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Spatial and moderating effects of greenspace on the association between air pollution and lung cancer incidence

4 Abstract

5 Lung cancer remains the primary cause of death globally. Studies have increasingly explored the role of greenspace in mitigating lung cancer risks, yet research gaps persist. 6 7 First, while the direct spatial effect of greenspace has received attention, its potential 8 spillover effects, driven by human mobility and air pollution dispersion, remain 9 underexamined. Second, despite prevalent assertions of greenspace as an air purifier, 10 the extent to which it moderates the air pollution-lung cancer association has yet to be 11 fully understood. Third, the evaluation of greenspace's effects, predominately analyzed 12 linearly a priori, demands exploration into their potential nonlinearity. We utilize three-13 year lung cancer datasets from 228 counties in China, to investigate greenspace's spatial, moderating, and threshold effects on lung cancer incidence in relation to air pollution. 14 15 Employing spatial econometric and threshold models, our findings indicate that 16 greenspace reduces lung cancer incidence in both local and neighboring counties. We 17 also observe a diminution in the detrimental impact of air pollution on lung cancer incidence in areas with higher greenspace, especially when the Normalized Difference 18 19 Vegetation Index surpasses a given threshold (0.38). These insights contribute to an 20 enhanced understanding of greenspace's role in lung cancer prevention and could 21 inform policies on greenspace expansion prioritization.

22

Keywords: greenspace; air pollution; lung cancer; spatial effect; moderating effect

23

24 **1 Introduction**

25 Lung cancer, the leading cause of death worldwide, necessitates an intensified focus on its preventive measures, underscoring its importance as a critical health 26 27 priority. In 2018, it accounted for 1.8 million deaths and 2.1 million new cases globally (WHO., 2020). China, as the most populous nation, contributes over 20% of new global 28 lung cancer cases and approximately 40% of deaths (Cao, Chen, Yu, Li, & Chen, 2021; 29 30 Y. Guo, et al., 2016; S. He, et al., 2020). This significant burden is further compounded 31 by rapid urban development; the country's swift urbanization in recent decades has led to several environmental and lifestyle issues, including air pollution, environmental 32 33 degradation, and decreased physical activity (Chung, et al., 2021; Sun, Bao, Zhao, Tang, 34 & Wang, 2021; L. Wang, et al., 2022; L. Wang, Zhao, Xu, Tang, & Jiang, 2016). As a consequence, projections suggest a rising lung cancer burden in China within the 35 36 coming two decades (Cao, et al., 2021).

37 Lung cancer risk factors include individual behaviors like smoking and alcohol 38 consumption, lifestyle factors such as unhealthy dietary choices and physical inactivity, 39 as well as the influence of socioeconomic status and environmental exposure to air 40 pollution (Barta, Powell, & Wisnivesky, 2019; Chung, et al., 2021; Sun, et al., 2021; L. 41 Wang, et al., 2022; L. Wang, et al., 2016). Notably, among these, ambient air pollutants 42 are recognized as one of the most critical determinants (Y. Guo, et al., 2016). 43 Biologically, these pollutants introduce viruses, bacteria, and harmful gases into the lungs, triggering chronic low-grade inflammation and oxidative stress, thereby 44 heightening lung cancer susceptibility (Loomis, et al., 2013; Turner, et al., 2020). 45 Numerous large-scale epidemiological studies consistently link ambient air pollution 46 47 with increased lung cancer incidence and mortality (Yuming Guo, et al., 2015; Raaschou-Nielsen, et al., 2013; L. Yang, et al., 2020). For example, a comprehensive
12-year cohort study in northern China associated long-term exposure to PM₁₀, SO₂,
and NO₂ with higher lung cancer mortality (Chen, et al., 2016). Another study across
75 Chinese communities found that PM_{2.5} and ozone exposure correlated with increased
lung cancer incidence (Y. Guo, et al., 2016).

53 The established link between ambient air pollution and increased lung cancer risk 54 has prompted investigations into mitigating factors of air pollution, notably greenspaces. 55 These areas, encompassing parks and forests, act as natural filters, efficiently capturing 56 airborne particulate matter (PM), dust, and pollen. Specific tree species, such as the 57 London plane tree, demonstrate a capacity for absorbing nitrogen dioxide (NO₂) 58 (Buccolieri et al. 2018), while broader vegetation effectively neutralizes sulfur dioxide 59 (SO₂) (Sarker et al. 2016). Greenspaces' shading effect also reduces high temperatures, 60 curbing the formation of ground-level ozone (O₃) in warm conditions (Knight et al. 2021). Empirically, the Normalized Difference Vegetation Index (NDVI), a metric 61 62 assessing vegetation density, is widely employed to measure greenspace exposure; a 63 close linkage between increased NDVI and air pollution reduction has been observed in the existing literature (Yu et al. 2021; Thiering et al. 2016). Within this context, the 64 formulation of health-oriented interventions aimed at augmenting residential 65 66 greenspace exposure is gaining recognition in mitigating lung cancer risks and lessening its associated health burdens (Loomis, et al., 2013). 67

Greenspaces, which facilitate physical activities (B. Xie, Lu, Wu, & An, 2021; Yu,
et al., 2023) and offer physical and mental health benefits (Lachowycz & Jones, 2013;
Bo Xie, Lu, & Zheng, 2022), have attracted considerable attention from scholars and
public health officials. In light of their potential as environmental interventions, a

72 growing body of research has explored the relationship between greenspace exposure 73 and lung cancer incidence (Markevych, et al., 2017; L. Wang, et al., 2022; Zare Sakhvidi, et al., 2022). While some suggest a protective role of greenspaces (Huang, et 74 al., 2022; Lei Yang, et al., 2021), others, including a comprehensive meta-analysis 75 76 (Coleman, et al., 2021), along with several case studies (Shao, et al., 2019; Sun, et al., 77 2021; Xu, Ren, Yuan, Nichol, & Goggins, 2017; Zare Sakhvidi, et al., 2021), report no 78 significant impact on lung cancer incidence or mortality. Despite these insights, critical 79 research gaps persist, which may contribute to such inconsistencies and hinder an in-80 depth understanding of this complex relationship.

