The Long-Term Effects of Outdoor Air Pollution in Urban Environments on Cardiovascular Health: A Global Review

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The effects of air pollution on health are listed as a significant cause of death worldwide. Slightly over 3 million deaths per year are due to outdoor air pollution. Studies have shown that short-term increases in exposure to particulate matter have increased the risk of cardiovascular diseases such as myocardial infarction, stroke, and heart failure. However, less is known about the longer-term effects of air pollution on various cardiovascular diseases. The American Heart Association formally recognized PM2.5 as a significant cardiovascular risk factor in 2010. Since then, more prolonged-term exposure to air pollution has been suggested to cause chronic cardiometabolic and cardiovascular problems. The effects of long-term (>1 year) air pollution are significant, but not as much is known about how location affects this exposure. Associations with cardiovascular diseases and their risk factors are often increased in urban settings, which is attributed to a higher concentrations of outdoor air pollution. Potential causes of long-term air pollution concentrations in cities or metropolitan areas come from traffic exposure and traffic intensity. The U.S. Environmental Protection Agency and United Nations have suggested changes in air quality standards, implementation plans, and ways to reduce vehicle emissions specifically to improve human health and reduce the adverse effects of air pollution; however, more work still needs to be done. This review assesses the impact of the global long-term (>1 year) air pollution

Keywords: Long-term air pollution, Cardiovascular disease, Pollution exposure

Introduction

Air pollution is a major global problem. The effect of air pollution has been directly associated with the exacerbated climate crisis as well as detrimental increases in our global public health. The threat of air pollution on health has been listed as a significant cause of death worldwide.^{1,2} Over 3 million deaths per year are due to particulate matter less than 2.5uM or PM₂.³ Air pollution will only increase over time due to the drastic increase in energy production, transportation, waste accumulation, and natural

disasters like forest fires^{2,3}, all of which stress the importance of addressing the health concerns associated with this pollution.

Air pollution is defined as a heterogeneous mixture of gases, liquids, and particulate matter (PM) that is mostly derived from combustion, specifically from fossil fuels.⁴ Location is a significant factor that must be considered when discussing air pollution. ⁵ Urbanized areas have a higher risk of increased air pollution concentrations largely due to

energy consumption and exhaust emissions.6,7,8 Therefore, the urban areas have a higher risk of various cardiovascular diseases (CVDs).4 The World Health Organization (WHO) has stated about 80% of people living in cities are breathing in air that is over the WHO guideline limit.6 Specifically, people from lower and middle-income countries suffer from the most exposure and, therefore, are at the highest risk of air pollution-related health risk.7 Within the U.S., 80.2% of Americans live in an urban area versus the 19.8% that live in a rural area. Interestingly, within the urban areas, 11.2% of Americans are in the lowest socioeconomic status (SES).9 Globally, the largest gap between the lowest and highest SES exists predominantly in urban settings.10

Air pollution can be created in indoor and outdoor environments, such as smoke in a house versus smog in a city, respectively. More specifically, outdoor or ambient air pollution has major implications on human health through the inhalation of toxic particles. The main sources of outdoor air pollution include particulate matter from fossil fuel combustion.¹¹ Additionally, changes in the composition in ground-level ozone (O₃) due to industrial combustion, traffic, and gasoline can play a major role in outdoor air pollution.¹¹PM and NO2 are usually the main contributors to outdoor air pollution; however, ultrafine particles (UFP) have also been associated with increased outdoor air pollution, often due to the combustion reactions12. While both outdoor and indoor air pollution are important in regards to global health, in this review, only outdoor air pollution-related to human cardiovascular health will be addressed.

