



# Article Association between Atrial Fibrillation Incidence and Temperatures, Wind Scale and Air Quality: An Exploratory Study for Shanghai and Kunming

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Abstract: As a common cardiovascular disease, atrial fibrillation has the characteristics of high morbidity, high disability, and high fatality rates, seriously endangering human health and sustainability. Some research has confirmed that environmental factors are related to the risk of illness and death from cardiovascular diseases (including atrial fibrillation), while there is still little comparison on the situation of the two cities in China. This research uses medical data in Shanghai and Kunming establishing, through two-step research, logistic models to compare the impacts on atrial fibrillation incidence to figure out the association between environmental factors (including air pollution, weather, temperature, and wind scales) and atrial fibrillation. Finally, this research shows that environmental impacts on atrial fibrillation prevalence have generality, regionality, and lagging characteristics. The result is significant for atrial fibrillation patients and provides a reliable medical theory basis for nursing measures. Besides, this research provides a prospective method of offering early warning for potential atrial fibrillation patients, helping to maintain human beings' sustainable development.

**Keywords:** atrial fibrillation; environmental factors; binary logistic model; sustainable development; early warning

# 1. Introduction

Atrial fibrillation (AF) is a disease that seriously endangers human health, recognized as a common complication of cardiovascular diseases. AF refers to the loss of routine and regular diastolic and contraction activities of the heart muscle, replaced by fast and uncoordinated weak peristalsis, causing the atrium to lose normal and effective contraction. Hence, blood tends to stagnate in the atrium and form a thrombus. The prevalence of AF in adults is about 3.0%, increasing with age and cardiovascular complications. In 2010, there were 33.5 million patients with AF worldwide. In the European Union and the United States, about one quarter of middle-aged people may suffer from AF [1]. In contrast, studies declare that AF's prevalence in China has increased by 20 times in the past ten years [2]. The total AF prevalence rate of people over 35 years old is 0.7%, and that of people over 80 is 7.5%, with the overall patient population reaching 4.87 million; however, 34.0% of them do not realize their AF history [3,4].

What is more, AF has seriously affected the sustainable development of human life. All-cause mortality caused by AF has increased by 1.5 times in men and 2 times in women [1]. Thromboembolic complications are the leading cause of death and disability in AF, and stroke is the most common manifestation, where strokes caused by AF account for 20.0% of all strokes [5–7]. Compared with non-AF-related strokes, AF-related strokes have



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**Copyright:** © 2021 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (https:// creativecommons.org/licenses/by/ 4.0/). severe symptoms, high disability, high mortality, and high recurrence. Its fatality rate is twice that of non-AF-related stroke [8].

As an essential part of the natural environment, climate profoundly impacts the global natural ecosystem and socio-economic system and even restricts all humankind's sustainable development. At present, many studies have confirmed that the lag-pattern impact of air pollutants may lead to the number of outpatients increasing, especially those who come to the hospital due to cardiovascular diseases, and an increase in the death rate of patients [9,10]. Besides, many governments have highlighted the importance of protecting the environment through sustainable development, wherein the air pollution index is vital [11,12]. With the increasing consciousness of air pollution's impact on human health [13,14], many scholars have focused on researching the association between air pollutant concentration and cardiovascular disease (CVD) prevalence. Nevertheless, as a sub-category of CVD, the association between environmental factors and AF are less concentrated. Therefore, this paper concentrates on the impact of environmental factors, researching the association between environmental factors and AF incidence to protect human health and provide early warnings.

# 2. Literature Review and Research Gap

Deeply researching and comprehending the impact of air pollution on the prevalence of AF is of significance. As AF is a sub-category of CVD, many scholars concentrate on researching the relationship between air pollution and CVD. Epidemiological studies have confirmed that air pollution impacts the cardiovascular system, and changes in air pollution concentrations are significantly related to the mortality and morbidity of CVD [15-18]. The previous research points out that every  $10 \,\mu g/m^3$  increase in the concentration of PM<sub>10</sub> in the air would increase the death rate from cardiovascular diseases by 2.4% [19], where  $PM_{10}$  denotes particulate matter (PM) with aerodynamic diameters  $\leq 10 \ \mu m$ , with the unit of concentration being  $\mu g/m^3$ . Similarly, a study evaluated the effects of long-term exposure to air pollution on Seoul residents' cardiovascular system, which carried out a 7-year follow-up of 136,094 participants in Seoul, asserting that the risk of cardiovascular events increased linearly with the increase in the average concentration of PM<sub>2.5</sub> [20], where FPM<sub>2.5</sub> denotes PM with aerodynamic diameters  $\leq 2.5 \,\mu$ m, recognized as a significant part of the total suspended particulate (TSP) as  $PM_{10}$  above, with the unit of concentration being  $\mu g/m^3$ . Besides, a prospective cohort study of 189,793 men over the age of 40 in 45 regions of China figured out that for every 10  $\mu$ g/m<sup>3</sup> increase in PM<sub>2.5</sub>, cardiovascular disease mortality increased by 9% [21]. Similarly, in 2010, the American Heart Association (AHA) published the scientific statement Air Pollution and Cardiovascular Diseases, stating that fine particulate matter exposure can lead to an increase in the incidence and mortality of CVD; therefore, fine particulate matter exposure is considered to be a form of controllable risk factor for the disease [21]. To sum up, it is of significance to develop the impact of air pollution on the prevalence of AF.

As a common complication of CVD, the prevalence of AF is considerably affected by air pollution, proven by many scholars. Studies have shown that in the general population of Asia, long-term exposure to  $PM_{2.5}$  is associated with an increased incidence of new-onset AF, and the situation for obese male subjects over 60 years of age with a history of hypertension or myocardial infarction is more serious [22]. It could be explained as ambient air pollution being positively correlated with elevated blood pressure and hypertension [23]. Furthermore, the Chinese Cardiovascular Health and Disease Report of 2019 points out, a series of studies based on daily data of air pollution and causes of death from 2013 to 2015 in 272 cities in China found that  $PM_{2.5}$ ,  $O_3$ ,  $SO_2$ , and  $NO_2$  increased by 10 µg/m<sup>3</sup>, and for every 1 mg/m<sup>3</sup> increase in CO, the risk of cardiovascular death increased by 0.3%, 0.3%, 0.7%, 0.9%, and 1.1%, respectively [24–28].

