

UTAH PHYSICIANS FOR A HEALTHY ENVIRONMENT

2017 REPORT ON AIR POLLUTION AND HEALTH RESEARCH



When UPHE was formed in 2007, we committed ourselves first and foremost to delivering a message on air pollution and public health based strictly on the science. As we searched through the research, we were all struck by the parallels between the health consequences of air pollution and those of cigarette smoking. The parallels between those two types of pollution exposure have only grown closer in the last 11 years. What follows is a summary of key medical research from the past year, and a reiteration and expansion of concepts that have been strengthened by this research.

* * *

We have often recited the evidence that there is no safe level of air pollution. Even at levels that fall far below national standards, indeed even the levels that approach our ability to measure them in the atmosphere, we see statistical and biologic evidence of impaired health.

The "no safe level of pollution" tenet and the parallel to cigarette smoking was solidified in a study published two weeks ago called a meta-analysis (i.e. a combining of the statistical findings of over 141 previous studies), which showed this remarkable result: Smoking one cigarette a day increases a person's risk for coronary artery heart disease 48-74% for men, and even more for women, 57-119%. The risks for strokes were a little less but showed a similar pattern. Overall the combined risks of smoking just one cigarette a day increases the risk of stroke and heart attack much more than we would have previously thought, specifically, almost half of the risk of smoking a full pack a day, i.e. 20 cigarettes.¹ Stated another way, the first few cigarettes smoked do far more damage per cigarette than the rest of the pack. The inevitable conclusion to be drawn is that reducing smoking is not enough to protect one's health, even one cigarette a day matters.



Like with cigarettes, the most recent air pollution research points in the same direction. There is a non-linear relationship between air pollution and impaired health. Per unit of air pollution, the health consequences are even greater at low levels. This fact has public policy implications. All air pollution matters, there is no "off season" for air pollution, and while current air quality standards are helpful, they are inadequate in protecting public health. All air pollution that can be removed from the Wasatch Front airshed will benefit public health regardless of the baseline air quality.

We have divided the past year's research into several categories below.

HEART, LUNGS, AND BLOOD VESSELS

Air pollution's effect on the heart and blood vessels is central to its overall impact on human health. While air pollution is only one of several risk factors, the cumulation of a person's risk factors are multiplicative rather than additive, which increases the urgency of reducing all of them.²

About 10 years ago it became well established that acute and chronic particulate pollution exposure were independent risk factors for heart disease and strokes because they inflamed the blood vessels. In the last 5 years we learned that acute particulate exposure can exert its effects through the microcirculation, the tiniest part of the blood vessel network, too small for interventions like surgery or angioplasties, and we learned that the diameter of these tiny blood vessels is almost immediately narrowed by acute air pollution inhalation. ³

The common denominators of damage to the blood vessels and probably the other organs as well, are twofold. One, chemical mediators of inflammation are triggered in the lung by inhaling harmful gases and particulate pollution and then distributed throughout the blood stream and eventually the entire body. Two, in the case of particulate pollution, the particles themselves, and in some cases atmospheric chemicals attached to those particles, are translocated from the lungs to the blood stream and then delivered to the rest of the body in like manner.

This past year we learned important details about what happens to air pollution nanoparticles once inhaled. Within 15 minutes these particles are detectable in the blood, liver and urine, and they can remain in the body for as long as three months or more. The very smallest particles, 5 nanometers, were the ones most likely to be distributed throughout the body.

Furthermore, when inhaled, particles preferentially accumulate in the lining of the blood vessels, specifically at sites of existing atherosclerosis and inflammation, which is likely to contribute further to the narrowing of critical blood vessels and increases the risk of plaques and debris rupturing and obstructing blood flow downstream. This new information provides a clear blueprint for the link between air pollution and heart attacks and strokes.⁴



We learned that even in healthy, college-age students, particulate air pollution at levels typical of our winter inversions, activates a complex interaction between the brain, pituitary gland, nervous system, adrenal glands, and kidneys which results in elevated blood pressure, production of stress hormones, inflammation, insulin resistance, and increased serum cholesterol and fatty acids. ⁵

No doubt this is a primary mechanism by which chronic exposure increases blood pressure and shortens life span.⁶

We learned more about air pollution's adverse impact on the pumping mechanism of the heart, ⁷ increasing vulnerability to heart failure, ⁸ and the compromising of the heart's ability to maintain a normal rhythm. ⁹

More studies were published showing air pollution increased rates of stroke ^{10,11} and increased rates of hospitalization for all causes, but especially for respiratory and heart problems. ¹² Like other studies on mortality, the rate of increase <u>per unit of exposure</u>, was even greater at PM 2.5 levels below 8 ug/m3. The current annual EPA standard is 12. Note that background levels of PM2.5 are about 5 ug/m3.

New clinical studies provided increased evidence of what these biological processes do to individual and public health. Multiple studies demonstrated that air pollution causes premature death.^{13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23} Research published in 2017 continues to expand the body of work demonstrating a strong connection between exposure to outdoor air pollution and adverse effects on lung function in infants, older adults, in both healthy individuals and those with asthma or smoking related lung disease (COPD).²⁴ Air pollution is directly toxic to the respiratory tract, degrading the inside lining of the nose, ²⁵ and causing DNA damage and cell death within the lungs. ²⁶

Overwhelming evidence from around the world shows that both long-term and short-term exposure to pollution at the levels frequently encountered along the Wasatch Front can both cause and exacerbate lung disease across all ages. Levels of pollution below current EPA standards still cause important health effects, including increased risk of death. These effects are seen even following short-term exposure (on the order of a few days) to PM2.5 levels below 25 mcg/m³.

