

REVIEW

Open Access

Climate change, air pollution and extreme events leading to increasing prevalence of allergic respiratory diseases

Gennaro D'Amato^{1*}, Carlos E Baena-Cagnani², Lorenzo Cecchi³, Isabella Annesi-Maesano⁴, Carlos Nunes⁵, Ignacio Ansotegui⁶, Maria D'Amato⁷, Gennaro Liccardi¹, Matteo Sofia⁷ and Walter G Canonica⁸

Abstract

The prevalence of asthma and allergic diseases has increased dramatically during the past few decades not only in industrialized countries. Urban air pollution from motor vehicles has been indicated as one of the major risk factors responsible for this increase.

Although genetic factors are important in the development of asthma and allergic diseases, the rising trend can be explained only in changes occurred in the environment. Despite some differences in the air pollution profile and decreasing trends of some key air pollutants, air quality is an important concern for public health in the cities throughout the world.

Due to climate change, air pollution patterns are changing in several urbanized areas of the world, with a significant effect on respiratory health.

The observational evidence indicates that recent regional changes in climate, particularly temperature increases, have already affected a diverse set of physical and biological systems in many parts of the world. Associations between thunderstorms and asthma morbidity in pollinosis subjects have been also identified in multiple locations around the world.

Allergens patterns are also changing in response to climate change and air pollution can modify the allergenic potential of pollens especially in presence of specific weather conditions.

The underlying mechanisms of all these interactions are not well known yet. The consequences on health vary from decreases in lung function to allergic diseases, new onset of diseases, and exacerbation of chronic respiratory diseases.

Factor clouding the issue is that laboratory evaluations do not reflect what happens during natural exposition, when atmospheric pollution mixtures in polluted cities are inhaled. In addition, it is important to recall that an individual's response to pollution exposure depends on the source and components of air pollution, as well as meteorological conditions. Indeed, some air pollution-related incidents with asthma aggravation do not depend only on the increased production of air pollution, but rather on atmospheric factors that favour the accumulation of air pollutants at ground level.

Considering these aspects governments worldwide and international organizations such as the World Health Organization and the European Union are facing a growing problem of the respiratory effects induced by gaseous and particulate pollutants arising from motor vehicle emissions.

Keywords: Airways hyper-responsiveness, Bronchial asthma, Climate change and allergy, Environment and respiratory allergy, Pollen allergy, Respiratory allergy, Urban air pollution

* Correspondence: gdamatomail@gmail.com

¹Division of Respiratory and Allergic Diseases, HighSpecialityHospital A. Cardarelli, Department of Respiratory Diseases, Napoli, Italy
Full list of author information is available at the end of the article

“He inhaled a breath of humid morning breeze and let in nitrogen, oxygen, argon, xenon & radon, steam, carbon monoxide, nitrogen dioxide, tetra-ethyl lead, benzene, some mould spores, a bacteria fleet, anonymous body hair, a pigeon ectoparasite, anemophilous pollen, a drop of sulphur dioxide flown from a distant factory, and a particle of dust carried by the night sirocco.

In other words he breathed air of the city”.
(Stefano Benni: *Achille piè veloce*. Italy: Mondadori; 2003.)

Introduction

The massive increase in emissions of air pollutants due to economic and industrial growth in the last century made air quality an environmental problem of first order in a large number of European and North American countries and it is now an emerging problem in other regions of the world [1-44].

Air pollution is convincingly associated with many signs of asthma aggravation (increased bronchial hyper-responsiveness, visits to emergency departments, hospital admissions, increased medication use etc.) [5,12,13,22-25]. Moreover, sensitive techniques to analyze time-series data have shown that there are clear adverse effects on mortality rates from current levels of air pollution [4,6,7].

Furthermore, several air pollutants, in particular carbon dioxide (CO₂) and ozone (O₃), are in the list of greenhouse gases which are involved in the global warming [6-8,11-13].

Global earth's temperature has markedly risen over the last 50 years due to the increase in greenhouse gas emissions, largely from anthropogenic sources. Changes are also occurring in the amount, intensity, frequency, and type of precipitation as well as the increase of extreme events, like heat waves, droughts, floods, thunderstorms and hurricanes and these are a real and daunting problem [5-15].

It is extremely unlikely that global climatic change can be explained without external forcing. As stated in the Working Group I Report of the Intergovernmental Panel on Climate Change (IPCC) “most of the observed increase in globally averaged temperatures since the mid-20th century is very likely due to the observed

increase in anthropogenic greenhouse gas concentrations” [6,7] (Table 1).

It has been also observed a rapid rise in the number of hot days and severe meteorological events [9,10,44] such as the 2003 and 2012 heat waves when temperatures reached or went beyond 35°C degrees resulting in excess deaths across Europe [9,10]. Moreover, climate scenarios for the next century predict that the warming will be associated with more frequent and more intense heat waves in wide areas of our planet with increased risk of wildfires and desertification. In urban areas the effects are higher since climate change influences outdoor air pollution because the generation and dispersion of air pollution is in strictly correlation with local patterns of temperature, wind and precipitation.

Climate change has led also to water deprivation in certain areas often associated with water degradation which potentially could result in population migration and the effects on health that result from mass population movement.

Atopy and asthma are more prevalent in developed and industrialized countries as compared with undeveloped and less affluent countries, and the effect of migration is age and time-dependent: early age and longer time increase the likelihood of developing atopy and asthma.

Climate changes will affect the acquisition of atopy and respiratory diseases [12,17].

A number of reports on time trends in allergic respiratory diseases and bronchial asthma have shown a substantial increase in prevalence since the early 1960s [1-3,5].

