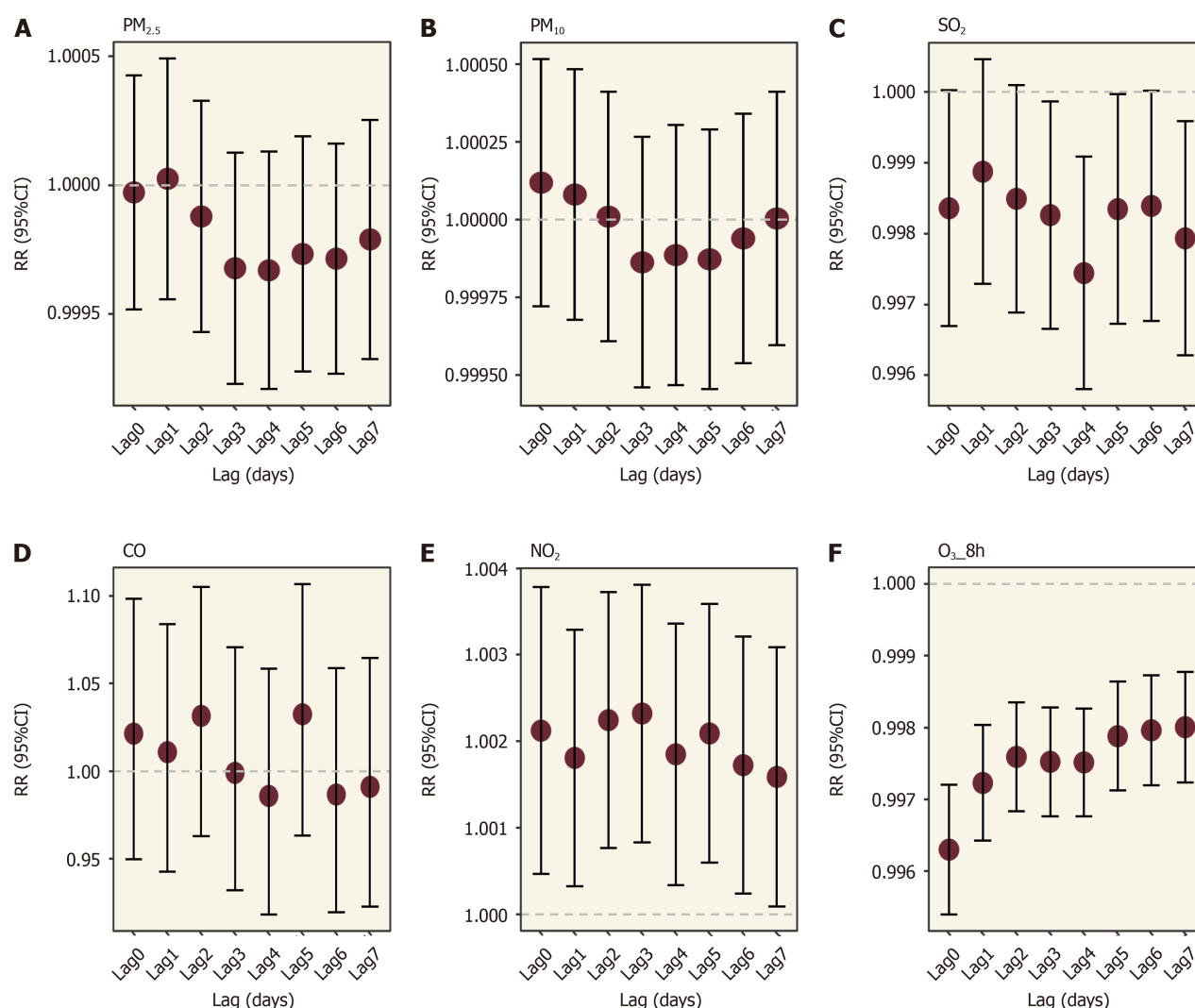
Figure 8 Plots of relative risk by relative humidity at lags 0-7. A: Relative risk (RR) by relative humidity (RH) at lag 0; B: RR by RH at lag 1; C: RR by RH at lag 2; D: RR by RH at lag 3; E: RR by RH at lag 4; F: RR by RH at lag 5; G: RR by RH at lag 6; H: RR by RH at lag 7. Reference at 67.9%. RR: Relative risk; RH: Relative humidity.

interaction effect with WS, indicating that most single effects of air pollutants were consistent at different levels of WS. Generally, when high WS and high concentrations of pollutants coexisted, hospital admissions for depressive disorders did not increase. Finally, the interactions between RH and air pollutants were analyzed, and the results are plotted in Figure 14. No significant interaction effects were found between any pair, indicating that the effects of air pollution were relatively stable across different levels of RH.

