JOINT OPINION

Taking a Stand Against Air Pollution—The Impact on Cardiovascular Disease

A Joint Opinion From the World Heart Federation, American College of Cardiology, American Heart Association, and the European Society of Cardiology

Ithough the attention of the world and the global health community specifically is deservedly focused on the COVID-19 pandemic, other determinants of health continue to have large impacts and may also interact with COVID-19. Air pollution is one crucial example. Established evidence from other respiratory viruses and emerging evidence for COVID-19 specifically indicates that air pollution alters respiratory defense mechanisms leading to worsened infection severity. Air pollution also contributes to comorbidities that are known to worsen outcomes among those infected with COVID-19, and air pollution may also enhance infection transmission due to its impact on more frequent coughing. Yet despite the massive disruption of the COVID-19 pandemic, there are reasons for optimism: broad societal lockdowns have shown us a glimpse of what a future with strong air pollution measures could yield. Thus, the urgency to combat air pollution is not diminished, but instead heightened in the context of the pandemic.

PROBLEM

Air pollution is a major contributor to the global burden of disease, with an estimated 12% of all deaths in 2019^{*} attributable to outdoor and household[†] air pollution. While the impacts of air pollution on respiratory diseases is widely recognized, 50% of the estimated 6.7 million deaths attributable to air pollution in 2019 are due to cardiovascular diseases.¹ Globally, nearly 20% of cardiovascular disease deaths were attributable to air pollution. Further, air pollution was the 4th highest ranking risk factor for mortality, with more attributable deaths than high LDL cholesterol, high body-mass index, physical inactivity, or alcohol use (Figure 1).

Globally air pollution contributes to an average loss of life expectancy of 20 months, nearly as high as that for tobacco use (22 months), with losses of 2.5 years in South Asia.² Despite increasing awareness of the impact of air pollution on population health, appreciation of air pollution as a modifiable risk factor is still limited among health care providers. With this joint opinion, and in accordance with the Sustainable Development Goals, the WHF, AHA, ESC and ACC call for action in research and policy development at a global scale. Further, we aim to generate awareness about the impact of air pollution on cardiovascular disease as a means toward reaching the World Heart Federation Goal of a 25% reduction in premature cardiovascular mortality by 2025.

AIR POLLUTION AND CVD: EVIDENCE

Air pollution is a complex and dynamic mixture of numerous compounds in gaseous and particle form, originating from diverse sources, subject to atmospheric transformation and varying over space and time. Three common air pollutants, Michael Brauer[®], ScD Barbara Casadei[®], MD, DPhil Robert A. Harrington, MD Richard Kovacs, MD Karen Sliwa, MD, PhD and the WHF Air Pollution Expert Group



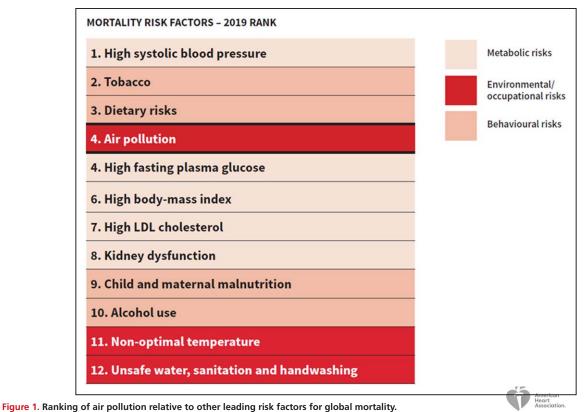
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*While other estimates exist and each have their uncertainties, all estimates of attributable disease burden are large and high-ranking in comparison to traditional and more widely recognized CVD risk factors.

tHousehold air pollution is mainly a concern in low income countries where polluting fuels (coal, wood, agricultural residue, animal dung) are used for cooking and heating.

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Mortality Risk Factors, Both sexes, all ages, 2019. Institute for Health Metrics and Evaluation. (Adapted from Institute for Health Metrics and Evaluation, 2020).

particulate matter (PM), ozone and nitrogen dioxide (NO_2) , are the focus of most monitoring programs, communication efforts,³ health impact assessments, and regulatory efforts.

