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# Greenness and chronic respiratory health issues: a systematic review and meta-analysis

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**Introduction:** The number of chronic respiratory disease (CRD) individuals worldwide has been continuously increasing. Numerous studies have shown that greenness can improve chronic respiratory health issues through different mechanisms, with inconsistent evidence. By quantitatively summarizing existing studies, our purpose is to determine the connection between greenness exposure and various chronic respiratory health.

**Methods:** We conducted a comprehensive search on PubMed, EMBASE, and Web of Science core databases to identify relevant studies on the correlation between greenness exposure and chronic respiratory health issues. Studies published up to January 2023 were included in the search. The study used the most frequent indicator (normalized difference vegetation index [NDVI]) as the definition of greenness exposure.

**Results:** We finally identified 35 studies for meta-analysis. We calculated pooled effects across studies using a random-effects model and conducted a subgroup analysis by age and buffer zones to discuss the effects on chronic respiratory health issues. This study showed that 0.1 increments in NDVI were significantly related to lower rates of asthma incidence, lung cancer incidence, and chronic obstructive pulmonary disease (COPD) mortality risk; the pooled RRs were 0.92 (95% CI: 0.85–0.98), 0.62 (95% CI: 0.40–0.95), and 0.95 (95% CI: 0.92– 0.99), respectively. For the age subgroup, the higher greenness exposure level was related to the incidence rate of asthma among teenagers aged 13–18years (RR: 0.91; 95% CI: 0.83–0.99). For the buffer subgroup, a positive relationship with greenness exposure and asthma incidence/prevalence at 200–300m and 800–1000m buffers, as well as the COPD mortality at 800–1000m buffer, the pooled RRs were 0.92 (95% CI: 0.86–0.98), 0.87 (95% CI: 0.81–0.93), and 0.93 (95% CI: 0.88–0.98), respectively. Evidence of publication bias was not detected in this study.

**Discussion:** Our study is the first global meta-analysis between greenness and various CRDs to report an inverse association. Further research is needed in order to determine the effect of greenness exposure on different CRDs. Therefore, when planning for green development, more consideration must be given to public health and green management as intervention measures.

https://www.crd.york.ac.uk/PROSPEROFILES/384029\_STRATEGY\_20230116.pdf

KEYWORDS

greenness, chronic respiratory disease, normalized difference vegetation index, meta-analysis, systematic review

## 1 Introduction

Chronic respiratory diseases (CRDs) have already become a non-negligible cause of death globally. The disability-adjusted life years (DALYs) counts of CRDs significantly increased from 1990 to 2017. Over the past 30 years, 544.9 million individuals worldwide have been diagnosed with CRDs (1). Among the most common CRDs are allergic rhinitis (AR), chronic obstructive pulmonary disease (COPD), lung cancer, asthma, non-cystic fibrosis bronchiectasis (NCFBE), cystic fibrosis (*CF*), and idiopathic pulmonary fibrosis (IPF) (2–4), which were the main causes of CRD-related incidence and mortality worldwide (5). CRD incidence is growing globally, posing a serious threat to human health and causing heavy financial burdens (6).

CRDs are caused by a variety of factors, including indoor and outdoor environmental exposure, interplays between genetic predispositions, and sociodemographic features throughout the lifespan (7). A series of environmental risk factors related to chronic respiratory diseases have been investigated, including outdoor air pollutants (8), urban heat islands (UHI) (9, 10), biodiversity loss (11), noise pollution (12, 13) indoor mold and humidity (14), tobacco smoke (15), renovation activities (16) and household pets (17). As urbanization continues to increase worldwide, more and more people are exposed to urban-related environmental hazards (18). Several studies have suggested that greenness can benefit chronic respiratory health issues through the purification of air, the alleviation of the heat island effect, the increase of microbial diversity, and the encouragement of physical activity (19–24).

A large number of studies have confirmed that greenness has positive effects on the prevention and treatment of CRD incidence and mortality, including asthma (25–27), AR (28), COPD (29, 30), and lung cancer (31, 32). A number of reviews and meta-analyses have been conducted to summarize the results of existing studies. Seven meta-analyses explored the correlation between greenness and chronic respiratory health outcomes, including asthma (33–36), AR (33–37), respiratory symptoms, diseases, allergies (38), and cancer incidence/prevalence/mortality (39). The relationship of greenness with asthma and AR was the most frequently investigated. However, the findings of the abovementioned reviews are inconsistent. Qiu et al. (34) reported that asthma and AR were significantly prevented by high-level residential greenness, while Parmes et al. (38) found that the odds of asthma and AR increased significantly by 5.9–13.0% for every 10% increase in greenness coverage. It has been recognized in recent years that research evidence regarding the influence of greenness exposure on health outcomes in CRDs has been growing; however, systematic reviews and meta-analyses concentrated only on one or two particular health outcomes in CRDs. Furthermore, no generalized systematic reviews and meta-analyses have been conducted to summarize studies on various CRDs, and no precise and global estimate has been made on the reduction in CRD incidence/prevalence and mortality rate correlated with greenness exposure. Therefore, it is necessary to conduct comprehensive research involving more CRD outcomes.

Millions of people of all ages suffer from CRDs, which are regarded as major causes of morbidity, death, economic burden, and social problems all over the world (40). Taking asthma as an example, asthma prevalence peaks between the ages of 5 and 9 (1) Some patients develop asthma when they are adults for the first time (adult-onset asthma) (41). Evidence of the connection between greenness exposure and CRDs has grown over the past decade, but systematic reviews and meta-analyses have focused on particular age groups, such as children and adolescents (33, 36–38). However, there is a lack of horizontal

comparisons between different age groups in regard to greenness exposure and CRDs.