The length of exposure to air pollution is an essential factor related to cardiovascular

Methods

This scoping literature review used articles found through the PubMed and Google Scholar databases to answer the initial question, "how does long-term outdoor air pollution found in urban environments affect cardiovascular health?" The initial search examined keywords displayed in Table 1 found

health as well. Studies have found that shortterm increases in exposure to increased concentrations of particulate matter have amplified the risk of cardiovascular diseases such as myocardial infarction, stroke, heart failure, etc..9 It is known that high amounts of ambient PM2.5 increase the risk of CVD, specifically in highly urbanized areas, even in short amounts of exposure time 13, indicating the necessity to fully understand the effect longterm. Specifically, a short-term exposure to outdoor air pollution was strongly associated with the risk factors of ischemic heart disease, specifically in cities¹⁴¹⁵ Other health risks may also intensify the risk of susceptibility to air pollution and CVDs, such as obesity. 16

However, until recently, less was known about the sizeable impact long-term air pollution has on cardiovascular health. The American Heart Association (AHA) formally recognized PM_{2.5} as a major cardiovascular risk factor in 2010.10 Since then, long-term exposure to air pollution has been suggested to cause chronic cardiometabolic problems such as diabetes.11 However, there is still a lack of information on the effect of long-term outdoor pollution on cardiovascular health, air specifically in urban locations such as cities or metropolitan areas with higher concentrations of air pollution.

In this review, short-term air pollution is described as less than one year of exposure, while long-term air pollution is three or more years of exposure. This review will outline the latest research from the last twenty years linking the extended exposure to air pollution in metropolitan areas to cardiovascular disease on a global scale.

in the title and abstract of retrieved papers as well as the index terms that describe the articles. All relevant reviews and articles were exported to Zotero for further examination and comparison. In the screen, all articles were scanned for titles based on the relevance of inclusion and exclusion criteria specified in

Table 1			
Search terms used in sys	tematic review of literature on long	g term outdoor air pollution found	in urban settings on
cardiovascular health, 2	000-2021		
Question	Cardiovascular Terms	Air Pollution Terms	Location Terms
How does long	"cardiovascular disease,"	"long-term air	"cities", "urban
term outdoor air	"heart," "myocardial	pollution," "outdoor air	setting", and
pollution found in	infarction," "heart	pollution," "ambient air	"metropolitan"
urban	failure," "ischemic heart	pollution" "particulate	
environments	disease," "hypertension"	matter," "UFP,"	
affect		"nitrogen dioxide,"	
cardiovascular		"ozone"	
health?			

Table 2, which also helped identify our Population, Exposure, and Outcome framework. The inclusion criteria consisted of studies that took place in defined cities or metropolitan areas studying outdoor air pollution effects on cardiovascular health for greater than one year. Exclusion criteria included separate, non-cardiovascular-related health effects such as renal function, lung function, and brain function as well as suburban or non-metropolitan specific locations. Any studies that were conducted for less than one year were considered short-term exposure and were not included.

Additionally, we searched for recent articles that researched over the last thirty years to emphasize the latest long-term studies; therefore, research studies conducted between 1990 and 2021 were included. We also examined

broader primary studies to help narrow our search for more recent and specific articles. We examined the reference list of these key papers to identify additional studies for more specific articles. Furthermore, to find grey literature on the topic, we individually searched for key authors in the field that were established in the original search, key agencies or organizations that often conduct relevant research, and key journals that have an index or database of related topics. Additionally, we referenced the PRISMA flowchart and checklist for a scoping literature review¹⁷. From this, a total of 9650 articles were found through database searching. After duplicates were removed and screened through titles and abstracts, approximately 173 articles were eligible. From this group, the articles were fully scanned and 59 were used for this review.

Question	Population/Location	review of literature Air	Health Effect	Include
-	-	Pollution		
How does long	Must be representative	Identifies	Addresses an	If it specifies
term outdoor air	of an urban setting	long term air	effect on heart or	the length of air
pollution found in		pollution	cardiovascular	pollution
urban		exposure	health	exposure in an
environments		(longer than		urban
affect		three		environment on
cardiovascular		months)		cardiovascular
health?				health

Results

Long-Term Air Pollution and CVD

Long-term exposure to PM2.5 and ozone (O3) has been connected with the progression of CVDs. Recently, long-term exposure to air pollution resulted in increased heart rate variability, blood pressure, vascular tone, blood coagulation, and atherosclerosis.18 A study through the AHA found that people exposed to PM2.5 and ozone over five years had poorer health one year following a myocardial infarction (MI). This was correlated with an added risk of death due to increased morbidity and mortality as well as worsened symptoms within the following five years.¹⁹ Using cohort studies, An et al.. summarized that long-term exposure to air pollution resulted in a significantly higher risk of CVD (1.24%) compared to short-term exposure, which was hypothesized to be because of the compounding effects of extended air pollution exposure. 18 Another study found that cardiopulmonary death was associated with fine PM (PM2.5) and sulfur dioxide (SO₂) pollution. Interestingly,