Different research figured out a similar result: air pollution's impact has a lag-effect pattern. For example, Kim et al. point out that the number of lag days for PM<sub>2.5</sub>'s effect on CVD depends on its chemical composition, and different lag times may vary with the

patient's health status and disease [29]. Moreover, the correlation between PM<sub>2.5</sub> and CVD mortality mainly occurred in the first 6 days, and the relationship between them lag from 0.5 to 3 days, and mortality was the largest [30].

Adding weather and wind scales into consideration, previous researchers have more specific conclusions. Y Ma et al. focused on the relationship between different air pollutants and the incidence of CVD on sandy and dusty days, and announced that  $PM_{10}$  and  $SO_2$  lag 1 day, and  $NO_2$  lags 2 days [31]. Research by Su Chang et al. asserts that the association between air pollutants and CVD incidence is more substantial in spring and winter, and the wind indirectly affects the incidence of CVD; meanwhile, the study also observed that the reduction in air pollution levels led to a reduction in the incidence of CVD [32]. Similarly, Ghanizadeh G et al. collected 1021 articles and pointed out that climate change parameters such as temperature, humidity, and air pollution significantly affect cardiorespiratory health [33]. What is more, it has been proven that meteorological fluctuations are most related to heart failure in the first 3 days of hospitalization, and that temperature and heart failure have a two-way relationship [34].

To sum up, although many studies have linked the increase of single or multiple air pollutants with adverse cardiovascular outcomes, the correlation between aggravated air pollution and AF has not been well investigated in China.

Therefore, this study explores the correlation between the concentrations and changes of different air pollutants (PM, O<sub>3</sub>, NO<sub>2</sub>, SO<sub>2</sub>, CO, et cetera) and AF by performing Holter monitoring on patients to provide a reliable medical theoretical basis to reduce the prevalence of AF, which is helpful for the assessment of the cause of AF in patients and the nursing measures for some high-risk groups. Those air pollutants are selected based on the WHO Air Quality Guidelines, which figured out that the four most common air pollutants are particulate matter (PM), ozone (O<sub>3</sub>), nitrogen dioxide (NO<sub>2</sub>), and sulfur dioxide (SO<sub>2</sub>).

#### 3. Materials and Methods

# 3.1. Data Collection and Resources

Shanghai MicroPort Medical (Group) Co., Ltd. (Shanghai, China) provided the research data containing the participants' test location, test time (date), and their health statuses (whether at a high risk of AF). All the data were collected through the investigation, and there was no private data of the participants. What is more, the duration of the investigations in Shanghai and Kunming were 1 year and 6 months, respectively. In addition, the related environmental factors were provided, including the current day and the previous 14 days' weather, temperature (with the unit of degrees Celsius), wind force scales, air quality indexes (AQI), and air pollution concentrations ( $PM_{2.5}$ ,  $PM_{10}$ ,  $NO_2$ ,  $SO_2$ ,  $O_3$ , CO), where the first five kinds of pollutants used the unit of  $\mu g/m^3$ , except for CO, with the unit of mg/m<sup>3</sup>. Therefore, there are 11 independent variables for each day, 15 days involved, and the AF status is the dependent variable.

In this research, two databases of Kunming and Shanghai were used to analyze the impact of the air pollution issue and the effect of other environmental factors. There were 3518 records collected, and after removing the incomplete records, 3160 records were kept (732 for Shanghai and 2428 for Kunming, respectively). Meanwhile, the high-risk AF (ICD-10 code I48) rate for Shanghai and Kunming participants were 9.3% and 2.8%, respectively. This research divided the data into two parts based on the AF status, as modeling parts (80%) and checking parts (20%), according to the locations.

#### 3.2. Research Logic and Data Curation

This paper establishes a two-step analysis process, and the variables are categorized into five types. The first step is to convert the parameters into binary variables and analyze whether they are significantly related to AF or not. The second step is to use the primary data to establish a logistic regression model to interrupt the effect quantitatively. In the first step, every related factors' differences were calculated to demonstrate the changes between the following days, shown in Equation 1 below.

$$\Delta(X)_{j,t}^{i,B} = \begin{cases} 1 & \text{when } (X)_{j,t}^{i} - (X)_{j,t+1}^{i} > 0 \\ 0 & \text{when } (X)_{j,t}^{i} - (X)_{j,t+1}^{i} \le 0 \end{cases}$$
(1)

where  $\Delta(X)_{j,t}^{i,B}$  is a binary variable recording the increase or decrease of the variable X between the day t and a day before for participant i, for which 1 means an increase and 0 means a decrease or equal.  $(X)_{j,t}^{i}$  denotes the value of the factor X on the day t for the tester i in city j. For instance,  $(AQI)_{5}^{i}$  represents the AQI index for the participant i on the day (5 days before the test day). Hence, if the  $\Delta(X)_{j,t}^{i,B}$  is positive, it means that the environment is getting worse from day t + 1 to day t.

In this research, all  $\Delta(X)_{j,t}^{i,B}$  are classified into the first-order difference category. What is more, to denote AF's situation, this paper use 1 to represent the patient being at a high risk of AF (over 50% chance of AF), and 0 for low risk (less than 50%). Furthermore, this research calculates the second-order differences, categorized into the second-order-difference type, to illustrate the fluctuating rate, shown in Equation (2) below.

$$\Delta'(X)_{j,t}^{i,B} = \begin{cases} 1 & \text{when } \Delta(X)_{j,t-1}^{i} - \Delta(X)_{j,t}^{i} > 0\\ 0 & \text{when } \Delta(X)_{j,t-1}^{i} - \Delta(X)_{j,t}^{i} \le 0 \end{cases}$$
(2)

where  $\Delta'(X)_{j,t}^{i,B}$  denotes whether the fluctuating speed of the difference of variable X between the day t – 1 and t and the difference between the day t and t + 1 for the participant i in city j. Based on the definition of the second-order-difference variables, if the value is zero, it means that the change is stable, keeping an unchanging tendency. However, if it is positive, it could represent one of the following three situations: (1) the value's decreasing rate is getting lower. (2) The value's increasing speed is getting faster. (3)  $(X)_{j,t}^{i}$  is the smallest compared with  $(X)_{j,t-1}^{i}$  and  $(X)_{j,t+1}^{i}$ . To sum up, if the second-order-difference variable is positive, it describes a more complicated environmental degradation.