What is now a landmark study, published in one of the most prestigious medical journals, found increased mortality in older adults following even modest increases in PM2.5 levels. These effects were particularly evident in those at the lower end of the socioeconomic spectrum. ²⁷ Similarly, an important study in the New England Journal of Medicine showed that chronic exposure to what we currently label as "acceptable" levels of PM2.5 or ozone was associated with increased risk of death for older individuals. ²⁸ These studies add confirmation to previous research that there is no safe level of air pollution.

Another important study evaluated the effect of taking a 2-hour walk in two different environments on symptoms and chemical markers of vascular health in older individuals with and without heart and lung



disease. Overall, walking in a low pollution environment had significant health benefits that were lost when walking in an urban environment with higher levels of pollution. ²⁹ There were quantitative relationships between levels of individual pollutants (such as PM2.5) and these effects. Thus, air pollution appears to counteract the beneficial effects of even mild exercise.

Physician researchers at the University of Utah and Intermountain Medical Center found that the risk of pneumonia and the severity of this illness (including ICU admission and need for critical care) was increased in response to a few days of increased PM2.5 levels.³⁰ Again, these effects were seen even at levels below EPA standards. These researchers identified the health care costs associated with these cases of severe pneumonia and estimate that improvements in air quality would both improve health and save millions of dollars in healthcare costs.

Ozone, our primary summer pollutant, increases the risk of hospitalization for heart attacks and has most of the same consequences as particulate pollution. ^{31, 32}

ENDOCRINE DISORDERS

Numerous new studies added to the evidence that air pollution contributes to metabolic disorders, especially type II diabetes. ^{33,34,35,36,37,38} Many of these studies found that pollution in utero can set the stage for obesity, type II diabetes and impaired glucose metabolism later in life. ^{39,40,41,42}

CANCER

The World Health Organization declared air pollution the most important environmental cause of cancer. Several new studies strengthened the relationship between air pollution and cancer, i.e. liver cancer, ⁴³ survival of liver cancer, ⁴⁴ breast cancer, ^{45, 46, 47, 48} lung and bladder cancer, ^{49,50} and malignant brain tumors. ⁵¹

REPRODUCTIVE HEALTH

New studies added to previous research showing air pollution impairs all aspects of human reproduction, likely via endocrine and hormonal impacts of chemicals attached to particulate matter. Sperm production and sperm quality are affected, ^{53, 54} in particular, a decrease in sperm carrying the y chromosome, which translates into a drop in the percentage of male newborns. ⁵⁵ Air pollution also affects menstruation in teenage girls. ⁵⁶

BRAIN

Beyond air pollution's connection to strokes, more and more studies are showing that air pollution has a profound adverse impact on the brain at every stage of life, from as early as the first trimester to the very elderly.



We now have a second study showing that tiny pollution nanoparticles from fossil fuel combustion end up inside our brains. Once there they can cause brain damage, disrupting cellular and intracellular architecture. This undoubtedly contributes to the many clinical studies that show impaired brain function, loss of memory, loss of intellectual abilities, behavior problems, and more degenerative brain diseases in people exposed to more air pollution. ⁵⁷

Two more studies showed a connection between air pollution and autism.^{58, 59} The first study showed that the affect could be mitigated by maternal extra consumption of folic acid during pregnancy. Attention span and memory in school children is impaired by the air pollution they breathe on the way to school. ^{60,61,62} Air pollution is associated with increases in the delinquent behavior of adolescents, much like lead, ⁶³ and we should be just as concerned about lead in the air our children breathe as we are about lead in the water they drink.⁷⁹ Air pollution and a more deprived social environment combine to contribute to lower intellectual scores in children. ⁶⁴

More studies showed air pollution's strong association with Alzheimer's, dementia, and the anatomic changes in the brain associated with these disorders. ^{65,66,67,68} Air pollution has been calculated to be responsible for about 20% of Alzheimer's disease overall, adding an especially strong risk for women, increasing their chance of the disease about 100%. ⁶⁹ There is emerging evidence that air pollution aggravates symptoms of Parkinson's disease. ⁷⁰

Air pollution is associated with higher risk for developing and seeking treatment for mental disorders like anxiety and depression and a much higher mortality risk for those with mental health and behavioral disorders, including suicide. ^{71,72,73,74,75,76}

CHILDREN

Most of the increased mortality from air pollution occurs in adults. Far fewer studies have been done exploring possible increased rates of infant mortality. A new study adds significantly to the evidence that air pollution also contributes to infant mortality. Almost 500,000 infants were tracked for over 6 years. Researchers found a very strong association between small increases in PM2.5 and total infant deaths, respiratory deaths, and SIDS. Specifically, 1.3 ug/m3 increases in PM2.5 increased the rate of these outcomes between 200% and 300%. ⁷⁷

This distressing outcome might be furthered by what we learned about how inhalation of particles is different depending on the age of the person. Particle deposition is highest in an infant, less in an older child, and lowest in an adult. ⁷⁸



PREGNANCY, FETAL DEVELOPMENT

Numerous studies added to the now overwhelming evidence that air pollution is a significant risk factor for just about every type of pregnancy complication and adverse outcome--premature birth, low birth weight, hypertension of pregnancy, intrauterine growth retardation, smaller placentas, reduction in the placental blood supply, still births and birth defects. ^{79,80,81,82,83,84,85,86,87,88,89,90,91,92,93,94} Even air pollution exposure of a mother prior to conception has been shown to cause an increase in certain types of birth defects, especially of the heart. ⁹⁵

Particulate pollution has been linked to 2.7 million preterm births per year, 18% of all pre-term births in the US.⁹⁶ This represents a substantial public health menace because it increases the risk of numerous adult onset diseases, impaired organ development, and an enormous economic liability.