There is also a link between climate changes and air pollution and an individual's response to air pollution depends on the source and components of the pollution, as well as on climatic agents [8,11-15]. Some air pollution-related episodes of rhinitis and asthma exacerbation are due to climatic factors that favour the accumulation of air pollutants, such as ozone, at ground level. However, the effects of air pollutants on lung function depend on the environmental concentration of the pollutant, the duration of pollutant exposure and the total ventilation of exposed persons.

A factor clouding the issue is that laboratory evaluations do not reflect what happens during natural exposition, when atmospheric pollution mixtures in polluted cities are inhaled. As a consequence, even if it is

Table 1 Global warming

-
- The global warming average air temperature has increased by 1.0±0.3°F (0.6±0.2°C) since the late 19th century
 - The average surface temperature of the earth is likely to increase by 2 to 11.5°F (1.1-6.4°C) by the end of the 21st century
 - The average rate of warming over each inhabited continent is very likely to be at least twice as large as that experienced during 20th century
-

plausible that ambient air pollution plays a role in the onset and in the increasing frequency of respiratory allergy, it is not easy to show that this happens at public health level. In addition, it is important to recall that an individual's response to pollution exposure depends on the source and components of air pollution, as well as meteorological conditions. Indeed, some air pollution-related incidents with asthma aggravation do not depend only on the increased production of air pollution but rather on atmospheric factors that favour the accumulation of air pollutants at ground level.

Greenhouse gas emissions

The key determinants of greenhouse gas emissions are energy production, transportation, agriculture and food production and waste management, and attempts at mitigating climate change will need to address each of these. However, while there is some uncertainty about predicting future meteorological trends, whatever interventions may be put in place to ameliorate climate change, it is likely that the world will experience more hot days, fewer frost days, and more periods of heavy rain and consequent flooding [6-8]. Paradoxically it is likely that there will be more periods of drought. A huge increase in CO₂ concentrations during the last two decades has been experienced (Figure 1). However, it is important to consider that after CO₂ emissions are reduced and atmospheric concentrations stabilized, surface air temperature continues to rise slowly for a century or more.

Urban air pollutants

The most abundant components of air pollution in urban areas are nitrogen dioxide, ozone and particulate matter. Sulphur dioxide is particularly abundant in industrial areas. It is estimated that more than 50% of the population of the United States live in areas where levels of ozone, nitrogen dioxide, sulphur dioxide, and particulates exceed current National Ambient Quality Standards, as monitored by the US Environmental Protection Agency [6,7]. With its particulate and gaseous emissions, road traffic is the main contributor to air pollution in most urban areas. Although associations between air pollution and respiratory diseases are complex, recent epidemiological studies have led to an increased recognition of the emerging importance of traffic-related air pollution in both developed and less-developed countries [6,7]. A number of experimental and epidemiological studies confirmed the negative effect of urban air pollution on human health and on allergic respiratory diseases [5,11-17] and projections of climate variability suggest these effects will increase in the next decades [6,7].

Nitrogen dioxide (NO₂)

Car and trucks exhausts, together with power plants, are the most significant sources of outdoor NO₂, which is a precursor of photochemical smog found in outdoor air in urban and industrial regions and, in conjunction with sunlight and hydrocarbons, results in the production of ozone. Like ozone, NO₂ is an oxidant pollutant, although

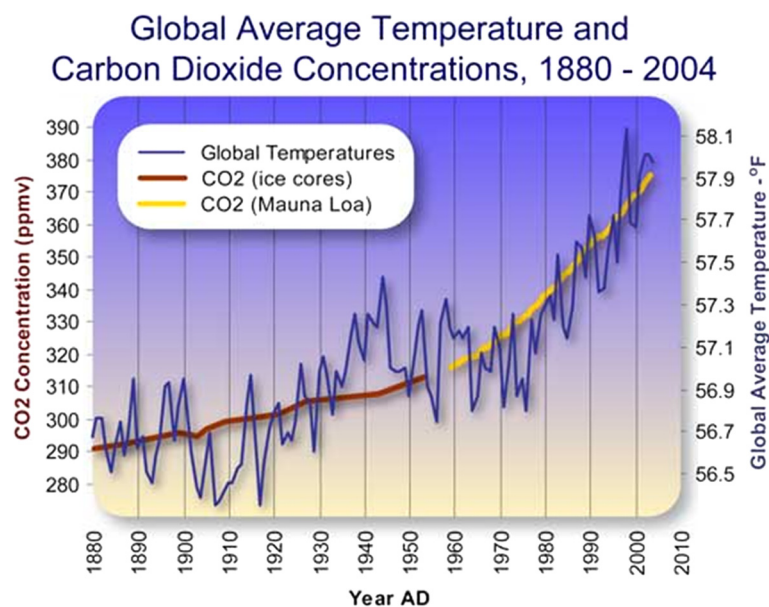


Figure 1 Global average temperature and carbon dioxide concentrations, 1980–2004.

it is less chemically reactive. NO₂ exposure is associated with increased emergency room visits, wheezing, and medication use among children with asthma [20]. Controlled exposure studies of asthmatics have found that NO₂ can enhance the allergic response to inhaled allergens, and NO₂ concentrations in ambient air are also reportedly associated with cough, wheezing and shortness of breath in atopic subjects [11-16]. A Norwegian study found no association between modeled NO₂ exposure at the birth address and doctor-diagnosed asthma in a large cohort of children [27].

Ozone (O₃)

Ozone is generated at ground level by photochemical reactions involving nitrogen dioxide, hydrocarbons, and UV radiation. Ozone inhalation induces epithelial damage and consequent inflammatory responses in the upper and lower airways, as shown by an increase in levels of inflammatory cells and mediators in nasal and bronchoalveolar lavage [20].