Therefore, there are 384 variables in this two-step research, divided into five categories (AF, primary pollution indexes, temperature, weather, and wind scale, first-order difference, and second-order difference, respectively). This research uses SPSS software (International Business Machines Corporation. Armonk, NY, USA. Version 25) to analyze the association between environmental factors and the incidence of AF and establish binary logistic models to elaborate on the relationship.

In the first step, all differences of variables (first-order difference and second-order difference) were converted into binary variables for modeling, 1 denoting the increase and 0 for the decrease.

#### 3.3. Binary Logistic Regression

This research established two models through binary logistic regression, one for Shanghai and one for Kunming (called model 1 and model 2). Many scholars have used this method to determine the association between the risk factors and CVD [31–33], while this research concentrates on the changing of environmental factors. With the binary logistic regression, this research established the models as:

$$p_{j}^{i} = \frac{e^{(c+D_{j}^{i}+U_{j,0}^{i}+U_{j,1}^{i}+U_{j,2}^{i})}}{1+e^{(c+D_{j,0}^{i}+U_{j,0}^{i}+U_{j,1}^{i}+U_{j,2}^{i})}}$$
(3)

$$U_{j,0}^{i} = \sum_{t} \beta_{X,t}^{j,0} \times (X)_{j,t}^{i}$$
(4)

$$\mathbf{U}_{j,1}^{i} = \sum_{\mathbf{t}} \beta_{X,\mathbf{t}}^{j,1} \times \Delta(\mathbf{X})_{j,\mathbf{t}}^{i}$$
(5)

$$U_{j,2}^{i} = \sum_{t} \beta_{X,t}^{j,2} \times \Delta \prime (X)_{j,t}^{i}$$
(6)

$$D_{j}^{i} = \begin{cases} 0 & \text{if } (X)_{j,t}^{1} = 0\\ \sum_{d,l} \beta_{d,t}^{l} & \text{if } (X)_{j,t}^{i} = 1 \end{cases}$$
(7)

where  $p_j^i$  denotes the probability that participant i has a high risk of AF, and  $U_{j,0}^i$ ,  $U_{j,1}^i$ , and  $U_{j,2}^i$  represents the utility of the sum of the primary factors (except for weather and wind scale), and the utilities of the sum of the first-order category factors and the second-order category factors, respectively.  $D_j^i$  is the sum of the dummy variables, representing the different situations of the weather and wind scales.  $\beta_{X,t}^{j,2}$  is the coefficients of the factor X in location j (Shanghai or Kunming, or the composition of the two cities) of the difference calculating order n (0 or 1 or 2) of the time t. Using the binary variables and the primary value of the factors, the  $p_i^i$  value could be calculated.

With 95% confidence interval (95% CI), this research first selected the statistically significant variables by checking their Wald statistic. What is more, this research reconsidered the correlations between every two variables to optimize the models.

With the binary logistic model results, the effect of the factors could be estimated by the coefficients' value. Besides, the odds ratio (OR) values were calculated to demonstrate the factors' impact, which equaled the exponential of  $\beta_{X,t}^{j,n}$ . If the OR value was larger than 1, it represented the specific factor's increase, positively related to the increase of the AF risk.

## 3.4. Model Tests

In this part, the models were tested by three methods, including chi-square tests, receiver operating characteristic curve (ROC curve) plotting, and applying to the new data sample to check the accuracy.

This research firstly analyzed the models' goodness of fit via omnibus tests and the Hosmer and Lemeshow test. They both use the chi-square value to check whether the model is further optimized to reject the null hypothesis. By comparing the chi-square values and the related *p*-values, it is persuasive to reject the null hypothesis evidently. Moreover, as many scholars have used the ROC-AUC approach in this field [34–37], this research used this method to examine this model's accuracy.

In conclusion, this research firstly checked the significance of all the variables and then used four methods to determine the models' statistical significance to determine the association between environmental factors and AF.

# 4. Results

This research establishes two models through the analysis methods above, and they could show three vital results about the environmental factors to AF, from the epidemical, geographical, and lag-affect pattern aspects.

# 4.1. Qualitative Models' Results

Analyzing the data using SPSS, this research figured out the factors that significantly affect the AF status qualitatively as the first step to establishing the models. The result is shown in Figures 1 and 2 (the qualitative model is in the Appendix A in Table A1).

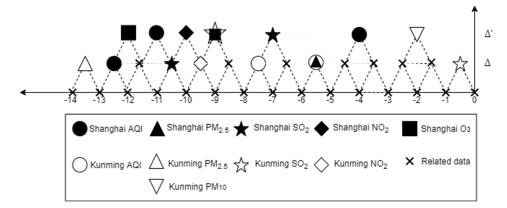


Figure 1. The impact of the air pollutants.

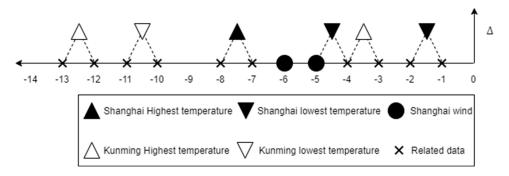


Figure 2. The impact of the weather, wind, and temperature difference.

As mentioned above, the  $\Delta$  and  $\Delta'$  in the two figures above denote the first and second-time differences. This step is used to figure out the significant factors and the related variables.

#### 4.2. Quantitative Models Results

Using SPSS 25, the following two models in Table 1 below exhibit the quantitative relationship between the specific environmental factors and AF.