Buffer zones of different radii were used to evaluate greenness exposure. The most commonly used indicator is the normalized difference vegetation index (NDVI) (42). However, different buffer zones of NDVI suggest different ways in which greenness affects health (10). NDVI within 500 m of a residential address can be used to indicate the immediate neighborhood greenness of a residence appropriate for physical activity (43, 44), decreasing noise and air pollution (45, 46). NDVI with smaller buffers (e.g., 200-300 m) indicates the visual impact of greenness seen from home (27). NDVI with a larger buffer (e.g., 800-1,000 m) may suggest that the area is suitable for recreational purposes and can be visited by people (47). Some studies have explored correlations between greenness exposure in different buffer zones and CRD risk (48-54). In larger buffers, Zeng et al. (54) observed stronger associations between asthma and school greenness, while Dadvand et al. (50) suggested higher relative prevalences of current allergic rhinoconjunctivitis and asthma for surrounding greenness. In addition, Xiao et al. (53) reported stronger correlations with the odds of COPD for NDVI in a 1,000 m buffer. Bereziartua et al. (49) found that increments of NDVI inside both a 300 m grid and a 1,000 m buffer were not associated with respiratory diseases. Fan et al. (55) discovered that neighborhood greenness had a significant positive correlation with COPD prevalence across different NDVI buffer sizes. Up to now, three systematic reviews and meta-analyses have carried out grouping analyses in different buffers of asthma and AR (34, 35, 37) A greater impact of residential greenness has been observed on the odds of respiratory system diseases in the larger buffer zones (e.g., asthma and AR) (34). Health outcomes for various CRDs are unknown as a result of different greenness buffer zones.

Therefore, we systematically conducted a meta-analysis of published studies to examine the correlation between greenness exposure and a broad range of chronic respiratory health issues. We also investigated the health impacts of different age groups and different greenness buffers, providing healthcare professionals and researchers with a more generalized and higher-quality piece of evidence on the connection between greenness exposure and chronic respiratory health issues.

# 2 Materials and methods

# 2.1 Search strategy and data sources

We followed standard protocols suggested by the PRISMA (Preferred Reporting Items for Systematic Reviews and Meta-Analyses) guidelines for the study (Supplementary Table S1) (56). In addition, our study protocol has been registered at PROSPERO (registered number: CRD42023384029). For the study, the Embase, Web of Science, and PubMed databases were searched to find epidemiological studies about greenness exposure and chronic respiratory health issues. The English-language search was conducted with the last update on 31 January 2023. We used combinations of terms concerning greenness (e.g., greenness, green space, and NDVI) and CRD incidence/prevalence/mortality (e.g., pulmonary phthisis, asthma, chronic obstructive pulmonary disease, and rhinitis). To find more potentially pertinent studies, we also conducted manual searches for reference lists of all pertinent systematic reviews and eligible

studies. The detailed search strategy and pertinent search outcomes are shown in Supplementary Table S2.

"moderate," "low," or "very low." The GRADE method for evaluating the quality of evidence refers to the report by Welsh et al. (60).

# 2.2 Study selection and eligibility criteria

Six criteria were required for studies to be eligible: (i) conducted among the general population; (ii) examined the connections between greenness and CRD incidence/prevalence or mortality; (iii) were cross-sectional, case-control, cohort, and ecological study designs; (iv) assessed greenness exposure with objective metrics based on continuous NDVI rather than classification based on measurement data (e.g., quartiles and tertiles). We chose an exposure index NDVI for comparisons between studies since it is an excellent target of overall greenness and is the most commonly used definition in the selection of studies; (v) evaluated the health outcomes of CRDs based on a clinical evaluation (e.g., International Classification of Primary Care[ICPC], International Classification of Diseases [ICD], self-reported physis diagnoses history, death records, and medical records related to respiratory situations); and (vi) written in English. We excluded reports that included greenness as a confounding factor or that lacked essential details. The references identified in three databases were imported into Endnote 20. The remaining articles were independently assessed through the title and abstract by two researchers (M.T. and F.L.) after the duplicates were removed, and disagreements were resolved by H.L. and W.L.

#### 2.3 Data extraction

The following data from eligible studies were extracted independently by two reviewers (M.T. and F.L.). Data extracted from each study included: year of publication, lead author, location (country/region), type of study, number and age range of samples, NDVI buffer, exposure unit reported, respiratory health issues assessed, results sourced, estimates of risk with a 95% CI, and confounders. During the review process, a third reviewer (H.L.) was consulted to address any inconsistencies in data extraction.

#### 2.4 Assessment of the risk of bias

The RoB of the included studies was assessed by two independent investigators (M.T. and W.L.), and any disagreement was addressed through discussion with two other investigators (F.L. and H.L.). In order to estimate the RoB of selected studies, we consulted a checklist developed by WHO (57) and van Kempen [(58), Tables S4, 5]. Based on the tool, we evaluated the RoB associated with exposure assessments, participants' selection, confounding, and health outcomes.

#### 2.5 Scientific evidence levels

The Grading of Recommendations, Assessments, Developments, and Evaluations (GRADE) standard was used to grade the credibility of the pooled evidence. Subsequently, each combination of exposure outcomes was assessed for five factors (59). We used GRADE Profiler 3.6 to grade the confidence rating into four descriptors: "high,"

# 2.6 Data standardization and meta-analysis methods

This study mainly focused on the connection between greenness exposure and CRD incidence and mortality, as well as the subgroup analysis of age groups and buffer groups. For the overall metaanalysis and age subgroup analysis, we selected the pooled effect obtained from the largest buffer (within 1,000 m buffer) in the study and estimated the associations between CRD outcomes with different age groups and greenness exposure in adults aged 0-7 years, 8-12 years, 13-18 years, 19-40 years, and over 40 years independently based on the classification of all age groups included in the study. Moreover, we estimated the associations between CRD outcomes and NDVI at 200-300 m, 400-500 m, and 800-1,000 m buffers. A meta-analysis of chronic respiratory health risk associations with greenness was conducted using all odds ratios (ORs), risk ratios (RRs), and hazard ratios (HRs). The outcomes of interest (OR, RR, and HR) were included in the same meta-analysis in accordance with previous practice (61). This is acceptable in this situation because the outcome of interest is relatively common, but its effect size is small (62). In this study, we evaluated the effects of exposure to greenness on CRD outcomes by calculating the pooled RR (63). In order to be able to compare between studies, the size of the effects used in the meta-analysis was normalized and scaled to the same size (NDVI increased by 0.1). Pooled effect estimates were calculated as follows:

 $OR_{(s)} = OR_{(o)}^{increment_{(s)}/increment_{(o)}}$  (substituting RR or HR as applicable).

where  $OR_{(s)}$  represented the standardized effect estimates,  $OR_{(o)}$  represented the original effect estimates,  $increment_{(s)}$  represented the standardized increases of NDVI (per 0.1-unit increase), and  $increment_{(o)}$  represented the original increases in NDVI (42).