DISCUSSION

Based on daily hospital admission data for depressive disorders from 2015 to 2022 in a hospital in Harbin, China, we obtained environmental data for the same period, including meteorological and ambient air pollution data, and conducted a time-series analysis to investigate the short-term exposure effects of these environmental factors on depressive disorders. Our results showed that meteorological factors, including AT and WS, could sharply influence daily hospital admissions, while AP, RH, and air pollutant NO₂ (immediate, lagged, or cumulatively lagged) had significant but moderate or small effects. We modeled the relationship between meteorological factors and air pollution and found significant interaction effects between AT and PM_{2.5}/PM₁₀/SO₂/CO/O₃-8 h, AP and PM_{2.5}/CO/NO₂/O₃-8 h, and WS and SO₂.

Specifically, we found that the risk of depressive disorder admission increases by 53% when an extremely low temperature (-29 °C, for Harbin) occurs. This finding was consistent with the results of a previous study conducted in Europe[8], but contrary to most similar studies[5,14,33-35], where increased or high AT was associated with increased depressive symptoms or higher hospital admissions. This inconsistency may have resulted from regional differences in the areas of interest selected by each study. Harbin City (45°25'-45°30'N, 126°20'-126°25'E) is located in Northeast China and has a typical continental monsoon climate. The highest temperature in the study period was 30.6 °C, indicating that it is never excessively hot in Harbin, unlike in Hong Kong[5] or Shanghai[33]. From reference[6], we know that both low and high temperatures might be important drivers of morbidity in mental disorders and that lower temperatures may have a more general and widespread effect on cause-specific morbidity. The result in Europe[8] even showed a protective



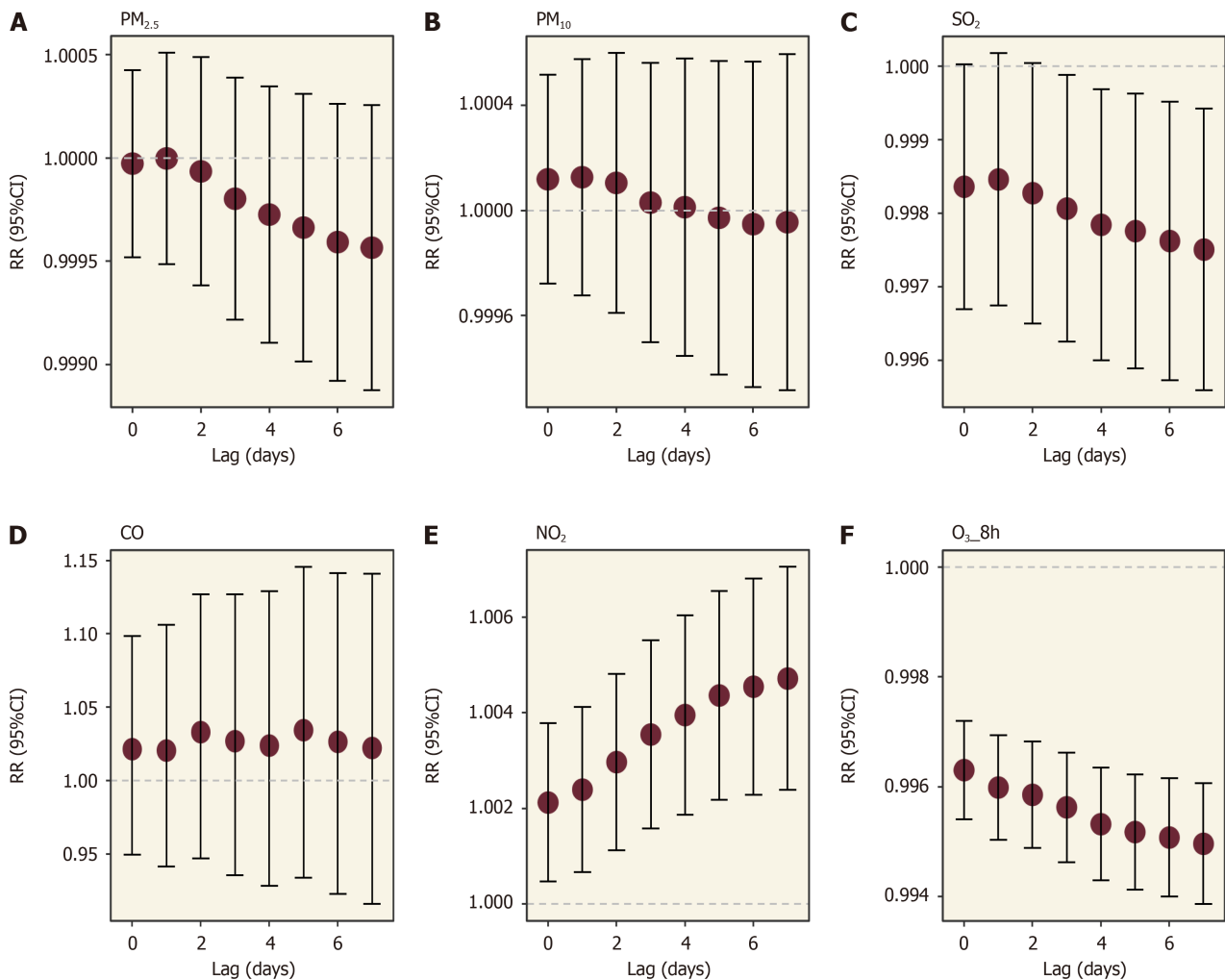
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Figure 9 Plots of relative risk by air pollutant concentrations at lags 0-7. A: Relative risk (RR) by fine particles concentrations at lags 0-7; B: RR by inhalable particles concentrations at lags 0-7; C: RR by sulfur dioxide concentrations at lags 0-7; D: RR by carbon monoxide concentrations at lags 0-7; E: RR by nitrogen dioxide concentrations at lags 0-7; F: RR by maximum 8-h average concentration of ozone concentrations at lags 0-7. PM_{2.5}: Fine particles; PM₁₀: Inhalable particles; SO₂: Sulfur dioxide; CO: Carbon monoxide; NO₂: Nitrogen dioxide; O_{3-8h}: Maximum 8-h average concentration of ozone; RR: Relative risk.