In the tables above, the OR value is the impact of per 1 unit increase of the particular variable on AF's incidence when other conditions are all kept unchanged, which equals to the  $exp(\beta)$ . The following Figures 3 and 4 plot the OR values with their lower and upper bounds, where the Y-axis shows the variables and the X-axis shows the corresponding OR values range with 95% CI and the expected OR value.

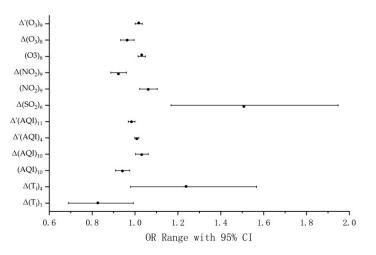


Figure 3. The odds ratio value in model 1.

	Variables			Sha	nghai					Kı	unming		
Description						OR <sup>4</sup> 95% CI						OR <sup>4</sup>	95% CI
		B <sup>1</sup>	Std. <sup>2</sup>	Sig. <sup>3</sup>	OR	Lower Bound	Upper Bound	β	Std.	Sig.	OR	Lower Bound	Upper Bound
	W <sub>5</sub> (1)	-1.645	0.649	0.011	0.193	0.054	0.689						
wind	W <sub>6</sub> (1)	1.121	0.426	0.009	3.067	1.330	7.076	-					
Temperature	$\begin{array}{c} (T_h)_3 \\ (T_h)_4 \\ (T_l)_{10} \\ (T_l)_{11} \\ (T_h)_{13} \\ \Delta(T_l)_1 \\ \Delta(T_l)_4 \end{array}$	-0.190 0.214	0.093 0.120	0.041 0.075	0.827 1.238	0.689 0.978	0.992 1.567	-0.207 0.200 0.347 -0.553 0.151	$\begin{array}{c} 0.072 \\ 0.080 \\ 0.153 \\ 0.170 \\ 0.062 \end{array}$	0.004 0.012 0.023 0.001 0.015	$\begin{array}{c} 0.813 \\ 1.221 \\ 1.416 \\ 0.575 \\ 1.163 \end{array}$	$\begin{array}{c} 0.706 \\ 1.044 \\ 1.048 \\ 0.412 \\ 1.030 \end{array}$	0.936 1.427 1.911 0.802 1.313
AQI	$\begin{array}{c} (AQI)_{10} \\ \Delta(AQI)_5 \\ \Delta(AQI)_7 \\ \Delta'(AQI)_4 \\ \Delta'(AQI)_{11} \end{array}$	-0.060 0.008 -0.016	0.018 0.005 0.008	0.001 0.089 0.047	0.942 1.008 0.984	0.909 0.999 0.968	0.975 1.018 1.000	0.041 0.029	0.011 0.013	0.000 0.030	1.042 1.029	1.019 1.003	1.066 1.056
TSP	$\Delta(PM_{2.5})_{13} \\ \Delta(PM_{10})_1$							0.125 0.009	0.033 0.017	0.000 0.602	1.133 1.009	1.062 0.976	1.208 1.043
SO <sub>2</sub>	$\begin{array}{c} \Delta(\mathrm{SO}_2)_0\\ \Delta(\mathrm{SO}_2)_6\\ \Delta'(\mathrm{SO}_2)_9 \end{array}$	0.411	0.130	0.002	1.508	1.168	1.947	-0.157 0.109	0.058 0.041	0.007 0.008	0.855 1.116	0.763 1.029	0.958 1.209
NO <sub>2</sub>	$(NO_2)_2$ $\Delta(NO_2)_9$	$0.060 \\ -0.080$	0.020 0.020	0.002 0.000	1.062 0.923	1.021 0.887	1.104 0.960	-0.014	0.023	0.531	0.986	0.943	1.031
O <sub>3</sub>	$(O_3)_8 \\ \Delta(O_3)_8 \\ \Delta'(O_3)_9$	$0.031 \\ -0.037 \\ 0.017$	0.009 0.016 0.009	0.000 0.024 0.044	1.031 0.964 1.017	1.014 0.933 1.000	1.049 0.995 1.034						
Constant	с	-4.008	0.874	0.000	0.018			-3.098	1.478	0.036	0.045		

**Table 1.** The results of the quantitative models.

<sup>1</sup> The β in this table represents the coefficients of the variables. <sup>2</sup> The Std. in this table represents the coefficients' standard deviation value. <sup>3</sup> The sig. in this table denotes the *p*-value of the variables. <sup>4</sup> The OR in this table denotes the odds ratio value.

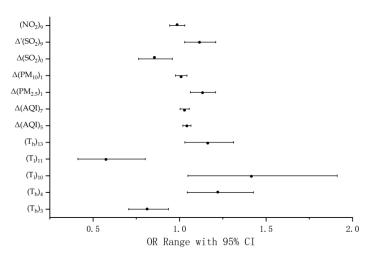


Figure 4. The odds ratio value in model 2.

For example, the  $\Delta(SO_2)_6$  in Table 1 model 1 is 1.508 (95% CI: 1.168, 1.947), larger than 1, showing that, with all other variables kept the same, for every 1 unit increase of the SO<sub>2</sub> index from the seventh day before the test date to the sixth day, the risk would increase by 1.508 times. In contrast, for the  $(T_h)_3$  (OR value as 0.813) in model 2, if the highest temperature occurred on the third day before the record increasing 1 degree, the patients in Kunming would have a lower probability of getting AF on the testing day, and the probability would decrease by about 20%. Therefore, the models' results could illustrate the relationship between environmental conditions changing and AF risk changing. What is more, with the descriptive statistics, the absolute value of the variables and related variables could be illustrated, showing the results more detailed and evidently.

As air pollution could last for days, and the air quality tendency is ordinarily stable in a particular city, the correlation between variables in this research is appropriately released. Hence, some variables may still be kept for the accuracy of the models. Moreover, by analyzing the data of air quality, wind scale, and temperature to assist in analyzing the association, this research used Figures 5 and 6 to illustrate the value of those selected variables. In the two figures, the boxes to the left of the vertical lines are for model 1, and others for model 2, and they describe the upper and lower bound of the value of the factors with their 25% to 75% range. The spots and the horizontal lines inside the boxes denote the mean and median values, respectively.

