

Systematic Review of Worldwide Variations of the Prevalence of Wheezing Symptoms in Children

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Abstract

Background

Considerable variation in the prevalence of childhood asthma and its symptoms (wheezing) has been observed in previous studies and there is evidence that the prevalence has been increasing over time.

Methods

We have systematically reviewed the reported prevalence and time trends of wheezing symptoms, in the past 12 months, among children, worldwide and within the same country over time. All studies comprising more than 1000 persons and meeting certain other quality criteria published over a 16-year period, between January 1990 and December 2005, are reported and a comparison of ISAAC (International Study of Asthma and Allergies in Childhood) and non-ISAAC studies is made, in part as a way of expanding the power to examine time trends (the older studies tend to be non-ISAAC), but also to examine possible methodological differences between ISAAC and non-ISAAC questions.

Results

A wide range of current prevalence of wheeze was observed between and within countries over time. The UK had the highest recorded prevalence of 32.2% in children aged 13-14 in 1994-5 and Ethiopia had the lowest prevalence, 1.7% in children aged 10-19 in 1996. All studies in Australia and the UK were compared using multiple logistic regression. ISAAC phase I and III studies reported significantly higher prevalence of current wheeze (OR=1.638) compared with non-ISAAC studies, after adjusting for various other factors (country, survey year, age of child, parental vs child response to the survey). Australia showed a significantly higher prevalence of current wheezing (OR=1.343) compared with the UK, there was a significant increase

in the prevalence odds ratio per survey year (2.5% per year), a significant decrease per age of child (0.7% per year), and a significantly higher response in current wheezing if the response was self-completed by the child (OR=1.290). These factors, when explored separately for ISAAC and non-ISAAC studies, showed very different results. In ISAAC studies, or non-ISAAC studies using ISAAC questions, there was a significant decrease in current wheezing prevalence over time (2.5% per year). In non-ISAAC studies, which tend to cover an earlier period, there was a significant increase (2.6% per year) in current wheezing prevalence over time. This is very likely to be a result of prevalence of wheezing increasing from the 1970s up to the early 1990s, but decreasing since then.

Conclusions

The UK has the highest recorded prevalence of wheezing and Ethiopia the lowest. Prevalence of wheezing in Australia and the UK has increased from the 1970s up to the early 1990s, but decreased since then and ISAAC studies report significantly higher prevalences than non-ISAAC studies.

Background

Considerable variation in the prevalence of childhood asthma and its symptoms (in particular, wheezing) has been observed in previous studies and there is evidence that the prevalence has been increasing over time. These differences may, in part, be due to geographical variations and due to methodological problems in defining asthma symptoms.

There are a multiplicity of endpoints used to define and diagnose asthma in an individual. For example, diagnosis is often based on a detailed medical history, including family health history, combined with examination of the upper respiratory tract.¹ Typically, this information is combined with information from laboratory tests. However, diagnostic criteria often differ between doctors in the same locality as well as between countries, and access to health care in different countries can also have an influence on the reported prevalence of doctor-diagnosed asthma.

Epidemiological studies have used different methods of measuring asthma prevalence and its symptoms in surveys. Questionnaires are administered, and depending on the wording of the questions asked, there has been variation in the symptoms elicited. The symptoms may not be present on a particular day, so a one-year period prevalence is often used in epidemiological studies to allow for seasonal variation.

In self-reported asthma, questions are usually asked about wheezing, chest tightness, breathlessness and coughing, but studies have shown that wheezing is the most important symptom for the identification of asthma in epidemiological studies.^{1,2} Some studies have shown that self-reported wheeze has reasonably good specificity and sensitivity for bronchial hyper-responsiveness both in children and adults.³⁻⁵ Wheeze is rarely a symptom of other diseases, in particular emphysema or chronic

bronchitis, which are rare in children, but it is very often indicative of acute viral infection, which is common in this age group.¹⁶³

Doctor diagnosed asthma has been shown to have a lower prevalence than the actual symptoms reported by the individual.^{1 6-8} Until the early 1990's, there was no standardized method of comparing asthma prevalence between countries. Only a small number of studies had used common protocols.⁸⁻¹² In 1991 the International Study of Asthma and Allergies in Childhood (ISAAC) was set up to achieve uniform diagnostic criteria.¹³ Their first worldwide epidemiological study, Phase I, was carried out in 1994-95. It included 56 countries and reported the prevalence of asthma symptoms in 6-7 year old children and in 13-14 year old adolescents.¹⁴ The Phase III study used the same research design as Phase I, but was carried out in 2002-03.¹⁵ The Phase II study comprised a much more detailed investigation of possible correlates of childhood asthma, in particular eczema, and in contrast to ISAAC Phase I and III used 9-11 year old children.^{161, 162} The ISAAC questionnaire is now widely used to assess self-diagnosed asthma by asking about the symptoms.¹⁶