impact of a higher AT (> 14.2 °C), which may serve as a reference for explaining the zero effects of a higher temperature in Harbin. A famous nickname for Harbin City is “Ice City,” indicating its freezing coldness in winter. Studies have shown that exercise interventions[15] and frequent visits to outdoor green spaces[17] effectively improve depression symptoms. Therefore, extremely low temperatures can negatively affect depressive symptoms by limiting outdoor exercises and visits to green spaces. Furthermore, low temperatures usually co-exist with insufficient sunlight in winter, and reduced sunlight exposure, both in intensity and duration, is related to an increased risk of depression, according to previous studies[22,36]. Despite this inconsistency with most similar studies, this study confirmed the importance of AT in depressive admissions.

In addition, we found a 58% increase in depressive disorder admissions when the WS approached zero. We propose two possible mechanisms underlying the effect of low WS. First, as coal-fired heating in the cold period always causes severe air pollution in Harbin[37,38], low WS would worsen the air pollution. Air pollution-induced inflammation of the respiratory airways is associated with depression in vulnerable individuals[39]. Severe air pollution may also decrease outdoor activities and sunlight exposure, thereby increasing the risk of depressive disorder[40]. Hence, low WS during cold winters may be associated with depressive symptoms caused by severe air pollution. Second, a previous study showed that during warmer periods, warm and dry winds have an increasing effect on the concentration of positive ions in the atmosphere, which increases blood and brain serotonin levels[8]. Low WS may not provide this improving effect on depressive symptoms.

We also found relatively small effects of AP, RH, and the air pollutant NO₂. These results are consistent with previous studies, where high RH levels seem to be pertinent to suicide risk[41] and nitrogen oxides increase the risk of mental health problems, probably *via* mechanisms of inflammation and neuronal injury[35].