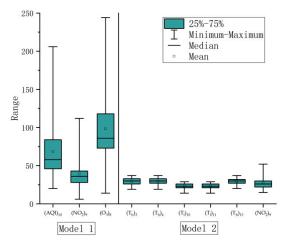


Figure 5. The description of the primary air quality and temperature data.

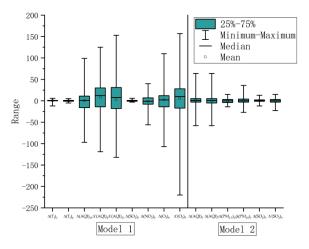


Figure 6. The description of the differential variables.

As most coefficients in the two models were from -1 to 1, with a larger value of the variable, this specific factor would cause a more significant impact. For instance, the (O<sub>3</sub>)<sub>8</sub> in model 1 is statistically significant, and the mean and median value of this variable were close to 100, larger than other factors'; therefore, the impact of the concentration of O<sub>3</sub> (lag of 8 days) could be evidently shown. Besides, judging by this graph, some insignificant variables could be explained, for example,  $\Delta(PM_{10})_1$  in model 2 (*p*-value is 0.602). In Figure 6, it is clear that the variable's value was close to 0, and half of them distributed close to 0. Therefore, the impact given by  $PM_{10}$ 's concentration changing was limited. Hence, compared with other factors, it could be statistically insignificant. In conclusion, these two figures above can explain the models' results by analyzing the distribution and value of the variables.

What is more, many variables were insignificant in the models because they could explain the first step's results. For example, the second-difference AQI value of four days before the test day in Shanghai (denoted as  $\Delta'$  (AQI)<sub>4</sub>) was insignificant, with 95% CI (*p*-value as 0.089), which is kept to show the impact of the AQI changing from the fifth day to the third day before the test day, which was found significant in the first step. Other insignificant variables played the same role. Besides, all insignificant variables' OR interval skipped 1, so it is hard to identify them to judge their impact confidently. Nevertheless, those variables are vital to support the conclusion achieved in the first step.

According to the tests of the models, the results of the goodness of fit test results are shown in Table 2. With the omnibus test's chi-square values and the significance values of all passing the test, it is evident that we can judge that the models are eloquent. As the *p*-values for the omnibus tests are all less than 0.5, this represents that the null hypothesis test could be rejected. Besides, the Hosmer and Lemeshow test results show that the information of the data has been fully extracted (all *p*-values are larger than 0.05), which means the models have good fitness. Therefore, from the results of these two tests of the models, all models have been proven to be relatively accurate.

Table 2. The results of models' goodness of fit tests.

Location	Omnibus Tests of	Model Co	efficients	Hosmer and Lemeshow Test				
	Chi-Square	df	Sig.	Chi-Square	df	Sig.		
Shanghai	63.89	14	0	15.394	8	0.052		
Kunming	44.41	12	0	10.039	8	0.262		

The ROC-AUC test results are shown in Table 3 below. As the AUC represents the degree or measure of separability of the model, it was chosen to be an approach to analyze the models' accuracy. Therefore, taking the AUC value for Shanghai's model as an example, which is 0.824 (95% CI: 0.771, 0.877), it represents excellent discrimination. Besides, the AUC value for model 2 is 0.756, denoting that the model is acceptable and has excellent abilities to separate AF's risk level.

Table 3. The description of the ROC curves and AUC.

T		0.1 5	A commutation Sig	Asymptotic 95% Confidence Interval				
Location	Area	Std. Error	Asymptotic Sig	Lower Bound	Upper Bound			
Shanghai	0.824	0.027	0	0.771	0.877			
Kunming	0.756	0.034	0	0.689	0.822			

Furthermore, by calculating the accuracy by checking the sample data, the accuracy was 88.8% and 97.3% for the two models mentioned above. As this research focused on the association between environmental factors and AF prevalence, the accuracy could prove the models' validity.

In conclusion, all models have shown great accuracy, separability, and information extraction through these three tests' results. Hence, these models were selected to analyze the association between AF and environmental factors.

#### 4.3. Epidemicical Results

The epidemical result is that three of the most common air pollutants mentioned by the WHO were positively correlated with the prevalence of AF, as PM, NO<sub>2</sub>, and SO<sub>2</sub>. Moreover, as the AQI is a synthetic factor in measuring the pollution level, the OR values of the AQI could demonstrate the impact of air pollution comprehensively.

Judging by the models, it is clear that with PM and SO<sub>2</sub> concentration increasing, it poses a higher risk for people getting AF, proven by the OR value of the concentration increase of  $PM_{2.5}$  13 days prior to the test day in Kunming (95% CI: 1.062, 1.208) and SO<sub>2</sub> 6 days prior in Shanghai (95% CI: 1.168, 1.947), respectively. Moreover, model 1 highlights that the concentration of O<sub>3</sub> had a negative impact, which was proven by the OR value

for the  $O_3$  concentration per 1 unit increase in fluctuation around the ninth day before the record date (95% CI: 1.000, 1.034). What is more, the AQI's increase 5 days (95% CI: 1.019, 1.066) and 7 days (95% CI: 1.003, 1.056) previously in model 2 prove that the comprehensive air pollution increase will cause the patients to get into a more dangerous environment.

Concentrating on the fluctuation of air pollutant concentrations (defined by the secondorder difference category), all coefficients were positive, except for one, the AQI fluctuation 11 days before the test day in model 1. However, by analyzing the primary data, it shows that the AQI index from the 12 days prior to the 10 days before the test day followed the increasing trend, while the increasing speed slowed down. Therefore, this research concludes that if the air pollution situation shows the tendency of going down initially and then immediately going up, there will be a higher risk of AF.

This paper uses the following Figures 7 and 8 to illustrate the impact of the pollutants and temperature changes.