81 First, research has predominately concentrated on the direct impact of greenspace 82 exposure, examining its correlation with lung cancer incidence within specific spatial areas (L. Wang, et al., 2016; Lei Yang, et al., 2021). However, the potential spillover 83 84 effects of greenspace exposure-its influence beyond immediate geographical boundaries due to human mobility and the diffusion of air pollution-remain largely 85 86 underexplored. This lack of attention to spatial spillover effects hampers the 87 development of a more comprehensive understanding of the overall health implications 88 of greenspaces, potentially leading to biased research outcomes (Elhorst, 2010).

Second, the majority of existing literature has viewed greenspace exposure as an air purifier, suggesting that it may reduce lung cancer risks through lowered levels of air pollution. However, beyond this mediating linkage, there is limited understanding of how greenspace moderates the relationship between air pollution and lung cancer risks. In essence, how air pollution affects lung cancer may vary with the amount of greenspace exposure. For example, studies have suggested that increased greenspace exposure may enhance lung function and immune response, thus bolstering resilience

against air pollutants (Zhang et al. 2023; Sun et al. 2023), which consequently affects 96 97 lung cancer risk. Evidence also suggests that greenspaces serve as communal areas, 98 fostering social interactions and community engagement (Lund 2003). Strong 99 community ties and collective efficacy enhance health information dissemination, 100 shaping individual responses to air pollution, such as increased awareness of its adverse 101 health effects and informed health choices to mitigate exposure like reducing outdoor 102 activities during periods of high pollution (see Ward et al. (2022) for a review). In 103 practice, investigating such moderating effect of greenspace helps inform targeted 104 greenspace planning to mitigate health disparities driven by variations in air pollution 105 exposure.

106 Third, while the existing research often assumed a linear relationship between 107 greenspace and lung cancer risks a priori (E. A. Richardson & Mitchell, 2010; E. A. 108 Richardson, et al., 2012), this assumption lacks comprehensive validation. Empirically, recent studies suggest a more complex, nonlinear association not only between 109 110 greenspace and lung cancer risk factors but also with various other health outcomes. 111 For example, Ai et al. (2023)'s study revealed that the expansion of greenspace 112 (measured by NDVI) only starts to reduce PM2.5 concentrations after exceeding a 113 specific threshold. Once this point is surpassed, the pollution-reducing impact 114 intensifies with additional greenspace, up until it reaches another threshold, where the 115 effects then plateau and remain relatively constant. Similar nonlinearity has also been 116 observed in the relationship between greenspace exposure and levels of physical activity (Klompmaker et al. 2018). Moreover, empirical evidence has revealed non-117 118 linear relationships between greenspace exposure and various other health outcomes, 119 such as general health (Huang et. al 2018), hypertension (Wensu, Wenjuan, Fenfen, Wen, & Li, 2022), and obesity (Ghimire et al. 2017). Therefore, it is reasonable to 120

explore the association between greenspace and lung cancer incidence within a nonlinear analytical framework. Against this backdrop, determining the presence and nature of threshold effects is crucial for implementing greenspace planning that is both effective and cost-efficient in mitigating lung cancer risks.

125 This study aims to examine the comprehensive effects of greenspace exposure on lung cancer incidence, utilizing panel data from 228 counties in China between 2013 126 and 2015. It seeks to examine: (1) the direct, spatial, and overall spillover effects of 127 128 greenspaces on lung cancer incidence; (2) the moderating role of greenspaces in the relationship between air pollution and lung cancer incidence; and (3) the presence of a 129 130 threshold effect in the greenspace-lung cancer incidence association. A conceptual 131 framework was developed to elucidate the interactions between greenspaces and lung cancer incidence (Fig. 1). Our findings could provide in-depth insights into the intricate 132 133 dynamics among greenspace, air pollution, and lung cancer incidence, serving as a 134 foundation for developing effective health-promoting greenspace strategies in 135 developing countries.



Direct effects
 Potential effects

136

137

Fig.1. Conceptual framework.

138 **2 Methods**

139 2.1 Study area

County-level administrative regions (counties or county-equivalent areas), which 140 141 are fundamental urban administrative units in China, served as the study area. 142 Considering the availability of data for all variables, we selected 228 sample counties 143 distributed across 30 municipalities, autonomous regions, and provinces (Fig. 2). On 144 average, new lung cancer cases reported in these selected counties covered 145 approximately 20% of the total new cases diagnosed in China from 2013 to 2015. Such 146 large case sizes in the study area ensured the generalizability of the findings. Moreover, 147 these sample counties consisted of 53% low-income counties and 47% high-income 148 counties with a balanced distribution of economic levels, which avoided biased results 149 caused by economic variation.



150 151

Fig. 2. Locations of the 228 sample counties in China.

152 **2.2 Data collection**

Four types of panel data over 3 years were used in this study: lung cancer incidence, air pollution (PM_{2.5}) exposure, greenspace exposure, and covariates. The summary and definition for all variables are presented in Table 1.

156 **2.2.1 Incidence of lung cancer**

157 The accurate medical information on lung cancer was gathered from the China 158 Cancer Registry Annual Report. Cancer registries in 31 provinces submit cancer 159 registration data to the National Cancer Center Registry of China (NCCR) annually. 160 These data are collected using active methods (registry personnel investigating the 161 sources of data) and passive methods (medical institution notification forms forwarded 162 to the registry, copies of abstracts for studies containing the necessary data). The NCCR 163 ensures the integrity and credibility of the submitted data by reviewing them against 164 the Guidelines for Chinese Cancer Registration and the applicable data quality criteria 165 outlined in the Cancer Incidence in Five Continents Volume published by IARC/IACR. 166 Quality problems were timely feedback to registries, who then revised and re-submitted 167 the data to the NCCR, forming a cancer reporting database. The database included the registry's name and the number of newly diagnosed cancer cases categorized by the 168 169 International Classification of Diseases (ICD-10) code at the county or equivalent 170 geographic unit level. Registries lacking complete lung cancer data for 2013 to 2015 171 were omitted. Consequently, 228 county-level cancer registries were incorporated into 172 the final analysis. From this dataset, we extracted data on the number of newly 173 diagnosed lung cancer cases, as defined by the ICD-10 code for lung cancer (C33-C34), 174 for each county or equivalent geographic unit. The incidence rate of lung cancer in each 175 of these areas was determined by dividing the new cases by the total population.