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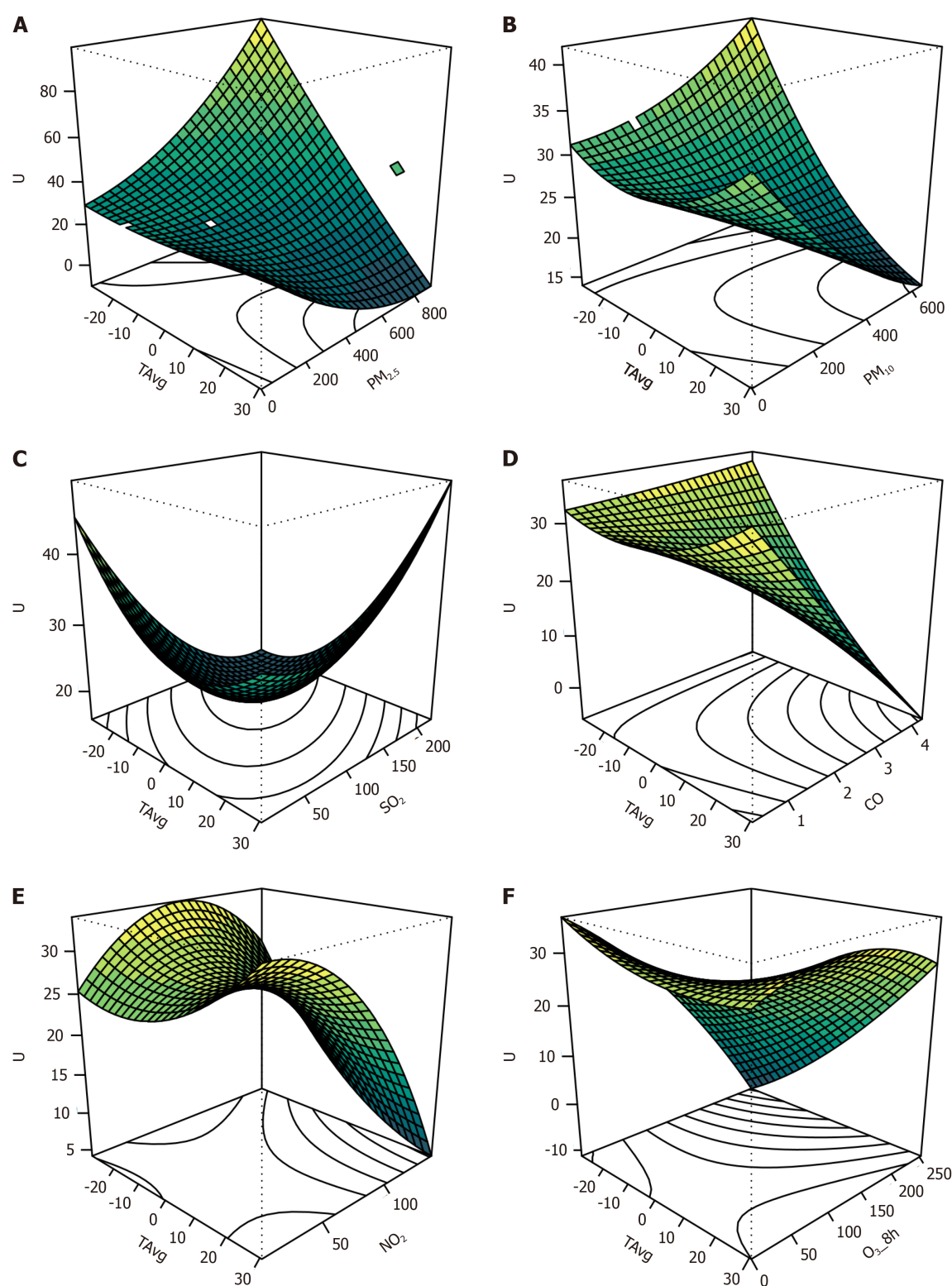
Figure 10 Plots of relative risk by cumulative air pollutant concentrations for lags 0-7. A: Relative risk (RR) by cumulative fine particles concentrations at lags 0-7; B: RR by cumulative inhalable particles concentrations at lags 0-7; C: RR by cumulative sulfur dioxide concentrations at lags 0-7; D: RR by cumulative carbon monoxide concentrations at lags 0-7; E: RR by cumulative nitrogen dioxide concentrations at lags 0-7; F: RR by cumulative maximum 8-h average concentration of ozone concentrations at lags 0-7. PM_{2.5}: Fine particles; PM₁₀: Inhalable particles; SO₂: Sulfur dioxide; CO: Carbon monoxide; NO₂: Nitrogen dioxide; O_{3_8h}: Maximum 8-h average concentration of ozone; RR: Relative risk.

The distinctive results of this study were the interactions between AT and PM_{2.5}/PM₁₀/SO₂/CO/O_{3_8h}, AP and PM_{2.5}/CO/NO₂/O_{3_8h}, and WS and SO₂. After carefully inspecting the patterns of these interactions, we assumed that most interactions corresponded to the relationship between the increased risk of depressive disorder and severe air pollution caused by coal-fired heating. In most significant interactions, typical meteorological factor levels in winter (including low AT and high AP) combined with high levels of air pollutants were associated with depressive symptoms. The interaction between WS and SO₂ was the same, as both high WS plus low SO₂ and low WS plus high SO₂ seemed unfavorable for outdoor activities and sunlight exposure. The increasing effect of low WS plus low SO₂ may appear during warmer periods, as low WS does not improve serotonin levels. The cases of O₃ were different from those of the others. We assumed that the reason for this inconsistency between O₃ and other air pollutants was seasonal differences. The O₃ concentration is typically high in the summer, whereas the concentrations of other pollutants are high in the winter.