Both chronic and acute ozone exposure during pregnancy increase the risk of still birth, as much as 39% even at levels below the EPA's standards. Even the ozone levels in the week prior to delivery increase the risk. Extrapolating from this research to Utah indicates about 100 still births a year occur in Utah due to our ozone, not counting the risk of PM2.5. ⁹⁷

There is growing evidence that the most potent toxin in urban air pollution, especially regarding pregnancy and fetal development, may be the chemicals attached to particulate matter, like PAHs (polycyclic aromatic hydrocarbons), rather than the particles themselves. ⁹⁸ This is more evidence that not all air pollution is created equal, and we should be paying much more attention to those sources that create high levels of PAH pollution--i.e. wood smoke, and industrial pollution.

TOXICITY TO GENES

Because almost all organs are at critical and vulnerable stages of development during pregnancy, in addition to altering pregnancy outcomes, even brief air pollution exposure during pregnancy can have lifelong consequences to a baby in utero, setting the stage for chronic diseases later on during childhood and adulthood, and a shortened life expectancy. The biological mechanism centers on the alteration of genes, and gene expression or "epigenetics," a term used to describe the microbiologic environment that surrounds the genes, and molecules that are attached to them, and can program their activity. Like genes, epigenetic changes can be passed on to subsequent generations. Numerous studies in 2017 added strong evidence for this phenomenon having real clinical significance.

Telomeres are repeating sequences of DNA at the ends of chromosomes that provide a marker of biological aging or life expectancy. Another new study showed that prenatal air pollution is associated with shorter telomere length in newborns, biologic evidence that their life expectancy has been shortened even before they are born. ⁹⁹

In utero exposure to air pollution, specifically diesel exhaust, has been shown in animals to lead to heart



failure in adulthood, and the likely mechanism is the alteration of genes that play a role in the function of heart cells. ¹⁰⁰ Air pollution exposure of a pregnant mother increases the likelihood of obesity later in childhood, ¹⁰¹ and can affect the function of the thyroid gland which is critical to normal growth and brain development. ¹⁰²

Exposure after birth can continue to alter gene and epigenetic function, providing a mechanism for impaired organ function and inflammation. ^{103, 104, 105, 106, 107, 108} The window of opportunity for air pollution to shorten telomere length and therefore life expectancy continues with exposure during childhood and adolescence ^{109 and} perhaps throughout adulthood.

MISCELLANEOUS EFFECTS

The most toxic type of particulate pollution is the ultrafine subset, i.e. less than 0.1 micron in size. Ultrafine pollution is associated with increased rates of inflammatory bowel disease, ¹¹⁰ and another study reveals at least one likely mechanism. When ingested, ultrafine pollution alters the microbial makeup of the bowel, and increased atherogenic lipid metabolites. ¹¹⁴

In 2017 we saw the first study we know of to show that osteoporosis is associated with air pollution exposure, ^{111 and} we saw the beginning of research that shows air pollution impairs kidney function. ¹¹² Given that inflammation is the biological pathway through which air pollution causes many diseases, for a new study to find an exacerbation of lupus with air pollution is not at all surprising. ¹¹³

A NEW RECOMMENDATION FROM THE RESEARCH

We'd like to finish our report on a positive note. There is sufficient research to recommend anti-oxidants as part of a healthy diet to help offset the inflammation triggered by air pollution. A new study in mice shows that an anti-oxidant in grapefruit reduces the DNA damage and oxidative stress in heart cells, and the tendency for blood clot formation caused by diesel exhaust. ^{115 This} is just one study, but we don't see any down side to eating more grapefruit, and it just might help.

UPHE is a 501(c)(3) tax-exempt, charitable organization. Tax ID# 80-0774496 423 W 800 S, Suite A108, Salt Lake City, UT 84101 Phone: 385.707.3677 www.uphe.org www.facebook.com/utahphysiciansforahealthyenvironment



REFERENCES

- 1. Hackshaw A, et al. Low cigarette consumption and risk of coronary heart disease and stroke: metaanalysis of 141 cohort studies in 55 study reports. BMJ 2018;360:j5855 http://dx.doi.org/10.1136/bmj.j5855
- 2. Stamler J, Neaton, Stamler R, Neaton DN. Blood Pressure, Systolic and Diastolic, and Cardiovascular Risks. Us Population Data. Arch Interm Med. 1993;153:598–615. [PubMed]
- 3. Louwies T, et al. Retinal Microvascular Responses to Short-Term Changes in Particulate Air Pollution in Healthy Adults. Environ Health Perspect. 2013 Sep; 121(9): 1011–1016.
- 4. Miller MR, et al. Inhaled Nanoparticles Accumulate at Sites of Vascular Disease. ACS Nano. 2017 Apr 26. doi: 10.1021/acsnano.6b08551. [Epub ahead of print]
- 5. Li H, et al. Particulate Matter Exposure and Stress Hormone Levels A Randomized, Double-Blind, Crossover Trial of Air Puri cation Circulation. 2017;136:618–627. DOI: 10.1161/CIRCULATIONAHA.116.026796
- 6. Magalhaes S, et al. Impacts of exposure to black carbon, elemental carbon, and ultrafine particles from indoor and outdoor sources on blood pressure in adults: A review of epidemiological evidence. Environ Res. 2017 Nov 28;161:345-353. doi: 10.1016/j.envres.2017.11.030. [Epub ahead of print]
- 7. Yang WY, et al. Left ventricular function in relation to chronic residential air pollution in a general population. Â Eur J Prev Cardiol. 2017 Jan 1:2047487317715109. doi: 10.1177/2047487317715109. [Epub ahead of print]
- 8. $S\tilde{A}_{s}$, rensen M_{s} , et al. Long-Term Exposure to Road Traffic Noise and Nitrogen Dioxide and Risk of Heart Failure: A Cohort Study. Environ Health Perspect. 2017 Sep 26;125(9):097021. doi: 10.1289/EHP1272.
- 9. Carll AP, et al. Inhaled ambient-level traffic-derived particulates decrease cardiac vagal influence and baroreflexes and increase arrhythmia in a rat model of metabolic syndrome. Part Fibre Toxicol. 2017 May 25;14(1):16. doi: 10.1186/s12989-017-0196-2.
- 10. Guo P, et al. Ambient Air Pollution and Risk for Ischemic Stroke: A Short-Term Exposure Assessment in South China. Â Int J Environ Res Public Health. 2017 Sep 20;14(9). pii: E1091. doi: 10.3390/ijerph14091091.
- 11. Guan T, et al. Differential Susceptibility in Ambient Particle-Related First-Ever Stroke Onset Risk: Findings From a National Case-Crossover Study. Am J Epidemiol. 2018 Jan 17. doi: 10.1093/aje/kwy007.