176 2.2.2 Air pollution exposure assessment

177 Research indicates PM2.5 as a key risk factor for non-communicable diseases, 178 notably lung cancer (Song, et al., 2017). The Global Burden of Disease (GBD) 2017 179 report (GBD Risk Factor Collaborators, 2018) shows that ambient PM_{2.5} air pollution is estimated to account for 14.1% of global lung cancer deaths, second only to tobacco 180 smoking. East Asia bore the heaviest regional burden of PM2.5 on lung cancer, 181 182 contributing to over 50% of the global disability-adjusted life years (DALYs) attributed 183 to PM2.5-induced lung cancer, with China being the most affected country (Yang et al. 184 2022). Therefore, the average annual PM_{2.5} was employed as a surrogate for assessing 185 air pollution exposure in this study. Other research suggests that exposure to other 186 ambient air pollutants such as SO₂, NO₂, and O₃ may also significantly contribute to 187 lung cancer risks (Yang et al. 2016). Therefore, we included SO₂ and NO₂ in our sensitivity analyses as adjustments to our model to ensure the robustness of our findings 188 (see, subsection 2.4.3). PM_{2.5} data were collected from the China High Air Pollutants 189 190 (CHAP) dataset (https://weijing-rs.github.io/product.html). These data are derived and 191 estimated using the Moderate Resolution Imaging Spectroradiometer (MODIS) Multi-192 Angle implementation of Atmospheric Correction (MAIAC) algorithm and Space-Time 193 Extra-Trees (STET) model, which achieved a cross-validation coefficient of 194 determination ranging from 0.80 to 0.92 (Wei, et al., 2020; Wei, et al., 2021). The PM_{2.5} 195 data, with a spatial resolution of 1 km, have been extensively utilized in studies related 196 to air pollution and public health (Feng, et al., 2023; D. He, Lu, Xie, & Helbich, 2022).

197 2.2.3 Greenspace assessment

Greenspace exposure estimates were based on the NDVI. We obtained NDVI from
Terra MODIS of the National Aeronautics and Space Administration (NASA); the

dataset provided comprehensive coverage of China, capturing spatial details at a
resolution of 250 m and temporal variations over 16 days from 2013 to 2015. The NDVI
values vary between -1 and 1, with higher positive values indicating greater vegetation
coverage and negative values corresponding to areas covered by clouds, water, or snow.
According to Kayyal-Tarabeia, Michael, Lensky, Blank, and Agay-Shay (2022), we
calculated the NDVI values for different months of the year to maximize NDVI contrast
and selected August, the greenest month, as the greenspace exposure.

207 2.2.4 Covariates

Four types of covariates were selected in this study: individual behavioral 208 209 covariates, meteorological covariates, socioeconomic covariates, and built environment 210 covariates (Table 1). The individual behavioral covariates used in this study were obtained from The China Health and Retirement Longitudinal Study (CHARLS) 211 212 (http://charls.pku.edu.cn/index.htm), published by the Institute of Social Science 213 Survey of Peking University, Beijing, China. CHARLS is a well-regarded longitudinal 214 survey capturing a broad range of data, from socioeconomic factors to health conditions 215 of individuals aged 45 and above in mainland China (Zhao, Hu, Smith, Strauss, & Yang, 216 2014). Launched in 2011 with biennial or triennial follow-ups, CHARLS has been 217 instrumental in lung cancer research, especially in integrating representative individual 218 health-related behavioral characteristics as covariates, enhancing the validity of 219 research findings (H. Guo, Chang, Wu, & Li, 2019; Huagui Guo, et al., 2021). In our 220 research, we extracted smoking and drinking rates from the CHARLS Wave 3 survey 221 as covariates. This survey, conducted in 2015, encompassed 150 cities across 28 of the 30 province-level administrative units in China, involving approximately 21,000 222 223 individuals (H. Guo, et al., 2019; Liu, Xu, & Yang, 2018). The data on smoking and alcohol consumption were obtained from the health status and function module of the survey. As CHARLS provides data at the prefectural city level, we assigned the same smoking and drinking information to all sample counties within the same prefectural city. We applied the provincial information to the counties for those sample prefecturelevel cities not covered by the target prefectural cities in CHARLS.

While CHARLS focused on individuals aged 45 and above, we argue that this may not significantly undermine the robustness of our findings. The reason is that lung cancer incidence in China predominantly increases after age 40 (S. Liu, et al., 2018), with individuals between 50-69 and over 70 accounting for over 80% of cases (Long, et al., 2023). This age distribution aligns closely with the CHARLS cohort, making it an ideal source for extracting representative covariates like smoking and drinking rates from its third wave in 2015.

Rainfall, temperature, and sunshine duration were included as meteorological
covariates in this study. Data were obtained from the National Meteorological Data
Center (<u>http://data.cma.cn/site</u>).

239 Socioeconomic covariates such as urbanization, gross domestic product (GDP), proportion of industrial production, and medical supply beds were included as 240 241 covariates. These variables were selected to control the impact factors of lung cancer 242 based on the urban development level, economic status, and medical conditions. Drawing on prior epidemiological studies (Ge, et al., 2021), we employed annual 243 244 average nighttime light as a proxy to assess urbanization. These data were obtained Visible 245 from the Infrared Imaging Radiometer Suite (https://ladsweb.modaps.eosdis.nasa.gov/), with higher values indicating higher levels 246 of urbanization. Data on the GDP, proportion of industrial production, and medical 247

supply beds were extracted from the China City Statistical Yearbook.

Built environments have significant effects on air pollution (Ferm & Sjoberg, 2015; Pant & Harrison, 2013) and can potentially affect the incidence of chronic noncommunicable diseases (Bo Xie, Jiao, An, Zheng, & Li, 2019). We selected population density and road density as built environment covariates; data were obtained from the China Statistical Yearbook at the county level.