The strength of our study lies in the precise effects we modeled and measured, especially the interaction between meteorological factors and ambient air pollution. Although our results indicate the effects of environmental factors on hospital admissions for depressive disorders, they are inconsistent with most similar studies and need to be replicated in other methodologically superior studies. The results cannot be generalized to the entire population of China because the sample came from only one northeastern city and urban population. Patient information was insufficient, which limited further analyses.

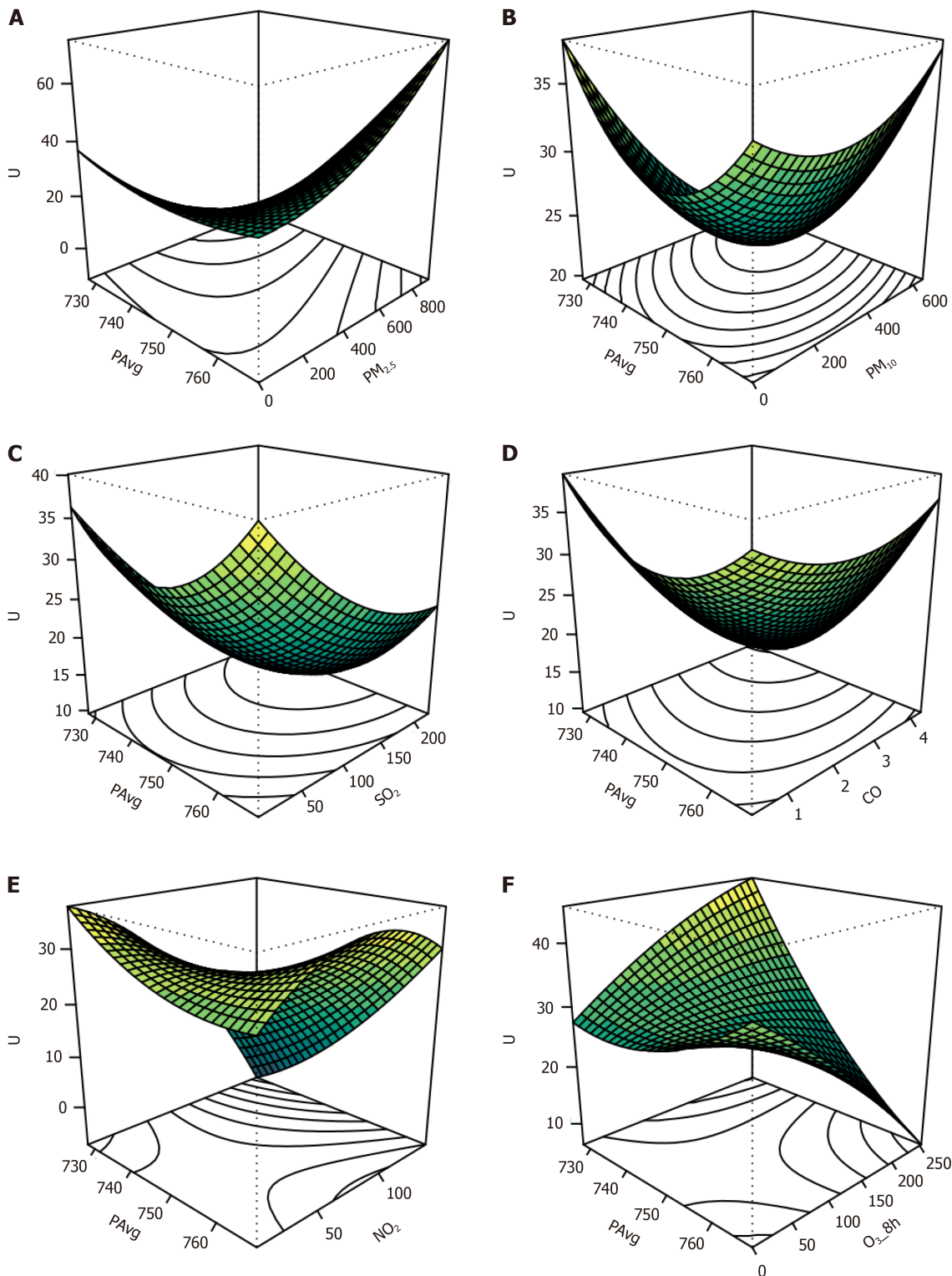
CONCLUSION

Our study suggests that meteorological factors and the air pollutant NO₂ affect daily hospital admissions for depressive disorders and that interaction effects exist between meteorological factors and ambient air pollution. Clinicians and



