

# *Deleterious Effects of Air Pollution on the Respiratory Epithelium.*

*Elnara Márcia Negri and Nilsa Regina Damaceno Rodrigues*

The air we breathe is becoming more and more toxic. Despite all studies and warnings from the scientific community on the subject, air pollution has been negligence as a morbidity factor and is making victims in megacity in the world. The worsening air quality in metropolitan areas is directly related to vehicular emissions, especially to fossil fuels vehicles.

In São Paulo City (Brazil), more than 6 million diesel-powered vehicles discharge approximately 1 ton of particulate matter into the air among other substances (Martins *et al.* 2017). These particles cause exacerbation of pre-existing diseases involving the airways such as COPD and asthma. In addition, the particulate matter, called PM 2.5 and PM 0.1, is able to penetrate the microcirculation, cross the alveolar barrier and cause cardiovascular, neurological and placental diseases with serious consequences on human health (Saldiva *et al.* 2002, Pope and Dockery 2006, Delfino *et al.* 2009 and Tsukue *et al.* 2010). In the atmosphere, more than 3000 different chemical components are identified, including a wide variety of mutagens and carcinogens, harmful to plants, animals and humans.

Several studies have been made pointing to pollution as one of the factors that cause cardiovascular diseases (Akinaga *et al.* 2009 and Santos *et al.* 2008), others associate the increased incidence of mutagenesis in plants and increased frequency of mortality due to cardiovascular diseases and cancer, to high levels of air pollution (Mariani *et al.* 2009 and Poirier, 2017).

Recent studies have pointed to air pollution as an isolated cause of lung cancer (Nafstad *et al.* 2003). Among other pollutants, SO<sub>2</sub>, NO and NO<sub>2</sub> are routinely measured parameters to describe air quality. Organic extracts took from particulate matter contain mutagenic chemical components of different origins, according to the site studied, and the concentration of NO is a strong indicator of the pollution emitted by vehicles and the mutagenic activity induced by air pollution. Nitrogen dioxide (NO<sub>2</sub>) is emitted directly into the atmosphere by vehicle exhausts, fertilizer industries, etc. NO<sub>2</sub> is an aggressive gas to the respiratory tract and its presence in the environment is related to cases of respiratory infections; in addition, it can be converted into the lungs into nitrosamines, some of which are known to be potentially carcinogenic. This gas can participate of a series of photochemical reactions in the atmosphere, for example, in the formation of the photochemical “smog”, which reduces visibility.

The Ozone (O<sub>3</sub>) is a colorless gas that is found in two distinct layers of the atmosphere: in the stratosphere, naturally formed by the photolysis of O<sub>2</sub>. It protects life on earth by absorbing the ultraviolet radiation from the sun that reach-

es this layer. Moreover, in the troposphere ozone is formed through a complex series of reactions involving volatile organic compounds and nitrogen oxides, in the presence of sunlight. In this layer, it affects the health of the populations, irritating the respiratory system and may cause worsening of asthma and chronic pulmonary obstructive diseases (COPD). Epidemiological studies show that daily variations of fine and ultrafine particles are associated with morbidity and mortality rates in the population, mainly causing toxic effects on respiratory and cardiovascular tract (Pinheiro *et al.* 2014 and Traboulsi *et al.* 2017).

Many studies have related pollution with increasing mortality rates in the elderly, newborns and people with chronic diseases (Lin *et al.* 1999, Braga *et al.* 2001, Lin *et al.* 2004 and Farhat *et al.* 2005). Particulate matter from the exhaust of diesel engines is formed by a complex and variable mixture of organic and inorganic components, the most important being polycyclic aromatic hydrocarbons (PAHs) and metals (Vasconcellos *et al.* 2003, Carvalho-Oliveira *et al.* 2005, Laks *et al.* 2008 and De Siqueira-Bueno *et al.* 2010). Metals are responsible for reactions that generate oxidizing substances and induce inflammation and methylation of DNA. PAHs, in addition to inducing oxidative stress and inflammation, have a chemical structure very similar to estrogens and can induce effects such as obesity, increased peripheral resistance to insulin and microbial alteration, in addition to pro-neoplastic effects (Calahan *et al.* 2017, Alderete *et al.* 2017).