254 2.3 Examination of spatial dependency

To avoid endogeneity bias in area-level estimations, we employed the Global Moran's I statistic to assess the spatial autocorrelation of lung cancer incidence within counties across the study area. The Global Moran's I value were 0.587, 0.636, and 0.663 in 2013, 2014, and 2015, respectively, illustrating that the incidence of lung cancer in a county presents a significant and strong spatial autocorrelation with that in adjacent counties. 261 Table 1

262 Summary statistics for all variables (N=228).

Variable	Description	Unit		Min	Ma	Mean	SD
				:	x		
The incidence of lun	ng Newly diagnosed lung cancer cases	New cases	per	0.715	23.1	5.137	2.13
cancer	/the total population in each county	100,000 person			14	7	
Air pollution	Annual average PM _{2.5} concentration	$\mu g/m^3$		16.42	112.	69.10	19.6
			4	(094 2	72	2
Greenspace	Average NDVI value in August			0.096	0.73	0.444	0.12
					4	2	

Individual factors							
Smoking	Smokers/total number of population in each county	%	0	25.80 3.	58.3 3 5	39.58 3	0.05
Alcohol drinking	The number of people who drink more than once a month/ total population in each county	%		4.918 93	56.4 3 6	35.58	0.07
Meteorological factors							
Rainfall	Annual average rainfall	mm	4	712.60 92	3422. 26 7'	1989. 72 3	513. 397
Temperature	Annual average temperature	°C		1.910	32.67	16.89	4.50

				5	0	9	
Sunshine duration	Annual average sunshine hours	hour	1.140	12	.65	5.761	1.98
				2		4	
Social-economic factors							
Urbanization	Annual average nighttime light		0.041	73	.12	11.23	14.5
i	ntensity			1	6	44	
GDP	Per capita GDP per year	10 thousand yuan	0.808	46	.77	5.652	3.97
	pe	er person		5		8	
Proportion of industrial	The proportion of industrial output in	%	0.061	5.:	595	1.662	0.81
oduction t	otal output per year					0	

Medical supply beds	Medical bed number per 1,000 person	counts per 1,000	11.897	241.6	49.91 26.2
	р	person	62	1	29
Built environment factors					
Population density	Population/total area of the counties	Person per km ²	79.000 .000	13971) 722	3710. 2616 .363
Road density	Road length/total area of the counties	km/km ²	0.002	3.902	0.346 0.53 2

263 Note: Min = minimum; Max = maximum; SD = standard deviation

264 2.4 Statistical analysis

265 2.4.1 Spatial econometric models for panel data

The existence of spatial dependence and autocorrelation violates the hypothesis that variables are independent of each other (LeSage & Pace, 2009). Therefore, we constructed spatial econometric models for the panel data to correct the biased and inconsistent estimations caused by the spatial spillover effect. A general spatial nesting model was constructed as follows:

271
$$Y_{it} = \rho W Y_{it} + \alpha I_n + \beta X_{it} + u, u = \lambda W_{\mu} + \varepsilon$$
(1)

272 where, for county *i* in year *t*, Y_{it} is the n \times 1 vector of lung cancer incidence; 273 X_{it} is the explanatory variable; β is the corresponding coefficients; *W* is an n \times n 274 spatial weight matrix; I_n is an N \times 1 vector which is associated with the constant term 275 parameter α ; ρ denotes the spatial autoregressive coefficient; λ denotes the spatial 276 autocorrelation coefficient; ε denotes a vector of disturbance terms; and W_{μ} denotes 277 the interaction effects arising from the disturbance terms across different spatial units.

Based on Eq. (1), when $\lambda = 0$, the model was transformed into a spatial Durbin model (SDM) containing the spatial lag terms of the dependent and independent variables:

281
$$Y_{it} = \rho W Y_{it} + \alpha I_n + \beta X_{it} + \theta W X_{it} + \varepsilon$$
(2)

Based on Eq. (2), when $\theta = 0$, the SDM could be simplified to a spatial lag model (SLM) containing the spatial lag terms of the dependent variable. If $\theta = -\rho\beta$, then $\lambda = \rho$, and we obtained the spatial error model (SEM), which only contained the spatial lag terms of the error term. To determine which model is more appropriate for describing data, the Lagrange multiplier (LM) test, robust Lagrange multiplier (robust
LM) test, and likelihood ratio (LR) test were used (for specific model selection methods,
see Elhorst (2012)). The Hausman test examined fixed effects when individual or time
effects were correlated with regressors (Lee & Yu, 2012).

290 The advantages of employing spatial econometric models are as follows. First, the 291 spatially lagged terms WY_{it} and WX_{it} can help explicitly reduce the endogeneity 292 bias caused by spatial dependence and spatial autocorrelation. Two main approaches 293 are commonly employed when constructing the spatial weight matrix: neighboring and 294 distance-based. As there were distances between some sample counties, the diffusion of air pollutants was not confined by administrative boundaries. Therefore, to account 295 296 for possible bias, the reciprocal of the Euclidean distance among counties was used as 297 an element in our distance weight matrix, according to the equation:

298
$$W = \begin{cases} \frac{1}{d} & i \neq j \\ 0 & i = j \end{cases}$$
(3)

where *d* represents the distance of the geometric center between county *i* and county *j*.

301 Second, spatial econometric models can be employed to calculate the spatial 302 spillover effects of greenspaces on lung cancer incidence. LeSage and Pace (2009) defined every average diagonal element of the WX_{it} matrix of WX_{it} as a direct effect. 303 304 This term refers to the influence of changes in the independent variable on the 305 dependent variable of local areas. Moreover, every non-diagonal average element is 306 defined as the spillover effect, which is interpreted as the impact of the independent variable on the dependent variable of neighboring areas. Finally, the total effect includes 307 both the average direct and spillover effects. 308

309 2.4.2 Panel threshold model

To explore the threshold effect of greenspace on lung cancer incidence, the panel threshold model, which can automatically identify the endogenous features of data, was employed in this study (Hansen, 1999).