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Figure 11 3D perspective plots of the interaction effects between air temperature and air pollutants on daily hospital admissions for depressive disorder. A: Interaction between air temperature (AT) and fine particles; B: Interaction between AT and inhalable particles; C: Interaction between AT and sulfur dioxide; D: Interaction between AT and carbon monoxide; E: Interaction between AT and nitrogen dioxide; F: Interaction between AT and maximum 8-h average concentration of ozone. AT: Air temperature; PM_{2.5}: Fine particles; PM₁₀: Inhalable particles; SO₂: Sulfur dioxide; CO: Carbon monoxide; NO₂: Nitrogen dioxide; O₃-8h: Maximum 8-h average concentration of ozone.



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Figure 12 3D perspective plots of the interaction effects between atmospheric pressure and air pollutants on daily hospital admissions for depressive disorder. A: Interaction between atmospheric pressure (AP) and fine particles; B: Interaction between AP and inhalable particles; C: Interaction between AP and sulfur dioxide; D: Interaction between AP and carbon monoxide; E: Interaction between AP and nitrogen dioxide; F: Interaction between AP and maximum 8-h average concentration of ozone. AP: Atmospheric pressure; PM_{2.5}: Fine particles; PM₁₀: Inhalable particles; SO₂: Sulfur dioxide; CO: Carbon monoxide; NO₂: Nitrogen dioxide; O_{3_8h}: Maximum 8-h average concentration of ozone.

patients are encouraged to pay more attention to the environmental effects on depressive symptoms. More stringent air quality control measures may be helpful to protect public mental health. Further studies involving more cities in China and a larger sample should be conducted to enable generalization. Thorough studies into the underlying mechanisms for environment impacting mental health are also warranted.