For this study, the "Review Manager 5.4" software was used, and a p-value <0.05 was taken as statistically significant. All results were presented as forest plots with 95% confidence intervals. In order to evaluate the impact of individual studies, every study was excluded one at a time during the sensitivity analysis. It was decided to use the random effect model in this study because it was a conservative method for high heterogeneity research (64).  $I^2$  statistics were used to evaluate the heterogeneity between studies ( $I^2$ =0–25% signified no heterogeneity;  $I^2$ =25–50% signified low heterogeneity;  $I^2$ =50–75% signified moderate heterogeneity;  $I^2$ =75–100% signified high heterogeneity) (65).

#### 3 Results

# 3.1 Retrieval of literature and characteristics of the study

As shown in Figure 1, according to our search strategy, a total of 6,691 relevant studies were retrieved from three databases, and another six studies were manually added to the reference list of review studies.

We assessed the titles and abstracts of 4,682 studies after removing duplicates. Next, the full texts of 118 studies were retrieved, and we further excluded 83 studies that were irrelevant. Finally, we selected 35 studies for meta-analysis, of which 20 had a pooled effect value of OR, 5 had a pooled effect value of RR, and 10 had a pooled effect value of HR.

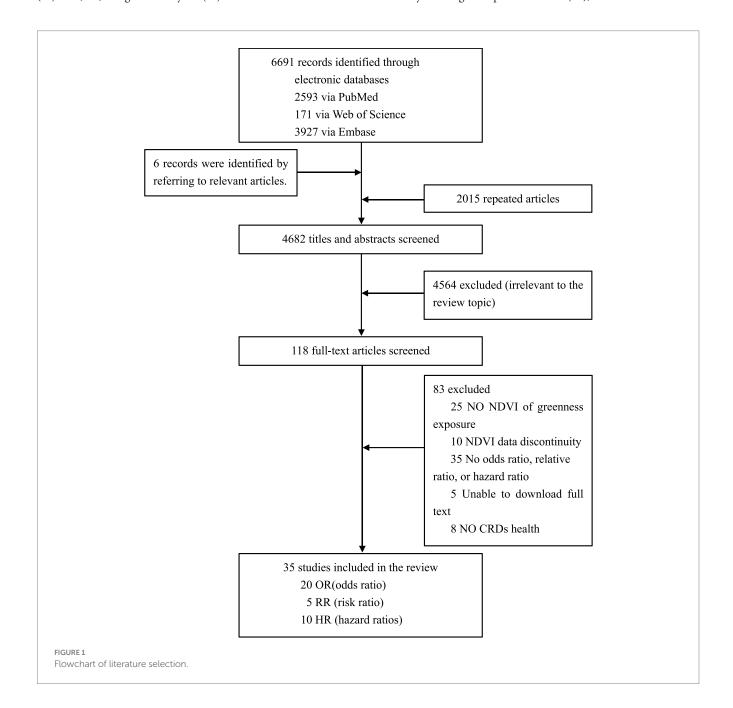
In Supplementary Table S3, the 35 studies were summarized according to their characteristics. The studies were all published in the last 10 years, indicating a growing interest in chronic respiratory health issues and greenness. The sample populations of this study were drawn from 20 countries, of which 15 (43%) were in Europe, 15 (43%) in Asia, and 5 (14%) in North America. There were 10 cross-sectional studies, 18 cohort studies, 4 ecological studies, and 3 case—control studies. There was a wide variation in the statistical sample size of these studies, from 478 children (66) to 10,481,566 aged over 30 years (31).

### 3.2 Primary outcomes

#### 3.2.1 NDVI and CRD incidence or prevalence

A total of 27 studies analyzed the connection between greenness exposure and CRD incidence/prevalence, including 14 cohort studies (1, 25, 26, 32, 52, 66–74), 10 cross-sectional studies (30, 45, 51, 53–55, 75–78), and 3 case–control studies (48, 79, 80).

Among these studies, 12 studies (44%) found that increasing NDVI and CRD incidence/prevalence in the overall samples had a significant beneficial effect, while 2 studies (7%) showed that an increment in NDVI was related to a slight increase in CRD incidence/prevalence. Eight studies found a stronger association between younger individuals (26, 54, 55), low urbanicity areas (25, 30, 81), low humidity and high-temperature areas (71), and lower household



income participants (25, 82). There were no statistically significant connections found in the remaining 13 studies (49%).

According to the pooled results, for every 0.1 increase in NDVI, the pooled RRs for asthma, AR, COPD, and lung cancer incidence were 0.92 (95% CI: 0.85–0.98, p=0.02,  $I^2$ =78%), 1.02 (95% CI: 0.97–1.08, p=0.41,  $I^2$ =65%), 0.92 (95% CI: 0.83–1.03, p=0.13,  $I^2$ =91%), and 0.62 (95% CI: 0.40–0.95, p=0.03,  $I^2$ =97%), respectively (Table 1 and Figure 2). The pooled RRs for asthma, AR, and COPD prevalence were 0.89 (95% CI: 0.74–1.08, p=0.23,  $I^2$ =54%), 0.91 (95% CI: 0.64–1.29, p=0.60,  $I^2$ =72%), and 1.00 (95% CI: 0.90–1.12, p=0.97,  $I^2$ =96%), respectively (Table 1 and Figure 2). High heterogeneity was observed for asthma incidence, COPD incidence/prevalence, and lung cancer incidence. Moderate heterogeneity was observed for asthma prevalence and AR incidence/prevalence. According to the GRADE system, asthma incidence and lung cancer incidence had "low" confidence, while the other CRDs had "very low" confidence in the pooled evidence (Supplementary Table S6).