[Epub ahead of print]

- 13. Ebenstein A, et al. New evidence on the impact of sustained exposure to air pollution on life expectancy from China's Huai River Policy. Proceedings of the National Academy of Sciences, 2017; 201616784 DOI: 10.1073/pnas.1616784114
- 14. Huang C, et al. Potential Cardiovascular and Total Mortality Benefits of Air Pollution Control in Urban China. Â Circulation. 2017 Sep 7. pii: CIRCULATIONAHA.116.026487. doi: 10.1161/CIRCULATIONAHA.116.026487. [Epub ahead of print.
- 15. Malley C, et al. Updated Global Estimates of Respiratory Mortality in Adults ≥30 Years of Age Attributable to Long-Term Ozone Exposure. Environ Health Perspect; DOI:10.1289/EHP1390
- 16. Blount, R, et al. Traffic-Related Air Pollution and All-Cause Mortality during Tuberculosis Treatment in California. Â Environ Health Perspect; DOI:10.1289/EHP1699
- 17. Zhang C, et al. Association between air pollution and cardiovascular mortality in Hefei, China: A timeseries analysis. Environ Pollut. 2017 Aug 7. pii: S0269-7491(16)32799-3. doi: 10.1016/j.envpol.2017.06.022. [Epub ahead of print]
- 18. Kim SE, et al. Associations between mortality and prolonged exposure to elevated particulate matter concentrations in East Asia. Â Environ Int. 2017 Oct 30. pii: S0160-4120(17)30909-1. doi: 10.1016/j.envint.2017.10.010. [Epub ahead of print]
- 19. Fang X, et al. Relationship between fine particulate matter, weather condition and daily non-accidental mortality in Shanghai, China: A Bayesian approach. PLoS One. 2017 Nov 9;12(11):e0187933. doi: 10.1371/journal.pone.0187933. eCollection 2017.
- 20. Holnicki P, et al. Burden of Mortality and Disease Attributable to Multiple Air Pollutants in Warsaw, Poland. Â Int J Environ Res Public Health. 2017 Nov 8;14(11). pii: E1359. doi: 10.3390/ijerph14111359.
- 21. Cappellari M, et al. Association between short- and medium-term air pollution exposure and risk of mortality after intravenous thrombolysis for stroke. J Thromb Thrombolysis. 2017 Nov 28. doi: 10.1007/s11239-017-1589-7. [Epub ahead of print]
- 22. Corrigan AE, et al. Fine particulate matters: The impact of air quality standards on cardiovascular



mortality. \hat{A} \hat{A} Environ Res. 2017 Nov 28;161:364-369. doi: 10.1016/j.envres.2017.11.025. [Epub ahead of print]

- 23. Parker JD, et al. Particulate Matter Air Pollution Exposure and Heart Disease Mortality Risks by Race and Ethnicity in the United States: 1997-2009 NHIS with Mortality Followup Through 2011. Circulation. 2017 Dec 13. pii: CIRCULATIONAHA.117.029376. doi: 10.1161/CIRCULATIONAHA.117.029376. [Epub ahead of print]
- 24. Panis L, Provost EB, Cox B, Louwies T, Laeremans M, Standaert A, Dons E, Holmstock L, Nawrot T, De Boever P. Â Short-term air pollution exposure decreases lung function: a repeated measures study in healthy adults. Environ Health. 2017 Jun 14;16(1):60. doi: 10.1186/s12940-017-0271-z.
- 25. Zhao R, et al. Nasal epithelial barrier disruption by particulate matter via tight junction protein degradation. J Appl Toxicol. 2017 Dec 13. doi: 10.1002/jat.3573. [Epub ahead of print]
- 26. de Oliveira Alves N, et al. Biomass burning in the Amazon region causes DNA damage and cell death in human lung cells. Scientific Reports, 2017; 7 (1) DOI: 10.1038/s41598-017-11024-3
- 27. Di, Q., et al., Association of Short-term Exposure to Air Pollution With Mortality in Older Adults. JAMA, 2017. 318(24): p. 2446-2456.
- 28. Di, Q., et al., Air Pollution and Mortality in the Medicare Population. N Engl J Med, 2017. 376(26): p. 2513-2522.
- 29. Sinharay, R., et al., Respiratory and cardiovascular responses to walking down a traffic-polluted road compared with walking in a traffic-free area in participants aged 60 years and older with chronic lung or heart disease and age-matched healthy controls: a randomised, crossover study. Lancet, 2017.
- 30. Pirozzi, C.S., et al., Short-Term Air Pollution and Incident Pneumonia: A Case-Crossover Study. Ann Am Thorac Soc, 2017.
- 31. Chiu HF, Weng YH, Chiu YW, Yang CY. Short-Term Effects of Ozone Air Pollution on Hospital Admissions for Myocardial Infarction: A Time-Stratified Case-Crossover Study in Taipei. J Toxicol Environ Health A. 2017 Jun 9:1-7. doi: 10.1080/15287394.2017.1321092. [Epub ahead of print]
- 32. Nuvolone D, Petri D, Voller F. The effects of ozone on human health. Environ Sci Pollut Res Int. 2017 May 25. doi: 10.1007/s11356-017-9239-3. [Epub ahead of print]
- 33. Mazidi M, et al. Ambient particulate air pollution (PM2.5) is associated with the ratio of type 2 diabetes to obesity. Sci Rep. 2017 Aug 22;7(1):9144. doi: 10.1038/s41598-017-08287-1.