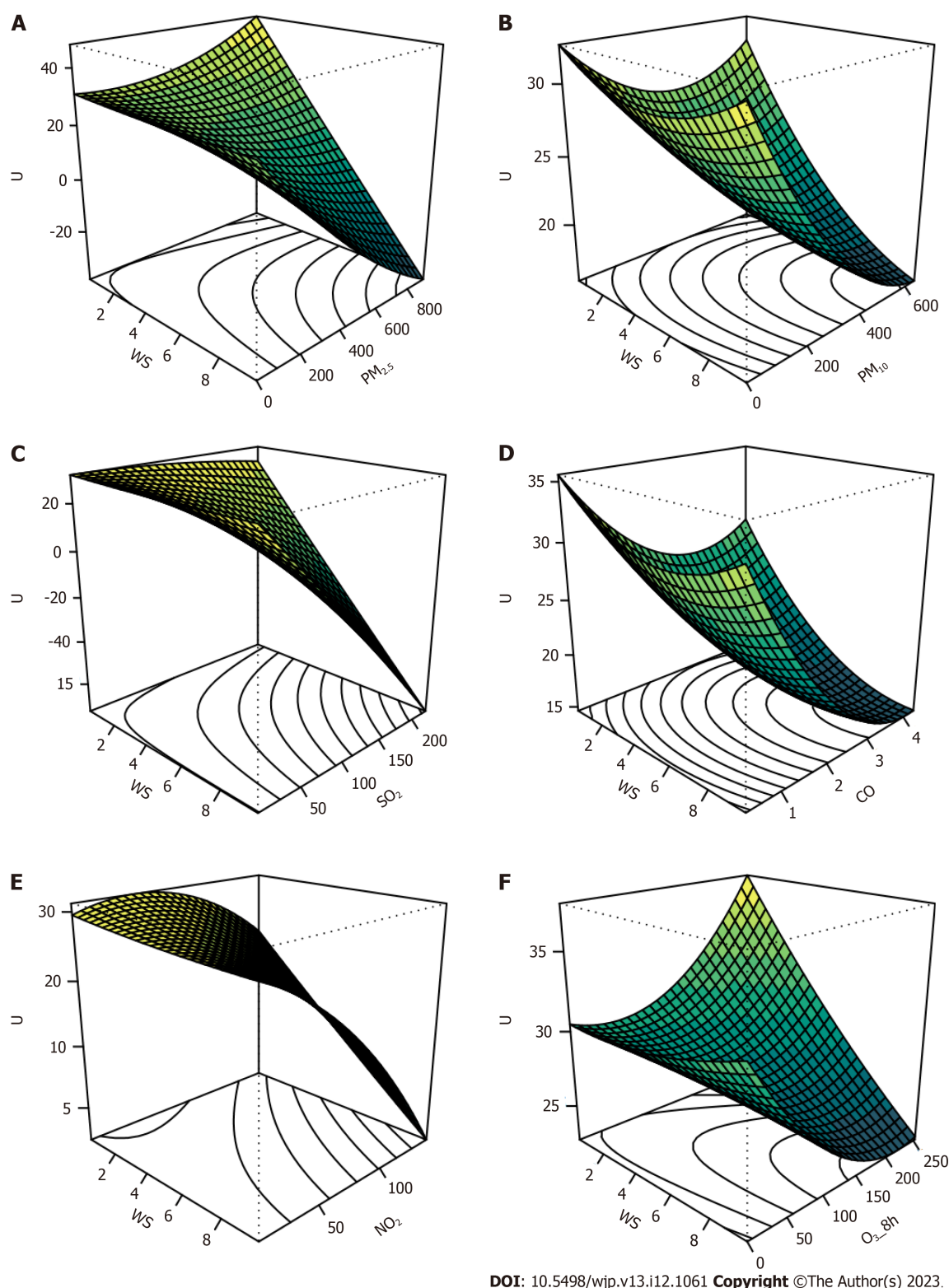


Figure 13 3D perspective plots of the interaction effects between wind speed and air pollutants on daily hospital admissions for depressive disorder. A: Interaction between wind speed (WS) and fine particles; B: Interaction between WS and inhalable particles; C: Interaction between WS and sulfur dioxide; D: Interaction between WS and carbon monoxide; E: Interaction between WS and nitrogen dioxide; F: Interaction between WS and maximum 8-h average concentration of ozone. WS: Wind speed; PM_{2.5}: Fine particles; PM₁₀: Inhalable particles; SO₂: Sulfur dioxide; CO: Carbon monoxide; NO₂: Nitrogen dioxide; O_{3_8h}: Maximum 8-h average concentration of ozone.