As a result of sensitivity analyses, we observed that most of the associations remained unchanged after any single study was excluded, which indicated that our findings were robust. However, the pooled estimates for COPD and lung cancer incidence became marginally significant after the exclusion of one study (Supplementary Table S9).

#### 3.2.2 NDVI and CRD mortality

Eight studies analyzed the connection between greenness exposure and CRD mortality, consisting of five cohort studies (31, 49, 83–85) and three ecological studies (86–88).

Three studies (38%) reported a significantly beneficial relationship between NDVI and CRD mortality. Two studies suggested that protective relationships appeared to be significant among women (85) and younger individuals (<65 years) (31, 85). No statistically significant association was reported in the remaining five (62%) studies.

The pooled RRs for COPD and lung cancer mortality (0.1 increments in NDVI) were 0.95 (95% CI: 0.92–0.99, p=0.01, I<sup>2</sup>=7%) and 0.98 (95% CI: 0.96–1.01, p=0.21, I<sup>2</sup>=88%), respectively (Table 2 and Figure 3). High heterogeneity emerged for lung cancer mortality, and no heterogeneity emerged for COPD mortality. According to the GRADE system, COPD mortality had "moderate" confidence and lung cancer mortality had "very low" confidence in the pooled evidence (Supplementary Table S6).

As a result of sensitivity analyses, we observed that most of the associations remained unchanged after any single study was excluded, which indicated that our findings were stable. However, COPD mortality estimates became marginally significant after excluding one study owing to the limited number of included studies (Supplementary Table S10).

# 3.3 Subgroup analysis

#### 3.3.1 Age subgroups

Considering the age characteristics of the subjects in each study, meta-analyses were performed in five age subgroups (aged 0–7 years, 8–12 years, 13–18 years, 19–40 years, and over 40 years) (Table 3). Sufficient data on asthma, AR, and COPD incidence/prevalence was available to perform age subgroup analyses.

Ten of the 13 studies on the relationship between greenness exposure and asthma incidence/prevalence conducted analyses in different age subgroups. There are four studies for preschool children under 7 years old, two for school-aged children aged 8–12 years old, two studies for teenagers aged 13–18, one study for youth aged 18–40, and one study for people over 40. The estimated RRs for subgroups of 0–7 years, 8–12 years, and 13–18 years were 0.98 (95% CI: 0.80–1.20, p=0.83, I<sup>2</sup>=51%), 0.91 (95% CI: 0.63–1.30, p=0.60, I<sup>2</sup>=63%), and 0.91 (95% CI: 0.83–0.99, p=0.03, I<sup>2</sup>=0%), respectively (Table 3 and Supplementary Figure S3). Moderate heterogeneity was observed for subgroups of 0–7 years and 8–12 years, and no heterogeneity was observed for 13- to 18-year subgroups.

Six of the seven studies on the relationship between greenness exposure and AR incidence/prevalence conducted analyses in different age subgroups. There are two studies for preschool children under seven, two for school-aged children aged 8–12 years old, one study for teenagers aged 13–18, and one study for youth aged 19–40. The estimated RRs for subgroups of 0–7 years and 8–12 years were 0.99 (95% CI: 0.83–1.19, p=0.93, I<sup>2</sup>=0%) and 0.91 (95% CI: 0.64–1.29, p=0.60, I<sup>2</sup>=72%), respectively (Table 3 and Supplementary Figure S4). Moderate heterogeneity emerged for 8- to 12-year subgroups, and no heterogeneity was observed for 0- to 7-year subgroups.

All four studies on the relationship between greenness exposure and COPD incidence/prevalence conducted analyses in different age subgroups. There are three studies for mid-life adults aged 40–65 and

Outcomes	Studies (n)	Continuous exposure		GRADE
		Heterogeneity I <sup>2</sup> (p-value)	Summary effect size (RR) (95% CI)	
Asthma incidence	9	78% (0.02)	0.92 (0.85, 0.98)	LOW
Asthma prevalence	4	54% (0.23)	0.89 (0.74, 1.08)	VERY LOW
AR incidence	6	65% (0.41)	1.02 (0.97, 1.08)	VERY LOW
AR prevalence	2	72% (0.60)	0.91 (0.64, 1.29)	VERY LOW
COPD incidence	2	91% (0.13)	0.92 (0.83, 1.03)	VERY LOW
COPD prevalence	2	96% (0.97)	1.00 (0.90, 1.12)	VERY LOW
Lung cancer incidence	5	97% (0.03)	0.62 (0.40, 0.95)	LOW

Weights are from random-effects analysis. When the 95% CI excludes 1 or 0, summary effect estimates are in bold. CRD, chronic respiratory disease; RR, relative risk;  $I^2$ , heterogeneity statistic; CI, confidence interval; NDVI, normalized difference vegetation index; AR, allergic rhinitis; COPD, chronic obstructive pulmonary disease.

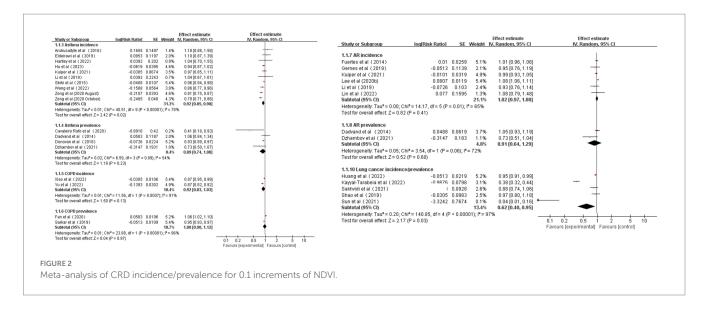


TABLE 2 Summary of meta-analyses of studies concerning NDVI and CRD mortality.

