

Review of Outdoor Air Pollution and Child Health in Hong Kong

SL LEE, H TINSLEY, J CHAU, HK LAI, TQ THACH, P IP, CB CHOW, AJ HEDLEY

Abstract

Our Air Quality Objectives (AQO) for seven air pollutants were established in 1987 under the Air Pollution Control Ordinance but have not been revised. Air pollution in Hong Kong has been continuously worsening and remains incompatible with acceptable standards of health protection for children. Although the government initiated a consultancy for AQO in 2007, revisions to the AQO will not be implemented for at least two more years, in 2014. Furthermore, the government does not accept that Hong Kong's AQO should follow the World Health Organization guidelines set in 2006, despite the high level of local and international evidence that poor quality air is detrimental to individual and population health and that the proposed new AQO will fail to provide adequate protection. We present a synopsis of the susceptibility of children to environmental pollution, international and local evidence of adverse health effects on children and call for support from paediatricians in urging the government to take actionable steps for the achievement of cleaner air to protect our child health.

Key words

Child health; Hong Kong; Outdoor air pollution

Introduction

Outdoor air pollution in Hong Kong has been a major environmental problem for more than two decades and

remains incompatible with acceptable standards of health protection for children. Our Air Quality Objectives (AQO) for seven air pollutants were established in 1987 under the Air Pollution Control Ordinance but were not revised. The government consultancy which began in 2007, following the World Health Organization (WHO) 2006 recommendations, has been criticised as a delaying tactic. However, any revision to the AQO will not be implemented by government for at least two more years, in 2014. The government does not accept that Hong Kong's AQO should follow those advised by WHO in 2006, despite the high level of evidence both locally and internationally showing that poor quality air is detrimental to individual and population health and that the proposed new AQO will not provide adequate protection.¹ What should be the roles of clinicians, in particular both paediatricians and public health practitioners, in this debate, since children are particularly vulnerable to environmental air pollution and the resulting harm can run a life-time course affecting susceptibility to illness, quality of life and life expectancy? In this report we provide a synopsis of updated evidence of the health impacts on child health attributable to outdoor air pollution and call for both paediatricians and public health experts to

Department of Paediatrics and Adolescent Medicine, LKS Faculty of Medicine, The University of Hong Kong, Pokfulam, Hong Kong, China

SL LEE (李素輪) *FHKCPaed, MPH*
H TINSLEY (丁詩妮) *FHKCPaed, FHKCCM*
P IP (葉柏強) *FHKCPaed, MRCP(UK)*
CB CHOW (周鎮邦) *FHKCPaed, FRCPC*

Department of Community Medicine and School of Public Health, LKS Faculty of Medicine, The University of Hong Kong, Pokfulam, Hong Kong, China

J CHAU (周宗欣) *MPhil, MSc*
HK LAI (黎克勤) *MSc, PhD*
TQ THACH *PhD*
AJ HEDLEY (賀達理) *FRCP, FHKCCM*

Correspondence to: Dr SL LEE

Received June 22, 2012

support and clearly state to the new Government of the HKSAR the mandatory and urgent steps required for cleaner air in Hong Kong. We earnestly urge the new government to take decisive actions in addressing and reversing, on the shortest possible time scale, the severe problem of poor air quality and its serious impact on the current and future health of children in Hong Kong.

Why Are Children More Vulnerable to Environmental Pollution?

Children are not miniature adults. Compared with adults, children have a particular vulnerability to environmental pollution (Table 1).^{2,3} First, exposure to air pollutants occurs at a critical window period from foetal to adolescent stage when tissues and organs are rapidly growing, developing

and differentiating until maturity. The resultant health effects on those exposed can be life-long. Next, a child's detoxification system is immature and less efficient in handling these toxic air pollutants. The incomplete development of the lung epithelium and frequent upper respiratory tract infection in childhood create increased permeability of the epithelial layer of the respiratory tract, resulting in increased damage for a given exposure.⁴ Molecular mechanism studies also suggest that the development of the immune system will skew towards the T-helper 2 phenotype, increasing proneness to the development of allergy in response to environmental exposure. Minute ventilation adjusted for body mass is greater in infancy (400 ml/min/kg) than adult (150 ml/min/kg) and children have greater activity levels than adults, so the resultant exposure of the lungs to air pollutants will be greater. The smaller peripheral airways in infants are also