Evidence for impacts on cardiovascular disease is most consistent for particulate matter, which is responsible for the vast majority of the disease burden via its impacts on ischemic heart disease,^{4–6} and stroke,⁷ as well as lung cancer, COPD, lower respiratory infections, Type 2 diabetes, pregnancy outcomes and related infant mortality.⁸ Time series studies conducted in hundreds of urban areas globally indicate a consistent association between shortterm variability in particulate matter and cardiovascular disease deaths,⁹ while large cohort studies from both high and lower income settings demonstrate increased cardiovascular disease incidence and mortality in association with PM levels.¹⁰ Further, particulate matter air pollution has been associated with progression of atherosclerosis.¹¹ Ozone is mainly associated with exacerbation of respiratory disease, with COPD incidence and mortality and with metabolic effects. NO, is often used as an indicator of traffic-related air pollution. Chronic exposure to NO₂ is associated with incident childhood asthma while short-term variability is associated with exacerbation of asthma and increased daily mortality counts.

A scientific statement from the American Heart Association provides a detailed description of the pathophysiologic mechanisms through which particulate

matter triggers cardiovascular events.⁴ Figure 2 summarises the likely dominant pathways including activation of oxidative stress/inflammation and autonomic imbalance, as well as translocation of components of the particulate matter mixture (ultrafine particles or specific constituents) into the systemic circulation.⁴ In turn, these alterations promote both subclinical cardiovascular disease (myocardial remodelling, atherosclerosis progression, systemic and pulmonary hypertension, enhanced vasoconstriction and coagulation) and thrombotic and nonthrombotic acute cardiovascular events (acute coronary syndromes, decompensated heart failure, stroke, life-threatening arrhythmias).4,5,12 The AHA statement suggests the existing evidence is consistent with a causal relationship between PM exposure and cardiovascular morbidity and mortality.

More recently, the European Society of Cardiology expert position paper⁵ updated the observational and mechanistic evidence and identified air pollution as one of several major modifiable risk factors relevant to the prevention and management of cardiovascular disease. The ESC paper identified a need for additional research regarding the role of air pollution in relation to hypertension and incident heart failure. This position paper also highlighted the role of health professionals, including cardiologists, in counseling their patients on the risks of air pollution while also supporting educational and policy initiatives to reduce air pollution exposure.

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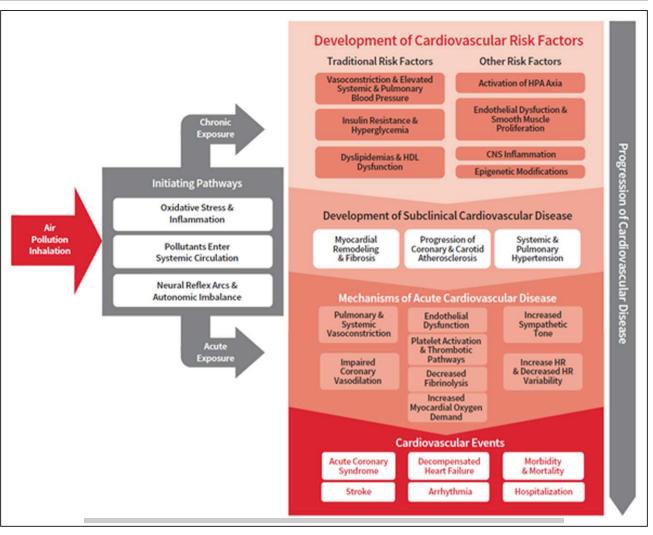


Figure 2. Biological pathways linking air pollution with cardiovascular disease. Mechanisms of cardiovascular disease attributable to air pollution exposures. (Adapted from Rajesh Vedanthan and Michael Hadley, 2019).

OPPORTUNITY FOR GLOBAL IMPACT

The threats posed by air pollution are substantial. Approximately 7 billion persons (92% of the global population), live in areas where the WHO Annual Air Quality Guideline for PM_{2.5} is exceeded and 3.6 billion people (47% of the global population) are exposed to household air pollution from the use of solid fuels for cooking. Recent research also indicates that air pollution impacts cardiovascular health even at very low levels below current WHO guidelines and most national standards.¹³ Further, climate change is leading to increased frequency and severity of wildfires, which leads to large smoke episodes and associated health impacts affecting major metropolitan areas.¹⁴

Despite these challenges, given its near-ubiquitous exposure, air pollution reduction offers a powerful opportunity to equitably reduce cardiovascular disease. Experience also indicates a clear path forward. In the United States, reductions in PM concentrations resulting from a diverse array of regulatory actions and technologies were responsible for up to 15% of the increase in US life expectancy observed over the past several decades.^{15,16} Similarly, programs such as India's Pradhan Mantri Ujjwala Yojana have made clean cooking fuels accessible to millions of households,¹⁷ helping to reduce inequalities in noncommunicable diseases.

ADDRESSING THE CHALLENGE: THE ROLE OF CARDIAC ORGANIZATIONS, SOCIETIES, AND FOUNDATIONS

The WHF, ACC, AHA, and ESC are united in our commitment to research, advocacy, and education to reduce the impacts of air pollution on cardiovascular health. Specifically, we will advocate for **further research on cardiovascular disease and air quality, disseminating findings** to our memberships and via other activities, such as World Heart Day. We will also work to **educate and raise awareness** among health care providers on the importance of reducing FRAME OF REFERENCE

air pollution and the cardiovascular benefits of air pollution mitigation. We will work with senior decisionmakers in national, regional and global governmental institutions to make air pollution–related heart disease a priority and to identify interventions to reduce air pollution and its impact on noncommunicable diseases. Finally, we will work with our members to increase the development and use of clinical guidelines on air pollution and cardiovascular disease to **ensure clinicians are educated** on the topic. In addition, we will strive to provide presentations on the links between air pollution and cardiovascular disease at our respective congresses, engagements, and events.

Structural actions to mitigate pollution emissions are ultimately necessary to reduce harmful exposures. Before mitigation is achieved, health care providers can play several important roles. First, clinicians can advocate for air pollution mitigation as a health measure. Second, clinicians can provide patients with personal measures to reduce exposures and associated risk at the individual level. For example, use of room air filtration can provide substantial improvements in PM levels within residences, schools and workplaces, with some evidence indicating improvements in blood pressure and measures of inflammation.¹⁸ Third, health care providers can integrate air pollution into disease management approaches. Communication tools such as various air quality indices focused on short-term variation in air quality can help patients adjust activities when air quality is poor.¹⁹ Detailed air pollution maps that provide information on long-term exposures and their impacts at neighborhood scales can also be used to target CVD treatment,²⁰ exposure reduction, and efforts to address other behavioural risks contributing to CVD in those living in locations with highest exposures.²¹ Finally, the health sector as a whole, which bears the impact of air pollution, can provide much-needed support for ministries of environment, energy, and transportation, which are traditionally responsible for mitigation efforts.

ARTICLE INFORMATION

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APPENDIX

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REFERENCES

- GBD 2019 Risk Factors Collaborators. Global burden of 87 risk factors in 204 countries and territories, 1990–2019: a systematic analysis for the Global Burden of Disease Study 2019. *Lancet*. 2020;396:1223–1249. doi: https://doi.org/10.1016/S0140-6736(20)30752-2
- Apte JS, Brauer M, Cohen AJ, Ezzati M, Pope CA. Ambient PM _{2.5} reduces global and regional life expectancy. *Environ Sci Technol Lett.* 2018;5:546– 551. doi: 10.1021/acs.estlett.8b00360
- Eagle KA, Riles E, Brook R. Clinical Effectiveness. The Air Quality Index: A Tool for Managing Patients with Cardiopulmonary Disease. Clincal Case Follow-Up. 2011. Accessed September 7: 2016;https://www.airnow.gov
 Brook RD, Rajagopalan S, Pope CA 3rd;***Brook JR, Bhatnagar A,
- 4. Brook RD, Rajagopalan S, Pope CA 3rd, Brook JR, Bhatnagar A, Diez-Roux AV, Holguin F, Hong Y, Luepker RV, Mittleman MA, et al; American Heart Association Council on Epidemiology and Prevention, Council on the Kidney in Cardiovascular Disease, and Council on Nutrition, Physical Activity and Metabolism. Particulate matter air pollution and cardiovascular disease: an update to the scientific statement from the American Heart Association. *Circulation*. 2010;121:2331–2378. doi: 10.1161/CIR.0b013e3181dbece1
- Newby DE, Mannucci PM, Tell GS, Baccarelli AA, Brook RD, Donaldson K, Forastiere F, Franchini M, Franco OH, Graham I, et al; ESC Working Group on Thrombosis, European Association for Cardiovascular Prevention and Rehabilitation; ESC Heart Failure Association. Expert position paper on air pollution and cardiovascular disease. *Eur Heart J.* 2015;36:83–93b. doi: 10.1093/eurheartj/ehu458
- Rajagopalan S, Al-Kindi SG, Brook RD. Air pollution and cardiovascular disease: JACC state-of-the-art review. J Am Coll Cardiol. 2018;72:2054– 2070. doi: 10.1016/j.jacc.2018.07.099
- Shah ASV, Lee KK, McAllister DA, Hunter A, Nair H, Whiteley W, Langrish JP, Newby DE, Mills NL. Short term exposure to air pollution and stroke: systematic review and meta-analysis. *BMJ*. 2015;350. doi: 10.1136/bmj.h1295
- Schraufnagel DE, Balmes JR, Cowl CT, De Matteis S, Jung SH, Mortimer K, Perez-Padilla R, Rice MB, Riojas-Rodriguez H, Sood A, et al. Air pollution and noncommunicable diseases: a review by the Forum of International Respiratory Societies' Environmental Committee, Part 2: air pollution and organ systems. *Chest.* 2019;155:417–426. doi: 10.1016/j. chest.2018.10.041
- Liu C, Chen R, Sera F, Vicedo-Cabrera AM, Guo Y, Tong S, Coelho MSZS, Saldiva PHN, Lavigne E, Matus P, et al. Ambient particulate air pollution and daily mortality in 652 cities. *N Engl J Med.* 2019;381:705–715. doi: 10.1056/NEJMoa1817364
- Yusuf S, Joseph P, Rangarajan S, Islam S, Mente A, Hystad P, Brauer M, Kutty VR, Gupta R, Wielgosz A, et al. Modifiable risk factors, cardiovascular disease, and mortality in 155 722 individuals from 21 high-income, middle-income, and low-income countries (PURE): a prospective cohort study. *Lancet.* 2020;395:795–808. doi: 10.1016/S0140-6736(19)32008-2
- Kaufman JD, Adar SD, Barr RG, Budoff M, Burke GL, Curl CL, Daviglus ML, Diez Roux AV, Gassett AJ, Jacobs DR Jr, et al. Association between air pollution and coronary artery calcification within six metropolitan areas in the USA (the Multi-Ethnic Study of Atherosclerosis and Air Pollution): a longitudinal cohort study. *Lancet.* 2016;388:696–704. doi: 10.1016/S0140-6736(16)00378-0

FRAME OF REFERENCE

- Münzel T, Gori T, Al-Kindi S, Deanfield J, Lelieveld J, Daiber A, Rajagopalan S. Effects of gaseous and solid constituents of air pollution on endothelial function. *Eur Heart J.* 2018;39:3543–3550. doi: 10.1093/eurheartj/ehy481
- Pappin AJ, Christidis T, Pinault LL, Crouse DL, Brook JR, Erickson A, Hystad P, Li C, Martin RV, Meng J, et al. Examining the shape of the association between low levels of fine particulate matter and mortality across three cycles of the Canadian Census Health and Environment Cohort. Environ Health Perspect. 2019;127:107008. doi: 10.1289/EHP5204
- Reid CE, Brauer M, Johnston FH, Jerrett M, Balmes JR, Elliott CT. Critical review of health impacts of wildfire smoke exposure. *Environ Health Per*spect. 2016;124:1334–1343. doi: 10.1289/ehp.1409277
- Bennett JE, Tamura-Wicks H, Parks RM, Burnett RT, Pope CA 3rd, Bechle MJ, Marshall JD, Danaei G, Ezzati M. Particulate matter air pollution and national and county life expectancy loss in the USA: a spatiotemporal analysis. *PLoS Med.* 2019;16:e1002856. doi: 10.1371/journal.pmed.1002856
- Pope CA 3rd, Ezzati M, Dockery DW. Fine-particulate air pollution and life expectancy in the United States. N Engl J Med. 2009;360:376–386. doi: 10.1056/NEJMsa0805646

- Kar A, Pachauri S, Bailis R, Zerriffi H. Using sales data to assess cooking gas adoption and the impact of India's Ujjwala programme in rural Karnataka. *Nat Energy.* 2019;4;806-814. doi: 10.1038/s41560-019-0429-8
- Chen R, Zhao A, Chen H, Zhao Z, Cai J, Wang C, Yang C, Li H, Xu X, Ha S, et al. Cardiopulmonary benefits of reducing indoor particles of outdoor origin: a randomized, double-blind crossover trial of air purifiers. J Am Coll Cardiol. 2015;65:2279–2287. doi: 10.1016/j.jacc.2015.03.553
- Riles E, Brook R. The air quality index: a tool for managing patients with cardiopulmonary disease. *Am J Med.* 2011;124:705–707. doi: 10.1016/j.amjmed.2011.02.034
- Radisic S, Newbold KB. Factors influencing health care and service providers' and their respective "at risk" populations' adoption of the Air Quality Health Index (AQHI): a qualitative study. *BMC Health Serv Res.* 2016;16:107. doi: 10.1186/s12913-016-1355-0
- Hadley MB, Baumgartner J, Vedanthan R. Developing a clinical approach to air pollution and cardiovascular health. *Circulation*. 2018;137:725– 742. doi: 10.1161/CIRCULATIONAHA.117.030377

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