This review has been carried out to assess and summarise the extent of the literature published on wheezing symptoms in children, which includes not only ISAAC but also all non-ISAAC studies that fulfilled specific quality criteria. There are many studies published which are not ISAAC and it is worthwhile to combine the published literature in a review such as this. We report the prevalence and time trends of current symptoms of childhood wheezing in the past 12 months in all studies, worldwide and within the same region over different time periods, and compare the results of ISAAC and non-ISAAC studies. A particular focus of parts of the analysis are studies in the UK and Australia, because of the large number of studies carried out in these two countries – we examine in some detail differences in time trends of

wheeze between the two countries. As we shall see, there are distinct, and perhaps surprising, differences between these two developed countries. In what follows one should note the distinction between the underlying medical condition, “asthma” and its principal symptom, “wheezing”; however, as above, we are referring in all cases to studies of wheezing symptoms.

Methods

Studies included in this systematic review had to satisfy the following requirements:

1. listed in Medline or Embase databases;
2. published in the period January 1990 to December 2005;
3. using the keywords: ‘prevalence’ AND
‘asthma OR wheeze OR wheezing’ AND
‘child OR children OR adolescent’;
4. full journal articles (rather than abstracts) published in English;
5. epidemiological studies of sample size greater than 1000;
6. prevalence of ‘current wheezing’ is reported.

In most epidemiological studies of the prevalence of asthma symptoms, two main types of questions are used.

- (i) ‘Current’ asthma/wheezing, which is normally a period prevalence, and where the question asked is often of the form “Have you had asthma/wheezing *in the past 12 months?*”
- (ii) ‘Lifetime’ asthma/wheezing, in which the question is often “Have you *ever* had asthma/wheezing at anytime in the past?”

Estimates of current prevalence are likely to be more reliable, although Kuehni *et al* have shown that retrospective recall of wheeze at age 8-13 years is a valid proxy measure for lifetime prevalence of wheeze.¹⁷ Questions which are similar to: “In the past 12 months, has the child had wheezing/whistling in the chest?” (from ISAAC questionnaire), i.e. current wheezing, will be used in this review (see footnote ‘Prevalence’ in Table 1). Examples of questions that were asked, and could not be included in this review were: doctor diagnosed asthma (as asthma diagnosis varies by countries); self-reported asthma or asthma attacks in the last 12 months; wheeze ever or occasional wheeze ever (for which there could be recall problems).

A minimum sample size of 1000 was used, as recommended by the ISAAC Steering Committee for small populations¹⁴, to obtain good estimates of wheezing prevalence.

For each selected article in this review, the following information was extracted and given in Tables 1-5:

<i>Country</i>	Country in which the study was carried out
<i>Reference</i>	Reference number of the journal article
<i>Survey year</i>	Year in which the survey was carried out, if reported, otherwise the year prior to publication year
<i>Area</i>	City / region
<i>N (Response rate)</i>	Number of questionnaires returned and response rate
<i>Age (years)</i>	Age range of children sampled (years)
<i>Ascertainment</i>	Method of ascertainment of wheeze (P=Parental-report, S=Self-report)
<i>Prevalence %</i>	The percentage of children (boys and girls combined) who responded ‘Yes’ to the particular wheezing question (see

footnote in Table 1 for different types of questions) out of the total number of children who answered that question, and type of question asked

95% CI

95% confidence interval for the prevalence (if unreported in the article then CI for a single proportion was calculated based on the prevalence and sample size)

In the ISAAC questionnaire, limited agreement has been shown between the written and video questionnaires of symptoms of asthma; the video questionnaire giving lower prevalence rates in 13-14 year old adolescents in two Canadian communities.¹⁸ In the studies reported here, if both methods are used then the written questionnaire results were reported.