How Extended Exposure to Outdoor Air Pollution Affects Cardiovascular Health

To better understand how to treat and prevent CVD associated with air pollution, it is essential to determine how air pollution causes an increase in disease. Some studies have explored how fine particulate air pollution causes a variety of CVDs through inflammatory mechanisms. One study examined increased pulmonary and systemic inflammation. This inflammation is associated with breathing in the particulate matter associated explicitly with air

Long-term Air Pollution in Urban Settings

The effect of long-term air pollution is significant, but not as much is known about how location affects this exposure. In cities, there is an escalation in traffic and motor emissions strongly associated with air pollution.²⁵ This kind of air pollution often consists of elemental carbon, ultrafine particles, and nitrogen oxides.²⁵ Pope et al. concluded that every ten microgram increase in $PM_{2.5}$ air pollution was associated with a 4% increased risk of cardiopulmonary mortality and, therefore, contributed to the adverse effects of long-term exposure on health.²⁰

A critical study by Miller et al. found that postmenopausal women had an amplified risk of CVD and death associated with cardiovascular issues because of long-term exposure to fine PM. In addition, they found that there were different risks and exposures to women in cities. ²¹ Another group found that the elderly community (65 years or older) are more at risk for the adverse effects of air pollution on cardiovascular health. A study of the elderly population in Taiwan found that added long-term exposure to ambient air pollution causes changes in blood pressure as well as blood sugar and blood lipids on older members of the population. ²²

pollution. This escalation in the inflammation from air pollution exacerbated atherosclerosis and decreased cardiac function.²³ Another group studied environmental PM in an atherosclerotic mouse model and found that long-term exposure to this air pollution changed the vascular tone and significantly increased vascular inflammation leading to a CVD such as atherosclerosis.²⁴

An initial study found that there was a strong association of increased fine PM air pollution that led to increased mortality in six different United States cities, including Steubenville, St. Louis, Harriman, Watertown, Topeka, and Portage²⁶. These cities were chosen because of the known air pollution in the area, with Steubenville being the worst and Portage being the best. Specifically, the authors found an increase in cardiopulmonary diseases in over eight thousand adults over 15 years.²⁶ Later in 2006, a similar study found that increased PM_{2.5} was associated with cardiovascular mortality, and when that exposure was reduced, there was a decrease in mortality.²⁷

High blood pressure or hypertension is a significant risk factor for CVD.²⁸ A study found that long-term exposure to PM led to an increase in arterial blood pressure. This association was significantly amplified in urban settings, which was attributed to an increased concentration of PM_{2.5}. This outcome was independent of ethnic groups or seasonal changes.²⁹

Some confounding variables have been suspected in outdoor air pollution studies. Specifically, changes in season and temperature have been thought to affect air quality.²⁴ A study examined the risk in cities over various seasons and found the risk of mortality related to outdoor air pollution was not significantly different in above-average or below-average temperatures. The risk was the same in cities in drier climates as well as humid climates. This indicated that the weather or temperature changes were not responsible for increased mortality related to air quality or condition.³⁰

As shown in Table 3, there are various pollutants commonly found in outdoor metropolitan environments that intensify cardiovascular problems. However, Cesaroni et al. suggested that air pollutants, specifically from traffic, including NO₂ and PM_{2.5}, cause the most severe and persisting cardiovascular problems. Specifically, traffic-related air pollution is defined as NO2 and PM2.5 that is uniquely augmented during rush hour periods, and is also found in vehicular emissions.³¹ In Rome, the long-term exposure to air pollution was attributed to increased exposure to NO2 and PM_{2.5}, which was associated with a greater number of incidents of ischemic heart disease.32 A similar result was also found in major cities in Greece, where CVD morbidity was associated with prolonged exposure to traffic-related air pollution.³³ In a cohort study from Norway and the United Kingdom, traffic air pollution was