Air pollution causes the destruction and malfunction of the mucociliary apparatus through changes in the functioning of the mitochondria and the cytoskeleton (Seriani *et al.* 2015). Such exposure may also affect gene expression and induce apoptosis and anti-apoptosis with impacts on the regeneration of the bronchial epithelium (Seriani *et al.* 2015).

In summary, there are numerous evidences of the deleterious effects of environmental pollution on our organism. We need to urgent action to improve air quality in large cities.

## References

1. Akinaga LM, Lichtenfels AJ, Carvalho-Oliveira R, Caldini EG, Dolhnikoff M, Silva LF, Bueno HM, Pereira LA, Saldiva PH, Garcia ML. 2009. Effects of chronic exposure to air pollution from Sao Paulo city on coronary of Swiss mice, from birth to adulthood. *Toxicol Pathol.* 37 (3): 306- 14.
2. Alderete TL, Jones RB, Chen Z, Kim JS, Habre R, Lurmann F, Gilliland FD, Goran MI. 2017. Exposure to traffic-related air pollution and the composition of the gut microbiota in overweight and obese adolescents. *Environ Res.* 161:472-478.
3. Braga AL, Saldiva PH, Pereira LA, Menezes JJ, Conceição GM, Lin CA, Zanobetti A, Schwartz J, Dockery DW. 2001. Health effects of air pollution exposure on children and adolescents in São Paulo, Brazil. *Pediatr Pulmonol.* 31(2):106-13.
4. Carvalho-Oliveira R, Saiki M, Pires-Neto RC, Lorenzi-Filho G, Macchione M, Saldiva PH. 2005. Anti-oxidants reduce the acute adverse effects of residual oil fly ash on the frog palate mucociliary epithelium. *Environ Res.* 98(3):349-54.