To investigate time trends within Australia and the UK, multiple logistic regression models were fitted using wheezing prevalence as the outcome variable. The standard log-linear logistic model was used, so that the probability of an individual in stratum i being affected by wheeze was assumed to be:

$$P[\text{individual in stratum } i \text{ has wheeze}] = \frac{\exp[\alpha_0 + \sum_i \alpha_i z_{ji}]}{1 + \exp[\alpha_0 + \sum_i \alpha_i z_{ji}]} \quad (1)$$

and where $(z_{ji})_j$ are the set of variables (country, year, ISAAC vs non-ISAAC study etc) associated with that stratum. The models were fitted via binomial maximum likelihood using SPSS and the odds ratios, that is to say the quantities, $\exp[\alpha_i]$, and 95% profile-likelihood confidence intervals are reported.¹⁹ For studies where the year of survey was not reported in the article for Australia and the UK, the year prior to the year of publication was used as survey year in the logistic regression analysis.

Results

From the literature search, 2,839 abstracts were listed in Medline, and 2,844 in Embase, from which 712 full articles were selected for further investigation after reviewing the abstracts. From these, 180 satisfied the above criteria. Some articles had referred to the same data set, thus there were 127 distinct studies reported in this review.

Prevalence

Tables 1-5 give the prevalence of current wheezing in children for the five continents. There is a very wide range of current prevalence of wheeze between and within different countries. The UK reported the highest prevalence of current wheeze in the world, 32.2%, in children aged 13-14 in 1994-1995.¹⁴ Ethiopia had the lowest prevalence, 1.7% in children aged 10-19 in 1996.²⁰

Studies in North and South America

Table 1 shows the studies carried out in North and South America; among these countries the USA had the largest number of published studies. A nationwide survey in the USA between 1988-1994 showed that the current prevalence of wheezing amongst 2-3 year olds was 26.4% and amongst 9-11 year olds was 13.4%.²¹ The highest prevalence rates were recorded in North Carolina, 26.1% in children aged 13-14 in 1999-2000.²²

In the rest of North America, Canada had recorded substantially higher prevalence rates in children aged 13-14 (30.6% in Hamilton and 24.0% in Saskatoon) than in children aged 6-7 (20.1% in Hamilton and 14.1% in Saskatoon).¹⁴ The study in Montreal²³ in 6-12 year olds showed very low prevalence of current wheezing, 5.1%; it has been shown that this is likely to be due to unsatisfactory translation of the term wheezing into French, in another study carried out in Quebec.²⁴ Mexico had the

lowest prevalence of current wheezing (<10%)^{14,25}. In Central America, both Costa Rica and Panama showed very high prevalence of wheezing (32.1% and 23.5% respectively) in 6-7 year old children in 1995.¹⁴

In studies carried out in South America high prevalence rates were observed in Chile¹¹ (17.2% in 15-yr-olds to 26.2% in 7-yr-olds), as early as 1990. In Brazil, the ISAAC Phase I study¹⁴ carried out in 1994-95 and the same ISAAC questionnaire methodology used in a study amongst 6-7 and 13-14 year olds carried out in 1999 in one of the same centres as the Phase I study²⁶, both showed higher prevalence of current wheezing than in the non-ISAAC study carried out in 1994, using the same ISAAC methodology in two non-ISAAC centres (iron-mining cities in a mountainous region).

Studies in Europe

Of the five continental groups Europe had the largest number of published studies overall (Table 2). The UK had the highest prevalence, of 32.2% in 1994-5, in the ISAAC Phase I study of 35,485 adolescents.^{14,27} Low prevalence rates, of less than 10%, were observed in Albania, Austria, Belgium, Cyprus, Estonia, Finland, France, Georgia, Greece, Hungary, Italy, Latvia, Malta, Romania, the Slovak Republic, and Switzerland in children aged 6-10, whereas Bulgaria (14.5%),²⁸ the Czech Republic (14.7%),²⁸ Ireland (17.4%),²⁹ and Norway (13.6%)³⁰ had markedly higher prevalence rates.

In the UK, national studies of the prevalence of asthma symptoms (wheezing) reported in 1986 that 6.6% of 16-year-olds had wheezing in the past year³¹ and by 1995 this had increased to 32.3% among 12-14 year old children¹⁴, using comparable questions. In the younger age group (6-10 years) in the UK, the current prevalence of

wheezing ranged from 7.6% in 1980³¹ to 20.2% in 1999³², using comparable questions.