Outcomes	Studies (n)	Continuous exposure		GRADE
		Heterogeneity I <sup>2</sup> (p-value)	Summary effect size (RR) (95% CI)	
COPD mortality	3	7% (0.01)	0.95 (0.92, 0.99)	MODERATE
Lung cancer mortality	6	88% (0.21)	0.98 (0.96, 1.01)	VERY LOW

Weights are from random-effects analysis. When the 95% CI excludes 1 or 0, summary effect estimates are in bold. CRD, chronic respiratory disease; RR, relative risk; I², heterogeneity statistic; CI, confidence interval; NDVI, normalized difference vegetation index; COPD, chronic obstructive pulmonary disease.

three studies for older adults aged over 65. The estimated RRs for subgroups of 40–65 years and over 65 years were 0.97 (95% CI: 0.91–1.05, p=0.46, I<sup>2</sup>=91%) and 0.91 (95% CI: 0.79–1.04, p=0.16, I<sup>2</sup>=86%), respectively (Table 3 and Supplementary Figure S5). High heterogeneity emerged for all the meta-analyses.

As a result of sensitivity analyses, we observed that most of the associations remained unchanged after any single study was excluded, which indicated that our findings were stable (Supplementary Table S11). According to the GRADE system, asthma incidence of 13–18 years had "moderate" confidence, AR incidence of 0–7 years had "low" confidence and other subgroups had "very low" confidence in the pooled evidence (Supplementary Table S7).

# 3.3.2 Buffer subgroups

Considering the buffer zones of greenness exposure characteristics of the subjects in each study, we classified the buffer zones into three groups (200–300 m, 400–500 m, and 800–1,000 m buffers) (Table 4). All CRD outcomes had sufficient data to perform subgroup meta-analyses. Concrete study information is shown in Supplementary Figures S7–S12.

Ten of the 13 studies on the relationship between greenness exposure and asthma incidence/prevalence conducted analyses in different buffer subgroups. The estimated RRs were 0.92 (95% CI: 0.86–0.98, p=0.02,  $I^2$ =69% at 200–300 m buffer), 0.93 (95% CI: 0.85–1.01, p=0.09,  $I^2$ =58%) at 400–500 m buffer, and 0.87 (95% CI: 0.81–0.93, p<0.0001,  $I^2$ =54%) at 800–1000 m buffer, respectively (Table 4 and Supplementary Figure S7). Moderate heterogeneity emerged for all the meta-analyses.

All eight studies on the relationship between greenness exposure and AR incidence/prevalence conducted analyses in different buffer subgroups. The estimated RRs were 1.02 (95% CI: 0.95–1.09, p=0.65,  $I^2$ =69%) at 200–300 m buffer, 0.99 (95% CI: 0.94–1.03, p=0.53,  $I^2$ =0%) at 400–500 m buffer, and 0.99 (95% CI: 0.91–1.06,  $I^2$ =0.70,  $I^2$ =26%) at 800–1000 m buffer, respectively (Table 4 and Supplementary Figure S8). Moderate heterogeneity emerged in the NDVI 200–300 m subgroup, low heterogeneity was observed in the 800–1,000 m buffer subgroup, and no heterogeneity was found in the 400–500 m buffer subgroup.

All four studies on the relationship between greenness exposure and COPD incidence/prevalence conducted analyses in different buffer subgroups. The estimated RRs were 0.95 (95% CI: 0.89–1.02, p=0.18,  $I^2$ =89%) at 400–500 m buffer and 0.92 (95% CI: 0.83–1.03, p=0.13,  $I^2$ =91%) at 800–1000 m buffer (Table 4 and Supplementary Figure S9). High heterogeneity emerged for all the meta-analyses.

Four of the five studies on the relationship between greenness exposure and lung cancer incidence conducted analyses in different buffer subgroups. The estimated RRs for lung cancer incidence were 0.70 (95% CI: 0.46–1.06, p=0.09; I<sup>2</sup>=98%) at 400–500 m buffer and 0.20 (95% CI: 0.01–4.48, p=0.31, I<sup>2</sup>=94%) at 800–1000 m buffer, respectively (Table 4 and Supplementary Figure S10). High heterogeneity emerged for all the meta-analyses.

Five of the six studies on the relationship between greenness exposure and lung cancer mortality conducted analyses in different buffer subgroups. The estimated RRs were 0.98 (95% CI: 0.94–1.01, p=0.19;  $I^2$ =80%) at 200–300 m buffer and 0.98 (95% CI: 0.94–1.03, p=0.43;  $I^2$ =85%) at 800–1000 m buffer, respectively (Table 4 and

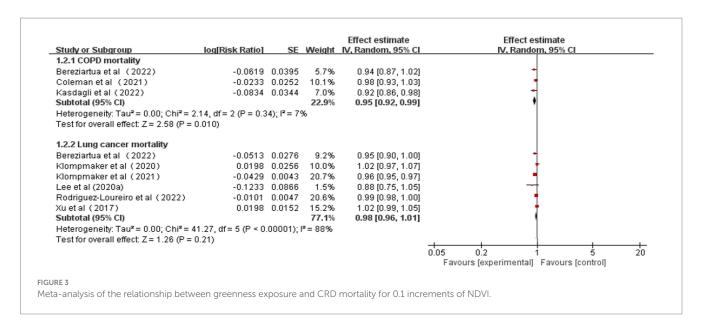


TABLE 3 Meta-analysis of age subgroups.