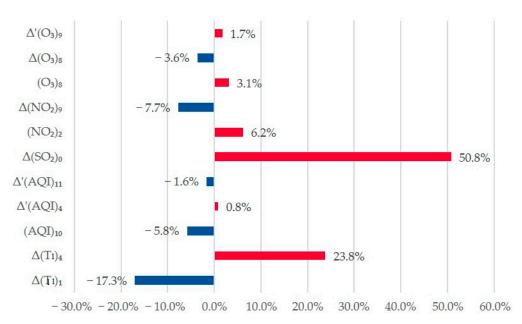


Figure 7. The odds change in Shanghai.

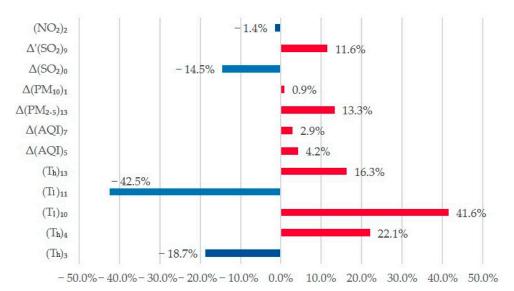


Figure 8. The odds change in Kunming.

The X-axis value is the odds change value, representing the changing of AF's incidence when the specific variable increases by 1 unit. From the two figures shown above, it is

clear that different cities are sensitive to different parameters. What is more,  $SO_2$  is more significant than the other variables, not only because the odds change is larger than for the others (close to 0.5), but also because both models contain this pollutant. Besides, as shown in Figures 7 and 8, it is clear that with the pollution getting severe, regardless of the absolute index or the polluting speeding up, people will have a higher risk of AF. For instance, the probability of AF for patients in Kunming will have 4% increase if there is 1 unit increase of the AQI, since from the sixth day to the fifth before the test day. The SO<sub>2</sub> increase 1 day previously played a protective role, because, in Kunming, the SO<sub>2</sub> concentration was relatively stable (mean and standard error for the increase is 0.21 and 0.057); if the concentration increases significantly, the impact could be significant.

Finally, both models' constant coefficients were negative, as a protecting factor from AF. There are few high-risk patients in the database (67 in Kunming and 68 in Shanghai) compared to low-risk people through the statistical analysis. Therefore, the constants here demonstrate that only a few people are at high risk in the initial stage.

To sum up, from the results related to the pollutants' concentrations, it is clear that air pollution may cause the prevalence of AF growth. Similarly, the second-order difference demonstrates that the fluctuating rate of the air pollutants' concentrations changing can significantly affect AF prevalence.

#### 4.4. Geographical Results

The geographical effect is clear from the models, mainly represented by the model variables involved. According to model 1 and model 2, it is clear that they contain different variables. The following Figure 9 illustrates the comparisons of the odds change of the temperature and wind on AF.

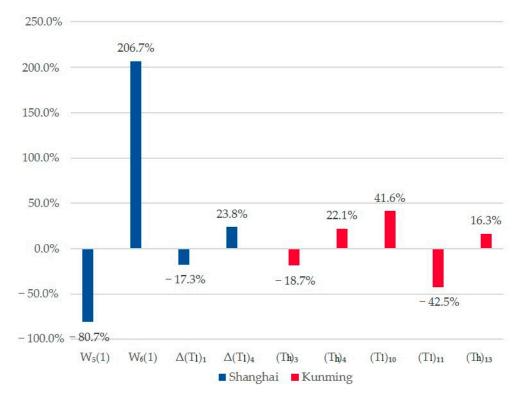


Figure 9. The odds change by the wind scale and temperature.

Nevertheless, different cities are sensitive to different days. From Figure 9, the AF incidence in Shanghai was not sensitive to the temperature, while it was closely related to the weather. In contrast, the situation in Kunming was the opposite. The OR values of the lowest temperature that occurred in Kunming (10 and 11 days lag for Kunming) were 1.416 (95% CI: 1.048, 1.911) and 0.575 (95% CI: 0.412, 0.802), showing that the different

days' temperature had a different impact. Although the temperature was usually kept at a relatively stable level, the impacts may have offset each other, and may have caused a considerable effect when the temperatures were changing a lot during those days.

Concentrating on the wind scales' impacts, it had a relatively complex influence. If the wind scale was above level 4 on the sixth day before the test, people had a higher probability of getting AF (95% CI: 1.330, 7.076) in Shanghai, while the wind on the five days lag decreased the chance (95% CI: 0.054, 0.689).

In conclusion, the models describe the difference caused by the location of the patients. The impact of geography is entire, while the negative impact of air pollution is the same.

# 4.5. Pollution's Lag-Pattern Impact

In this research, most of the variables were not directly related to the test date, with a lag of 3 to 12 days. Therefore, it is reasonable to conclude that the risk of AF is associated with the environment and air pollutions. Comparing the models, nearly all the significant variables had at least three days lag, representing that the impact of the pollution could not immediately cause AF.

# 5. Discussion

## 5.1. The Influence of Air Pollution Factors on AF

There has been some research concentrating on the medical mechanism of the impact given by environmental factors on CVD. For example, Bhatnagar [38] in 2004 asserted that, as important components of  $PM_{2.5}$ , metals in the environment deposit in the heart and blood vessels, and their toxicity increases the risk of CVD. Besides, some gaseous pollutants can mediate or modify PM, which may cause ischemic heart diseases. Similarly, it has been proven that  $O_3$  is highly related to pulmonary inflammation and edema, but it is still unclear whether it is associated with CVD [39]. Besides, other research figured out that environmental factors (especially air pollution) can cause CVD, such as endothelial dysfunction and the role of oxidative stress [40], which proves this paper's results. What is more, PM can cause atherosclerosis via promoting vascular dysfunction and alternating the vasoactive mediators' responsiveness [41]. These results explain the medical mechanism of CVD related to environmental factors, and they highlight the significant impact of environmental factors, which shows the importance of this paper's research.