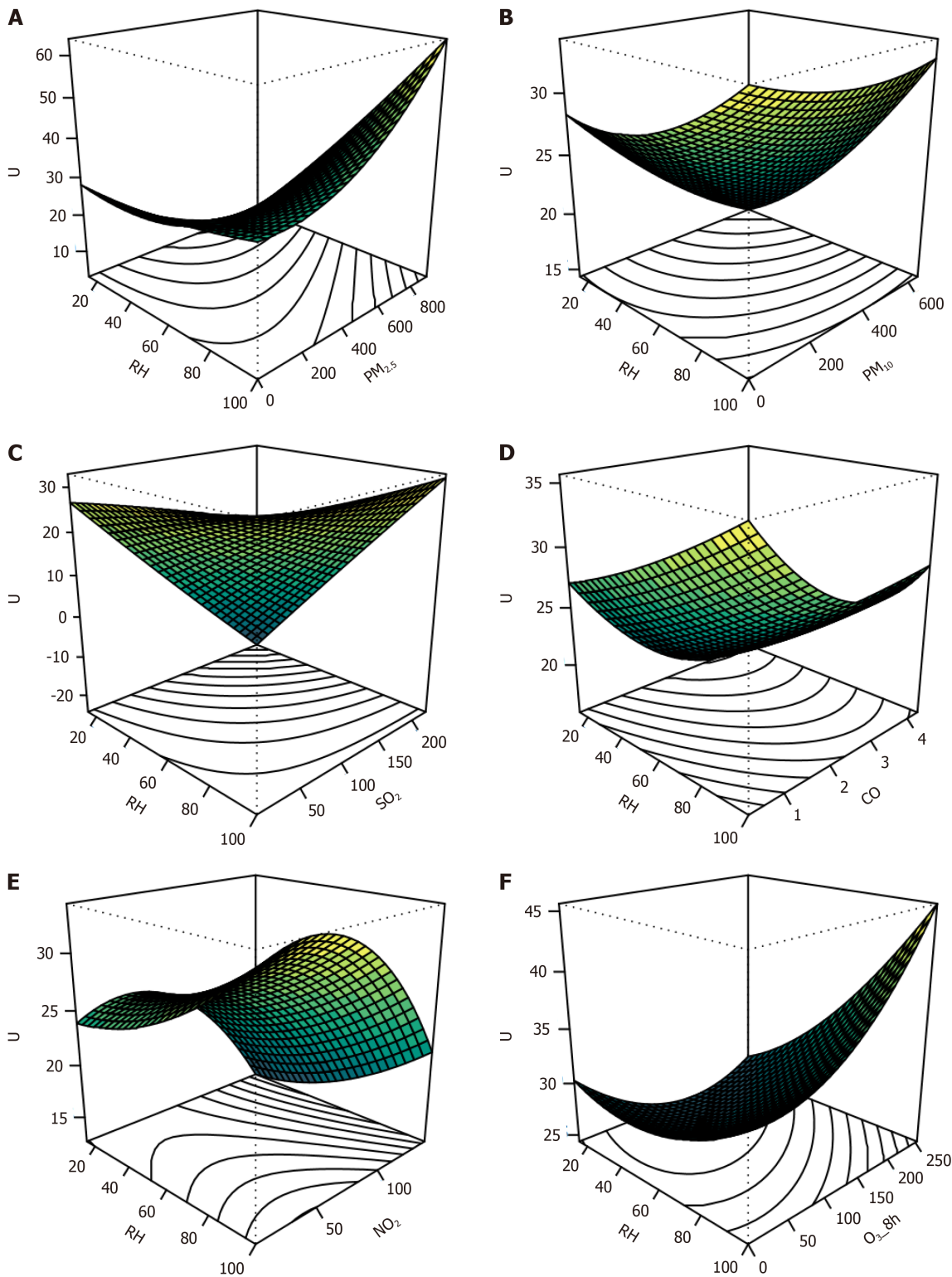


Figure 14 3D perspective plots of the interaction effects between relative humidity and air pollutants on daily hospital admissions for depressive disorder. A: Interaction between relative humidity (RH) and fine particles; B: Interaction between RH and inhalable particles; C: Interaction between RH and sulfur dioxide; D: Interaction between RH and carbon monoxide; E: Interaction between RH and nitrogen dioxide; F: Interaction between RH and maximum 8-h average concentration of ozone. RH: Relative humidity; $PM_{2.5}$: Fine particles; PM_{10} : Inhalable particles; SO_2 : Sulfur dioxide; CO: Carbon monoxide; NO_2 : Nitrogen dioxide; O_3_{8h} : Maximum 8-h average concentration of ozone.

ARTICLE HIGHLIGHTS

Research background

Environmental factors have been shown to affect individual mental health; however, the relationships and effects of different environmental factors in relatively cold areas have not yet been clarified.

Research motivation

We studied the effects of environmental factors, including their interaction effects, on daily hospital admissions for depressive disorders, aiming to discover the patterns of interaction between various environmental factors and provide regional references for this research field.

Research objectives

To investigate the single and interaction effects of environmental factors, we modeled, measured, and plotted these potential effects and added evidence to explore the complex relationship between environmental factors and human mental health.

Research methods

We performed a retrospective study using data from Harbin, China between 2015 and 2022, including daily hospital admissions for depressive disorders, meteorological data, and air pollution data. A time-series analysis was conducted using generalized additive models with quasi-Poisson regression to measure the effect of each single factor on depression. A bivariate response surface model was used to model the interaction effects of different factors. Our data came from a higher-latitude area, offering an opportunity to investigate regional differences in relationship patterns and the effects of environmental factors.

Research results

Our results showed that air temperature and wind speed influenced daily hospital admissions. Extremely low temperatures and wind speeds could increase daily hospital admissions for depressive disorders by approximately 50%. atmospheric pressure, relative humidity, and the air pollutant nitrogen dioxide had significant but moderate effects. Interaction effects between three meteorological factors and six air pollutants were discovered.

Research conclusions

Meteorological factors and air pollutants have single and interaction effects on daily hospital admissions for depressive disorders, and the effect pattern may be related to coal-fired heating in winter.

Research perspectives

Specific mechanisms behind the complex relationships of environmental factors are to be studied.

FOOTNOTES

Author contributions: Hu T designed and performed the research and wrote the paper; Guo BB designed the research and supervised the report; and Xu ZY, Wang J, and Su Y provided clinical advice.

Institutional review board statement: This study was reviewed and approved by the Ethics Committee of The First Psychiatric Hospital of Harbin.

Informed consent statement: This paper is a retrospective study that collects information on cases that have already occurred in the past, is analysed anonymously, does not expose patients' private information, does not cause harm to patients, and successful application for waiver of informed consent has therefore been made.

Conflict-of-interest statement: We have no financial relationships to disclose.

Data sharing statement: No additional data are available.

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Country/Territory of origin: China

ORCID number: Ting Hu 0009-0002-5352-7650; Zhao-Yuan Xu 0009-0000-7134-0422; Jian Wang 0009-0005-4386-6765; Yao Su 0009-0007-8178-7595; Bing-Bing Guo 0009-0006-2100-319X.

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