Outcomes	Studies (n)	Continuous exposure		GRADE
		Heterogeneity I <sup>2</sup> (p-value)	Summary effect size (RR) (95% CI)	
Aged 0–7 years				
Asthma incidence/prevalence	4	51% (0.83)	0.98 (0.80, 1.20)	VERY LOW
AR incidence	2	0% (0.93)	0.99 (0.83, 1.19)	LOW
Aged 8–12 years				
Asthma prevalence	2	63% (0.60)	0.91 (0.63, 1.30)	VERY LOW
AR prevalence	2	72% (0.60)	0.91 (0.64, 1.29)	VERY LOW
Aged 13-18 years				
Asthma incidence	2	0% (0.03)	0.91 (0.83, 0.99)	MODERATE
AR incidence	1	-	0.93 (0.76, 1.14)	_
Aged 19-40 years				
Asthma incidence	1	-	0.97 (0.85, 1.11)	-
AR incidence	1	-	0.99 (0.93, 1.05)	_
Aged 40+ years				
Asthma incidence	1	-	0.86 (0.77, 0.96)	_
COPD incidence/prevalence (aged 40–65 years)	3	91% (0.46)	0.97 (0.91, 1.05)	VERY LOW
COPD incidence/prevalence (aged over 65 years)	3	86% (0.16)	0.91 (0.79, 1.04)	VERY LOW

Weights are from random-effects analysis. Significant estimates are bolded. RR, relative risk;  $I^{\prime}$ , heterogeneity statistic; CI, confidence interval; AR, allergic rhinitis; COPD, chronic obstructive pulmonary disease.

Supplementary Figure S11). High heterogeneity emerged for all the meta-analyses.

All three studies on the relationship between greenness exposure and COPD mortality conducted analyses in different buffer subgroups. The estimated RR was 0.93 (95% CI: 0.88–0.98, p=0.04, I<sup>2</sup>=0%) at 800–1000 m buffer (Table 4 and Supplementary Figure S12). No heterogeneity emerged for the meta-analyses.

As a result of sensitivity analyses, we observed that most of the associations remained unchanged after any single study was excluded, which indicated that our findings were stable (Supplementary Table S12). According to the GRADE system, AR incidence/prevalence at

400–500 m and 800–1,000 m buffers and COPD mortality at 800–1000 m buffer had "low" confidence; other subgroups had "very low" confidence in the pooled evidence (Supplementary Table S8).

# 3.4 Study RoB assessment

In Supplementary Tables S4, S5, we summarize the RoB valuations for studies exploring the connection between greenness exposure and chronic respiratory health issues. There was a single study that was deemed to have a "high risk of bias" (67). As far as confounding

TABLE 4 Meta-analysis of buffer subgroups.

Outcomes	Studies (n)	Continuous exposure		GRADE
		Heterogeneity I <sup>2</sup> (p-value)	Summary effect size (RR) (95% CI)	
NDVI 200-300 m				
Asthma incidence/prevalence	9	69% (0.02)	0.92 (0.86, 0.98)	VERY LOW
AR incidence/prevalence	5	69% (0.65)	1.02 (0.95, 1.09)	VERY LOW
COPD prevalence	1	-	1.05 (1.02, 1.08)	-
Lung cancer incidence	1	-	0.90 (0.77, 1.06)	-
Lung cancer mortality	5	80% (0.19)	0.98 (0.94, 1.01)	VERY LOW
COPD mortality	1	-	0.96 (0.90, 1.02)	-
NDVI 400-500 m				
Asthma incidence/prevalence	9	58% (0.09)	0.93 (0.85, 1.01)	VERY LOW
AR incidence/prevalence	7	0% (0.53)	0.99 (0.94, 1.03)	LOW
COPD incidence/prevalence	4	89% (0.18)	0.95 (0.89, 1.02)	VERY LOW
Lung cancer incidence	3	98% (0.09)	0.70 (0.46, 1.06)	VERY LOW
Lung cancer mortality	1	-	0.98 (0.97, 0.99)	-
NDVI 800-1,000 m				
Asthma incidence/prevalence	8	54% (<0.0001)	0.87 (0.81, 0.93)	VERY LOW
AR incidence/prevalence	4	26% (0.70)	0.99 (0.91, 1.06)	LOW
COPD incidence/prevalence	3	91% (0.13)	0.92 (0.83, 1.03)	VERY LOW
Lung cancer incidence	2	94% (0.31)	0.20 (0.01, 4.48)	VERY LOW
Lung cancer mortality	4	85% (0.43)	0.98 (0.94, 1.03)	VERY LOW
COPD mortality	2	0% (0.04)	0.93 (0.88, 0.98)	LOW

Weights are from random-effects analysis. Significant estimates are bolded. RR, relative risk; P, heterogeneity statistic; CI, confidence interval; NDVI, normalized difference vegetation index; AR, allergic rhinitis; COPD, chronic obstructive pulmonary disease.

assessments and unblinded outcome assessments are concerned, all studies were deemed to have a "low risk of bias." Nine studies were deemed to have a "high risk of bias" due to the bias in selecting the participants. Three studies were appraised to have a "high risk of bias" for health outcome assessment. Based on these indicators, no studies were excluded.

According to the funnel plot (Supplementary Figures S1, S2, S4, S11, S12), there was no evidence of published bias in this meta-analysis.

#### 4 Discussion

This study identified 35 articles, consisting of presumably 32 million participants from 20 countries. A comprehensive evaluation was conducted according to the currently available data on the connection between greenness exposure and chronic respiratory health outcomes. Our study observed that the increment in greenness was connected with a reduction in asthma incidence, lung cancer incidence, and COPD mortality. Estimates of lung cancer incidence detected significant heterogeneity. The meta-analyses of age subgroups discovered that greenness exposure was connected with a low risk of asthma incidence in teenagers aged 13–18 years. A further meta-analysis of the buffer zone subgroups indicated that greenness exposure was positively correlated with asthma incidence/prevalence at 200–300 m and 800–1,000 m

buffers, and COPD mortality was associated with greenness exposure at 800–1000 m buffers.

#### 4.1 Comparison with other studies

Seven previously published reviews summarized the relationship between greenness and CRDs, which focused on one or two particular chronic respiratory health diseases (34). Our study assessed the relevance between greenness exposure and various CRD health outcomes. In line with this study, three previous meta-analyses indicated no significant overall association between greenness exposure and AR (33, 35, 37).

Three reviews and meta-analyses on asthma discovered no relationship between greenness exposure and asthma incidence (35–37) which is indirectly inconsistent with our findings (RR = 0.92, 95% CI: 0.85–0.98). Qiu et al. (34) combined 11 articles and reported asthma and rhinitis incidence and found that high-level residential greenness can significantly contribute to preventing respiratory diseases (OR = 0.95, 95% CI: 0.92–0.98). Our meta-analyses on different age subgroups observed that greenness exposure was correlated with a low risk of asthma incidence in teenagers aged 13–18 years (RR = 0.91, 95% CI: 0.83–0.99), but no connection was indicated in other age groups. According to Cao et al. (37) and Ye et al. (36), there is no significant correlation between greenness exposure and asthma in children and adolescents (aged under 19). It is thus

clear that more detailed results for the age subgroup were provided in our study. Compared to children over 12 years old, Zeng et al. (54) found that greenness and asthma had a more positive relationship with children under 12 years old. In preschool years, Sbihi et al. (26) indicated a reduced asthma risk with every interquartile increase in NDVI (OR=0.96, 95% CI: 0.93–0.99), but no similar relationships were observed during the school-age period. However, the finding was contrary to ours.

Our meta-analysis of the buffer subgroups concluded that greenness exposure was positively correlated with asthma incidence/prevalence at 200–300 m and 800–1,000 m buffers, and beneficial associations between greenness exposure at 800–1000 m buffer and asthma incidence/prevalence were stronger (RR=0.87, 95% CI: 0.81–0.93). Inconsistent results were reported by a previous meta-analysis (35), which indicated no significant relationships between the NDVI and current asthma in the 0–100 m group, 100–300 m group, and 500–1,000 m group. Two cross-sectional studies found similar evidence to ours that stronger associations for greenness exposure in larger buffers (45, 54). The case–control study conducted in Kaunas confirmed the opposite finding that an IQR increment in NDVI-100 m was statistically significantly associated with an increase in asthma risk (OR=1.43, 95% CI: 1.10–1.85) (48).

Our meta-analysis on lung cancer incidence and mortality was conducted separately, which indicated a significant positive correlation between greenness exposure and lung cancer incidence (RR=0.62, 95% CI: 0.40–0.95). Sakhvidi et al. (39) combined nine studies on greenspace exposure and lung cancer incidence /mortality and observed no correlation between greenspace exposure and lung cancer outcomes. By contrast, we provided more detailed meta-analysis results.

In this study, we carried out the first meta-analysis of the connection between greenness exposure and COPD incidence/prevalence and mortality. We found that increased greenness exposure was related to a reduced risk of COPD mortality (RR=0.95, 95% CI: 0.92–0.99), but not to COPD incidence and prevalence. However, three of four studies indicated the productive effect of greenness exposure on COPD (30, 53, 74). The higher prevalence of COPD has been linked to higher greenness levels across a countywide cross-sectional study (OR=1.19, 95% CI: 1.13–1.27) (55). Possible reasons might be the method used to diagnose COPD, the different periods used to estimate NDVI, and the different dominant types of vegetation in different regions (55).

Greenness exposure was positively associated with COPD mortality at  $800-1000\,\mathrm{m}$  buffer (RR=0.93, 95% CI: 0.88–0.98) in our study. There is inconsistency in the study results included in our meta-analysis. An ecological study in Greece, consisting of 1,035 municipal units, reported that greenness with a 1,000 m buffer was protective for COPD mortality (86). A cohort study containing 14 sub-cohorts from Europe discovered no association between NDVI in 300 m and 1,000 m buffers and COPD mortality (49). Various types and populations of studies could contribute to the differences in estimate sizes across studies.

As compared to previously published reviews, our study specifically focuses on the role of greenness in various CRDs. To the best of our knowledge, this is the first study to explore quantitatively the relationship between chronic respiratory diseases among different ages. As a result, our review provides a deeper understanding of the epidemiologic relationships between greenness and CRDs than previous studies.

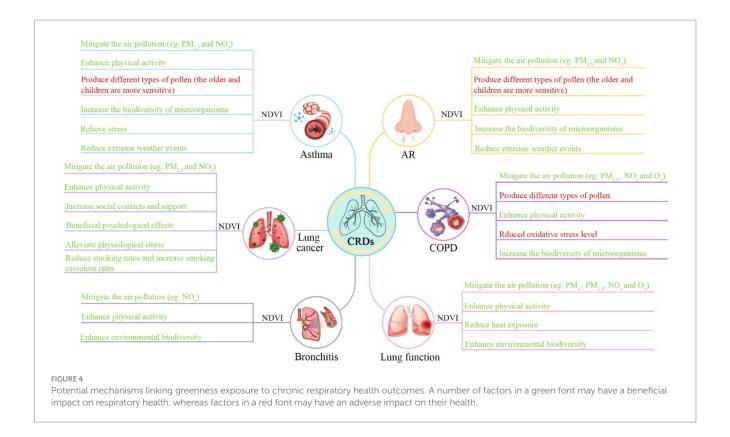
#### 4.2 Potential mechanisms

Exposure to greenness remains unclear in terms of its mechanisms of action. The benefits of greenness on chronic respiratory health issues have been explained by several mechanisms. First, green vegetation can reduce heat exposure by absorbing solar radiation (89, 90). High temperatures are correlated with airway drying, which may trigger bronchoconstriction and affect respiratory health (91, 92). Second, greenness can attenuate air pollution. Based on the results of some studies, the most important symptoms of ozone (O<sub>3</sub>), particle matter (PM<sub>10</sub>), sulfide dioxide (SO<sub>2</sub>), nitrogen dioxide (NO<sub>2</sub>), polyaromatic hydrocarbons (PAHs), and metals increase respiratory morbidity and mortality (93-96). Vegetation and trees may reduce air pollution-related chronic respiratory health issues through dispersion, deposition, and modification of air pollutant concentrations (97, 98). Moreover, greenness can promote physical activity by providing places for play or exercise, as well as a place for walking or cycling (99). The evidence suggests that physical activity could serve as an elastic mechanism that can improve emotional regulation (100) and reduce stress reactivity (101). This can encourage well-being and reduce the odds of chronic respiratory health complications (102). Additionally, greenness can reduce inflammatory responses through exposure to a wider variety of microbes, which may contribute to the relationship between greenness and health (103). As a result of increased exposure to a more diverse microbial environment, immune tolerance may be enhanced, and CRDs may be prevented (104). The potential mechanisms of each specific CRD are integrated into Figure 4.

However, greenness may have problems with respiratory health as they are sources of emissions, VOC, molds, aerosols, and pollen, which have been indicated to create AR problems (105–108). Despite the fact that COPD is generally regarded as a non-allergic respiratory disease, some studies have observed that higher levels of pollen in green areas are connected with higher rates of hospitalization for COPD (76). This may be why our meta-analysis showed no significant association between AR and COPD incidence/prevalence.

Our study indicated that greenness exposure has a beneficial effect on asthma incidence in teenagers aged 13–18 years. As teenagers spend most of their time in school, inhalation rates are higher when they are engaged in medium- to intense physical activity compared to when they are sleeping at home or resting at home (109). The planting of roadside trees has been shown to reduce ambient levels of particulate matter, such as ultrafine particles (110), and decrease levels of heavy metals in PM2.5 (111). Thus, the downregulated air pollutants in the larger buffer zone from greenness exposure may be large enough to see a meaningful diminution in asthma risk. The relatively small buffer zone had a positive effect on asthma, which may be due to the visual impact of greenness on health seen from home (112), and living close to greener areas is likely to encourage more regular and active participation in physical activity (113).

In addition to age and buffer zones, there are also many potential factors related to greenness that can affect respiratory health, including atmospheric environment (71, 81), educational attainment of mothers (26), urban–rural differences (69, 70, 83), gender (85, 88), underweight or premature birth (25, 30, 81), and household income (25, 82). A more comprehensive analysis of different potential factors is necessary in the future, which may assist in reducing socio-economic health inequalities.



# 4.3 Strengths and limitations

Our study included comprehensive and up-to-date evidence on greenness exposure and various CRD incidence/prevalence and mortality. An important strength of our study was the assessment of greenness exposure across age groups and buffer groups with subgroup meta-analysis to explore associations with chronic respiratory health issues. Moreover, we evaluated the RoB and rated the quality of the evidence using the GRADE method to improve the credibility of the study.

However, our study also had a number of limitations. First, we were constrained by the number of studies. This makes it impossible to conduct meta-analyses on various types of CRDs, such as bronchiectasis and bronchitis. Second, the greenness exposure indicator we selected was NDVI, which cannot characterize the structure, quality, and accessibility of greenness. Third, studies with different potential confounders were included, and there was no uniformity among them. Thus, we cannot rule out confounding effects when estimating the effects of greenness exposure. Fourth, due to a lack of studies, a high degree of heterogeneity, and a RoB, it is difficult to draw any absolute conclusions about the relationship between greenness and CRDs. Furthermore, providing important estimates of impact, adjusting covariates as necessary, and explaining bias issues in detail will improve the quality of reports in the future. Fifth, pooling OR, HR, and RR results together could produce imprecise estimates of the influence of greenness exposure on chronic respiratory health outcomes. Finally, some subgroup analyses combine prevalence and incidence because of the limited number of studies, which may impair the quality of the results of a meta-analysis. Moreover, as a language restriction, we only included English-language manuscripts, which is also a limitation.

#### 5 Conclusion

There was a correlation between higher greenness exposure and a reduced risk of some chronic respiratory health issues in this study, especially for asthma incidence, lung cancer incidence, and COPD mortality. Our results also indicate that age and buffer zone differences have potential implications for the correlation between greenness exposure and chronic respiratory health issues. According to this study, an increase in greenness exposure may be an appropriate measure to take precautions against some particular CRDs, potentially attracting the attention of policymakers and city managers to the public health issue. In order to provide detailed and specific recommendations for planners and policymakers, further research is necessary, focusing especially on the quality of greenness.

# Data availability statement

The original contributions presented in the study are included in the article/Supplementary material, further inquiries can be directed to the corresponding author.

#### **Author contributions**

MT: Conceptualization, Data curation, Investigation, Software, Writing – original draft. WL: Conceptualization, Formal analysis, Methodology, Visualization, Writing – original draft. HL: Resources, Supervision, Validation, Visualization, Writing – review & editing. FL: Formal Analysis, Funding

acquisition, Project administration, Resources, Supervision, Writing – review & editing.

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# Conflict of interest

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

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# Supplementary material

The Supplementary material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fpubh.2023.1279322/full#supplementary-material

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# Glossary

CRD	chronic respiratory disease	
DALY	disability-adjusted life-year	
COPD	chronic obstructive pulmonary disease	
AR	allergic rhinitis	
CF	cystic fibrosis	
NCFBE	non-cystic fibrosis bronchiectasis	
IPF	idiopathic pulmonary fibrosis	
UHI	urban heat island	
NDVI	normalized difference vegetative index	
PRISMA	Preferred Reporting Items for Systematic Reviews and Meta-Analyses	
RoB	risk of bias	
OR	odds ratio	
RR	risk ratio	
HR	hazard ratio	
PM	particle matter	
O <sub>3</sub>	Ozone	
NO <sub>2</sub>	nitrogen dioxide	
SO <sub>2</sub>	sulfide dioxide	
PAH	polyaromatic hydrocarbon	