313 The threshold model is expressed by Eq. (4) and Eq. (5) as follows:

314
$$Y_{it} = \mu_i + \beta_1 X_{it} + \varepsilon_{it}, \quad q_{it} \le \gamma$$
(4)

315
$$Y_{it} = \mu_i + \beta_2 X_{it} + \varepsilon_{it}, \quad q_{it} > \gamma$$

316 (5)

For county *i* in year *t*, Y_{it} is lung cancer incidence, X_{it} is the independent variable, and q_{it} denotes the threshold variable; μ_i represents the individual effect. Depending on the threshold γ , the observations are categorized into two stages. Each stage represents a distinct regime with its regression slope, either β_1 or β_2 .

321 2.4.3 Sensitivity analysis

322 To enhance the robustness of our findings, we performed two types of sensitivity 323 analyses. First, we employed an alternative weight matrix structure, specifically the 324 queen contiguity matrix, to ascertain the stability of our model given its distinct weight 325 matrix configuration. Second, we recalibrated the primary model, incorporating adjustments for SO2 and NO2 concentrations, to explore the potential confounding 326 327 effects of these gaseous pollutants on lung cancer incidence. The data for these pollutants, with a 10km resolution, were sourced from the Comprehensive Air-quality 328 Prediction (CHAP) dataset (https://weijing-rs.github.io/product.html). 329

330 3 Results

331 **3.1 Descriptive analysis**

According to the data presented in Table 2, there was no significant variation in the incidence of lung cancer from 2013 to 2015, whereas the variations in PM_{2.5} and NDVI differed significantly. The mean PM_{2.5} concentration significantly decreased by $36 \ \mu g/m^3$ in absolute terms and 41.2% in percentage terms. The mean NDVI value in 2015 were slightly higher than those in 2013 (0.46 vs. 0.42).

In terms of spatial distribution, a higher incidence of lung cancer was observed in counties in the eastern region (Fig. 3). Moreover, the concentration of $PM_{2.5}$ was found to be higher in the eastern region when compared to both the central and western regions within the study area (Fig. 4). Fig. 5 shows that the distribution of the NDVI differed significantly among the three regions in China, with the central region holding the highest level, followed by the eastern and western regions.

343 Table 2

344 The temporal variations of lung cancer incidence, PM_{2.5} concentration, and the345 NDVI.

Inc cancer	vidence	of lung	PM ₂	.5		ND	VI	
2	2	2	20	20	201	20	20	20
014-	015-	015-	14-	15-	201 5 2014	14-	15-	15-
2013	2013	2014	2013	2013	5-2014	2013	2013	2014

-									
Z	-	-		-	-	-	-	-	-
value	1.074 ^b	1.489 ^b	601 ^b	11.887°	13.083°	13.067°	9.012 ^b	10.101 ^b	2.194 ^b
Р	0.	0.	0.	0.0	0.0	0.0	0.	0.0	0.0
value	283	136	548	00	00	00	000	00	28

346 Note: The Wilcoxon signed-rank test was used to examine the temporal variations of variables in

each county from 2013 to 2015. b was based on negative ranks. c was based on positive ranks.

348





349

350



a. 2013

351

c. 2015



b. 2014





353



356

Fig. 4. Spatial distribution of PM_{2.5} concentrations in 2013-2015.





359a. 2013b. 2014c. 2015

Fig. 5. Spatial distribution of the NDVI in 2013-2015.

361 **3.2 Results of basic models**

Table 3 presents the estimated results of greenspaces' effects on lung cancer incidence. The statistical values of the LM and LR tests indicated that the SDM was deemed more suitable compared to the SLM and SEM in the present study. Furthermore, according to the Hausman test, fixed effects were more suitable for our research. Therefore, all models employed in our study incorporate the time-fixed effect. The results showed that the direct effect of the NDVI on the incidence of lung cancer was significantly negative at the 5% significance level. A one-unit increase in NDVI caused a 2.793% decrease in the incidence of lung cancer. Furthermore, we found a negative spatial spillover effect of NDVI on the incidence of lung cancer. Every one-unit increase in NDVI decreased the incidence of lung cancer in the surrounding counties by 0.593%.

Regarding covariates, the incidence of lung cancer was significantly influenced by higher rates of smoking and alcohol consumption in counties. Moreover, rainfall and the proportion of industrial production had direct and spatial spillover effects on lung cancer incidence. We also found that road density was negatively related to lung cancer incidence in both local and nearby counties.

Table 3

Dependent	SDM				
Variable: Incidence of	Main	Wx	Direct	Indirect	Total
lung cancer	Walli	WA	effect	effect	effect
NDVI	-	-	-	-0.593*	-
	2.805**	3.350*	2.793**	(0.3191)	3.386**
	(1.091	(1.893	(1.133		(1.40
	8)	0)	1)		02)

Smoking		7.730*		5.317*		7.748*	1.631***		9.379
	**		*		**		(0.5937)	***	
		(1.515		(2.747		(1.446			(1.82
	5)		2)		0)			03)	
Alcohol drinking		5.573*		6.899*		5.738*	1.218**		6.956
	**		**		**		(0.4759)	***	
		(1.079		(2.119		(1.156			(1.49
	5)		4)		8)			08)	
Rainfall		0.455*		0.711*		0.449*	0.096*		0.545
		(0.239		(0.402	*		(0.0636)	**	
	1)		4)			(0.241			(0.29
					0)			75)	
Sunshine duration		0 390		1 263		0.403	0.087		0 490
Sunshine duration		0.390		1.205		0.403	0.087		0.490
		(0.388		(0.623		(0.385	(0.0921)		(0.47
	1)		7)		3)			17)	
Temperature		-0 397		-0 464		-0 390	-0.080		_
Temperature		0.377		0.101		0.370	0.000	0.4'	70
		(0.305		(0.431		(0.298	(0.0673)	0.4	10
	0)		0)		7)				(0.35
								98)	

GDP		0.003		0.008		0.004	0.001		0.005
		(0.024		(0.045		(0.026	(0.0059)		(0.03
	2)		3)		1)			18)	
Urbanization		-0.001		-0.018		-0.001	-0.001		-
		(0.006		(0.011		(0.005	(0.0013)	0.0	02
	0)		2)		7)				(0.00
								69)	
Proportion of		0.475*		0.049*		0.485*	0.103**		0.588
industrial production	**			(0.188	**		(0.0400)	***	:
		(0.100	7)			(0.100			(0.12
	6)				2)			78)	
Medical supply-		-0.001		-0.001		-0.001	-0.001		-
beds		(0.000		(0.001		(0.000	(0.0001)	0.0	02
	5)		5)		5)				(0.00
								06)	
Population density		0.200		0.747		0.200	0.043		0.243
		(0.399		(0.726		(0.406	(0.0929)		(0.49
	5)		6)		0)			45)	
Road density		-		-0.396		-	-0.074*		-

		0.3	55**		(0.285	0.3	50**	(0.0452)	0.424**
			(0.170	9)			(0.179		(0.21
		2)				9)			92)
	ρ		0.195*						
		**							
	σ		3.805*						
		**							
	R ²		0.128						
	Ν		228						
	SEM-LM		0.009*						
	SEM-Robust LM		2.158*						
	LR test (SDM &		20.790						
SEI	M)	**							
	SLM-LM		2.191*						
		**							
	SLM-Robust LM		4.340						
	LR test (SDM &		21.330						

SLM)

Hausman test 40.010

*

Note: *p <0.1; **p <0.05; ***p <0.01. In parentheses denotes the standard error. "Main" refers to the non-spatial regression coefficient; "Wx" refers to the spatial regression coefficient. "Direct effect" represents the impacts of variables on the incidence of lung cancer in local areas; "Indirect effect" represents the impacts of variables on the incidence of lung cancer in nearby areas. The spatial correlation coefficient of the dependent variable is denoted as ρ , and the standard error is defined as σ .

387 **3.3 Moderating effect of greenspace**

388 To examine the moderating effect of greenspace on the relationship between PM_{2.5} 389 and lung cancer incidence, we added PM_{2.5} and the interaction term (PM_{2.5} \times NDVI) 390 into the basic model. Table 4 shows that the negative direct and spatial spillover effects of NDVI remained significant after controlling for covariates. In addition, there was a 391 392 positive direct and spillover effect of PM_{2.5} on the incidence of lung cancer. Furthermore, we observed that the interaction term of PM2.5 and NDVI had a significant 393 394 negative direct effect on the incidence of lung cancer. That is, a one-unit increase in the 395 NDVI is associated with a 0.050% decrease in the impact of PM_{2.5} on the incidence of lung cancer. However, no spillover effect of the interaction term was observed in this 396 397 model. Our sensitivity analyses revealed that after recalibrating the model with the 398 queen contiguity matrix and incorporating adjustments for SO₂ and NO₂, the results

demonstrated substantial consistency (Table S1 in Supplementary Material).

400 Table 4

401 Tests for the moderating effect of greenspace.

Dependent	SDM				
Variable: Incidence	;				
of lung cancer	Main	Main Wx Direct effect		Indirect	Total
				effect	effect
NDVI	_	-	_	-0.646*	_
	3.056***	3.587*	3.047**	(0.3266)	3.693**
	(1.090	(1.929	(1.132		(1.40
	8)	9)	2)		21)
PM _{2.5}	0.020*	0.002*	0.020*	0.004**	0.024
	**	*	**	(0.0019)	***
	(0.006	(0.010	(0.006		(0.00
	8)	5)	6)		81)
PM _{2.5} ×NDVI	-	-0.034	-	-0.010	-
	0.053*	(0.067	0.050*	(0.0077)	0.060*
	(0.034	8)	(0.033		(0.04
	4)		2)		01)

	Covariates		\checkmark
	ρ	**	0.194*
	σ	**	3.755*
	R ²		0.169
	Ν		228
	SEM-LM		0.035*
	SEM-Robust LM		2.838*
SEN	LR test (SDM & M)	*	21.180
	SLM-LM		3.215*
	SLM-Robust LM	*	6.018*
SLN	LR test (SDM & M)	*	21.360

Hausman test 31.940

402 Note: *p < 0.1; **p < 0.05; ***p < 0.01. In parentheses denotes the standard error.

403 **3.4 Threshold of moderating effects**

We developed two models to examine whether there was a significant threshold effect on the incidence of lung cancer. Model 1 in Table 5 shows that the total effect of the NDVI did not exhibit a significant threshold. However, the single and double thresholds in Model 2 passed the 5% significance test, suggesting significant double thresholds for NDVI's moderating effect.

409 The NDVI moderated the relationship between PM_{2.5} and the incidence of lung 410 cancer in the three stages (Table 6). When the NDVI value was less than 0.38, the interaction term did not significantly impact lung cancer incidence. When the NDVI 411 412 was between threshold values of 0.38 and 0.4, the interaction term was negatively 413 associated with the incidence of lung cancer. Specifically, with a one-unit increase in NDVI, the positive effect of $PM_{2.5}$ on lung cancer incidence decreased by 0.139%. 414 415 Finally, when the NDVI was higher than a threshold value of 0.4, the negative effect of 416 the interaction term on lung cancer incidence remained significant. However, the extent of the moderating effect decreased significantly compared with that in the second stage. 417

418 Table 5

419 Tests for the threshold effect of the NDVI.

d	Threshol	F statistic	P value		10%		5%		1%	
	Threshold test of the total effect (model 1)									
	Single	14.870	0.15 0	1	16.52	9	18.55	5	24.45	
	Double	6.770	0.58 6	6	14.63	0	16.16	7	19.96	
	Triple	4.820	0.83	5	16.10	7	19.74	6	25.13	
	Threshold to	est of the mode	rating effec	et (moo	lel 2)					
	Single	13.140*	0.07		11.730	8	15.01	4	19.04	
	Double	17.540* *	0.02 6	7	12.27	9	15.58	7	19.72	
	Triple	14.100	0.25	7	20.78	8	29.95	0	32.98	

420 Note: *p <0.1; **p <0.05; ***p <0.01.

421 Table 6

Dependent Variable: Incidence of lung cancer	Threshold	Coefficient	T statistic
	NDVI≤0.38	0.002	0.260
PM _{2.5} ×NDVI	0.38 < NDVI ≤0.4	-0.139***	- 4.080
	NDVI >0.4	-0.013***	- 2.820

The threshold value and parameter estimation of the moderating effects of 422 greenspace. 423