Table 1 Vulnerability of children to environmental pollution

| Factors | Resultant effects |
|---|--|
| Biological | |
| Foetal to adolescent stage is a critical window for growth | Resultant effect due to exposure to air pollutants can be life-long |
| Immature detoxification system in foetus and young child | Inefficient in handling toxic air pollutants |
| Lung development has not completed | Increased permeability of epithelia layer of respiratory tract and increased damage per a given exposure |
| Frequent upper respiratory tract in childhood | May skew towards T-helper 2 phenotype in response to environmental exposure to air pollutants |
| Maturing immune system | |
| Physiological | |
| Minute ventilation adjusted for body mass greater in infancy | Exposure of lungs to air pollutants greater |
| Children have greater activity levels | |
| Smaller peripheral airways | More susceptible to airway obstruction |
| Proneness to mouth breathing due to enlarged adenoid and tonsils or allergic rhinitis | Loss of nasal filtering and greater deposition of air pollutants to lower airway |
| Genetic | |
| Genetic susceptibility to air pollution induced lung injury and repair | More prone to develop symptoms with exposure to air pollutants |
| Behavioural | |
| Spend more time in outdoor activities than adults | More likely to be exposed to air pollutants and develop more severe symptoms |
| Less likely to pay heed of warnings of poor air quality | |
| Less likely to stop activities when symptoms develop due to exposure to air pollution | |
| Child Protection | |
| No choice on where to live or attend school | Passive to exposure to air pollutants |

more susceptible to airway obstruction by inflammatory secretion and bronchospasm.⁵ Children, who are prone to mouth breathing due to enlarged adenoids and tonsils or allergic rhinitis, experience a loss of nasal filtering and greater deposition of air pollutants in the lower respiratory tract.⁶ There is also evidence of genetic susceptibility to air pollutant-induced lung injury and repair. Glutathione-S-transferases (GST) catalyse the conjugation of glutathione to secondary oxidation products produced by exposure of lipids to environmental oxidants such as ozone. Asthmatic children with GST M1 null and GST P1 valine/valine genotypes appear more susceptible to developing respiratory symptoms related to ozone exposure.^{7,8} Polymorphism in the genotype of tumour necrosis factor (TNF), an inflammatory cytokine, also influences the lung function response to ozone. Those with TNF-308 GG genotype had a significantly reduced risk of bronchitic symptoms with low-ozone exposure.⁹ Children's behaviour also increases their personal exposure to air pollutants as they spend more time in outdoor activities than adults. These risks to health are especially marked in the cooler months of the year when ozone, particulates and nitrogen dioxide levels are highest and there is an important interaction between seasonal and air pollutant health effects. Most importantly, children usually have no choice where they live or attend school and there is no good evidence that restricting activity on selected days in a generally high pollution environment confers any overall health protection.

Current Evidence for Adverse Health Effects of Poor Quality Air on Children

There have been abundant time series studies that captured relationships between air pollution and more severe health impacts such as hospital admissions or mortality. They have provided strong evidence to support the significant association of ambient air pollutants with hospital admission for asthma, especially in children.^{10,11} Patterns of O₃ and fine particles (PM_{2.5}) in particular were associated with severe asthma attacks requiring admission for general or intensive care in children aged 6-18 years.¹² Longitudinal studies are scarce as they are more resource demanding but they can also explore the less severe but important health impacts of air pollution that have often been overlooked. A recent longitudinal study carried out in California showed that wheeze was significantly associated with short-term exposures to NO₂ and the coarse particulate fraction of PM_{10-2.5} in children aged 6-11 years,

especially those who were sensitised to cat or common fungi, and in boys with mild intermittent asthma.¹³ Modest but consistent associations were also found between NO₂, PM_{2.5} and wood-smoke and otitis media in a large birth cohort exposed to relatively low levels of ambient air pollution.¹⁴ In recent years, there has been increasing concern about the effect of traffic-related air pollution. There is consistent evidence that living near traffic sources is associated with asthma incidence and exacerbations.¹⁵ Even in areas with good regional air quality, a cross-sectional study showed associations between current asthma in children in grade 3 to 5 and residential proximity to traffic, with highest risk for those living within 75 metres of freeways/highways.¹⁶ A hospital based longitudinal study showed that exposure to traffic-related air pollution within 300 metres of residence increased asthma severity and hospital utilisation in children younger than 18 years.¹⁷ In Hong Kong, exposure to traffic emissions is generally unavoidable in children living in urban areas.

It is noteworthy that air pollution does not just cause acute health effects but can induce long term pathophysiological damage to the developing lung in children and adolescents.¹⁸ Residential traffic-related air pollution exposure is associated with reduced lung function in schoolchildren.^{19,20} Longitudinal studies indicate that traffic exhaust contributes to the development of respiratory illness in childhood. There is substantial evidence for the development of asthma,²¹⁻²³ in addition to the aggravation of symptoms in those with asthma, and many studies have shown that early life exposure to traffic-related air pollutants is associated with new onset childhood asthma. The associations between traffic-related pollution with other allergic diseases such as allergic rhinitis and eczema were initially inconsistent but were strengthened with the findings in more recent studies.^{24,25} In addition, there is a growing body of evidence indicating the adjuvant effect of traffic-related air pollutants, including diesel exhaust particles and NO₂, in promoting allergic sensitisation to inhaled allergens.²⁶⁻³¹ There are increasing numbers of observational studies which suggest an association between traffic-related air pollution and children's cognitive function^{32,33} and neurobehavioural function^{34,35} although the underlying mechanisms await further elucidation. Traffic is also a source of toxic environmental exposures, including benzene, which may be associated with childhood leukemia. A national case-control study in France showed that acute leukaemia in children was significantly associated with higher estimates of traffic NO₂ concentrations at the place of residence (>27.7 µg/m³ versus <21.9 µg/m³) and with

the presence of a heavy-traffic road within 500 metres. There was also a significant association between acute leukaemia and a high density of heavy-traffic roads within 500 metres, with a significant positive linear trend in the association of acute leukaemia with the total length of heavy-traffic road within 500 metres.³⁶

Evidence for the adverse health effects of chronic exposure in pregnant women and young infants is accumulating. Babies born to mothers who were exposed to high levels of ambient air pollutants showed increased evidence of intrauterine growth retardation, low birth weight and preterm delivery.^{3,37-39} Increased micronuclei and bulky DNA adducts in cord blood after maternal exposures to traffic-related air pollution have been found, demonstrating that these transplacental environmental exposures induce DNA damage in newborns.⁴⁰ Meta-analysis showed an association between ambient air pollution and congenital anomalies, notably congenital heart disease, including NO₂ and SO₂ exposures with coarctation of the aorta and Tetralogy of Fallot, and PM₁₀ exposure with atrial septal defects.⁴¹ Two studies showed an association with omphalocele and PM₁₀ exposure⁴² and unspecified nervous system anomalies with black smoke.⁴³ A Korean birth cohort study demonstrated a relationship between gestational exposures to particulate matter and infant mortality for all-causes and respiratory mortality in normal birth weight infants.⁴⁴

There are numerous good quality local studies to show the adverse population health effects of air pollution, some of which are particularly related to child health. There is evidence of reduced oxygen uptake during exercise in children living in Kwun Tong and Shatin that led to impairment of physical performance during sports activities.⁴⁵ Air pollution (NO₂, PM₁₀, PM_{2.5} and O₃) in Hong Kong increases paediatric asthma admissions to hospital. This is also the first study to show the association between fine particles PM_{2.5} with increased asthma admissions to hospital.¹¹

Cross sectional and cohort studies have shown how air quality controls can lead to a reduction in respiratory symptoms and doctor visits. Bronchial hyper-responsiveness provoked by a histamine challenge in primary school children who were not symptomatic at the time of the initial and follow-up examinations, was reduced after air quality improvement. These health gains were more marked in the more polluted areas after implementation of the legislation to restrict fuel sulphur levels in July 1990.^{46,47} The legislation also led to a significant reduction in mortality in individuals aged 46 and older especially due to

respiratory and cardiovascular causes in the subsequent 5-year period.⁴⁸

High concentrations of air pollutants (NO₂, SO₂, PM₁₀, O₃) are related to higher excess risks of mortality and hospitalisation, mainly from cardiopulmonary disease.^{49,50} The damaging effect of Hong Kong's air pollution on environmental justice and health inequity has been convincingly demonstrated by the fact that those in lower socio-economic groups are most affected.^{51,52} These are landmark studies and the findings in Hong Kong and mainland China were adopted in the consensus statement which established the WHO 2006 Air Quality Guideline.⁵³

Current Situation in Hong Kong

Despite the strong evidence for the adverse health effects related to air pollution provided by both local and international studies, the Government of the HKSAR has failed to take sufficient action over the past two decades to improve air quality. Except for the ambient sulphur dioxide SO₂ level, the levels of other major criteria air pollutants have continuously deteriorated (Figure 1). As a matter of fact, Hong Kong's air quality is regarded as poor by most international standards. Even the successful dramatic reduction of ambient sulphur dioxide SO₂ level due to implementation of legislation to reduce fuel sulphur level in mid 1990 was not sustained with predictable damage to child health. There was a continuous increase in SO₂ levels from the late 1990s followed by a slow decline to a point at least 100% above the WHO limit for safer air quality (Figure 2). The 2011 roof-top SO₂ level was 13.5 µg/m³, which is 170% higher than the 5 µg/m³ level predicted if we had complied with the WHO 24 hour limit of 20 µg/m³. The corresponding roadside figure was 12 µg/m³, 140% higher than the predicted level achievable through compliance with the short term limit.

It is recognised that the Hong Kong Air Pollution Index (API), which has been used for the past 20 years, needs revision. Even using this seriously outdated index, the number of days when roadside stations regularly recorded an index of greater than 100 was very high. Due to the 'canyon effect' of buildings, the levels of air pollutants measured at roadside monitoring stations are usually much higher than those measured at general (roof top) stations. Using internationally accepted 2006 WHO guidelines, air pollutants in Hong Kong are continuously above safer levels for PM₁₀, NO₂, and frequently for SO₂ in certain locations, with important consequences for illness, health care and

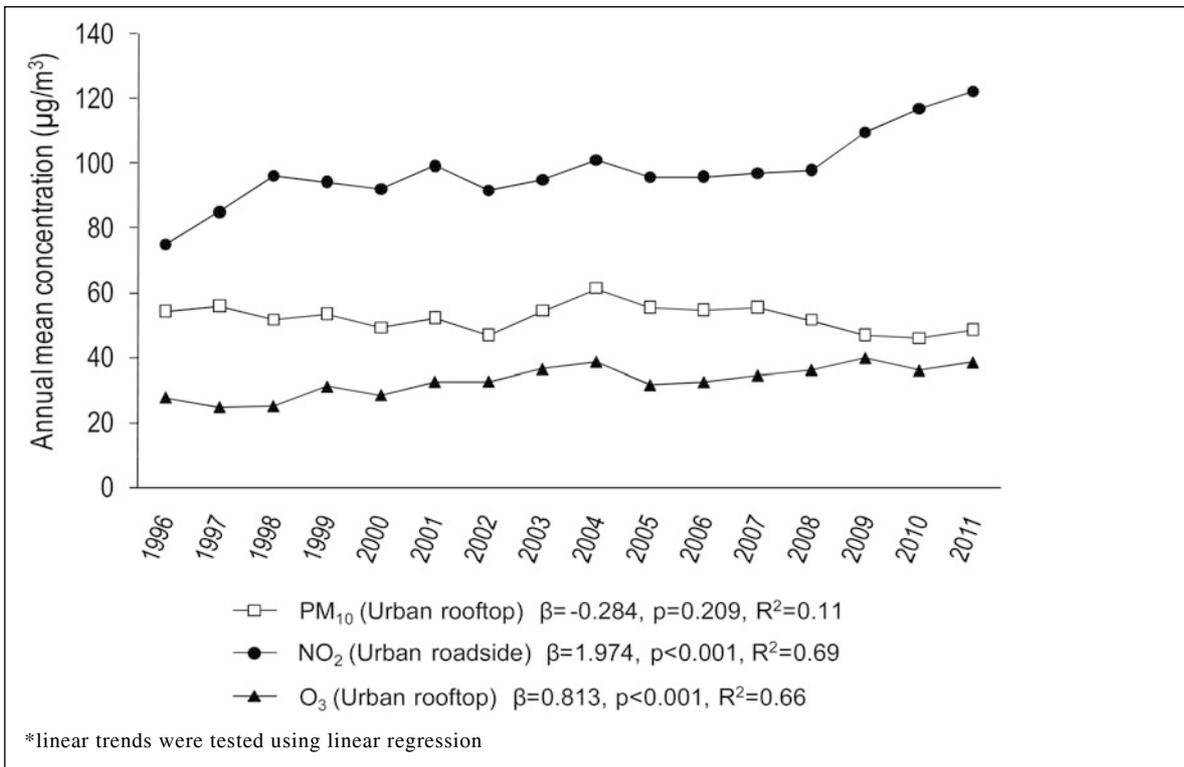


Figure 1 Trends of annual mean concentrations of PM₁₀, NO₂ and O₃ from 1996 to 2011.

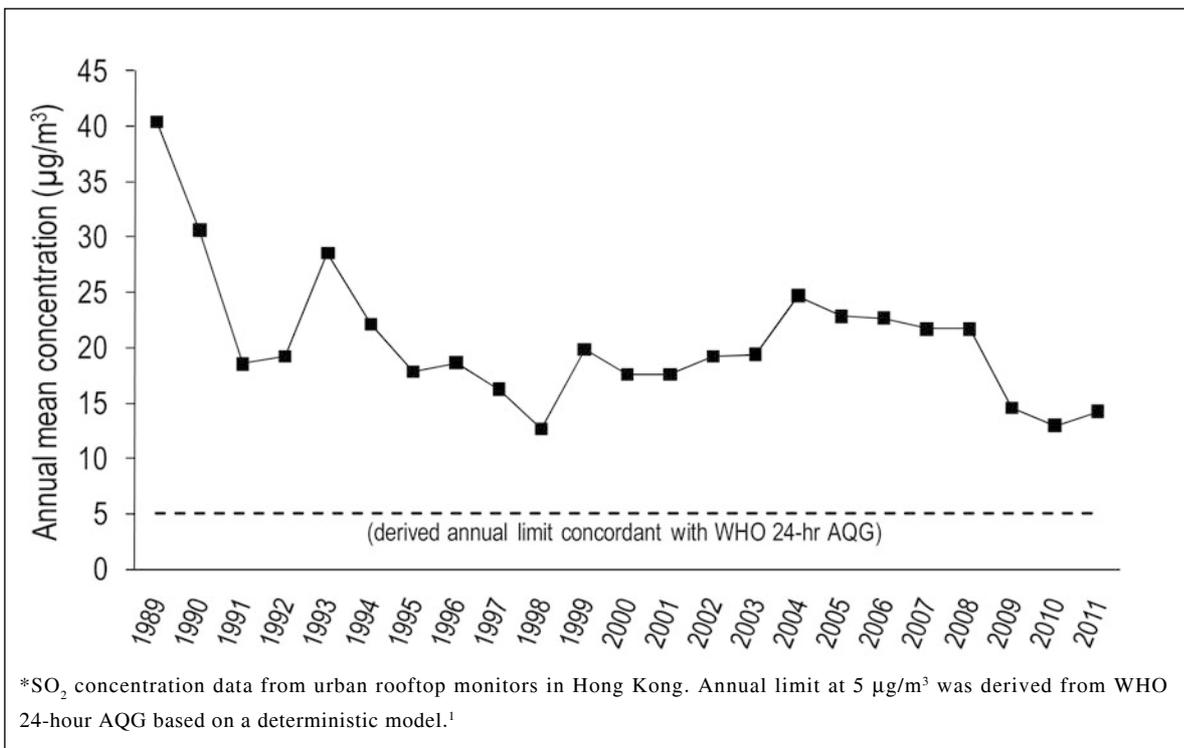


Figure 2 Annual mean SO₂ concentration from 1989 to 2011 compared with the annual limit derived from WHO 24-hour Air Quality Guideline (AQG).

community costs as demonstrated on a continuing basis by the Hedley Environmental Index. (<http://hedleyindex.sph.hku.hk/pollution/home.php#s>).

A high API has resulted in advice to restrict outdoor sport and leisure activities, especially for vulnerable groups, for example children with pre-existing respiratory and cardiac diseases, but these advisories come from the Environmental Protection Department with no specialist public health or medical input. They are meaningless in terms of public health protection. The Child Health Survey 2005 to 2006 suggested that only around 15% of children aged 6-14 years in Hong Kong had moderate to severe vigorous activities each day, in contrast to nearly 60% who spent at least two hours in screen time at home.⁵⁴ If air pollution contributes to the adoption of an even more sedentary lifestyle in our local children population, then it may also contribute to the emerging epidemic of childhood obesity in addition to the many other associated medical problems. The loss of daily visibility, landscapes and horizons degrades a child's natural environment but its impact on health related quality of life has not yet been assessed. In Hong Kong, myopia was the commonest chronic health problem reported in the Child Health Survey 2005 to 2006.⁵⁴ There is concern that the deterioration in visibility due to air pollution over the past 40 years,⁵⁵ with less exposure to greater depth of field and less usage blur than with higher outdoor light intensities, may be an additional contributing factor in the genesis of myopia in this region.

While exposure to air pollutants in residential areas is a major concern for child health, another priority area is the location of most of the schools in Hong Kong. There are a total of around 650 primary schools in Hong Kong of which 20% are situated close to a main road, as defined by the Transport Department, at a mean distance of 20.5 with standard deviation of 24.7 metres⁵⁶ and traffic related air pollution must therefore be regarded as a major health threat to school children. It is clear that the current neglect of air quality in Hong Kong and acceptance of the status quo will have a significant impact on quality of life, life expectancy and community costs of health care for our children. The Government of the HKSAR has made considerable efforts in the past to improve our child health through many interventions, for example the establishment of the Maternal and Child Health Service and Student Health Service, and provision of universal immunisation. However the question remains as to whether there is the political will, organisation and expertise to accept primacy of the need to improve air

quality for protection of the health of our children, and indeed of the whole population, now and in the future. The government consultancy for new air quality objectives which began in 2007 has resulted in an extremely lax limit for pollutants which does not conform to the evidence guidelines of WHO. These proposed new AQO will not be effective instruments to support air quality regulation and improvement. We can show that their adoption will likely lead to predictable and measurable continuing adverse health effects in children.¹ Hong Kong should adopt the full WHO 2006 Air Quality Guidelines as the basis for risk assessment and risk communication. It should be recognised that there is no discernible threshold for the health effects of air pollutants. While it may be argued that the creation of intermediate compromised standards could possibly lead to some mitigation of the threat to child health, given Hong Kong's extremely high levels of pollution, the impact on avoidable morbidity and morbidity will be relatively small and unacceptable as a public health approach to the problem.

We have suggested mandatory actionable steps for the Government of the HKSAR, local authorities and other organisations to clean the air of Hong Kong. The government should acknowledge that air quality is an urgent public health problem with serious implications for the current and future child health in the community. Most importantly the government has a clear duty to adopt immediate legislative and administrative measures to reduce the impact of our current urban environment and transport policies on air quality, and develop evidence-based strategies for rapid improvements. These can include adopting advanced technology to reduce emissions from power plants, traffic and marine sources, clean transportation options and modern vehicle fleets, infrastructure planning including establishing low emission zones in urban areas,⁵⁷ protecting kindergartens, schools and sports facilities from intense pollutant sources, planting vegetation in street canyons⁵⁸ and promoting building designs like urban wide conversion of black roof to green roof.⁵⁹ Local councils should press the government for immediate effective action to protect children in inner conurbations, consistent with the Department of Health's promotion of the WHO Healthy City concept to District Councils. The government should fund scientific-based population health research on the long term implications of toxic air on child health, to support policies for health protection. Food and Health Bureau should take the lead in monitoring the health impact of air pollution and use

information from expert groups in Hong Kong and overseas, including paediatric and public health expertise, to provide evidence-based advice to relevant sectors of our government to strengthen policy decisions.

Conclusion

Children need our continuing commitment to their future health and wellbeing. Good health care in a nurturing environment, with safe water, food and air are all important for our children now and in the future. The Government of the HKSAR is a signatory to the Convention of Rights of the Child adopted by the United Nations. We urge all paediatricians to join to press the government to support these principles by taking effective and immediate action, on the shortest possible timescale, to minimise environmental pollution, and prevent further serious harm to Hong Kong's young people. Failure to take a comprehensive, effective and precautionary approach will result in large scale detriment to health related quality of life among our children for decades to come.

Acknowledgements

The authors would like to thank Professor YL Lau, Doris Zimmern Professor in Community Child Health and Chair Professor of Paediatrics, Department of Paediatrics & Adolescent Medicine, The University of Hong Kong and Professor TH Lam, Sir Robert Kotewall Professor in Public Health, Director of School of Public Health, The University of Hong Kong for their helpful advice and critiques for this review.

References

- Lai HK, Wong CM, McGhee SM, Hedley AJ. Assessment of the health impacts and economic burden arising from proposed new Air Quality Objectives in a high pollution environment. *Open Epidemiol* 2011;4:106-22.
- Finkelstein JN, Johnston CJ. Enhanced Sensitivity of the Postnatal Lung to Environmental Insults and oxidant stress. *Pediatrics* 2004;113:1092-6.
- Salvi S. Health effects of ambient air pollution in children. *Paediatr Respir Rev* 2007;8:275-80.
- Schwartz J. Air pollution and children's health. *Pediatrics* 2004; 113:1037-43.
- Stocks J. Respiratory physiology during early life. *Monaldi Arch Chest Dis* 1999;54:358-64.
- Abreu RR, Rocha RL, Lamounier JA, Guerra AF. Etiology, clinical manifestations and concurrent findings in mouth-breathing children. *J Pediatr (Rio J)* 2008;84:529-35.
- Romieu I, Sienna-Monge JJ, Ramirez-Aguilar M, et al. Genetic polymorphism of GSTM1 and antioxidant supplementation influence lung function in relation to ozone exposure in asthmatic children in Mexico City. *Thorax* 2004;59:8-10.
- Islam T, Berhane K, McConnell R, et al. Glutathione-S-transferase (GST) P1, GSTM1, exercise, ozone and asthma incidence in school children. *Thorax* 2009;64:197-202.
- Lee YL, McConnell R, Berhane K, Gilliland FD. Ambient ozone modifies the effect of tumor necrosis factor G-308A on bronchitic symptoms among children with asthma. *Allergy* 2009;64: 1342-8.
- Giovannini M, Sala M, Riva E, Radaelli G. Carbon monoxide and nitrogen dioxide may be associated with emergency hospital admissions for respiratory conditions among children in Southwest Milan. *Acta Paediatr* 2010;99:1180-5.
- Lee SL, Wong WH, Lau YL. Association between air pollution and asthma admission among children in Hong Kong. *Clin Exp Allergy* 2006;36:1138-46.
- Silverman RA, Ito K. Age-related association of fine particles and ozone with severe acute asthma in New York City. *J Allergy Clin Immunol* 2010;125:367-73.e5.
- Mann JK, Balmes JR, Bruckner TA, et al. Short-term effects of air pollution on wheeze in asthmatic children in Fresno, California. *Environ Health Perspect* 2010;118:1497-502.
- MacIntyre EA, Karr CJ, Koehoorn M, et al. Residential air pollution and otitis media during the first two years of life. *Epidemiology* 2011;22:81-9.
- Salam MT, Islam T, Gilliland FD. Recent evidence for adverse effects of residential proximity to traffic sources on asthma. *Curr Opin Pulm Med* 2008;14:3-8.
- Kim JJ, Huen K, Adams S, et al. Residential traffic and children's respiratory health. *Environ Health Perspect* 2008;116:1274-9.
- Chang J, Delfino RJ, Gillen D, Tjoa T, Nickerson B, Cooper D. Repeated respiratory hospital encounters among children with asthma and residential proximity to traffic. *Occup Environ Med* 2009;66:90-8.
- Gauderman WJ, Avol E, Gilliland F, et al. The effect of air pollution on lung development from 10 to 18 years of age. *New Engl J Med* 2004;351:1057-67.
- Gauderman WJ, Vora H, McConnell R, et al. Effect of exposure to traffic on lung development from 10 to 18 years of age: a cohort study. *Lancet* 2007;369:571-7.
- Rosenlund M, Forastiere F, Porta D, De Sario M, Badaloni C, Perucci CA. Traffic-related air pollution in relation to respiratory symptoms, allergic sensitisation and lung function in schoolchildren. *Thorax* 2009;64:573-80.
- Clark NA, Demers PA, Karr CJ, et al. Effect of early life exposure to air pollution on development of childhood asthma. *Environ Health Perspect* 2010;118:284-90.
- McConnell R, Islam T, Shankardass K, et al. Childhood incident asthma and traffic-related air pollution at home and school. *Environ Health Perspect* 2010;118:1021-6.
- Carlsten C, Dybuncio A, Becker A, Chan-Yeung M, Brauer M. Traffic-related air pollution and incident asthma in a high-risk birth cohort. *Occup Environ Med* 2011;68:291-5.
- Heinrich J, Wichmann HE. Traffic related pollutants in Europe and their effect on allergic disease. *Curr Opin Allergy Clin*

- Immunol 2004;4:341-8.
25. Morgenstern V, Zutavern A, Cyrus J, et al. Atopic diseases, allergic sensitization, and exposure to traffic-related air pollution in children. *Am J Respir Crit Care Med* 2008;177:1331-7.
 26. Bråbäck L, Forsberg B. Does traffic exhaust contribute to the development of asthma and allergic sensitization in children: findings from recent cohort studies. *Environ Health* 2009;8:17.
 27. Riedl M, Diaz-Sanchez D. Biology of diesel exhaust effects on respiratory function. *J Allergy Clin Immunol* 2005;115:221-8.
 28. Samuelsen M, Nygaard UC, Løvik M. Allergy adjuvant effect of particles from wood smoke and road traffic. *Toxicology* 2008;246:124-31.
 29. de Haar C, Hassing I, Bol M, Bleumink R, Pieters R. Ultrafine but not fine particulate matter causes airway inflammation and allergic airway sensitization to co-administered antigen in mice. *Clin Exp Allergy* 2006;36:1469-79.
 30. Bevelander M, Mayette J, Whittaker LA, et al. Nitrogen dioxide promotes allergic sensitization to inhaled antigen. *J Immunol* 2007;179:3680-8.
 31. Hodgkins SR, Ather JL, Paveglio SA, et al. NO₂ inhalation induces maturation of pulmonary CD11c+ cells that promote antigenspecific CD4+ T cell polarization. *Respir Res* 2010;11:102.
 32. Suglia SF, Gryparis A, Wright RO, Schwartz J, Wright RJ. Association of black carbon with cognition among children in a prospective birth cohort study. *Am J Epidemiol* 2008;167:280-6.
 33. Guxens M, Aguilera I, Ballester F, et al. Prenatal exposure to residential air pollution and infant mental development: modulation by antioxidants and detoxification factors. *Environ Health Perspect* 2012;120:144-9.
 34. Wang S, Zhang J, Zeng X, Zeng Y, Wang S, Chen S. Association of traffic-related air pollution with children's neurobehavioral functions in Quanzhou, China. *Environ Health Perspect* 2009;117:1612-8.
 35. Siddique S, Banerjee M, Ray MR, Lahiri T. Attention-deficit hyperactivity disorder in children chronically exposed to high level of vehicular pollution. *Eur J Pediatr* 2011;170:923-9.
 36. Amigou A, Sermage-Faure C, Orsi L, et al. Road Traffic and Childhood Leukemia: The ESCALE Study (SFCE). *Environ Health Perspect* 2011;119:566-72.
 37. Mattison DR. Environmental exposures and development. *Curr Opin Pediatr* 2010;22:208-18.
 38. Darrow LA, Klein M, Strickland MJ, Mulholland JA, Tolbert PE. Ambient air pollution and birth weight in full-term infants in Atlanta, 1994-2004. *Environ Health Perspect* 2011;119:731-7.
 39. van den Hooven EH, Pierik FH, de Kluizenaar Y, et al. Air pollution exposure during pregnancy, ultrasound measures of fetal growth, and adverse birth outcomes: a prospective cohort study. *Environ Health Perspect* 2012;120:150-6.
 40. Pedersen M, Wichmann J, Autrup H, et al. Increased micronuclei and bulky DNA adducts in cord blood after maternal exposures to traffic-related air pollution. *Environ Res* 2009;109:1012-20.
 41. Vrijheid M, Martinez D, Manzanares S, et al. Ambient air pollution and risk of congenital anomalies: A systematic review and meta-analysis. *Environ Health Perspect* 2011;119:598-606.
 42. Dolk H, Armstrong B, Lachowycz K, et al. Ambient air pollution and risk of congenital anomalies in England, 1991-1999. *Occup Environ Med* 2010;67:223-7.
 43. Rankin J, Chadwick T, Natarajan M, Howel D, Pearce MS, Pless-Mullooli T. Maternal exposure to ambient air pollutants and risk of congenital anomalies. *Environ Res* 2009;109:181-7.
 44. Son JY, Bell ML, Lee JT. Survival analysis of long-term exposure to different sizes of airborne particulate matter and risk of infant mortality using a birth cohort in Seoul, Korea. *Environ Health Perspect* 2011;119:725-30.
 45. Yu TS, Wong TW, Wang XR, Song H, Wong SL, Tang JL. Adverse effects of low-level air pollution on the respiratory health of schoolchildren in Hong Kong. *J Occup Environ Med* 2001;43:310-6.
 46. Peters J, Hedley AJ, Wong CM, et al. Effects of an ambient air pollution intervention and environmental tobacco smoke on children's respiratory health in Hong Kong. *Int J Epidemiol* 1996;25:821-8.
 47. Wong CM, Lam TH, Peters J, et al. Comparison between two districts of the effects of an air pollution intervention on bronchial responsiveness in primary school children in Hong Kong. *J Epidemiol Community Health* 1998;52:571-8.
 48. Hedley AJ, Wong CM, Thach TQ, Ma S, Lam TH, Anderson HR. Cardiorespiratory and all-cause mortality after restrictions on sulphur content of fuel in Hong Kong: an intervention study. *Lancet* 2002;360:1646-52.
 49. Wong CM, Ma S, Hedley AJ, Lam TH. Effect of air pollution on daily mortality in Hong Kong. *Environ Health Perspect* 2001;109:335-40.
 50. Wong CM, Atkinson RW, Anderson HR, et al. A tale of two cities: Effects of air pollution on hospital admissions in Hong Kong and London compared. *Environ Health Perspect* 2002;110:67-77.
 51. Ou CQ, Hedley AJ, Chung RY, et al. Socioeconomic disparities in air pollution-associated mortality. *Environ Res* 2008;107:237-44.
 52. Wong CM, Ou CQ, Chan KP, et al. The effects of air pollution on mortality in socially deprived urban areas in Hong Kong. *Environ Health Perspect* 2008;116:1189-94.
 53. WHO. Air Quality Guidelines. Global update 2005. Geneva: World Health Organization, 2006. http://www.euro.who.int/_data/assets/pdf_file/0005/78638/E90038.pdf.
 54. Department of Health. The Child Health Survey 2005 to 2006. http://www.chp.gov.hk/files/pdf/chs_0506_eng.pdf.
 55. Thach TQ, Wong CM, Chan KP, et al. Daily visibility and mortality: assessment of health benefits from improved visibility in Hong Kong. *Environ Res* 2010;110:617-23.
 56. Transport Department. The annual traffic census, 2009, TTSD Publication No. 10CAB1. Hong Kong SAR, 2009. http://www.td.gov.hk/filemanager/en/content_4379/annual%20traffic%20census%202009.pdf
 57. Kelly F, Armstrong B, Atkinson R, et al. The London low emission zone baseline study. *Res Rep Health Eff Inst* 2011;163:3-79.
 58. Pugh TA, Mackenzie AR, Whyatt JD, Hewitt CN. Effectiveness of green infrastructure for improvement of air quality in urban street canyons. *Environ Sci Technol* 2012;46:7692-9.
 59. Susca T, Gaffin SR, Dell'osso GR. Positive effects of vegetation: urban heat island and green roofs. *Environ Pollut* 2011;159:2119-26.