About 40-60% of inhaled O₃ is absorbed in the nasal airways, the remainder reaching the lower airways. Exposure to increased atmospheric levels of O₃ induces reduction of lung function, increased airway hyperreactivity to bronchoconstrictor agents and is related to an increased risk of asthma exacerbations in asthmatic subjects [12-16]. Epidemiologic studies have provided evidence that high ambient concentrations of this air pollutant are associated with an increased rate of asthma exacerbations, increased hospital admissions, and/or emergency department visits for respiratory diseases, including asthma. Furthermore, several studies suggest that O₃ increases asthma morbidity by enhancing airway inflammation and epithelial permeability [11-16].

O₃ exposure significantly increases levels of inflammatory cells (in particular neutrophils) and mediators, such as IL-6, IL-8, granulocyte-macrophage colony-stimulating factor (GM-CSF) and fibronectin, in bronchoalveolar lavage fluid (BALF) of asthmatic subjects [16,17].

It has long been speculated that O₃ and other pollutants may render allergic subjects more susceptible to the antigen they are sensitized [11-13]. It has been observed that the incidence of new diagnoses of asthma is associated with heavy exercise in communities with high concentrations of O₃, thus, air pollution and outdoor exercise could contribute to the development of asthma [16,17].

However, it is important to take into account that physical exercise in polluted areas results in greater deposition of air pollutants, including allergen-carrying particles, in the lower airways.

The acute health effects of exposure to ambient ozone have been examined in many geographical regions. Potential adverse effects include decrease in lung

function, airway inflammation, symptoms of asthma, increases in hospitalization due to respiratory diseases, and excess mortality. Ozone exposure has both a priming effect on allergen-induced responses and an intrinsic inflammatory action in the airways of allergic asthmatics [8,13-16].

In the long term, continuous exposure to high ozone levels impairs respiratory function and causes or exacerbates airway inflammation in healthy subjects and atopic asthmatics. At the population level, long-term exposure to ozone may reduce lung function in school children and adults and increase the prevalence of asthma and asthmatic symptoms [16,17]. In addition, studies have long shown that asthma can be exacerbated by ozone, as measured by increased visits to emergency departments on days with higher levels of ozone and other pollutants [11-13]. Moreover, it has been observed an increased asthma prevalence in milder climatic areas, possibly due to the positive effect of air temperature on ozone synthesis [12-15].

Traffic-related air pollution, particulate matter and diesel exhaust particles

There is evidence that living near high-traffic roads is associated with impaired respiratory health including asthma [22-25]. First, McConnell et al. [17] observed that the incidence of newly diagnosed asthma in children is associated with physical exercise in areas with high concentrations of ozone and particulate matter. Since then, other prospective cohort studies have indicated that long-term exposure to traffic pollution could contribute to the development of asthma-like symptoms and allergic sensitization in children [24,25]. Potential long-term effects of traffic exhaust on the development of allergic sensitization were only assessed in the four European birth cohorts.

Long-term exposure to outdoor air pollutants had no association with sensitization in ten-year-old schoolchildren in Norway [27]. These studies, however, are flawed by the fact that no objective assessment of air pollution concentrations was available. Individual exposure was estimated by the distance from the highways.

Particulate matter is a mixture of organic and inorganic solid and liquid particles of different origins, size, and composition. It is a major component of urban air pollution and has the greatest effect on health. Penetration of the tracheobronchial area is related to particle size and the efficiency of airway defence mechanisms. Ultrafine particulate matter (UFP), with diameters of 0.1 µm or less, is a major component of emissions from vehicles. These particles accumulate into larger fine PM with a diameter of ≤2.5 µm (PM_{2.5}), within short distances from the point of release. PM₁₀ (particulate matter with a diameter of 10 µm or less) consists of PM_{2.5}

and larger particles of mainly crustal or biological origin, including many aeroallergens. On the basis of epidemiological and laboratory studies, $PM_{2.5}$ appears to be more potent agent for the development of respiratory and cardiovascular disease compared with PM_{10} . [21-23] PM_{10} can penetrate the lower airways, and $PM_{2.5}$, is thought to constitute a notable health risk since it can be inhaled more deeply into the lungs at the alveoli level). While human lung parenchyma retains $PM_{2.5}$, particles larger than 5 μm and smaller than 10 μm reach the proximal airways only, where they are eliminated by mucociliary clearance if the airway mucosa is intact [11-13].

A large portion of urban particulate matter originates from diesel engines, the source of diesel exhaust particles (DEPs) which includes other components such as polycyclic aromatic hydrocarbons [PAH]. DEPs account for up to 90% of airborne particulate matter in the world's largest cities and are composed of fine particles (2.5-0.1 μm) and ultrafine (0.1 μm) particles, which can also coalesce to form aggregates of varying sizes [21].

PM_{10} levels have been associated with early respiratory exacerbations in children with persistent asthma and with higher prevalence rates even after having considered the dispersion of the particles. Although there is compelling evidence that ambient air pollution exacerbates existing asthma, the link with the development of the asthma syndrome is still less well established, as few studies provide extensive exposure data. Researches have elucidated the mechanisms whereby fine particles induce adverse effects; they appear to affect the balance between antioxidant pathways and airway inflammation. Gene polymorphisms involved in antioxidant pathways can modify responses to air pollution exposure. Acute exposure to diesel exhaust causes specific effects, like irritation of nose and eyes, headache, lung function abnormalities, respiratory changes, fatigue, and nausea in acute exposure, while chronic exposure is associated with cough, sputum production, and diminished lung function [21-25].

Studies have demonstrated inflammation in the airways of healthy individuals after exposure to diesel exhaust and DEPs, and elevated expression and concentrations of inflammatory mediators have similarly been observed in the respiratory tract after diesel exhaust and DEP exposure [21,25]. Even if an increased sensitivity of asthmatic individuals to the pro-inflammatory effects of DEPs has not been confirmed, some studies showed a bigger effect of exposure to high-traffic roads in asthmatics compared to non asthmatic subjects, also accompanied by increases in levels of biomarkers of neutrophilic inflammation.

Effect of climate change on allergic respiratory diseases

A body of evidence suggests that major changes involving the atmosphere and the climate, including global

warming induced by human activity, have an impact on the biosphere and human environment [6,7].

Current knowledge on the worldwide effects of climate change on respiratory allergic diseases is provided by epidemiological and experimental studies on the relationship between asthma and environmental factors, like meteorological variables, airborne allergens and air pollution. However, we can try to summarize the correlation between climate changes, allergenic plants and pollen distribution [12-15] in the following points:

- (i) increase and faster plant growth;
- (ii) increase the amount of pollen produced by each plant;
- (iii) increase the amount of allergenic proteins contained in pollen,
- (iv) increase the start time of plant growth and therefore the start of pollen production and
- (v) earlier and longer growing pollen seasons.

Pollen allergy is frequently used to study the interrelationship between air pollution and allergic respiratory diseases (rhinitis and asthma). Epidemiologic studies have demonstrated that urbanization, high levels of vehicle emissions and westernized lifestyle are correlated to an increase in the frequency of pollen-induced respiratory allergy prevalent in people who live in urban areas compared to those who live in rural areas [12-15].

Meteorological factors (temperature, wind speed, humidity, thunderstorms etc.) along with their climatic regimes (warm or cold anomalies and dry or wet periods, etc.), can affect both biological and chemical components of this interaction. In addition, by inducing airway inflammation, air pollution overcomes mucosal barrier priming allergen-induced responses.

Climate changes might induce negative effects on respiratory allergic diseases favouring the increased length and severity of pollen season, the higher occurrence of heavy precipitation events and the increasing frequency of urban air pollution episodes.

Climatic factors (temperature, wind speed, humidity, thunderstorms, etc.) can affect both components (biological and chemical) of this interaction.

However, the relationship between air pollution, pollen exposure and respiratory allergy is based on an individual's response to air pollution, which depends on the source and components of the pollution, as well as on climatic agents.

Thunderstorm-related allergic respiratory diseases and bronchial asthma in pollinosis subjects

There are observations that thunderstorms occurring during pollen season can induce severe asthma attacks in pollinosis patients [45-69].

Table 2 Weather changes with climate change

<ul style="list-style-type: none">• More extreme weather patterns, such as increase in thunderstorm• High number of thunderstorms in spring and summer at the same time as high pollen counts• Pollen grain rupture with thunderstorm with higher level of respirable allergens; also increased in zone• More asthma outbreaks

-UK, Australia and Italy

According to current climate change scenarios, there will be an increase in intensity and frequency of heavy rainfall episodes, including thunderstorms, over the next few decades, which can be expected to be associated with an increase in the number and severity of asthma attacks both in adults and in children (Table 2).

From [46,50,57,62] mod.

Associations between thunderstorms and asthma morbidity have been identified in multiple locations around the world [45-69]. So called "thunderstorm asthma" is characterized by asthma outbreaks possibly caused by the dispersion of more respirable allergenic particles derived from pollen and spores by osmotic rupture.

The most prominent hypotheses for thunderstorm-related asthma are linked with bioaerosols, and involve the role of rainwater in promoting the release of respirable particulate matter [47,59,63].

After hydration and rupture by osmotic shock during the beginning of a thunderstorm, pollen grains may release in atmosphere part of their cytoplasmic content, including inhalable, allergen-carrying paucimicronic particles such as starch granules and other cytoplasmic components .

Thunderstorm-related asthma outbreaks have been described in various geographical zones. One of the first observations regarding thunderstorms and asthma outbreaks was provided by Packe and Ayres at the East Birmingham Hospital (Birmingham, UK) on July 6 and 7, 1983 [45]. These authors described a remarkable increase in the number of asthma emergency department admissions during the hours of a thunderstorm. In a 36-h period, 26 asthma cases were treated in the emergency department, compared with a daily average of two or three cases in the days preceding the outbreak.

Another asthma outbreak occurred in London, UK, coinciding with a heavy thunderstorm on June 24, 1994, when a large increase in the number of visits for asthma at the emergency departments of London and the southwest of England was observed. Several patients who were examined, who were not known to be asthmatics or were affected only by seasonal rhinitis, experienced an asthma attack [50-52]. During a 30-h

period from 6 p.m. on June 24, 1994, 640 patients with asthma or other airways disease (283 of whom were not known to be asthmatic and 403 were affected only by seasonal rhinitis) attended several emergency departments, nearly 10 times the expected number of 66 patients. In total, 104 patients were admitted (including five to an intensive care unit) (574 patients attributable to the thunderstorm). Not all affected patients attended a hospital and this epidemic was the largest outbreak ever recorded.

Other asthma outbreaks during thunderstorms have been described in Australia. In Melbourne, two large asthma outbreaks (rapid increase in hospital or general practitioner visits for asthma) coincided with thunderstorms [46]. In WaggaWagga, 215 asthmatic subjects attended the local emergency department, 41 of whom required admission to hospital [57].

In south eastern Australia, Marks et al. [59] observed that the incidence of excess hospital attendances for asthma during late spring and summer was strongly linked to the occurrence of thunderstorm outflows and demonstrated that the arrival of a thunderstorm outflow was accompanied by a large increase in the concentration of ruptured pollen grains in ambient air.

Thunderstorm-related asthma was observed in Naples, Italy, on June 3 and 4, 2004 [62,63], when six adults and one child received treatment in emergency departments. One patient was admitted to an intensive care unit for a very severe bronchial obstruction and acute respiratory insufficiency following a sudden thunderstorm. All individuals were outdoors when the thunderstorm struck. In one severe case, a female sensitized only to *Parietaria* pollen allergen, soon began to show symptoms of intense dyspnoea, which gradually worsened. She was taken to hospital where she was intubated and given high intravenous doses of corticosteroids. She was discharged a few days later. This patient had previously suffered from seasonal asthma but had been asthma-free for the past few years and did not need continuous therapy. None of the other six subjects took anti-allergic and/or anti-asthma drugs regularly. All seven patients were sensitized with allergic respiratory symptoms upon exposure to *Parietaria* pollen, but were not sensitized to grasses. *Parietaria* is an *Urticacea* that is widespread in the Naples area of Italy with a spring and summer pollen season that is, in part, coexistent with that of grasses. During the thunderstorm, the concentration of airborne *Parietaria* pollen grains was particularly high, with a peak of 144 grains/m³ being recorded on June 3, 2004 [62,63]. Air pollution levels for both gaseous and particulate components based on the hourly concentrations of nitric dioxide, ozone and respirable particulate matter were not particularly high in Naples on June 3 and 4, 2004. Subjects with sensitization to *Parietaria*

Table 3 The evidence about thunderstorm-related epidemics of rhinitis and asthma exacerbations

- 1) The occurrence of epidemics is closely linked to thunderstorm.
- 2) The thunderstorm-related epidemics are limited to late spring and summer when there are high levels of airborne pollen grains.
- 3) There is a close temporal association between the arrival of the thunderstorm, a major rise in the concentration of pollen grains and the onset of epidemics.
- 4) Subjects with pollen allergy, who stay indoors with window closed during thunderstorm, are not involved.
- 5) There are not high levels of gaseous and particulate components of air pollution during outbreaks.
- 6) There is a major risk for subjects who are not under anti-asthma correct treatment.

who were indoors in Naples with the windows closed during the night between June 3 and 4, 2004, did not experience asthma attacks. No moulds or viruses were involved in the Naples epidemics. Recently, Losappio et al. [69] observed 20 patients with allergic sensitization to Olea pollen who presented to an emergency department in Barletta, Italy, for sudden and severe asthmatic symptoms in May 2010 following a thunderstorm. A similar phenomenon has been suggested for moulds [65] after the observation of a possible key role of sensitization to *Alternaria* species in thunderstorm related asthma. To date, among pollens, only grass, *Parietaria* and olive pollen have been suggested as possible key factors in thunderstorm-related asthma [63,66,69]. However, there is a risk of relapse of thunderstorm-related asthma [44,70].

Although thunderstorm-associated asthma outbreaks are not frequent, it is possible to observe in clinical practice single cases of patients with deterioration of the allergic respiratory symptoms during a thunderstorm (Table 3).

In summary the occurrence of these epidemics is closely linked to thunderstorm and they are limited to late spring and summer when there are high levels of airborne pollen grains. There is a close temporal association between the arrival of the thunderstorm, a major rise in the concentration of pollen grains and the onset of epidemics. As a consequence, subjects affected by pollen allergy should be alert to the danger of being outdoors during a thunderstorm in the pollen season.

Conclusions

In conclusion, strategies to reduce climate changes and air pollution are political in nature, but citizen and in particular health professionals and societies must raise their voices in the decision process to give strong support for clean policies on both national and international levels [71,72].

Endnotes

Manuscript from a lecture in XXI World Congress of Asthma (Organized by Interasma Global Asthma Association) held in Quebec City in August 18–21 2012.

Competing interests

The authors declare that they have no competing interests.

Author details

¹Division of Respiratory and Allergic Diseases, HighSpecialityHospital A. Cardarelli, Department of Respiratory Diseases, Napoli, Italy. ²Pediatric Hospital, Santa Rosa 381, Research Centre for Respiratory Medicine, Faculty of Medicine, Catholic University Córdoba, Córdoba, Argentina. ³Allergy and ClinicalImmunologySection, Azienda Sanitaria di Prato, Prato, Italy. ⁴INSERM, UMR S 707, EPAR, Paris, France. ⁵Centro de ImunoAlergologia de Algarve, Porto, Portugal. ⁶Department of Allergy and Immunology, Hospital QuirónBizkaiaErandio, Erandio-Bilbao, Spain. ⁷Division of Pneumology, University Hospital Dei Colli-Monaldi and Medical School Federico II, Napoli, Italy. ⁸DIMI Allergy and Respiratory Diseases, University of Genova, Genova, Italy.

Received: 15 January 2013 Accepted: 15 January 2013

Published: 11 February 2013

References

1. Burney PGJ: Evidence for an increase in atopic disease and possible causes. *Clin Exp Allergy* 1993, **23**:484–492.
2. Woolcock AJ, Peat JK: Evidence for the increase in asthma worldwide. In *"The rising trend in asthma"*. Ciba Foundation Symposium 206. Chichester, UK: John Wiley & Sons; 1997:122–139.
3. The International Study of Asthma and Allergy in Childhood (ISAAC), Steering Committee: Worldwide variation in prevalence of symptoms of asthma, allergic rhinoconjunctivitis and atopic eczema. *Lancet* 1998, **351**:1225–1232.
4. Wilkinson P, Smith KR, Beevers S, Tonne C, Oreszczyn T: Energy, energy efficiency, and the built environment. *Lancet* 2007, **370**:1175–1187.
5. D'Amato G, Liccardi G, D'Amato M, Holgate ST: Environmental risk factors and allergic bronchial asthma. *Clin Exp Allergy* 2005, **35**:1113–1124.
6. Hegerl GC, Zwiers FW, Braconnot P, Gillett NP, Luo Y, Marengo JA, et al: Understanding and attributing climate change. In *Climate change 2007: the physical science basis. Contribution of the Working Group I to the Fourth Assessment Report of the Intergovernmental Panel on Climate Change*. Edited by Solomon S, Qin D, Manning M, Chen Z, Marquis M, Averyt KB, et al. Cambridge, UK and New York: Cambridge University Press; 2007:663–746.
7. Parry ML, Canziana OF, Palutikof JP, Adger N, Agarwal P, Agrawala S, et al: Technical summary. In *Climate change 2007: impacts, adaptation, and vulnerabilities. Contribution of Working Group II to the Fourth Assessment Report of the Intergovernmental Panel on Climate Change*. Edited by Parry ML, Canziana OF, Palutikof JP, van der Linden PJ, Hansen CE. Cambridge, U K: Cambridge University Press; 2007:23–78.
8. Shea KM, Truckner RT, Weber RW, Peden DB: Climate change and allergic disease. *J Allergy Clin Immunol* 2008, **122**(3):443–453.
9. Stafoggia M, Forastiere F, Agostini D, Caranci N, de' Donato F, Demaria M, Michelozzi P, Miglio R, Rognoni M, Russo A, Perucci CA: Factors affecting in-hospital heat-related mortality: a multi-city case-crossover analysis. *J Epi Comm Health* 2008, **62**:209–215.
10. Michelozzi P, Accetta G, De Iorio M, D'Ippoliti D, Marino C, Baccini M, Biggeri A, Anderson HR, Katsouyanni K, Ballester F, Bisanti L, Cadum E, Forsberg B, Forastiere F, Goodman PG, Hojs A, Kirchmayer U, Medina S, Paldy A, Schindler C, Sunyer J, Perucci CA, PHEWE Collaborative Group: High temperature and hospitalizations for cardiovascular and respiratory

- causes in 12 European cities. *Am J Respir Crit Care Med* 2009, **179**(5):383–389.
11. Ayres JG, Forberg B, Annesi-Maesano I, Dey R, Ebi KL, Helms PJ, Medina-Ramon M, Menne B, Windt M, Forestiere F: **Climate change and respiratory disease. European Respiratory Society position paper on behalf of the Environment & Human Health Committee.** *Eur Respir J* 2009, **34**:295–302.
 12. D'AG E: **Outdoor air pollution, climate and allergic respiratory diseases: evidence of a link.** *Clin Exp Allergy* 2002, **32**:1391–1393.
 13. D'Amato G, Liccardi G: **Allergenic pollen and urban air pollution in the Mediterranean area.** *Allergy Clin Immunol Int* 2003, **15**:73–78.
 14. D'Amato G: **Outdoor air pollution in urban areas and allergic respiratory diseases.** *Monaldi Arch Chest Dis* 2000, **6**:471–474.
 15. D'Amato G, Cecchi L: **Effects of climate change on environmental factors in respiratory allergic diseases.** *Clin Exp Allergy* 2008, **38**(8):1264–1274.
 16. McDonnell WF, Abbey DE, Nishino N, Lebowitz MD: **Long-term ambient ozone concentration and the incidence of asthma in nonsmoking adults: the AHSMOG Study.** *Environ Res* 1999, **80**:110–121.
 17. McConnell R, Berhane K, Gilliland F, London SJ, Islam T, Gauderman WJ, Avol E, Margolis HG, Peters JM: **Asthma in exercising children exposed to ozone: a cohort study.** *Lancet* 2002, **359**:386–391.
 18. Schelegle ES, Morales CA, Walby WF, Marion S, Allen RP: **Hour Inhalation of Ozone Concentrations from 60 to 87 Parts per Billion in Healthy Humans.** *Am J Respir Crit Care Med* 2009, **180**:265–272.
 19. Gent JF, Triche EW, Holford TR, Belanger K, Bracken MB, Beckett WS, Leaderer BP: **Association of low-level ozone and fine particles with respiratory symptoms in children with asthma.** *JAMA* 2003, **290**(14):1859–1867.
 20. Gauderman WJ, Avol E, Lurmann F, Kuenzli N, Gilliland F, Peters J, McConnell R: **Childhood asthma and exposure to traffic and nitrogen dioxide.** *Epidemiology* 2005, **16**:737–743.
 21. Riedl M, Diaz Sanchez D: **Biology of diesel exhaust effects on respiratory function.** *J Allergy Clin Immunol* 2005, **115**:221–228.
 22. Sacks JD, Stanek LW, Luben TJ, Johns DO, Buckley BJ, Brown JS, Ross M: **Particulate matter-induced health effects: who is susceptible?** *Environ Health Perspect* 2011, **119**:446–454.
 23. Ristovski ZD, Miljevic B, Surawski NC, Morawska L, Fong KM, Goh F, Yang IA: **Respiratory health effects of diesel particulate matter.** *Respirology* 2012, **17**(2):201–212.
 24. Wjst M, Reitmar M, Dold S, Wulff A, Nicolai T, von Loeffelholz-Colberg E, von Mutius E: **Road traffic and adverse effects on respiratory health in children.** *BMJ* 1993, **307**:596–600.
 25. 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.
 26. Laumbach RJ, Kipen HM: **Respiratory health effects of air pollution: update on biomass smoke and traffic pollution.** *J Allergy Clin Immunol* 2012, **129**.
 27. Oftedal B, Brunekreef B, Nystad W, Nafstad P: **Residential outdoor air pollution and allergen sensitization in schoolchildren in Oslo, Norway.** *Clin Exp Allergy* 2007, **37**:1632–1640.
 28. Bernstein JA, Alexis N, Barnes C, Leonard Bernstein I, Bernstein JA, Nel A, Peden D, Diaz-Sanchez D, Tarlo SM, Brock Williams P: **Health effects of air pollution.** *J Allergy Clin Immunol* 2004, **114**:1116–1123.
 29. D'Amato G, Liccardi G, D'Amato M, Cazzola M: **Outdoor air pollution, climatic changes and allergic bronchial asthma.** In *The impact of air pollution on respiratory health*, European Respiratory Monograph, Volume 21. Edited by D'Amato G, Holgate S; 2002:30–51.
 30. Beggs PJ: **Impacts of climate change on aeroallergens: past and future.** *Clin Exp Allergy* 2004, **34**:1507–1513.
 31. D'Amato G: **Urban air pollution and plant-derived respiratory allergy: a review.** *Clin Exp Allergy* 2000, **30**:628–636.
 32. Ariano R, Canonica GW, Passalacqua G: **Possible role of climate changes in variations in pollen seasons and allergic sensitizations during 27 years.** *Ann Allergy Asthma Immunol* 2010 Mar, **104**(3):215–222.
 33. Cecchi L, Morabito M, Domeneghetti MP, Crisci A, Onorari M, Orlandini S: **Long-distance transport of ragweed pollen as a potential cause of allergy in central Italy.** *Ann Allergy Asthma Immunol* 2006, **96**:86–91.
 34. D'Amato G, Liccardi G: **Pollen-related allergy in the European Mediterranean area.** *Clin Exp Allergy* 1994, **24**:210–219.
 35. Melén E, Nyberg F, Lindgren CM, Berglund N, Zucchelli M, Nordling E, Hallberg J, Svartengren M, Morgenstern R, Kere J, Bellander T, Wickman M, Pershagen G: **Interactions between glutathione S-transferase P1, tumor necrosis factor, and traffic-related air pollution for development of childhood allergic disease.** *Environ Health Perspect* 2008, **116**:1077–1084.
 36. Morgenstern V, Zutavern A, Cyrys J, Brockow I, Gehring U, Koletzko S, Bauer CP, Reinhardt D, Wichmann HE, Heinrich J: **Respiratory health and individual estimated exposure to traffic-related air pollutants in a cohort of young children.** *Occup Environ Med* 2007, **64**:8–16.
 37. Brauer M, Hoek G, Smit HA, de Jongste JC, Gerritsen J, Postma DS, Kerkhof M, Brunekreef B: **Air pollution and development of asthma, allergy and infections in a birth cohort.** *Eur Respir J* 2007, **29**:879–888.
 38. Traidl-Hoffmann C, Kasche A, Menzel A, Jakob T, Thiel M, Ring J, Behrendt H: **Impact of pollen on human health: more than allergen carriers?** *Int Arch Allergy Immunol* 2003, **131**:1–13.
 39. D'Amato G, Cecchi L, Bonini S, Nunes C, Annesi-Maesano I, Behrendt H, Liccardi G, Popov T, van Cauwenbergh P: **Allergenic pollen and pollen allergy in Europe.** *Allergy* 2007, **62**:976–990.
 40. D'Amato G, Spiekma FTM, Liccardi G, Jager S, Russo M, Kontou-fili K, Nikkels H, Wutrich B, Bonini S: **Pollen-related allergy in Europe. Position paper of the European Academy of Allergology and Clinical Immunology.** *Allergy* 1998, **53**:567–578.
 41. Islam T, Gauderman WJ, Berhane K, McConnell R, Avol E, Peters JM, Gilliland FD: **Relationship between air pollution, lung function and asthma in adolescents.** *Thorax* 2007, **62**:957–963.
 42. D'Amato G, Liccardi G, D'Amato M, Cazzola M: **Outdoor air pollution, climatic changes and allergic bronchial asthma.** *Eur Resp J* 2002, **21**:12–26.
 43. D'Amato G, Ruffilli A, Ortolani C: **Allergenic significance of Parietaria (pellitory-of-the-wall) pollen.** In *Allergenic Pollen and Pollinosis in Europe*. Edited by D'Amato G, Spiekma F, Bonini S. Oxford: Blackwell Scientific Publications; 1991:113–118.
 44. D'Amato G, Cecchi L, Annesi-Maesano I: **A trans-disciplinary overview of case reports of thunderstorm-related asthma outbreaks and relapse.** *Eur Respir Rev* 2012, **21**(124):82–7.
 45. Packe GE, Ayres JG: **Asthma outbreak during a thunderstorm.** *Lancet* 1985, **ii**:199–204.
 46. Bellomo R, Gigliotti P, Treloar A, Holmes P, Suphioglu C, Singh MB: **Two consecutive thunderstorm associated epidemics of asthma in Melbourne.** *Med J Aust* 1992, **156**:834–837.
 47. Knox RB: **Grass pollen, thunderstorms and asthma.** *Clin Exp Allergy* 1993, **23**:354–356.
 48. Murray V, Venables K, Laing-Morton T, Partridge M, Williams D: **Epidemic of asthma possibly related to thunderstorms.** *BMJ* 1994, **309**:131–132.
 49. Wallis DN, Davidson AC, Welch J, et al: **Clinical and immunological characteristics of patients with thunderstorm asthma.** *Eur Respir J* 1995, **8**:500s. abstr.
 50. Thames Regions Accident and Emergency Trainer Association, Davidson AC, Emberlin J, Cook AD, Venables KM: **A major outbreak of asthma associated with a thunderstorm.** *BMJ* 1996, **312**:601–604.
 51. Celenza A, Fothergill J, Kupek E, Shaw RJ: **Thunderstorms associated asthma: A detailed analysis of environmental factors.** *BMJ* 1996, **312**:604–607.
 52. Venables KM, Allitt U, Collier CG, Emberlin J, Greig JB, Hardaker PJ, Highham JH, Laing-Morton T, Maynard RL, Murray V, Strachan D, Tee RD: **Thunderstorm-related asthma - epidemic 24/25 June 1994.** *Clin Exp Allergy* 1997, **27**:725–736.
 53. Antò JM, Sunyer J: **Thunderstorms: a risk factor for asthma attacks.** *Thorax* 1997, **52**:669–670.
 54. Burney PGJ, Malmberg E, Chinn S, Jarvis D, Luczynska C, Lai E: **The distribution of total and specific serum IgE in the European Community Respiratory Health Survey.** *J Allergy Clin Immunol* 1997, **99**:314–322.
 55. Suphioglu C: **Thunderstorm asthma due to grass pollen.** *Int Arch Allergy Immunol* 1998, **116**:253–260.
 56. D'Amato G, Liccardi G, D'Amato M: **Environmental risk factors (outdoor air pollution and climatic changes) and increased trend of respiratory allergy.** *J Invest Allergol Immunol* 2000, **10**:123–128.
 57. Girgis ST, Marks GB, Downs SH, Kolbe A, Car GN, Paton R: **Thunderstorm-associated asthma in an inland town in southeast Australia. Who is at risk?** *Eur Respir J* 2000, **16**:3–8.
 58. D'Amato G: **Airborne paucimicronic allergen-carrying particles and seasonal respiratory allergy (Editorial).** *Allergy* 2001, **56**:1109–1111.
 59. Marks GB, Colquhoun JR, Girgis ST, Koski MH, Treloar AB, Hansen P, Downs SH, Car NG: **Thunderstorm outflows preceding epidemics of asthma during**

- spring and summer. *Thorax* 2001, **56**:468–471.
60. Taylor PE, Hagan R, Valenta R, Glovsky MM: **Release of allergens in respirable aerosols: A link between grass pollen and asthma.** *J Allergy Clin Immunol* 2002, **109**:51–56.
 61. Villeneuve PJ, Leech J, Bourque D: **Frequency of emergency room visits for childhood asthma in Ottawa, Canada: the role of weather.** *Int J Biometeorol* 2005, **50**:48–56.
 62. D'Amato G, Liccardi G, Gilder JA, et al: **Thunderstorm-associated asthma in pollinosis patients.** *BMJ* 2005, <http://www.bmj.com/rapid-response/2011/10/30/thunderstorm-associated-asthma-pollinosis-patients>.
 63. D'Amato G, Liccardi G, Frenguelli G: **Thunderstorm asthma and pollen allergy.** *Allergy* 2007, **62**:11–16.
 64. Haines A, Smith KR, Anderson D, Epstein R, McMichael AJ, Roberts I, Wilkinson P, Woodcock J, Woods J: **Policies for accelerating access to clean energy, improving health, advancing development, and mitigating climate change.** *Lancet* 2007, **370**:1264–1281.
 65. Pulimood TB, Corden JM, Bryden C, Sharples L, Nasser SM: **Epidemic asthma and the role of the fungal mold *Alternaria alternata*.** *J Allergy Clin Immunol* 2007, **120**:610–617.
 66. D'Amato G, Cecchi L, Liccardi G: **Thunderstorm-related asthma: not only grass pollen and spores.** *J Allergy Clin Immunol* 2008, **120**:530–532.
 67. Grundstein A, Sarnat SE, Klein M, Shepherd M, Naeher L, Mote T, Tolbert P: **Thunderstorm associated asthma in Atlanta, Georgia.** *Thorax* 2008, **63**:659–660.
 68. Nasser SM, Pulimood TB: **Allergens and thunderstorm asthma.** *Curr Allergy Asthma Rep* 2009, **9**:384–390.
 69. Losappio L, Heffler E, Contento, Cannito C, Rolla G: **Thunderstorm-related asthma epidemic owing to *OleaEuropea* pollen sensitization.** *Allergy* 2011, **66**:1510–1511.
 70. D'Amato G, Corrado A, Cecchi L, et al: **A relapse of near-fatal thunderstorm-asthma in pregnancy resolved with magnesium sulphate.** *Eur Ann Allergy Clin Immunol* 2013, in press.
 71. Pawankar R, Canonica GW, Holgate ST, et al: *WAO White Book on Allergy*. Milwaukee: World Allergy Organization; 2011. Available from: www.worldallergy.org/UserFiles/file/WAOWhite-Book-on-Allergy_web.pdf.
 72. D'Amato G, Holgate ST: *The impact of air pollution on respiratory Health*, European Respiratory Monograph, Volume 21. Sheffield UK; 2002.

doi:10.1186/2049-6958-8-12

Cite this article as: D'Amato et al.: Climate change, air pollution and extreme events leading to increasing prevalence of allergic respiratory diseases. *Multidisciplinary Respiratory Medicine* 2013 **8**:12.

Submit your next manuscript to BioMed Central and take full advantage of:

- Convenient online submission
- Thorough peer review
- No space constraints or color figure charges
- Immediate publication on acceptance
- Inclusion in PubMed, CAS, Scopus and Google Scholar
- Research which is freely available for redistribution

Submit your manuscript at
www.biomedcentral.com/submit