After adjusting for other gaseous pollutants, the significance of PM<sub>2.5</sub>, PM<sub>10</sub>, SO<sub>2</sub>, NO<sub>2</sub>, and O<sub>3</sub> in research results is highlighted, suggesting that they are currently the main pollutants in Shanghai and Kunming. There is a positive correlation between the risk of AF for people in Shanghai and Kunming and the increase in PM concentration [42]. Other studies have evaluated the impact of six major environmental pollutants on the risk of AF in patients with cardiac implantable electronic devices (CIED). With a 10  $\mu$ g/m<sup>3</sup> increase in PM<sub>2.5</sub> and PM<sub>10</sub> concentrations, the risk of AF increased by 3.8% and 2.7%, respectively; however, the correlation with other gaseous pollutants (SO<sub>2</sub>, NO<sub>2</sub>, CO, O<sub>3</sub>) was not statistically significant [43,44], which is similar to this research's results. Furthermore, this research emphasizes the impact of air pollution fluctuations in Shanghai and Kunming, filling the blanks of previous research.

On the other hand, statistics show that the increase in the number of visits for arrhythmia in the emergency room was significantly related to  $PM_{2.5}$  on warm days (>23 °C) and cold days (<23 °C), and high levels of  $PM_{2.5}$  would increase the risk of emergency visits due to arrhythmia [45]. In this research, the average highest temperature in Shanghai on the seventh and eighth day before was 23.3 °C, and 22.8 °C (standard error as 0.334 °C and 0.312 °C), respectively, and the lowest temperature in Kunming was close to 23 °C as well, which meets the conclusion. A large number of studies have explained this impact from the perspective of pathology, as certain substances can cause oxidative stress, cardiovascular inflammation, endothelial dysfunction, high C-reactive protein level, myocardial ischemia, and right atrial pressure [46–50], all of which are related to atrial remodeling and AF pathophysiology [51–54]. Therefore, it is persuasive to confirm that air pollution has the risk of increasing the probability of arrhythmia or other acute cardiac events. Hence, patients at high risk of AF should be educated to monitor the local air quality index and follow the recommendations to reduce exposure and reduce outdoor activities. What is more, this research helps prevent AF's high-risk situation and can give an early warning to the number of outpatients increasing through the lag-affect analysis of air pollution and wind. Therefore, this research can further strengthen the theoretical basis of urban air pollution control, establish air pollution control measures, and optimize community and hospital diagnoses and treatment configuration.

# 5.2. AF Risk Factors Are Variable in Different Cities

According to this research, for the results of model 1 and model 2, air pollution particles' impact on AF incidence was similar, while the impact of temperature factors showed opposite laws. Besides, the risk of AF in Shanghai was more sensitive to the effects wind scale, while that in Kunming was not.

Many studies have shown that the effects of seasons and temperature on the incidence and mortality of cardiovascular diseases are different in different cities [55,56]. The results of an epidemiological study containing 16 cities in China showed that  $PM_{10}$  was significantly associated with deaths from cardiopulmonary system diseases, and the association is still statistically significant after adjusting for other gaseous pollutants; additionally, the relationship between  $PM_{10}$  and death risk in different cities differed due to the different levels of pollutants in each city [57]. It explains the reason why the impact of  $PM_{10}$ 's concentration increase has different lag periods.

The seasons' effect on the risk of AF is another consideration. K Spengos et al. analyzed the symptom onset pattern of more than 300 patients with acute cardiogenic stroke in Greece for the first time due to AF, figuring out that the patients' symptoms were cyclically distributed, with the peak season in winter and the incidence in summer declining [58]. Similarly, J Ahn et al.'s study of the seasonal changes in AF incidence in Seoul also showed that the frequency of AF in summer was significantly lower than in other seasons [59]. Moreover, a national statistical study in Germany showed that when the outdoor temperature was between 0 and 10 °C, the hospital admission rate for arrhythmia, including AF, reached its peak [60]; however, some studies have shown that in some temperate countries, cardiovascular diseases in winter are lower than in summer [61]. Although most participants took the test in autumn in this research, it is hard to show the seasonal impact, and the different impacts of changing temperature, weather, and wind scale on AF risk have been pointed out.

In summary, different cities have different effects on the incidence of AF due to the intensity, mode of action, duration of the effect of cold and heat, and maybe the local residents' physical fitness. Therefore, different locations may have their own specific models to estimate the risk of AF, which might be another way to describe whether a city is livable or not.

#### 5.3. The Impact's Lag-Affect Pattern

The lag-effect of environmental factors' impact on AF is shown by the variables' OR values, most of which show a three to twelve days lag. For example, the increase of the previous six day's  $SO_2$ 's concentration is statistically significant in Shanghai. Although the four kinds of air pollutants ( $PM_{2.5}$ ,  $PM_{10}$ ,  $SO_2$ , and  $O_3$ ) could significantly affect AF incidence, they all show lag-affect influence.

Many studies have also confirmed that the lag of air pollutants may affect the occurrence of CVD and the mortality rate. The correlation between  $PM_{2.5}$  and CVD mortality mainly occurred in the first 6 days, with the fourth day being the most significant [62]. Additionally, LC Martins and other scholars have asserted that air pollutants' delayed effects (CO,  $PM_{10}$ ,  $O_3$ ,  $NO_2$ , and  $SO_2$ ) have different effects on different genders. Besides heart failure inducing diseases, the delayed effects have more noticeable effects on women for other cardiovascular diseases [63]. Similarly, M Dastoorpoor et al. announced that for people over age 60 and under 18, after a delay of 3 days and 13 days, there was also a significant relationship between the increase in the interquartile range (IQR) of particles below 10  $\mu$ m and cardiovascular death. For those under 18 years old (lag 11) and over 60 years old (lag 9), there was a significant relationship between the increase in IQR of NO<sub>2</sub> and CO and cardiovascular death, respectively [64]. Kim et al. pointed out that the number of lag days for the impact of PM<sub>2.5</sub> on CVD depended on its chemical composition, and the different lag times may vary with the patient's health status and disease [65].