Note: *p <0.1; **p <0.05; ***p <0.01. 424

4 Discussion 425

4.1 Associations between greenspace, PM_{2.5}, and lung cancer 426

427 This study found that greenspace exposure is a protective factor against lung cancer. To date, only a few studies have reported the preventive effects of greenspace on lung 428 429 cancer. A cohort study from Taiwan, China, revealed a significant relationship between an increment of 0.1 units in NDVI and a hazard ratio (HR) of 0.95 in lung cancer risks 430 (Huang, et al., 2022). Another cohort study conducted in Tel Aviv, Israel, over a 21-year 431

follow-up period, involving 144,427 participants, revealed a beneficial correlation
between higher residential greenness and lower incidence of lung cancer (KayyalTarabeia, et al., 2022). We obtained consistent results at the national scale, as noted in
previous studies. This finding provides evidence that greenspace intervention is a useful
avenue for reducing the incidence of lung cancer.

437 The effects of greenspace on health outcomes in previous ecological studies are 438 often simplified to a direct one-to-one relationship at the area level, which might report 439 biased findings. For example, a comprehensive ecological study from the United 440 Kingdom found no relationship between greenspace and lung cancer mortality (E. A. Richardson & Mitchell, 2010). Similarly, an urban study conducted in New Zealand 441 442 failed to provide evidence that greenspace influences lung cancer mortality (E. 443 Richardson, Pearce, Mitchell, Day, & Kingham, 2010). Although these studies observed 444 consistent results at the local level across different countries, they do not fully capture the comprehensive health impact of greenspace, especially when considering the 445 446 presence of spillover effects that align with the direction of the direct effects (Benjamin-447 Chung, et al., 2018). In our study, we go beyond previous studies that typically consider 448 the direct effects of greenspace on lung cancer and adopt spatial econometric models to 449 investigate the presence and extent of spatial spillover effects. The results from the 450 SDM showed that the total protective effect of greenspace involved both direct and 451 indirect effects (i.e., spatial spillover effect). These findings reveal that exposure to greenspaces can mitigate the risk of lung cancer not only in local counties but also in 452 453 neighboring ones. Remarkably, the protective effects of greenspace from nearby 454 counties account for 20% of the benefits derived from local greenspace. Increasing 455 greenspace not only provides health benefits to residents who lived in local counties 456 but also extends its impact to individuals who do not directly receive the greenspace

457 intervention. Our findings highlight the importance of synergistically scaling up or458 subsidizing greenspace interventions nationally.

459 Two potential reasons have been proposed in this study to explain the spatial effects of greenspace on lung cancer incidence. A plausible explanation is that greenspaces in 460 461 local counties may deposit and filter $PM_{2.5}$ diffused from adjacent counties by chemical, 462 biological, and physical effects (Markevych, et al., 2017; Zare Sakhvidi, et al., 2022), thereby decreasing the harmful effects of PM_{2.5}. Moreover, most existing studies on 463 464 cancer incidence failed to consider the greenspace exposure changes caused by human 465 mobility across the geographical boundary (Gailey, McElroy, Benmarhnia, & Bruckner, 466 2021; Namin, Zhou, Neuner, & Beyer, 2021). In China, straddle and circle mobility 467 among counties (such as inter-city commuting and hukou-based migrant) is widespread. This large-scale individual mobility involving approximately 18% of the country's total 468 469 population may lead to continuous variations in residents' exposure to greenspace (Mai & Wang, 2022; Namin, et al., 2021). Under this scenario, the health benefits derived 470 471 from greenspace may not solely originate from the local area but should encompass the 472 cumulative effect of local and nearby counties. Our findings on spatial spillover effects 473 emphasize the need for future studies to estimate greenspace exposure using the mobility-based approach in similar countries with high internal mobility levels to 474 475 provide more comprehensive insights into the health effects of greenspace.

476 Second, recent epidemiological studies have shown that greenspace may moderate 477 the effect of $PM_{2.5}$ on health outcomes. For example, a nationwide modeling study from 478 China suggested that areas with higher levels of greenspace exhibited stronger 479 protective effects against tuberculosis in the presence of $PM_{2.5}$ compared to areas with 480 lower levels of greenspace (Zhu, et al., 2022). Similarly, an ecological study conducted in Greece reported that areas with greater greenness had lower PM_{2.5} effects on
cardiovascular mortality (Kasdagli, Katsouyanni, de Hoogh, Lagiou, & Samoli, 2021).
Our estimated results suggest that greenspace can moderate the relationship between
PM_{2.5} and lung cancer incidence, which is in accordance with previous studies
(Coleman, et al., 2021; Kasdagli, et al., 2021). In other words, the positive effects of
PM_{2.5} on the incidence of lung cancer decreased in greener counties.

487 However, a moderating greenspace effect was observed only after the NDVI 488 exceeded a certain threshold. According to Table 6, when the NDVI exceeded 0.38, 489 greenspace was conducive to reducing the positive effect of PM_{2.5} on the incidence of lung cancer. Furthermore, when the NDVI increased to 0.4, the strength of the 490 491 moderating effect decreased. Recent studies conducted in developed countries have observed similar nonlinear effect trends. For example, evidence from a national cohort 492 493 of Canadian adults showed no significant relationship between PM2.5 and non-494 accidental and cardiovascular mortality in the two greenest quintiles (Crouse, et al., 495 2019). Research comprising 5.5 million cancer patients and survivors across 14 states 496 and metropolitan regions in the United States has found that an increase in county-level 497 PM_{2.5} is associated with a heightened risk of cardiopulmonary mortality in regions with low levels of NDVI. In contrast, areas with high levels of greenness were almost 498 499 immune to any variations in PM_{2.5} levels and did not pose any relative risk (Coleman, 500 et al., 2021). Our findings differed slightly from those of previous studies. Results in our study emphasize the existence of a certain threshold value that may trigger the 501 502 potential emergence of the moderating effect of greenspace. This finding implies that 503 compared to developed countries, local governments in developing countries need to 504 prioritize policy interventions focusing on expanding greenspaces to generate positive 505 health effects even in counties with high air pollution.