Studies in the Eastern Mediterranean and Africa

Apart from the ISAAC studies conducted in 1994-95, very few countries had carried out epidemiological studies of asthma in the Eastern Mediterranean and Africa, reported in English (Table 3).

In Africa, very low prevalence rates were observed in Ethiopian rural communities (2.0% in 0-9 year olds, 1.7% in 10-19 year olds),²⁰ intermediate levels of wheeze prevalence (5%-14%) were observed in Algeria, Kenya, Morocco and Nigeria and the highest rates were in South Africa, 26.8% in 7-8 year olds in 1993.³³

In the Eastern Mediterranean, Iran, Oman and Palestine (West Bank) had the lowest prevalence of wheeze (<11%), while the highest rates were observed in Israel (17.9%)³⁴, Kuwait (16.1%)³⁵, and Malta (16.0%)¹⁴, amongst 13-14 year old adolescents. In Turkey, many of the studies had not used the ISAAC question and the prevalence of wheeze was low.

Studies in Asia

Amongst studies conducted in Asia, low prevalence rates (<9%) were observed in China, Hong Kong, India, Indonesia and Malaysia while Japan (17.3%)¹⁴, Korea (13.6%)³⁶ and Singapore (15.7% in 1994 and 10.2% in 2001)^{14 37} had higher prevalence rates in 6-7 year olds (Table 4). The majority of the studies reported had used the ISAAC questions relating to current wheeze.

Studies in Australasia

Australia, New Zealand and Fiji had a very high prevalence of current wheezing with the majority of the studies showing the prevalence of current wheezing in the range 18% - 30% (Table 5). The highest prevalence of 30.2% was observed in New

Zealand¹⁴ amongst a very large sample of 13-14 year olds in 1992-93, followed by Australia⁸ which observed a prevalence of 29.7% amongst 12-15 year olds in 1991. Fiji³⁸ reported a prevalence of 21.0% in 1990 amongst 9-10 year old children.

Subgroup analysis: Studies in Australia and the UK

Australia and the UK had the largest number of studies carried out and published (14 and 25 publications respectively over the 16-year period), and these are investigated further to assess differences in prevalence and trends in prevalence between the two countries.

The overall trend of the current wheeze prevalence, by calendar year for both Australia and UK can be seen in Figure 1. This shows overall (over the period 1990-2005) a clear increasing trend for both countries. Table 6 shows the findings of multiple logistic regression analysis of the effects of country (UK versus Australia), year of survey, age of child, parental- or self-report questionnaire, and if the study was an ISAAC study or not, on the current prevalence of wheeze. This allows each factor to be assessed while adjusting for the other factors. When using the data from all studies, Australia showed a statistically significant increase in prevalence compared with the UK, OR=1.343 [95% CI 1.318, 1.369] allowing for all other factors. There was also a statistically significant increase in the prevalence per survey year (2.5% per year; 95% CI 2.3%, 2.6%), and a significant decrease per year of age of child (0.7% per year; 95% CI 0.3%, 1.1%). If the study was an ISAAC study, phase I or III, then the odds of wheezing was significantly higher compared with non-ISAAC studies [OR=1.638; 95% CI 1.598, 1.678], after adjusting for the other factors. Similar results were obtained when comparing ISAAC question with a non-ISAAC question (OR=1.331; 95% CI 1.304, 1.359) after adjusting for all other factors.

When analysis was restricted to ISAAC studies there was a statistically significant decrease in the prevalence per survey year (2.5% per year; 95% CI 2.1%, 2.9%), after allowing for age of child and country (neither of which were significant).

If the analysis is restricted to non-ISAAC studies, even if the study had used the ISAAC question, then there was a statistically significant increase in prevalence of wheezing in Australia compared with the UK [OR=1.470; 95% CI 1.438, 1.502], a significant increase in the prevalence per survey year (2.6% per year; 95% CI 2.4%, 2.8%), a significant decrease in the prevalence per age of child (1.1% per year; 95% CI 0.7%, 1.5%), and a significant increase in wheezing if the questionnaire was self-report compared with parental-report [OR=1.242; 95% CI 1.197, 1.287]. This shows that the results of ISAAC studies are very different from non-ISAAC studies.

The majority of studies carried out since ISAAC phase I, have used the ISAAC question on wheezing, even if the study was not a phase I or phase III ISAAC study. So the analysis was then restricted only to studies which had used the ISAAC question on wheezing. The results showed that there was no significant difference in the prevalence of wheezing between UK and Australia and age of child was also not significant, after controlling for year of survey and parental- or self-report questions. However there has been a statistically significant decrease in the prevalence per survey year (2.3% per year; 95% CI 1.9%, 2.7%) and a significant increase in reported wheezing if the response was self- compared with parental-report (OR=1.739; 95% CI 1.624, 1.862).

As the rate of increase in the trends for wheezing prevalence over time looked different in UK and Australia (Figure 1), an extra interaction terms was fitted in the multiple logistic regression model using all studies (Table 7). The country by year interaction effect shows that Australia has a significantly lower rate of increase in

wheezing (6.2% per calendar year; 95% CI 5.7%, 6.7%) compared with the UK, after allowing for all other factors.

Discussion

In this review we have reported the prevalence of current wheezing in children, published in all epidemiological studies comprising more than 1000 persons and meeting certain other quality criteria (see the Methods), over a 16-year-period, between 1990 and 2005, and further investigated the differences in reported symptoms of wheeze between ISAAC and non-ISAAC studies, in the UK and Australia.

Overall, the highest prevalence rates of current wheezing were reported in the UK, Australia and New Zealand, and the lowest prevalence was found in Albania, China, Ethiopia, Indonesia and Turkey, which gives an indication of the difference between developed and developing countries. The pattern in Africa and Asia also supports this. However, this is not supported in America, where Chile, Costa Rica and Peru had equally high wheezing prevalence as the US and Canada. Chile and Costa Rica are relatively developed countries, that may have similar characteristics in relation to development of wheezing in childhood as fully developed countries such as the US and Canada. However, this apparent inconsistency (in relation to Peru) requires further research.

Within the UK there was slightly higher prevalence of wheezing in adolescents in Scotland compared with England but there were no other substantial geographic variations, suggesting no major impact of climate, diet or outdoor environment.³⁹ Also, prevalence of wheezing was lower in children born outside the UK but currently residing in the UK, suggesting a role of the environment in infancy and possibly

heritable genetic factors.^{39 40} However, although genetic factors are important risk factors for individuals with symptoms of asthma, migrant studies indicate that they are unlikely to be responsible for the large variations in asthma symptoms that exist between populations, and cannot be responsible for the increasing prevalence of asthma within populations.⁴¹ Environmental factors are likely to be more important and offer the greatest opportunities for prevention.

The cross-sectional ISAAC phase I study, carried out in 1994-1995, was a major achievement, and repeated in 2002-2003, the phase III study.^{14 15} However, the selected ISAAC centres were most commonly an urban area (a city) and therefore may not be representative of the country. This is illustrated, for example, in Brazil where in the ISAAC phase I centres, which were all major cities, the prevalence of current wheezing in 6-7 year olds was higher than in non-ISAAC centres, which were iron-mining cities and mountainous regions (23% vs 14% respectively). The ISAAC studies also have the disadvantage of reporting wheezing symptoms only amongst two age groups (6-7 yr and 13-14 yr), and at only 2 time points (phase I and III), whereas this review shows the results of all studies of all age groups.

This review has shown that differing rates of asthma symptoms are observed in developed and developing countries. The validity of the question on wheezing in the questionnaire is likely to have varied across cultures as some languages do not have an equivalent word for “wheezing” as understood by English speakers. However, large variations in the prevalence of wheezing across the countries and over time, found in these studies are unlikely to be explained by methodological factors alone. When making comparisons of the prevalence of wheeze or asthma between different studies, it is necessary to critically assess the content of the question. There is, as yet, no accepted definition of asthma and identification of asthma by questionnaire

remains a contentious issue.⁴² One question is whether the everyday meaning of the word wheezing has changed over time. Do better educated parents use this word more freely for symptoms in their children? The threshold of observing mild respiratory symptoms could be lower now than previously and health campaigns may have increased parental awareness of symptoms in their children. Another interesting hypothesis is the loss of protective effect of respiratory infection in early childhood, the “hygiene hypothesis”.⁴³ This confirms the importance of the ISAAC phase II data collection, which was completed in 2003, and in which objective measures of pulmonary function and bronchial responsiveness are recorded in conjunction with other factors, so that further study of possible aetiological factors common to different countries can be investigated.

Australia and the UK had the most published studies on wheezing prevalence in children and were investigated in much more detail (Tables 6, 7). In addition, in these countries the ISAAC studies reported significantly higher wheezing prevalence than non-ISAAC studies and this was not due to an increase over time, nor age of child, both of which (and various other variables) were adjusted for in the analyses. It is possible that this is due to the selection of the ISAAC centres, which were all major cities. There are various possible explanations for the differences in current prevalence of wheezing including susceptibility to environmental stimuli and changes in exposure to environmental factors, genetic susceptibility, diet, low birth weight and young maternal age⁴⁵⁻⁵⁰.

Using all the studies in UK and Australia, we find that there was a significantly higher odds of wheezing in Australia than the UK, but the rate of increase in Australia is significantly lower than the UK, as shown by the highly significant interaction between country and year. The multiple logistic regression analysis we performed for

the prevalence of wheezing adjusted for age, time period, type of study (ISAAC vs non-ISAAC) and type of response (parental or self report) and does not appear therefore to result from methodological bias (e.g., confounding by ISAAC status). These differences indicate some significant discrepancy in early life environment between the two countries over the last 20 or so years.

If only ISAAC studies are investigated then there was no difference in prevalence between the two countries, after adjusting for time and age of the child, whereas non-ISAAC studies show significantly higher odds of wheezing in Australia. This is very likely because the ISAAC studies in these countries were carried out at two similar time points and for two age groups. The non-ISAAC studies span a much larger time period, use a wider range of ages and include many more study groups, and in particular are not restricted to large conurbations.

If time trends are explored in all studies in the UK and Australia, then overall there is a significant increase in the odds of wheezing over time, but only ISAAC studies show a significant decrease, which was also reported in the recent results of the phase I in 1995 and phase III in 2002, ISAAC study comparisons¹⁵. This is almost certainly a result of the different time periods covered by these two sorts of survey. The ISAAC studies cover the period from the early 1990s onwards, whereas other studies tend to cover earlier years, some as early as 1975. This is consistent with the trend of wheezing prevalence increasing since the 1970's but levelling out in the most recent 10-15 years, as shown in Figure 1. This is also confirmed in other reports; evidence from many repeat surveys shows that the prevalence has increased over the past 3 decades⁵¹, but in studies between 1991 to 1998 the increase was confined to minor symptoms of asthma⁵².

A decrease in reported symptoms of wheezing, as the child gets older, was observed in the UK and Australia, which confirms previous reports⁵³. In ISAAC studies the age effect is not significant, perhaps because only two age groups were studied. Self-report of wheezing was significantly higher than parental-report, in the UK and Australia, again confirming previous work⁴⁴.

The ISAAC questionnaire has become almost ubiquitous since the early 1990's and we have shown that using an ISAAC question or the results of the ISAAC studies give similar results within Australia and the UK.

Wheeze may indicate undiagnosed asthma in some patients. Some studies have shown that self-reported wheeze has reasonably good specificity and sensitivity for bronchial hyper-responsiveness both in children and adults.³⁻⁵ Wheeze is rarely a symptom of emphysema or chronic bronchitis in children, but it is very often indicative of acute viral infection, which is common in this age group.¹⁶³ Doctor-diagnosed asthma tends to be reported in only a small proportion (about 40%) of persons reporting wheeze,¹⁶⁴ so that the possibility of selection or information bias in studies of asthma or wheeze cannot be discounted in general.

In summary, the strength of this review is the reporting of the prevalence of all studies of more than 1000 persons, providing a full description of the scale and distribution of asthma symptoms (wheeze in the past year), worldwide and over time within each country. We have documented a clear increase in the prevalence over time within Australia and the UK, with a levelling off or even decline in prevalence in more recent years. ISAAC studies show higher prevalence rates of wheezing than non-ISAAC studies.

Competing Interests

The authors declare that they have no competing interests.

Authors Contributions

SP performed database searches, MPL determined and oversaw the statistical analyses, and all three authors contributed substantially to the conception, design and writing of the paper.

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There is no conflict of interest.

Table 1. Studies of wheeze prevalence in North and South America

Country	Reference	Survey Year	Area	N (Response rate)	Age (years) / ascertainment	Prevalence %	95% CI
Argentina	14	'94-'95	Buenos Aires, Rosario	6,012 [†] (>80%):	6-7: P	16.4 ^{IS}	15.5, 17.3*
				5,374 (89.5%):	13-14: S	10.9 ^{IS}	10.1, 11.7*
Brazil	54	'94	Santa Maria & