

Environmental Pollution: An Under-recognized Threat to Children’s Health, Especially in Low- and Middle-Income Countries

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SUMMARY: Exposures to environmental pollutants during windows of developmental vulnerability in early life can cause disease and death in infancy and childhood as well as chronic, non-communicable diseases that may manifest at any point across the life span. Patterns of pollution and pollution-related disease change as countries move through economic development. Environmental pollution is now recognized as a major cause of morbidity and mortality in low- and middle-income countries (LMICs). According to the World Health Organization, pollution is responsible for 8.9 million deaths around the world each year; of these, 94% (8.4 million) are in LMICs. Toxic chemical pollution is growing into a major threat to children’s health in LMICs. The disease and disability caused by environmental pollution have great economic costs, and these costs can undercut trajectories of national development. To combat pollution, improved programs of public health and environmental protection are needed in countries at every level of development. Pollution control strategies and technologies that have been developed in high-income countries must now be transferred to LMICs to assist these emerging economies to avoid the mistakes of the past. A new international clearinghouse is needed to define and track the health effects of pollution, quantify the economic costs of these effects, and direct much needed attention to environmental pollution as a risk factor for disease.

Introduction

The worst industrial accident in the world occurred in 1984 in a pesticide plant in Bhopal, India, where 200,000 people were exposed to methyl isocyanate. The gas leak and explosion at the plant caused more than 6,000 deaths, and another 50,000 people suffered long-term health effects (Dhara et al. 2002).

Exposures to ambient air pollution, toxic chemicals, and pesticides generally have been problems that cause disease in high-income countries (HICs). Was the Bhopal disaster an anomaly? Or was it an early indicator of an emerging global pattern in which environmental pollution and toxic chemicals are becoming a greater source of health risk in low- and middle-income countries (LMICs)?

To explore these questions, the authors considered data on patterns of environmental exposure and disease in 12 countries that are at different levels of development (United Nations Development Programme 2015): Australia, Brazil, Canada, China, Ghana, Iran, Mexico, South Africa, South Korea, Switzerland, Thailand, and the United States. These data were collected from official sources in each country, from data compiled through the Global Burden of Disease (GBD) project (Lim et al. 2012; Lozano et al. 2012; Murray et al. 2012, 2015), and from World Health Organization sources (WHO 2005, 2009, 2014a, 2014b, 2014c, 2014d) that include physicians and scientists who are members of the WHO Collaborating Centres for Children’s Environmental Health (CEH; <http://www.niehs.nih.gov/research/programs/geh/partnerships/network/index.cfm>). The WHO has established these centers over the past decade in countries at all levels of industrial and economic development. The WHO Collaborating Centres (WHO CCs) are now forming into a network to ensure effective collaboration and coordination of research efforts (Sly et al. 2014). To date, the CEH WHO CCs are in Australia, Japan, Mexico, Republic of Korea, Thailand, United States, and Uruguay. A proposed WHO CC in Brazil is currently in the application process. The network is formally coordinated by the WHO CC

at the National Institute of Environmental Health Sciences (<http://www.niehs.nih.gov/research/programs/geh/partnerships/>).

Discussion

Patterns of environmental pollution and of the diseases caused by pollution vary greatly from country to country. National income and level of development are critical factors responsible for these sharp differences (Barreto 2004; De Maio 2011).

High-Income Countries

The principal pediatric diseases seen today in HICs are chronic, non-communicable diseases (NCDs) (WHO 2009). These diseases include birth defects—a leading cause of infant death in the United States (CDC 2006)—and asthma, which has been increasing in prevalence since 1980 (Akinbami 2002; Moorman et al. 2012). In addition, the reported prevalence of neurodevelopmental disorders, including dyslexia, mental retardation, attention deficit/hyperactivity disorder, and autism, have also increased sharply (CDC 2014), as have leukemia and brain cancer among children (Siegel et al. 2016). Childhood obesity more than doubled from 1980 to 2012 (CDC 2015), and its sequel, type 2 diabetes, has become epidemic (Hu et al. 2015; Beller 2000). The major diseases of adults in these countries are also NCDs—heart disease, stroke, cancer, diabetes, and chronic lung disease (Kelly et al. 2012).

Low- and Middle-Income Countries

In contrast, indoor air pollution and contaminated drinking water have been the major environmental risk factors for disease in LMICs, as clearly demonstrated in recent analyses in Ghana and South Africa (WHO 2005, 2014a, 2014d). Malnutrition and the high incidence and prevalence of parasitic and vector-borne diseases have also been major threats to human health (Pronczuk et al. 2011; WHO 2009, 2014a). These health problems are still major contributors to the burden of disease in LMICs.

Toxic Chemicals and Pesticides

Toxic chemicals and pesticides have long been important environmental pollutants in HICs, and thousands of these substances have been disseminated widely into the environment over the past century (Fischetti 2010). This long-standing concentration of toxic synthetic chemicals in the environment in HICs reflects the geographic origins of the chemical manufacturing industry, which began in Western Europe in the late 19th and early 20th centuries and then spread in the 20th century to North America, Japan, and Australia (Aftalion 2001). Many of the synthetic chemicals in widest use in these countries have never undergone any safety testing, and their potential toxicity is not known (Landrigan and Goldman 2011). Only about 20% have been screened for developmental toxicity (Goldman 1998). Scientists know even less about the possible synergistic effects of simultaneous exposures to multiple untested synthetic chemicals. National biomonitoring surveys conducted in the United States have documented several hundred synthetic chemicals in detectable quantities in the bodies of virtually all Americans of all ages (CDC 2009).

Toxic chemicals have been linked to numerous diseases through toxicological and epidemiological studies, and the list is growing as research into environmental causes of non-communicable disease

continues. The likelihood is high that there are additional diseases and disabilities caused by widely used synthetic chemicals where the etiologic associations have not yet been recognized (Grandjean and Landrigan 2014). Chemicals of particular concern include organic chemicals that persist in the environment long after their production and use have been stopped, such as polychlorinated biphenyls, and non-persistent chemicals to which individuals are constantly exposed, such as the plastic components and plasticizers bisphenol A, other bisphenols, and phthalates. Exposures to these organic chemicals have been associated with increased risk of diabetes (Lee et al. 2007, 2012, 2014), hypertension (Goncharov et al. 2011), cardiovascular disease (Lang et al. 2008; Lind et al. 2012), obesity (Newbold 2010; Trasande et al. 2012), and cancer (Lauby-Secretan et al. 2013).

Global Spread of Toxic Chemicals and Pesticides

In the past decade, with the globalization of trade, the spread of the Western life style, and the increasing globalization of the chemical manufacturing industry, toxic chemicals, highly hazardous pesticides, and chemical wastes, which previously were found only in developed countries, have been infiltrating LMICs with increasing rapidity (Spitz 2003). The manufacture and use of chemicals are shifting to LMICs, where labor costs are low and environmental and public health protections are few (Cole and Elliott 2005; Cole et al. 2010; Cole 2004; Kearsley and Riddell 2010). Chemical and pesticide pollution are increasing in LMICs, and hazardous wastes, including electronic waste, are accumulating (Grant 2014; Heacock et al. 2015; Luzardo et al. 2014; Perkins et al. 2014). At the same time, pollution-related chronic diseases such as asthma, heart disease, stroke, and cancer are becoming epidemic in countries where they were previously seldom seen (De Maio 2011; Kelly et al. 2012; Landrigan and Fuller 2014; Lim et al. 2012; Murray et al. 2015). The once separate patterns of disease in LMICs and HICs are converging (Dhara et al. 2002).

Tragic episodes of occupational and environmental exposure to toxic chemicals have resulted from the movement of the chemical industry to LMICs and have caused great damage. These include the devastating events such as the Bhopal disaster in India, and chronic, slowly unfolding tragedies such as the exposure of more than 1 million people to chrysotile asbestos in China, South and Southeast Asia, and sub-Saharan Africa (Frank and Joshi 2014). The new reality in global health is that NCDs are becoming major health problems in all countries around the world (Lim et al. 2012; Murray et al. 2015). In LMICs, especially those that are undergoing rapid industrialization, high risk of NCDs results in a double burden of disease, adding new threats to such age-old problems as infectious disease, inadequate clean drinking water, and poor nutrition (Pronczuk et al. 2011; Suk et al. 2003).

Toxic Chemicals and Children's Health in LMICs

A shift toward pollution-related diseases in developing countries presents a special problem for children who are already vulnerable because of inadequate nutrition and lack of access to clean drinking water. For children, exposures to environmental pollutants can be especially dangerous, as they are more sensitive than are adults to pesticides and other chemicals (reviewed in Landrigan and Goldman 2011). They intake more food, air, and water per pound than adults, and thus have greater exposures to toxic chemicals for their body weight. In addition, their metabolic pathways are immature, their early developmental processes are easily disrupted, and they have more time to develop chronic diseases than do adults (Landrigan and Goldman 2011). The concept that adverse environmental exposures in early life increase risk for disease outcomes in later life is not new and has been formalized in the developmental origins of health and disease (DOHAD) concept (Barker 2004). There are now many well-studied

examples of DOHAD in the literature, including the increased risk of obesity and cardiovascular disease in adult life that were the consequence of maternal malnutrition during pregnancy during the Dutch Famine. Cancers and non-cancerous lung diseases also have been shown to result from prenatal exposure to high concentrations of arsenic (reviewed in Boekelheide et al. 2012).

Such toxic environmental exposures before birth or in early postnatal life can cause short-term death from acute disease in infancy and childhood as well as chronic NCDs that can manifest at any point across the human lifespan (Boekelheide et al. 2012; Grandjean and Landrigan 2014). Indeed, low-dose exposure occurring during developmental windows of susceptibility—brief, precisely timed periods in embryonic, fetal, and early postnatal life when vital organs are sculpted through highly choreographed and tightly scheduled developmental processes—have been shown to have far greater effects on health than high-dose exposures to the same chemicals among adults (Ho et al. 2012). Major acute diseases associated with environmental pollution in early life include pneumonia (Darrow et al. 2014; Fuertes et al. 2014) and diarrheal disease (Lanata et al. 2013). Chronic NCDs associated with environmental exposures in early life include neurobehavioral disorders (Bouchard et al. 2011; Braun et al. 2009; Engel et al. 2010; Jacobson and Jacobson 1996; Rauh et al. 2011; Grandjean and Landrigan 2014), adult and pediatric asthma (Gauderman et al. 2004), hypertension, obesity, diabetes, cardiovascular disease (Barker 2004), and cancer (Smith et al. 2012).

Global Climate Change and Health

Global climate change could further exacerbate health risks from toxic environmental exposures, especially in LMICs, by increasing concentrations of many chemicals in water, air, and sediment (Noyes et al. 2009), as well as by imposing additional stress to individuals' immune, endocrine, and neurological systems that may leave some even more sensitive to the pollutants they encounter (Hooper et al. 2013).

Cost of Pollution

The diseases caused by pollution impose great economic costs on countries around the world—direct medical costs, opportunity costs reflecting the diminished productivity of populations damaged by pollution, and costs to health care systems (Jacobson and Jacobson 1996; Engel et al. 2010; Rauh et al. 2011; Bouchard et al. 2011; Trasande and Liu 2011; Landrigan and Fuller 2014).

The widespread pollution in a number of rapidly industrializing LMICs can result in adverse health effects, including damage to the brains, lungs, and other organ systems, for large numbers of persons (Laborde et al. 2015; Lanphear et al. 2000). This damage can result in diminished economic productivity of entire countries (Grosse et al. 2002; Landrigan and Fuller 2014; Trasande and Liu 2011; WHO 2010). For example, it is estimated that persons exposed to lead in countries that used leaded gasoline suffered widespread low-grade lead poisoning that resulted in cognitive impairment and reduced the population mean IQ in those countries by about five points (Lanphear et al. 2000; WHO 2010). A downward shift in cognitive function of this magnitude across an entire population has the effect of reducing, by more than 50%, the number of persons with IQ scores above 130, while at the same time increasing by more than 50% the number of persons with IQ scores below 70 (Weiss 1982). Such widespread cognitive impairment can reduce the intelligence and lifelong economic productivity of entire generations, thus undermining the developmental trajectory of whole societies (Landrigan and Fuller 2014).

Worldwide, environmental pollution is insufficiently appreciated and inadequately quantified as a cause of disease. It is estimated that pollution (the joint effects of household air pollution and ambient air pollution) is responsible for 7 million deaths per year (WHO

2014a, 2014d). This total is greater than the number of deaths due to HIV/AIDS, malaria, and tuberculosis combined (WHO 2014b, 2014c). Moreover, the overwhelming majority of the deaths attributed to ambient air pollution (88%) occurred in LMICs, and almost all of the deaths attributable to household air pollution were in LMICs (WHO 2014a).

Need for a Global Pollution Control Strategy

Despite these sobering facts, pollution receives far less attention in national and international health and development assistance programs than do AIDS, malaria, and tuberculosis (Landrigan and Fuller 2014). Pollution, as a risk factor for health effects, is often undercounted because of the very high standard of proof required in the public policy arena to establish etiologic associations between pollution and illness and to bring about effective remedial action. Examples are provided by the protracted debates over whether lead in gasoline causes subclinical lead poisoning in children that preceded the removal of lead from gasoline (Needleman and Landrigan 1981), and the long-running controversy over whether particulate air pollution is responsible for cardiovascular disease and lung cancer (Hamra et al. 2014).

A second reason for the relative lack of attention paid to pollution as a cause of disease and death is that there is a lack of reliable exposure data at the population level, especially exposures in early life that may have occurred years or decades ago (Briggs 2003). Exposures to environmental toxicants can occur at low doses and are frequently ongoing from ingestion of food or water or from inhalation of pollutants in air. These factors make proving causation very difficult, especially when the adverse health outcome from prenatal and early-life exposures occurred many years or decades later.

Yet another reason for the lack of attention paid to pollution is that the various components of pollution such as air pollution, water pollution, asbestos, and lead have been counted separately, one at a time (Landrigan and Fuller 2014). This disaggregated analytical approach is typical of environmental health research, and it is consistent with the structure of most public health and environmental protection agencies, which have separate bureaus for air, water, and solid waste. An unintended consequence of this fragmentation is that it minimizes the total impact of pollution and thus fails to give pollution the attention it deserves in planning and policy making.

Conclusions

Patterns of disease are changing rapidly in LMICs. Pollution-related chronic diseases are becoming more common. This shift presents a particular problem for children, who are proportionately more heavily exposed than are adults to environmental pollutants and for whom these exposures are especially dangerous. Better quantification of environmental exposures and stepped-up efforts to understand how to prevent exposures that cause disease are needed in LMICs and around the globe.

To confront the global problem of disease caused by pollution, improved programs of public health monitoring and environmental protection are needed in countries at every level of economic development. Pollution control strategies and technologies that have been developed and successfully deployed in HICs need to be transferred to LMICs, and their implementation must be adequately funded. Pollution control strategies in HICs have succeeded by controlling exposures at the source. For instance, lead has been removed from gasoline (Grosse et al. 2002), asbestos use has been sharply curtailed and banned in some countries (Frank and Joshi 2014), and air and water pollution have been reduced. Highly toxic pesticides have been replaced. These actions have produced tangible benefits for human health and the environment. Such strategies can also succeed in LMICs, as evidenced by experience with reducing lead in gasoline, with resulting declines in children's blood lead levels in countries such as the Philippines, India, and Pakistan (Suk et al.

2003). Such pollution control strategies can be highly cost effective and can lift the economies of entire nations. For example, the removal of lead from gasoline has resulted in much lower body lead burdens in children. Current evidence suggests that lowering lead exposure has a direct effect on reducing the incidence of impaired cognitive development and increasing economic productivity (Grosse et al. 2002). In the United States, removing lead from gasoline has returned approximately \$200 billion to the U.S. economy each year since 1980 (Grosse et al. 2002).

Constructing strong public health and environmental protection programs will require several elements at the country level. Tracking systems to monitor environmental pollution, actual exposures, and disease provide an essential foundation for these systems. Training physicians and other health care providers to recognize and manage diseases caused by environmental pollution is a second essential need. Legally mandated safety testing of new chemicals before they are introduced to commercial markets and of existing chemicals is a third essential pillar of chemical control (Landrigan and Goldman 2011). Assessment of toxicity must be followed by governmental regulation, as voluntary controls appear to be of little value (Ashford and Caldwell 2008).

The international development agenda must put a higher priority on pollution control, with a level of attention at least equal to that assigned to HIV/AIDS, malaria, and tuberculosis control (Landrigan et al. 2015). The lack of attention given to pollution in the program priorities of major international organizations is striking (Landrigan et al. 2015; Landrigan and Fuller 2014), especially given the substantial impact and favorable cost-benefit ratio of many pollution control programs such as the removal of lead from gasoline (Grosse et al. 2002), the installation of stack scrubbers on coal-fired power plants, and national bans on asbestos (Frank and Joshi 2014, Trasande and Liu 2011). Pollution protections are urgently needed in LMICs, where chemical production and use are rapidly increasing and environmental and occupational safeguards are too few. Current legal structures in many countries fail to adequately protect workers, children, and other vulnerable populations against environmental threats to health (Landrigan and Goldman 2011).

At the international level, an argument can be made for the formation of a new international clearinghouse focused on tracking the global movement of highly toxic pollutants and on defining the health effects of environmental pollution (Grandjean and Landrigan 2014). This new agency could be modeled on the International Agency for Research on Cancer and be placed in a position to:

- Assess industrial chemicals and other forms of pollution for potential health effects using a precautionary approach that emphasizes prevention and does not require absolute proof of toxicity.
- Facilitate and coordinate epidemiological and toxicological studies.
- Lead the urgently needed global programs for pollution prevention.

In summary, the adverse health consequences of exposure to environmental toxicants constitute a large and rapidly growing global problem, yet they receive insufficient attention in the global health and international development agendas. It is time to focus the world's attention on the great and growing global problem of environmental pollution. We must set stringent, but feasible numerical targets for pollution control.

Pollution deserves as much attention as infectious diseases. And, the global response to pollution deserves the same degree of rigor as has been applied to such infectious diseases as AIDS, tuberculosis, and malaria. Focus by the international community on environmental pollution can save the lives of millions, cost effectively and predictably. The need is great. The time is now.

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REFERENCES

- Aftalion F. 2001. A History of the International Chemical Industry. Philadelphia, PA:Chemical Heritage Press.
- Akinbami LJ, Schoendorf KC. 2002. Trends in childhood asthma: prevalence, health care utilization, and mortality. *Pediatrics* 110(2 Pt 1):315–322.
- Ashford NA, Caldart CC. 2008. Environmental Law, Policy, and Economics: Reclaiming the Environmental Agenda. Cambridge, MA:MIT Press.
- Barker DJ. 2004. The developmental origins of adult disease. *J Am Coll Nutr* 23(6 suppl):588S–595S.
- Barreto ML. 2004. The globalization of epidemiology: critical thoughts from Latin America. *Int J Epidemiol* 33(5):1132–1137.
- Beller GA. 2000. President's page: the epidemic of type 2 diabetes and obesity in the U.S.: cause for alarm. *J Am Coll Cardiol* 36(7):2348–2350.
- Boekelheide K, Blumberg B, Chapin RE, Cote I, Graziano JH, Janesick A, et al. 2012. Predicting later-life outcomes of early-life exposures. *Environ Health Perspect* 120(10):1353–1361, doi: 10.1289/ehp.1204934.
- Bouchard MF, Chevrier J, Harley KG, Kogut K, Vedar M, Calderon N, et al. 2011. Prenatal exposure to organophosphate pesticides and IQ in 7-year-old children. *Environ Health Perspect* 119(8):1189–1195, doi: 10.1289/ehp.1003185.
- Braun JM, Yolton K, Dietrich KN, Hornung R, Ye X, Calafat AM, et al. 2009. Prenatal bisphenol A exposure and early childhood behavior. *Environ Health Perspect* 117(12):1945–1952, doi: 10.1289/ehp.0900979.
- Briggs D. 2003. Environmental pollution and the global burden of disease. *Br Med Bull* 68(1):1–24, doi: 10.1093/bmb/ldg019.
- CDC (Centers for Disease Control and Prevention). 2006. Improved national prevalence estimates for 18 selected major birth defects—United States, 1999–2001. *MMWR Morb Mortal Wkly Rep* 54(51):1301–1305.
- CDC. 2009. Fourth National Report on Human Exposure to Environmental Chemicals. Atlanta, GA:U.S. Department of Health and Human Services, CDC. Available: <http://www.cdc.gov/exposurereport/pdf/fourthreport.pdf> [accessed 14 September 2015].
- CDC. 2014. Prevalence of autism spectrum disorder among children aged 8 years—autism and developmental disabilities monitoring network, 11 sites, United States, 2010. *MMWR Surveill Summ* 63(2):1–21.
- CDC. 2015. Childhood Obesity Facts. Available: <http://www.cdc.gov/healthyschools/obesity/facts.htm> [accessed 20 May 2015].
- Cole MA. 2004. Trade, the pollution haven hypothesis and the environmental Kuznets curve: examining the linkages. *Ecol Econ* 48(1):71–81.
- Cole MA, Elliott R. 2005. FDI and the capital intensity of “dirty” sectors: a missing piece of the pollution haven puzzle. *Rev Dev Econ* 9(4):530–548.
- Cole MA, Elliott R, Okubo T. 2010. Trade, environmental regulations and industrial mobility: an industry-level study of Japan. *Ecol Econ* 69(10):1995–2002.
- Darrow LA, Klein M, Flanders WD, Mulholland JA, Tolbert PE, Strickland MJ. 2014. Air pollution and acute respiratory infections among children 0–4 years of age: an 18-year time-series study. *Am J Epidemiol* 180(10):968–977.
- De Maio FG. 2011. Understanding chronic non-communicable diseases in Latin America: towards an equity-based research agenda. *Global Health* 7(1):36, doi: 10.1186/1744-8603-7-36.
- Dhara VR, Dhara R, Acquilla SD, Cullinan P. 2002. Personal exposure and long-term health effects in survivors of the Union Carbide disaster at Bhopal. *Environ Health Perspect* 110(5):487–500.
- Engel SM, Miodovnik A, Canfield RL, Zhu C, Silva MJ, Calafat AM, et al. 2010. Prenatal phthalate exposure is associated with childhood behavior and executive functioning. *Environ Health Perspect* 118(4):565–571, doi: 10.1289/ehp.0901470.
- Fischetti M. 2010. The great chemical unknown: a graphical view of limited lab testing. *Sci Am* 303(5):92.
- Frank AL, Joshi TK. 2014. The global spread of asbestos. *Ann Glob Health* 80(4):257–262.
- Fuertes E, MacIntyre E, Agius R, Beelen R, Brunekreef B, Bucci S, et al. 2014. Associations between particulate matter elements and early-life pneumonia in seven birth cohorts: results from the ESCAPE and TRANSPHORM projects. *Int J Hyg Environ Health* 217(8):819–829.
- Gauderman WJ, Avol E, Gilliland F, Vora H, Thomas D, Berhane K, et al. 2004. The effect of air pollution on lung development from 10 to 18 years of age. *N Engl J Med* 351(11):1057–1067.
- Goldman LR. 1998. Chemicals and children's environment: what we don't know about risks. *Environ Health Perspect* 106(suppl 3):875–880.
- Goncharov A, Pavuk M, Foushee HR, Carpenter DD. 2011. Blood pressure in relation to concentrations of PCB congeners and chlorinated pesticides. *Environ Health Perspect* 119(3):319–325, doi: 10.1289/ehp.1002830.
- Grandjean P, Landrigan PJ. 2014. Neurobehavioural effects of developmental toxicity. *Lancet Neurol* 13(3):330–338.
- Grant K, Goldizen FC, Sly PD, Brune MN, Neira M, van den Berg M, et al. 2014. Health consequences of exposure to e-waste: a systematic review. *Lancet Glob Health* 1(6):e350–e361.
- Grosse SD, Matte TD, Schwartz J, Jackson RJ. 2002. Economic gains resulting from the reduction in children's exposure to lead in the United States. *Environ Health Perspect* 110(6):563–569.
- Hamra GB, Guha N, Cohen A, Laden F, Raaschou-Nielsen O, Samet JM, et al. 2014. Outdoor particulate matter exposure and lung cancer: a systematic review and meta-analysis. *Environ Health Perspect* 122(9):906–911, doi: 10.1289/ehp.1408092.
- Heacock M, Kelly CB, Asante KA, Birnbaum LS, Bergman AL, Bruné MN, et al. 2015. E-waste and harm to vulnerable populations: a growing global problem. *Environ Health Perspect*, doi: 10.1289/ehp.1509699.
- Ho S-M, Johnson A, Tarapore P, Janakiram V, Zhang X, Leung Y-K. 2012. Environmental epigenetics and its implication on disease risk and health outcomes. *ILAR J* 53(3-4):289–305.
- Hooper MJ, Anklely GT, Cristol DA, Maryoung LA, Noyes PD, Pinkerton KE. 2013. Interactions between chemical and climate stressors: a role for mechanistic toxicology in assessing climate change risks. *Environ Toxicol Chem* 32(1):32–48.
- Hu FB, Satija A, Manson JE. 2015. Curbing the diabetes pandemic: the need for global policy solutions. *JAMA* 313(23):2319–2320.
- Jacobson JL, Jacobson SW. 1996. Intellectual impairment in children exposed to polychlorinated biphenyls in utero. *N Engl J Med* 335(11):783–789.
- Kearsley A, Riddell M. 2010. A further inquiry into the pollution haven hypothesis and the environmental Kuznets curve. *Ecol Econ* 69(4):905–919.
- Kelly BB, Narula J, Fuster V. 2012. Recognizing global burden of cardiovascular disease and related chronic diseases. *Mt Sinai J Med* 79(6):632–640.
- Laborde A, Tomasina F, Bianchi F, Brune MN, Buka I, Comba P, et al. 2015. Children's health in Latin America: the influence of environmental exposures. *Environ Health Perspect* 123(3):201–209, doi: 10.1289/ehp.1408292.
- Lanata CF, Fischer-Walker CL, Olascoaga AC, Torres CX, Aryee MJ, Black RE. 2013. Global causes of diarrheal disease mortality in children < 5 years of age: a systematic review. *PLoS One* 8(9):e72788, doi: 10.1371/journal.pone.0072788.
- Landrigan PJ, Fuller R. 2014. Environmental pollution: an enormous and invisible burden on health systems in low- and middle-income countries. *World Hosp Health Serv* 50(4):35–40.
- Landrigan PJ, Fuller R, Horton R. 2015. Environmental pollution, health, and development: a Lancet-Global Alliance on Health and Pollution—Icahn School of Medicine at Mount Sinai Commission. *Lancet* 386(10002):1429–1431.
- Landrigan PJ, Goldman LR. 2011. Children's vulnerability to toxic chemicals: a challenge and opportunity to strengthen health and environmental policy. *Health Aff (Millwood)* 30(5):842–850.
- Lang IA, Galloway TS, Scarlett A, Henley WE, Depledge M, Wallace RB, et al. 2008. Association of urinary bisphenol A concentration with medical disorders and laboratory abnormalities in adults. *JAMA* 300(11):1303–1310.
- Lanphear BP, Dietrich K, Auinger P, Cox C. 2000. Cognitive deficits associated with blood lead concentrations <10 microg/dL in U.S. children and adolescents. *Public Health Rep* 115(6):521–529.
- Lauby-Secretan B, Loomis D, Grosse Y, El Ghissassi F, Bouvard V, Benbrahim-Tallaa L, et al. 2013. Carcinogenicity of polychlorinated biphenyls and polybrominated biphenyls. *Lancet Oncol* 14(4):287–288.
- Lee DH, Lee IK, Jin SH, Steffes M, Jacobs DR, Jr. 2007. Association between serum concentrations of persistent organic pollutants and insulin resistance among nondiabetic adults: results from the National Health and Nutrition Examination Survey 1999–2002. *Diabetes Care* 30(3):622–628.
- Lee DH, Lind L, Jacobs DR Jr, Salihovic S, van Bavel B, Lind PM. 2012. Associations of persistent organic pollutants with abdominal obesity in the elderly: the Prospective Investigation of the Vasculature in Uppsala Seniors (PIVUS) study. *Environ Int* 40(4):170–178, doi: 10.1016/j.envint.2011.07.010.
- Lee DH, Porta M, Jacobs DR Jr, Vandenberg LN. 2014. Chlorinated persistent organic pollutants, obesity, and type 2 diabetes. *Endocr Rev* 35(4):557–601.
- Lim SS, Vos T, Flaxman AD, Danaei G, Shibuya K, Adair-Rohani H, et al. 2012. A comparative risk assessment of burden of disease and injury attributable to 67 risk factors and risk factor clusters in 21 regions, 1990–2010: a systematic analysis for the Global Burden of Disease Study 2010. *Lancet* 380(9859):2224–2260.
- Lind PM, van Bavel B, Salihovic S, Lind L. 2012. Circulating levels of persistent organic pollutants (POPS) and carotid atherosclerosis in the elderly. *Environ Health Perspect* 120(1):38–43, doi: 10.1289/ehp.1103563.

- Lozano R, Naghavi M, Foreman K, Lim S, Shibuya K, Aboyans V, et al. 2012. Global and regional mortality from 235 causes of death for 20 age groups in 1990 and 2010: a systematic analysis for the Global Burden of Disease Study 2010. *Lancet* 380(9859):2095–2128.
- Luzardo OP, Boada LD, Carranza C, Ruiz-Suárez N, Henríquez-Hernández LA, Valerón PF, et al. 2014. Socioeconomic development as a determinant of the levels of organochlorine pesticides and PCBs in the inhabitants of Western and Central African countries. *Sci Total Environ* 1:(497–498):97–105.
- Moorman JE, Akinbami LJ, Bailey CM, Zahran, HS, King ME, Johnson CA, et al. 2012. National Surveillance of Asthma: United States, 2001–2010. National Center for Health Statistics. *Vital Health Stat* 3(35). Available: http://www.cdc.gov/nchs/data/series/sr_03/sr03_035.pdf [accessed 14 September 2015].
- Murray CJ, Barber RM, Foreman KJ, Abbasoglu Ozgoren A, Abd-Allah F, et al. 2015. Global, regional, and national disability-adjusted life years (DALYs) for 306 diseases and injuries and healthy life expectancy (HALE) for 188 countries, 1990–2013: quantifying the epidemiological transition. *Lancet* 386(10009):2145–2191.
- Murray CJ, Ezzati M, Flaxman AD, Lim S, Lozano R, Michaud C, et al. 2012. GBD 2010: a multi-investigator collaboration for global comparative descriptive epidemiology. *Lancet* 380(9859):2055–2058.
- Needleman HL, Landrigan PJ. 1981. The health effects of low-level exposure to lead. *Annu Rev Public Health* 2:277–298.
- Newbold RR. 2010. Impact of environmental endocrine-disrupting chemicals on the development of obesity. *Hormones (Athens)* 9(3):206–217.
- Noyes PD, McElwee MK, Miller HD, Clark BW, Van Tiem LA, Walcott KC, et al. 2009. The toxicology of climate change: environmental contaminants in a warming world. *Environ Int* 35(6):971–986.
- Perkins DN, Brune Drisse MN, Nxele T, Sly PD. 2014. E-waste: a global hazard. *Ann Glob Health* 80(4):286–295, doi: 10.1016/j.aogh.2014.10.001.
- Pronczuk J, Bruné MN, Gore F. 2011. Children's environmental health in developing countries. In: *Encyclopedia of Environmental Health* (Nriagu JO, ed). Burlington, MA:Elsevier, 601–610.
- Rauh V, Arunajadai S, Horton M, Perera F, Hoepner L, Barr DB, et al. 2011. Seven-year neurodevelopmental scores and prenatal exposure to chlorpyrifos, a common agricultural pesticide. *Environ Health Perspect* 119(8):1196–1201, doi: 10.1289/ehp.1003160.
- Siegel RL, Miller KD, Jemal A. 2016. Cancer statistics, 2016. *CA Cancer J Clin* 66(1):7–30, doi:10.3322/caac.21332.
- Sly PD, Neira M, Collman G, Carpenter DO, Landrigan PJ, Van Den Berg M, et al. 2014. Networking to advance progress in children's environmental health. *Lancet Glob Health* 2(3):e129–e130.
- Smith AH, Marshall G, Liaw J, Yuan Y, Ferreccio C, Steinmaus C. 2012. Mortality in young adults following in utero and childhood exposure to arsenic in drinking water. *Environ Health Perspect* 120(11):1527–1531, doi: 10.1289/ehp.1104867.
- Spitz P. 2003. *Chemical Industry at the Millennium: Maturity, Restructuring, and Globalization*. Philadelphia, PA:Chemical Heritage Foundation.
- Suk WA, Ruchirawat KM, Balakrishnan K, Berger M, Carpenter D, Damstra T, et al. 2003. Environmental threats to children's health in Southeast Asia and the Western Pacific. *Environ Health Perspect* 111(10):1340–1347.
- Trasande L, Attina TM, Blustein J. 2012. Association between urinary bisphenol A concentration and obesity prevalence in children and adolescents. *JAMA* 308(11):1113–1121.
- Trasande L, Liu Y. 2011. Reducing the staggering costs of environmental disease in children, estimated at \$76.6 billion in 2008. *Health Aff (Millwood)* 30(5):863–870.
- United Nations Development Programme. 2015. *Human Development Report 2015*. Available: http://hdr.undp.org/sites/default/files/2015_human_development_report_1.pdf [accessed 15 January 2016].
- Weiss B. 1982. Food additives and environmental chemicals as sources of childhood behavior disorders. *J Am Acad Child Psychiatry* 21(2):144–152.
- WHO (World Health Organization). 2005. *Indoor Air Pollution, Health, and the Burden of Disease. Indoor Air Thematic Briefing 2*. Geneva, Switzerland:WHO. Available: <http://www.who.int/indoorair/info/briefing2.pdf> [accessed 25 January 2016].
- WHO. 2009. *Global Health Risks: Mortality and Burden of Disease Attributable to Selected Major Risks*. Geneva, Switzerland:WHO. Available: http://www.who.int/healthinfo/global_burden_disease/GlobalHealthRisks_report_full.pdf [accessed 5 October 2015].
- WHO. 2010. *Childhood lead poisoning*. Available: <http://www.who.int/ceh/publications/leadguidance.pdf> [accessed 17 January 2016].
- WHO. 2014a. *Burden of Disease from Household Air Pollution for 2012: Summary of Results*. Geneva, Switzerland:World Health Organization. Available: http://www.who.int/phe/healthinfo/outdoorair/databases/FINAL_HAP_AAP_BoD_24March2014.pdf [accessed 14 September 2015].
- WHO. 2014b. *Global Health Observatory (GHO) Data Repository. Estimated Deaths, Data by Region*. Geneva:World Health Organization. Available: <http://apps.who.int/gho/data/view.main.14117?lang=en> [accessed 18 January 2016].
- WHO. 2014c. 2011. *Global Health Sector Strategy on HIV/AIDS 2011–2015*. Available: http://apps.who.int/iris/bitstream/10665/44606/1/9789241501651_eng.pdf [accessed 18 January 2016].
- WHO. 2014d. *7 million premature deaths annually linked to air pollution*. WHO Media Centre [news release], 25 March 2014. Geneva, Switzerland:World Health Organization. Available: <http://www.who.int/mediacentre/news/releases/2014/air-pollution/en/> [accessed 18 January 2016].