also associated with increased cardiovascular mortality. Specifically, in high traffic areas, the long-term exposure to particulate air pollution was connected to more cases of CVD. Additionally, even air pollution levels lower than the current European air quality standards were linked to CVD risk.³⁴ In general, air pollution in major European cities is caused by traffic pollution, which is resulting in elevated risks of CVDs. Finally, Beelen et al. created a multicenter program called European Study of Cohorts for Air Pollution Effects (ESCAPE), which assessed long-term air pollution on mortality. This study assessed large cities in Europe, some of which include Oslo, Augsburg, Stockholm, Rome, and San Sebastian, and found an increase in risk factors for cardiovascular diseases. Overall, this study summarized the massive negative effect on heart health due to air pollution in cities, specifically in Europe.35

In the Netherlands, extended exposure to traffic-related air pollution was linked to an added risk of cardiovascular mortality. The study measured black smoke, nitric oxide, sulfur dioxide, and PM2.5. They measured local or home outdoor exposure compared to urban exposure based on geographic information systems to determine the intensity of traffic exposure in the respective areas. They discovered that traffic intensity, specifically near the home, was associated with a 10ug/m³ increase in black smoke, NO, and PM2.5, which was related to cardiovascular mortality and lung cancer. Downward et al. found that long-term exposure to ultrafine particles (UFP) was strongly associated with an increase in CVD. Specifically, the authors found that UFP was often underestimated, due to the current emphasis on PM. This long-term exposure study in Amsterdam found increased incidents of myocardial infarction as well as heart failure associated with UFP12.

There is an apparent exacerbated risk of CVD because of outdoor urban air pollution, but there are also populations that are more at risk, which must be considered. For example, a study found that long-term air pollution exposes the elderly to an increased hospital admission rate for cardiac disease.³⁷ A study focused on

Comparison of outdoor air pollutants commonly found in cities20Adverse Health EffectsWHO guideline levelPollution found in CitiesComponents of sulfates, nitrates, ammonia, sodium chloride, black carbon, mineral dust and water suspended in the air, often from combustionPM10 can enter lungs causing cardiovascular and respiratory problems. barrier and blood system causing cardiovascular and respiratory problemsPM25 pM25 meanNitrogen Dioxide (NO2)Aerosols, component of PM25, emission from combustion for heat,Aerosols, component of pM25 causes inflammation of airways, bronchitis in asthmatic children, atchildren, 200 µg/m3 1-hour
Outdoor Air Pollution found in CitiesComponents of sulfates, nitrates, ammonia, sodium chloride, black carbon, mineral dust and water suspended in the air, often from combustionPM10 can enter lungs through inhalation causing cardiovascular and respiratory problems.PM10 20 µg/m³ annual meanNitrogen Dioxide (NO2)Aerosols, component of PM2.5, emission fromPM20 causes inflammation of airways, bronchitis inPM40 20 µg/m³ annual mean
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combustion for heat, asthmatic children, $200 \ \mu g/m^3 1$ -hour
power, engines of vehicles reduced lung function mean
and ships
Sulfur DioxideProduced fromReduced lung function, $20 \ \mu g/m^3 24$ -hour
(SO ₂) combustion of fossil fuels eye irritation, mean
(coal and oil) mostly for inflammation of $500 \mu\text{g/m}^3 10$ -minute
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increased risk of
cardiovascular disease
Ozone (O ₃) Ground level component Causes breathing $100 \mu\text{g/m}^3$ 8-hour
of photochemical smog problems, asthma, mean
created when sunlight reduced lung function,
reacts with other air enhanced lung diseases
pollutants

metropolitan areas of Greece found that longterm exposure to traffic-related air pollution caused an added risk of fatal and nonfatal ischemic heart disease and stroke. Importantly, they found that this CVD risk was higher in women and younger children.³³

Urban, traffic, and industrial-related air pollution is the biggest issue in large cities in Asia.³⁸ In Tianjin, Shenyang, Taiyuan, and Rizhao, air pollution levels are significantly higher than even the upper limits of the WHO guidelines.³⁹²⁰ These increases in PM_{2.5} are associated with higher risks of CVD mortality. Importantly, the more significant impact on CVD mortality was associated with a longer duration of exposure.⁴⁰ Air quality has not only been associated with a decline in human health but also in food instability in China. In six sectors of China, the PM_{2.5} and the O₃ pollutants were found to be detrimental to human health as well as crop production leading to an overall reduction in health.⁴¹ Additionally, a study from Seoul, South Korea found that long-term exposure to PM_{2.5}, specifically a 1µg/m³ increase over an average of seven years caused a 36% increase in cardiovascular events⁴². Importantly,

these cardiovascular events, which included heart failure, myocardial infarction, and stroke, occurred in individuals who had no previous cardiovascular issues, emphasizing the staggering implication of long-term air pollution on heart health.⁴²

Furthermore, India has a considerably higher prevalence of elevated levels of urban air pollution. This is associated with the country's increased motor vehicle activity as well as A group analyzed 59 industrialization. independent studies of short (hours, days, weeks) and long-term (months, years) ambient air pollution cases in India. They determined that long-term air pollution was strongly associated asthma, with heart attack, cardiovascular mortality, and premature mortality. The majority of health problems related to air pollution were caused by PM_{2.5}.⁴⁴

In the United States, outdoor air pollution has been associated with increased cardiovascular death, specifically in urbanized areas.⁴⁵ A comprehensive study examined 90 U.S. cities for PM₁₀ and found an increase in mortality, specifically cardiovascular and respiratory mortality.⁴⁶ Another study analyzed 88 of the largest urban areas in the United States for over eight years and analyzed all causes of mortality. They found that variations of PM₁₀ were attributed to cardiovascular and respiratory mortality compared to all other causes of

death.⁴⁷ The same group examined five outdoor air pollutants relative to mortality in 20 of the largest cities in the United States over the same eight-year period. They found that the PM_{10} was the air pollutant most strongly associated with cardiovascular mortality. In fact, for every 10µg per cubic meter increase in PM_{10} , there is a rise in cardiovascular-related death.⁴⁸ Another study analyzed data from 13.2 million U.S. residents within a six-mile radius of a PM_{2.5} monitor. They found that in eastern and central U.S. urban centers, a 10µg per cubic meter increase of PM_{2.5} over six years was associated with a 6.8% increase in mortality.⁴⁹ Even with this information, the United States Environmental Protection Agency (US EPA) standards are not enough. limiting А study examined hospitalization and congestive heart failure cases in seven U.S. cities over 13 years. They found that even air pollution levels below the EPA standard still resulted in additional hospital admission rates for congestive heart failure in people above the age of 65.5°

Across the globe, it is clear that increased long-term exposure to air pollution has a significant impact on cardiovascular health. However, there is still not much understood about how to reverse or lessen the adverse health effects of air pollution.

Discussion

How to Reduce Long-term Air Exposure Risks in Urban Settings

Long-term exposure in urban areas is a significant risk factor for various CVDs. The United States Environmental Protection Agency (US EPA) created an annual National Ambient Air Quality Standard (NAAQS) in 1997 specifically for PM_{2.5} that would require a decrease of 1µg per cubic meter in PM_{2.5}. A study examined the rate of cardiovascular mortality based on changes in the PM_{2.5} before and after the changes in the NAAQS guidelines in over 180 countries across the globe over 20 years. Interestingly, they found that for every 1µg per cubic meter decrease in PM_{2.5} lead to

about a 1.10% decrease in cardiovascular-related deaths. 51

Langrish et al., found that reducing personal exposure to PM by wearing masks that have a polypropylene filter that filters airborne PM and does not change ambient gas did improve the health of patients with coronary heart disease.⁵² Car filters have also been suggested as a possible recirculatory way to reduce airborne nanoparticles. The filters help recirculate the air to minimize nanoparticle concentrations below dangerous levels. The filters can reduce the concentration of pollutants less than nanoparticle concentration accumulated after three minutes of driving in heavy traffic.⁵³ Even low-efficiency filters have been shown to protect from traffic PM exposure effectively.⁵³ Exhaust PM traps on specific car engines have also been successful at decreasing PM exposure. These traps have even been used as a biomarker for CVD improvement.⁵⁴