This research selected the medical records in Shanghai and Kunming for statistical research, and the conclusion showed the similarity with the previous study. Notably, the pollutants' delayed impact on AF in the two cities was different, which may be due to the study cities' different characteristics, such as indoor air quality, local climate type, residents' sensitivity to pollutants (such as economic level, population age, and smoking rate), pollutant concentration levels, and differences in pollutant composition.

#### 5.4. Research Significance, Limitations and Prospects

This research attempted to figure out the association between environmental factors and AF, which is of significance to the government, medical research, and patients. As it emphasizes the hysteresis of air pollutions and weather changes' impact, the government and hospitals can use these conclusions to improve and optimize weather reports and forecasting. Meanwhile, this research uses the data of two cities in China, comparing and demonstrating the different impacts caused by the different locations. Therefore, other researchers can follow this direction to determine the overall regularity, boosting AF and CVD comprehension. Furthermore, patients with a history of AF could be alerted by this research to pay more attention to air pollution and take self-protection more seriously. On the other hand, medical workers may also remind the patients to remember the air pollution conditions in order to protect them in the following days.

However, there were still some limitations to this research. This research used the average value of each monitoring site in each city as the population exposure level of pollutants. However, each monitoring site's pollutant measurement methods may differ, so the environmental monitoring results and the individual level of pollutant exposure might not be accurate enough. Hence, this study cannot calculate the precise individual exposure in each city, affecting the analysis results. Meanwhile, as the participants' gender and age level were often misrecorded in the records, this research does not contain those two vital factors to optimize the model. What is more, as most of the data were collected in the summer and autumn, some contingents might exist in this paper's results. Therefore, more data are being collected by a more extended duration investigation and tracking surveys, which can help to improve the results. In the subsequent research, more detailed data will be used to optimize the models and provide more conclusions.

#### 6. Conclusions

Increasing environmental pollution will increase the incidence of atrial fibrillation, and the concentration of the increase of the two major air pollutants ( $PM_{10}$  and  $SO_2$ ) will raise the risk significantly. Meanwhile, the environmental impact of each region is regional and lagging. Besides, the acceleration of air pollution will also increase the probability of atrial fibrillation. Therefore, continuously monitoring the environmental indexes are of importance for warning and protecting potential patients. What is more, different cities may have their unique model to evaluate and estimate the significant factors to improve the local environmental departments' working process, as integrating the previous environmental indexes into the daily weather reports has a strong impact on the sustainability of AF.

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Z.C., M.D, and W.Y.; validation, Q.Z. and W.Y. All authors have read and agreed to the published version of the manuscript.

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**Conflicts of Interest:** We have no conflict of interest to declare. All the authors have seen and approved the manuscript being submitted.

# Appendix A

	Variables	Shanghai							Kunming					
Description						OR <sup>4</sup> 95% CI						OR <sup>4</sup> 9	95% CI	
		β 1	Std. <sup>2</sup>	Sig. <sup>3</sup>	OR	Lower Bound	Upper Bound	β	Std.	Sig.	OR	Lower Bound	Upper Bound	
Wind	W <sub>5</sub> (1) W <sub>6</sub> (1)	-2.900 2.498	0.760 0.552	0.000 0.000	0.055 12.162	0.012 4.119	0.244 35.917							
Temperature	$\begin{array}{c} \Delta(T_{l})_{1} \\ \Delta(T_{l})_{4} \\ (T_{h})_{7} \\ \Delta(T_{h})_{3} \\ \Delta(T_{l})_{10} \\ \Delta(T_{h})_{12} \end{array}$	-1.213 1.999 1.662	0.439 0.455 0.454	0.006 0.000 0.000	0.297 7.378 5.267	0.126 3.022 2.162	0.703 18.013 12.833	-0.906 1.095 -1.983	0.370 0.336 0.541	0.014 0.001 0.000	0.404 2.988 0.138	0.196 1.545 0.048	0.835 5.777 0.397	
AQI	$\begin{array}{c} \Delta(AQI)_5\\ \Delta(AQI)_7\\ \Delta(AQI)_{12}\\ \Delta'(AQI)_4\\ \Delta'(AQI)_{11} \end{array}$	-1.857 1.935 -1.834	0.453 0.445 0.477	0.000 0.000 0.000	0.156 6.925 0.160	0.064 2.893 0.063	0.379 16.578 0.407	0.763 -1.105	0.309 0.346	0.014 0.001	2.144 0.331	1.170 0.168	3.928 0.652	
TSP	$\begin{array}{c} \Delta(PM_{2.5})_5 \\ \Delta(PM_{2.5})_{13} \\ \Delta'(PM_{10})_2 \end{array}$	0.926	0.422	0.028	2.525	1.105	5.770	0.827 0.770	0.316 0.345	0.009 0.026	2.285 2.160	1.230 1.098	4.246 4.248	
SO <sub>2</sub>	$\begin{array}{c} \Delta(SO_2)_0\\ \Delta(SO_2)_{10}\\ \Delta'(SO_2)_7\\ \Delta'(SO_2)_9 \end{array}$	1.305 1.960	0.424 0.416	0.002 0.000	3.686 7.102	1.606 3.142	8.457 16.056	-1.156 0.763	0.372 0.345	0.002	0.315	0.152	0.652	
NO <sub>2</sub>	$\Delta(NO_2)_9 \\ \Delta'(NO_2)_{10}$	-1.419	0.430	0.001	0.242	0.104	0.562	-1.264	0.349	0.000	0.283	0.143	0.560	
O <sub>3</sub>	$\Delta'(O_3)_9 \ \Delta'(O_3)_{12}$	0.913 -1.363	0.425 0.437	0.032 0.002	2.491 0.256	1.083 0.109	5.726 0.603							
Constant	с	-4.222	0.854	0.000	0.015			-3.773	0.579	0.000	0.023			

Table A1. The qualitative model results.

 $^{1}$  The  $\beta$  in this table represents the coefficients of the variables.  $^{2}$  The Std. in this table represents the coefficients' standard deviation value.  $^{3}$  The sig. in this table denotes the *p*-value of the variables.  $^{4}$  The OR in this table denotes the odds ratio value.

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