5. de Siqueira Bueno HM, de Souza Leão Martins R, Pannuti C, dos Santos RN, Sowmy T, Barbosa Junior F, Carvalho-Oliveira R, Garcia MLB, Saldiva PHN. 2010. Metal embryotoxicity from urban particles in Sao Paulo city: An experimental study in chicken embryos. *Ecotoxicol Environ Saf.* 73(6):1385–90.
6. Delfino RJ, Staimer N, Tjoa T, Gillen DL, Polidori A, Arhami M, Kleinman MT, Vaziri ND, Longhurst J, Sioutas C. 2009. Air pollution exposures and circulating biomarkers of effect in a susceptible population: clues to potential causal component mixtures and mechanisms. *Environ Health Perspect* 117:1232–8.
7. Farhat SC1, Paulo RL, Shimoda TM, Conceição GM, Lin CA, Braga AL, Warth MP, Saldiva PH. 2005. Effect of air pollution on pediatric respiratory emergency room visits and hospital admissions. *Braz J Med Biol Res.* 38(2):227-35.
8. Laks D, de Oliveira RC, de André PA, Macchione M, Lemos M, Faffe D, Saldiva PH, Zin WA. 2008. Composition of diesel particles influences acute pulmonary toxicity: an experimental study in mice. *Inhal Toxicol* 11:1037–42.
9. Lin CA, Martins MA, Farhat SC, Pope CA 3rd, Conceição GM, Anastácio VM, Hatanaka M, Andrade WC, Hamaue WR, Böhm GM, Saldiva PH. 1999. Air pollution and respiratory illness of children in São Paulo, Brazil. *Paediatr Perinat Epidemiol.* 13(4):475-88.
10. Lin CA, Pereira LA, Nishioka DC, Conceição GM, Braga AL, Saldiva PH. 2004. Air pollution and neonatal deaths in São Paulo, Brazil. *Braz J Med Biol Res.* 37(5):765-70.
11. Mariani RL, Jorge MP, Pereira SS, Melione LP, Carvalho-Oliveira R, Ma TH, Saldiva PH. 2009. Association between micronuclei frequency in pollen mother cells of *Tradescantia* and mortality due to cancer and cardiovascular diseases: a preliminary study in Sao José dos Campos, Brazil. *Environ Pollut.* 157(6):1767-70.
12. Martins MHRB, Muramoto CA et al. 2017. Qualidade do ar no estado de São Paulo 2016/CETESB - São Paulo. CETESB, 2017. Disponível em: <<http://ar.cetesb.sp.gov.br/publicacoes-relatorios/>>. ISBN 978-85-9467-013-7. Acessado em novembro de 2017.
13. Nafstad P, Haheim LL, Oftedal B, Gram F, Holme I, Hjerermann I, Leren P. 2003. Lung cancer and air pollution: a 27 year follow up of 16 209 Norwegian men *Thorax* 58:1071–6.
14. Pope CA 3rd, Dockery DW. 2006. Health effects of fine particulate air pollution: lines that connect. *J Air Waste Manag Assoc.* 56(6):709-42.
15. Pinheiro Sde L, Saldiva PH, Schwartz J, Zanobetti A. 2014. Isolated and synergistic effects of PM10 and average temperature on cardiovascular and respiratory mortality. *Rev Saude Publica.* 48(6):881-8.
16. Poirier AE, Grundy A, Khandwala F, Friedenreich CM, Brenner DR. Cancer incidence attributable to air pollution in Alberta in 2012 *CMAJ Open.* 2017; 5(2): E524–E528.
17. Saldiva PH, Clarke RW, Coull BA, Stearns RC, Lawrence J, Murthy GG, Diaz E, Koutrakis P, Suh H, Tsuda A, Godleski JJ. 2002. Lung inflammation induced by concentrated ambient air particles is related to particle composition. *Am J Respir Crit Care Med.* 165:1610–7.
18. Santos UP, Terra-Filho M, Lin CA, Pereira LA, Vieira TC, Saldiva PH, Braga AL. 2008. Cardiac arrhythmia emergency room visits and environmental air pollution in Sao Paulo, Brazil. *J Epidemiol Community Health.* 62(3):267-72.
19. Seriani R, Junqueira MS, Toledo AC, Corrêa AT, Silva LF, Martins MA, Saldiva PH, Mauad T, Macchione M. 2015. Organic and inorganic fractions of diesel exhaust particles produce changes in mucin profile of mouse trachea explants. *J Toxicol Environ Health A* 78(4):215–25.
20. Seriani R, Junqueira MS, de Toledo AC, Martins MA, Seckler M, Alencar AM, Negri EM, Silva LF, Mauad T, Saldiva PH, Macchione M. 2015. Diesel exhaust particulates affect cell signaling, mucin profiles, and apoptosis in trachea explants of Balb/C mice. *Environ Toxicol.* 30(11):1297-308.

21. Seriani R, Junqueira MS, Carvalho-Sousa CE, Arruda AC, Martinez D, Alencar AM, Garrippo AL, Brito JM, Martins MA, Saldiva PH, Negri EM, Mauad T, Macchione M. 2015. Enriched inorganic compounds in diesel exhaust particles induce mitogen-activated protein kinase activation, cytoskeleton instability, and cytotoxicity in human bronchial epithelial cells. *Exp Toxicol Pathol.* 67(4):323-9.
22. Traboulsi H, Guerrina N, Iu M, Maysinger D, Ariya P, Baglolle CJ. 2017. Inhaled Pollutants: The Molecular Scene behind Respiratory and Systemic Diseases Associated with Ultrafine Particulate Matter. *Int J Mol Sci.* 18(2). pii: E243.
23. Tsukue N, Okumura H, Ito T, Sugiyama G, Nakajima T. 2010. Toxicological evaluation of diesel emissions on A549 cells. *Toxicol in Vitro* 24(2):363–9.
24. Vasconcellos PC, Zacarias D, Pires MAF, Pool CS, Carvalho LRF. 2003. Measurements of polycyclic aromatic hydrocarbons in airborne particles from the metropolitan area of São Paulo City, Brazil. *Atmos Environ* 37(21):3009–18.