As of 2019, there are studies designed to understand how outdoor air pollution affects cardiovascular health in specific populations. The Reducing Air Pollution in Detroit

Addressing Issues Regarding Outdoor Air Pollution

There is a dire need for change when it comes to PM and other air pollutants. The US EPA has suggested changes in air quality standards, statewide implementation plans, and ways to reduce vehicle emissions⁵⁶, but more work still needs to be done. For example, improvements have been made relative to traffic pollution emission through controls. Specifically, there are requirements for fewer tailpipe emissions as well as reduced crankcase emissions (combustion-related gases from the engine) through mandatory vehicle emission testing in the US25, however, universally these measures are still lacking.

Globally, the WHO Air Quality Guidelines (AQGs) are updated annually to be used as a baseline and reference for world leaders and lawmakers to make decisions for air quality limits and management.⁵⁷ The primary air pollutants the AQGs address included the particulate matter of various sizes, ozone, nitrogen dioxide, and sulfur dioxide. More recently, the guidelines have taken the length of exposure more seriously by including the impact of the duration of exposure in the AQGs as a way to emphasize the drastic need to update laws and educate the public.⁵⁷

Conclusions

Overall, the long-term exposure of outdoor ambient air pollution in urban settings is an increased hazard and risk for global health,

Intervention Study (RAPIDS) is analyzing a senior citizen community in Detroit for CVDs, pressure, increased blood and vasoconstriction⁵⁵⁵⁴. This area has been strongly linked to high levels of outdoor air pollution, specifically PM2.5. The group is using low efficiency and high-efficiency air filters in various conditions to assess if they can help improve cardiovascular health outcomes for high-risk populations.54 Continued studies such as this would be beneficial for an understanding of cardiovascular global health relative to outdoor air pollution.

These guidelines are meant to aid and advise world leaders; however, the impact and use of the guidelines are not followed equally across the globe. Joss et al. discovered that air quality standards for ambient outdoor air pollutants from 170 countries out of the 194 analyzed had varied levels of air quality standards, most not complying with the WHO guidelines. Moreover, 57 countries did not have any air quality standards at all.⁵⁸ This indicated not only a need for a better global air quality plan but also a need for better universal approaches to improve air quality.

More recently, the United Nations General Assembly created the global 2030 Agenda for Sustainable Development in 2015. One of the main goals created by this group was to adequately address air pollution and emphasize the detrimental effect on human health.⁵⁹ The current seventeen Sustainable Development Goals (SDGs) do not have an individual goal for air standards or air pollution. So, this 2030 Agenda was put forth to create a global air quality standard by focusing on international law about the atmosphere and global air pollution.⁵⁹

specifically for cardiovascular health. There is significant evidence of the cardiovascular risk associated with outdoor air pollution in urban

environments. Specifically, long-term exposure (more than three months) to outdoor air pollution leads to an amplified potential for a variety of CVDs and their risk factors. Cities and metropolitan areas have an increased susceptibility to the negative effects of outdoor air pollution because of added exposure to traffic and motor emission pollution, which leads to increased concentrations of dangerous air pollutants. Mechanistically, an increase in this outdoor air pollution has been shown to escalate inflammation in the heart and vasculature, which causes exacerbated effects of CVD. Yet, the detailed mechanism and side effects are still unknown.

There are still many questions that need to be answered in regards to the specific concentrations of various air pollutants and

their effects cardiovascular on health, specifically in vulnerable populations worldwide. A comprehensive understanding of all the various outdoor air pollutants, their concentrations in urban environments globally, and their toxicities, would provide insight into why outdoor air pollution is so harmful in certain areas and what can be done to prevent further damage. There are still many unknowns regarding the extended exposure to outdoor air pollution on cardiovascular health,.38 A better understanding of the mechanism could allow for more targeted interventions for preventing or lessening the adverse effects on human health. Finally, there is still a need for universal policies on air quality standards that are implemented and followed globally.

Conflicts of Interest

The authors have no conflicts to disclose.

Statement of Contributions

LK had the idea for the article under the guidance of HM. LK did the literature review and writing. HM provided critical revisions and oversight.

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