506 Third, comprehensively controlled for individual, meteorological, we 507 socioeconomic, and built environment factors, thus avoiding potential biases and giving 508 more realistic results. Smoking and alcohol consumption were seldom incorporated in 509 previous ecological studies, although they are crucial contributors to respiratory diseases like lung cancer (Lin, Murray, Cohen, Colijn, & Ezzati, 2008). In our research, 510 511 these two factors based on data derived from CHARLS were introduced to the models. 512 The results showed that they were positively correlated with the incidence of lung 513 cancer, which is consistent with the results of previous studies (WHO., 2020). 514 Regarding meteorological factors, we found that rainfall was positively related to the 515 incidence of lung cancer, perhaps owing to condensation nuclei in microdroplets of 516 rainfall water that carried air pollutants and damaged human lungs (Clauss, Mayes, 517 Hilton, & Lawrenson, 2005; Javorac, et al., 2021). Furthermore, our findings indicate a 518 positive correlation between the proportion of industrial production and the incidence 519 of lung cancer, which is consistent with previous studies (López-Cima, et al., 2013; 520 Lopez-Cima, et al., 2011). For built environment factors, the results suggest a negative correlation between road density and the incidence of lung cancer, which is inconsistent 521 522 with the findings of other studies (Bechle, Millet, & Marshall, 2011; Chawinska, 523 Tukiendorf, & Miszczyk, 2014; Sun, et al., 2021). A plausible explanation is that high 524 road density in Chinese counties may decrease per capita emissions by discouraging 525 car usage and promoting walking as a mode of transportation (Brownstone & Golob, 2009), thereby reducing the risk of lung cancer. 526

527 4.2 Policy implications

528 This study has several policy implications that can facilitate greenspace 529 construction and lung cancer prevention in China and other developing countries. First, urban planners should note the strong spillover effects of greenspace, which can reap the health benefits that transcend the local population. Consequently, there is a pressing need to prioritize and allocate resources to support the expansion of greenspace at a national level, aiming to optimize health outcomes for a broader population.

534 Second, we found a moderating effect of greenspaces on the association between 535 PM_{2.5} and lung cancer incidence, providing new evidence and insights for central and local governments to enhance the understanding of greenspace-related health benefits. 536 537 Although the government has taken proactive measures to control air pollution, PM_{2.5} 538 will likely continue to contribute significantly to the high burden of lung cancer. 539 Therefore, policymakers and urban planners should pay attention to the mitigating 540 function of greenspaces in lung cancer associated with air pollution. Specifically, 541 policymakers can use NDVI as a crucial quantitative indicator and incorporate it into 542 lung cancer resilience strategies and urban planning policies. The critical greenspace 543 value, where the moderating effect was significant, could be considered the basic 544 requirement for regional greenspace construction.

545 Third, to improve our understanding of the threshold value of the moderating effect 546 of greenspace, we calculated the proportion of counties in which the NDVI value was 547 less than 0.38, even in the greenest season, and presented their spatial distribution (see 548 Fig. S1 and Table S2). In over 20% of the counties, NDVI was found to have no 549 moderating effect on the relationship between PM_{2.5} and lung cancer incidence. 550 Therefore, the policymakers should provide more resources (e.g., financial investment 551 to expand greenspace and mitigate air pollution) for these counties to reduce lung 552 cancer risks at both the local and national levels.

553 4.3 Strengths and limitations

554 This study makes three contributions to the existing literature. First, this study 555 considered greenspaces' direct and spatial spillover effects simultaneously, avoiding the 556 estimated bias caused by ignoring spatial autocorrelation in previous studies. More importantly, our study provides insights for policymakers to develop broader regional 557 policies to improve greenspaces. Second, we explored the moderating impact of 558 559 greenspace on the link between PM_{2.5} and lung cancer incidence after controlling for 560 drinking and alcohol drinking factors, extending our knowledge of how residing in 561 greener areas may contribute to better health outcomes. In most developing countries, 562 people are simultaneously exposed to greenspaces and high levels of air pollution. 563 Given this widespread issue, our findings can help local governments in China and 564 similar developing countries realize that greenspace intervention effectively prevents 565 health problems caused by high air pollution concentrations. Third, this study examined 566 the nonlinear trend in the health effects of greenspaces, which could guide the implementation of greenspace interventions to improve health outcomes. 567

This study is subject to several limitations. First, due to data constraints and privacy 568 569 laws in China, we did not include detailed sociodemographic details such as age and 570 sex for individuals with lung cancer. Second, while our reliance on smoking and alcohol 571 consumption rates from the CHARLS datasets, targeting those aged 45 and above, aligns with the age bracket most affected by lung cancer in China, it may not fully 572 573 represent younger demographics, potentially affecting the generalizability of our findings. Also, our method of assigning uniform health-related behavior information to 574 575 counties within the same prefectural cities or provinces might not account for local behavioral differences. Third, although we investigated the relationship between PM_{2.5}, 576

577 greenspace exposure, and lung cancer, and enhanced our findings by adjusting for SO₂ 578 and NO₂, our approach might not capture all air pollutant factors influencing lung 579 cancer. Addressing these data limitations in future studies would be beneficial. Fourth, 580 our study did not account for the cumulative effects of greenspace and PM_{2.5} exposure 581 on lung cancer incidence, potentially introducing uncertainty regarding threshold levels. 582 It is advisable for future research to re-examine our findings using longitudinal data 583 over an extended period, which would better facilitate the modeling of cumulative 584 exposure to greenspace and $PM_{2.5}$.

585

5 Conclusions

586 This study provides robust evidence to explore the spatial effect of greenspaces on 587 the incidence of lung cancer in developing countries, using panel data from 2013 to 588 2015. The results of spatial econometric models showed that greenspace has a negative 589 direct effect and a spatial spillover effect on the incidence of lung cancer. Moreover, a negative moderating effect of greenspace on the relationship between PM_{2.5} and the 590 591 incidence of lung cancer was identified. Furthermore, the moderating effect of 592 greenspaces became significant when it was greater than 0.38. This study suggests that 593 investment in greenspaces can be an effective intervention strategy for reducing the 594 incidence of lung cancer in China.

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598 References

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