- 34. Strak M, et al. Long-term exposure to particulate matter, NO2 and the oxidative potential of particulates and diabetes prevalence in a large national health survey. Environ Int. 2017 Sep 5;108:228-236. doi: 10.1016/j.envint.2017.08.017. [Epub ahead of print]
- 35. Khafaie MA, et al. Particulate matter and markers of glycemic control and insulin resistance in type 2 diabetic patients: result from Wellcome Trust Genetic study. J Expo Sci Environ Epidemiol. 2017 Dec 21. doi: 10.1038/s41370-017-0001-1. [Epub ahead of print]
- 36. Dendup T, et al. Environmental Risk Factors for Developing Type 2 Diabetes Mellitus: A Systematic Review. Int J Environ Res Public Health. 2018 Jan 5;15(1). pii: E78. doi: 10.3390/ijerph15010078.
- 37. Bai L, et al. Exposure to Ambient Ultrafine Particles and Nitrogen Dioxide and Incident Hypertension and Diabetes. Epidemiology. 2018 Jan 9. doi: 10.1097/EDE.000000000000798. [Epub ahead of print]
- 38. Tan C, et al. Long-term high air pollution exposure induced metabolic adaptations in traffic policemen. Environ Toxicol Pharmacol. 2018 Jan 5;58:156-162. doi: 10.1016/j.etap.2018.01.002. [Epub ahead of print]
- 39. Chen M, et al. Prenatal Exposure to Diesel Exhaust PM2.5 Causes Offspring & Cell Dysfunction in Adulthood. Am J Physiol Endocrinol Metab. 2017 Dec 26. doi: 10.1152/ajpendo.00336.2017. [Epub ahead of print]
- 40. Madhloum N, et al. Cord plasma insulin and in utero exposure to ambient air pollution. Environ Int. 2017 May 22. pii: S0160-4120(16)30886-8. doi: 10.1016/j.envint.2017.05.012. [Epub ahead of print]
- 41. Mao G, et al. Individual and Joint Effects of Early-Life Ambient PM2.5 Exposure and Maternal Prepregnancy Obesity on Childhood Overweight or Obesity. Environ Health Perspect; DOI:10.1289/EHP261
- 42. Alderete TL, et al. Prenatal traffic-related air pollution exposures, cord blood adipokines and infant weight. Pediatr Obes. 2017 Nov 3. doi: 10.1111/ijpo.12248. [Epub ahead of print]
- 43. Pedersen M, et al. Ambient air pollution and primary liver cancer incidence in four European cohorts within the ESCAPE project. Environ Res. 2017 Jan 17;154:226-233. doi: 10.1016/j.envres.2017.01.006. [Epub ahead of print]
- 44. Deng H, et al. Particulate matter air pollution and liver cancer survival. Int J Cancer. 2017 Jun 7. doi: 10.1002/ijc.30779. [Epub ahead of print]
- 45. Sifaki-Pistolla D, Lionis C, Koinis F, Georgoulias V, Tzanakis N; On behalf of the Cancer Registry of



- 46. White AJ, et al. Indoor Wood-Burning Stove and Fireplace Use and Breast Cancer in a Prospective Cohort Study. Environ Health Perspect. 2017 Jul 18;125(7):077011. doi: 10.1289/EHP827.
- 47. Goldberg MS, et al. The association between the incidence of postmenopausal breast cancer and concentrations at street-level of nitrogen dioxide and ultrafine particles. Environ Res. 2017 Jun 5;158:7-15. doi: 10.1016/j.envres.2017.05.038. [Epub ahead of print]
- 48. Large C, Wei Y. Geographic variations in female breast cancer incidence in relation to ambient air emissions of polycyclic aromatic hydrocarbons. Environ Sci Pollut Res Int. 2017 Jun 14. doi: 10.1007/s11356-017-9395-5. [Epub ahead of print]
- 49. Collarile P, et al. Residence in Proximity of a Coal-Oil-Fired Thermal Power Plant and Risk of Lung and Bladder Cancer in North-Eastern Italy. A Population-Based Study: 1995-2009.
- 50. Lamichhane DK, et al. Lung Cancer Risk and Residential Exposure to Air Pollution: A Korean Population-Based Case-Control Study. Yonsei Med J. 2017 Nov;58(6):1111-1118. doi: 10.3349/ymj.2017.58.6.1111.
- 51. Andersen Z, et al. Long-term Exposure to Ambient Air Pollution and Incidence of Brain Tumor: the European Study of Cohorts for Air Pollution Effects (ESCAPE). Neuro Oncol. 2017 Aug 31. doi: 10.1093/neuonc/nox163. [Epub ahead of print]
- 52. Jedrychowski WA, Majewska R, Spengler JD, Camann D, Roen EL, Perera FP. Prenatal exposure to fine particles and polycyclic aromatic hydrocarbons and birth outcomes: a two-pollutant approach. Int Arch Occup Environ Health. 2017 Feb 7. doi: 10.1007/s00420-016-1192-9. [Epub ahead of print]
- 53. Qiu L, et al. Exposure to Concentrated Ambient PM2.5 Compromises Spermatogenesis in a Mouse Model: Role of Suppression of Hypothalamus-Pituitary-Gonads Axis. Toxicol Sci. 2017 Nov 20. doi: 10.1093/toxsci/kfx261. [Epub ahead of print]
- 54. Zhou N, et al. Exposures to Atmospheric PM10 and PM10-2.5 Affect Male Semen Quality: Results of MARHCS Study. Environ Sci Technol. 2018 Jan 11. doi: 10.1021/acs.est.7b05206. [Epub ahead of print]
- 55. Radwan M, et al. Air Pollution and Human Sperm Sex Ratio. Am J Mens Health. 2018 Jan 1:1557988317752608. doi: 10.1177/1557988317752608. [Epub ahead of print]
- 56. Mahalingaiah S, et al. Perimenarchal air pollution exposure and menstrual disorders. Human



Reproduction, 2018; DOI: 10.1093/humrep/dey005

- 57. González-Maciel A, Reynoso-Robles R, Torres-Jardón R, Mukherjee PS, Calderón-Garcidueñas L. Combustion-Derived Nanoparticles in Key Brain Target Cells and Organelles in Young Urbanites: Culprit Hidden in Plain Sight in Alzheimer's Disease Development. J Alzheimers Dis. 2017 Jun 3. doi: 10.3233/JAD-170012. [Epub ahead of print]
- 58. Goodrich AJ, et al. Joint effects of prenatal air pollutant exposure and maternal folic acid supplementation on risk of autism spectrum disorder. Autism Res. 2017 Nov 9. doi: 10.1002/aur.1885. [Epub ahead of print]
- 59. Li K, et al. Early postnatal exposure to airborne fine particulate matter induces autism-like phenotypes in male rats. Toxicol Sci. 2017 Nov 7. doi: 10.1093/toxsci/kfx240. [Epub ahead of print]
- 61. Forns J, et al. Longitudinal association between air pollution exposure at school and cognitive development in school children over a period of 3.5 years. Environ Res. 2017 Aug 28;159:416-421. doi: 10.1016/j.envres.2017.08.031. [Epub ahead of print]
- 62. Alvarez-Pedrerol M, et al. Impact of commuting exposure to traffic-related air pollution on cognitive development in children walking to school. Environmental Pollution, 2017; 231: 837 DOI: 10.1016/j.envpol.2017.08.075
- 63. Younan D, et al. Longitudinal Analysis of Particulate Air Pollutants and Adolescent Delinquent Behavior in Southern California. Journal of Abnormal Child Psychology, 2017; DOI: 10.1007/s10802-017-0367-5
- 64. Lett LA, et al. The Combined Influence of Air Pollution and Home Learning Environment on Early Cognitive Skills in Children. Â Int J Environ Res Public Health. 2017 Oct 26;14(11). pii: E1295. doi: 10.3390/ijerph14111295.
- 65. Hullmann M, et al. Diesel engine exhaust accelerates plaque formation in a mouse model of Alzheimer's disease. Part Fibre Toxicol. 2017 Aug 30;14(1):35. doi: 10.1186/s12989-017-0213-5.
- 66. Chen H, et al. Exposure to ambient air pollution and the incidence of dementia: A population-based cohort study. Environment International. Volume 108, November 2017, Pages 271-277



- 67. Chen H, et al. Living near major roads and the incidence of dementia, Parkinson's disease, and multiple sclerosis: a population-based cohort study. Published: 04 January 2017 DOI: http://dx.doi.org/10.1016/S0140-6736(16)32399-6
- 68. Linares C, et al. Short-term association between environmental factors and hospital admissions due to dementia in Madrid. Environ Res. 2016 Oct 27;152:214-220. doi: 10.1016/j.envres.2016.10.020. [Epub ahead of print]
- 69. Cacciottolo M, et al. Particulate air pollutants, APOE alleles and their contributions to cognitive impairment in older women and to amyloidogenesis in experimental models. Translational Psychiatry (2017) 7, e1022; doi:10.1038/tp.2016.280 Published online 31 January 2017.
- 70. Lee H, Myung W, Kim DK, Kim SE, Kim CT, Kim H. Short-term air pollution exposure aggravates Parkinson's disease in a population-based cohort. Sci Rep. 2017 Mar 16;7:44741. doi: 10.1038/srep44741.
- 71. Ho HC, et al. Spatiotemporal influence of temperature, air quality, and urban environment on cause-specific mortality during hazy days. Environment International Volume 112, March 2018, Pages 10-22
- 72. Jia Z, et al. Exposure to Ambient Air Particles Increases the Risk of Mental Disorder: Findings from a Natural Experiment in Beijing. Int J Environ Res Public Health. 2018 Jan 19;15(1). pii: E160. doi: 10.3390/ijerph15010160.
- 73. Oudin A, et al. The association between daily concentrations of air pollution and visits to a psychiatric emergency unit: a case-crossover study. Environ Health. 2018 Jan 10;17(1):4. doi: 10.1186/s12940-017-0348-8.
- 74. Casas L, et al. Does air pollution trigger suicide? A case-crossover analysis of suicide deaths over the life span. European Journal of Epidemiology. November 2017, Volume 32, Issue 11, pp 973–981
- 75. Lin H, et al. Exposure to air pollution and tobacco smoking and their combined effects on depression in six low- and middle-income countries. Br J Psychiatry. 2017 Aug 10. pii: bjp.bp.117.202325. doi: 10.1192/bjp.bp.117.202325. [Epub ahead of print]
- 76. Pun VC, et al. Association of Ambient Air Pollution with Depressive and Anxiety Symptoms in Older Adults: Results from the NSHAP Study. Environ Health Perspect; DOI:10.1289/EHP494
- 77. Son JY, et al. Pregnancy and Lifetime Exposure to Fine Particulate Matter and Infant Mortality in Massachusetts, 2001-2007. Am J Epidemiol. 2017 Nov 7:1-9. doi: 10.1093/aje/kwx015. [Epub ahead of



print]

- 78. Deng Q, et al. Particle deposition in tracheobronchial airways of an infant, child and adult. Sci Total Environ. 2017 Aug 27;612:339-346. doi: 10.1016/j.scitotenv.2017.08.240. [Epub ahead of print]
- 79. Gonzalez-Maciel A, Reynoso-Robles R, Torres-Jardan R, Mukherjee PS, Calderon-Garcidueas L. Combustion-Derived Nanoparticles in Key Brain Target Cells and Organelles in Young Urbanites: Culprit Hidden in Plain Sight in Alzheimer's Disease Development. J Alzheimers Dis. 2017 Jun 3. doi:10.3233/JAD-170012. [Epub ahead of print]
- 80. Liu C, et al. Different exposure levels of fine particulate matter and preterm birth: a meta-analysis based on cohort studies. Environ Sci Pollut Res Int. 2017 Jun 15. doi: 10.1007/s11356-017-9363-0. [Epub ahead of print]
- 81. Hettfleisch, K, et al. Short-Term Exposure to Urban Air Pollution and Influences on Placental Vascularization Indexes. Environ Health Perspect; DOI:10.1289/EHP300
- 82. Zhu Y, et al. Ambient air pollution and risk of gestational hypertension. Am J Epidemiol. 2017 May 4. doi: 10.1093/aje/kwx097. [Epub ahead of print]
- 83. Basu R, et al. Association between PM2.5 and PM2.5 Constituents and Preterm Delivery in California, 2000-2006. Paediatr Perinat Epidemiol. 2017 Jul 21. doi: 10.1111/ppe.12380. [Epub ahead of print]
- 84. Clemens T, Turner S, Dibben C. Maternal exposure to ambient air pollution and fetal growth in North-East Scotland: A population-based study using routine ultrasound scans. Environ Int. 2017 Jul 25;107:216-226. doi: 10.1016/j.envint.2017.07.018. [Epub ahead of print]
- 85. Blum J, et al. Exposure to Ambient Particulate Matter during Specific Gestational Periods Produces Adverse Obstetric Consequences in Mice. Environmental Health Perspectives, July 2017 DOI: 10.1289/EHP1029
- 86. Soto SF, et al. Exposure to fine particulate matter in the air alters placental structure and the reninangiotensin system. PLoS One. 2017 Aug 18;12(8):e0183314. doi: 10.1371/journal.pone.0183314. eCollection 2017.
- 87. Reis MMD, et al. Air pollution and low birth weight in an industrialized city in Southeastern Brazil, 2003-2006. Rev Bras Epidemiol. 2017 Apr-Jun;20(2):189-199. doi: 10.1590/1980-5497201700020001.
- 88. Clemens T, et al. Maternal exposure to ambient air pollution and fetal growth in North-East Scotland: A population-based study using routine ultrasound scans. Environ Int. 2017 Oct;107:216-226.



doi: 10.1016/j.envint.2017.07.018. Epub 2017 Jul 25.

- 89. Kingsley SL, et al. Maternal residential air pollution and placental imprinted gene expression. Â Environ Int. 2017 Sep 5;108:204-211. doi: 10.1016/j.envint.2017.08.022. [Epub ahead of print]
- 90. Wang L, et al. The association between cooking oil fume exposure during pregnancy and birth weight: A prospective mother-child cohort study. Sci Total Environ. 2017 Sep 4;612:822-830. doi: 10.1016/j.scitotenv.2017.08.031. [Epub ahead of print]
- 91. Wu H, et al. Exposure to fine particulate matter during pregnancy and risk of term low birth weight in Jinan, China, 2014-2016. Int J Hyg Environ Health. 2017 Oct 28. pii: S1438-4639(17)30399-1. doi: 10.1016/j.ijheh.2017.10.013. [Epub ahead of print]
- 92. Ji Y, et al. Association between exposure to particulate matter during pregnancy and birthweight: a systematic review and a meta-analysis of birth cohort studies. J Biomed Res. 2017 Nov 1. doi: 10.7555/JBR.31.20170038. [Epub ahead of print]
- 93. Dutta A, et al. Household air pollution and chronic hypoxia in the placenta of pregnant Nigerian women: A randomized controlled ethanol Cookstove intervention. Sci Total Environ. 2017 Nov 14;619-620:212-220. doi: 10.1016/j.scitotenv.2017.11.091. [Epub ahead of print]
- 94. Smith R, et al. Impact of London's road traffic air and noise pollution on birth weight: retrospective population based cohort study. BMJ, 2017; j5299 DOI: 10.1136/bmj.j5299
- 95. Ren S, et al. Periconception Exposure to Air Pollution and Risk of Congenital Malformations. The Journal of Pediatrics, 2017; DOI: 10.1016/j.jpeds.2017.09.076
- 96. Malley CS, et al. Preterm birth associated with maternal fine particulate matter exposure: A global, regional and national assessment. Environment International, 2017; DOI: 10.1016/j.envint.2017.01.023
- 97. Mendola P, Ha S, Pollack AZ, Zhu Y, Seeni I, Kim SS, Sherman S, Liu D. Chronic and Acute Ozone Exposure in the Week Prior to Delivery Is Associated with the Risk of Stillbirth.Int J Environ Res Public Health. 2017 Jul 6;14(7). pii: E731. doi: 10.3390/ijerph14070731.
- 98. Jedrychowski WA, Majewska R, Spengler JD, Camann D, Roen EL, Perera FP. Prenatal exposure to fine particles and polycyclic aromatic hydrocarbons and birth outcomes: a two-pollutant approach. Int Arch Occup Environ Health. 2017 Feb 7. doi: 10.1007/s00420-016-1192-9. [Epub ahead of print]
- 99. Martens DS, et al. Prenatal Air Pollution and Newborns' Predisposition to Accelerated Biological Aging. JAMA Pediatr. 2017 Oct 16. doi: 10.1001/jamapediatrics.2017.3024. [Epub ahead of print]



- 100. Goodson J, et al. In utero exposure to diesel exhaust particulates is associated with an altered cardiac transcriptional response to transverse aortic constriction and altered DNA methylation. The FASEB Journal, 2017; fj.201700032R DOI: 10.1096/fj.201700032R
- 101. Mao G, et al. Individual and Joint Effects of Early-Life Ambient PM2.5 Exposure and Maternal Prepregnancy Obesity on Childhood Overweight or Obesity. Environ Health Perspect; DOI:10.1289/EHP261
- 102. Janssen BG, et al. Fetal Thyroid Function, Birth Weight, and in Utero Exposure to Fine Particle Air Pollution: A Birth Cohort Study. Environ Health Perspect; DOI:10.1289/EHP508
- 103. Huang Q, et al. Fine particulate matter 2.5 exerted its toxicological effect by regulating a new layer, long non-coding RNA. Sci Rep. 2017 Aug 24;7(1):9392. doi: 10.1038/s41598-017-09818-6.
- 104. de Oliveira A, et al. Biomass burning in the Amazon region causes DNA damage and cell death in human lung cells. Sci Rep. 2017 Sep 7;7(1):10937. doi: 10.1038/s41598-017-11024-3.
- 105. Wang C, et al. Acute Inflammation Following Personal Exposure to Fine-particulate Air Pollution. Â Am J Epidemiol. 2017 Aug 17. doi: 10.1093/aje/kwx277. [Epub ahead of print]
- 106. Lai CH, et al. Exposure to fine particulate matter causes oxidative and methylated DNA damage in young adults: A longitudinal study. Sci Total Environ. 2017 Apr 23;598:289-296. doi: 10.1016/j.scitotenv.2017.04.079. [Epub ahead of print]
- 107. Ding R, et al. Dose- and time- effect responses of DNA methylation and histone H3K9 acetylation changes induced by traffic-related air pollution. Sci Rep. 2017 Mar 3;7:43737. doi: 10.1038/srep43737.
- 108. Chen R, et al. Fine Particulate Air Pollution and the Expression of microRNAs and Circulating Cytokines Relevant to Inflammation, Coagulation, and Vasoconstriction. Environ Health Perspect. 2018 Jan 17;126(1):017007. doi: 10.1289/EHP1447.
- 109. Lee E, et al. Â Traffic-Related Air Pollution and Telomere Length in Children and Adolescents Living in Fresno, CA. Journal of Occupational and Environmental Medicine, 2017; 59 (5): 446 DOI: 10.1097/JOM.0000000000000996 Â
- 110. van der Sloot KWJ, Amini M, Peters V, Dijkstra G, Alizadeh BZ. Inflammatory Bowel Diseases: Review of Known Environmental Protective and Risk Factors Involved. Inflamm Bowel Dis. 2017 Aug 2. doi: 10.1097/MIB.000000000001217. [Epub ahead of print]



- 111. Prada, D, Zhong, J, Colicino, E et al. Association of air particulate pollution with bone loss over time and bone fracture risk: analysis of data from two independent studies. Lancet Planet Health. 2017; 1: e337–e347
- 112. Bowe B, et al. Particulate Matter Air Pollution and the Risk of Incident CKD and Progression to ESRD. J Am Soc Nephrol. 2017 Sep 21. pii: ASN.2017030253. doi: 10.1681/ASN.2017030253. [Epub ahead of print]
- 113. Alves AGF, et al. Influence of air pollution on airway inflammation and disease activity in childhood-systemic lupus erythematosus. Clin Rheumatol. 2017 Nov 2. doi: 10.1007/s10067-017-3893-1. [Epub ahead of print]Â
- 114. Li R, et al. Ambient Ultrafine Particle Ingestion Alters Gut Microbiota in Association with Increased Atherogenic Lipid Metabolites. Sci Rep. 2017 Feb 17;7:42906. doi: 10.1038/srep42906.
- 115. Nemmar A, et al. Thrombosis, systemic and cardiac oxidative stress and DNA damage induced by pulmonary exposure to diesel exhaust particles, and the effect of nootkatone thereon. Am J Physiol Heart Circ Physiol. 2018 Jan 5. doi: 10.1152/ajpheart.00313.2017. [Epub ahead of print]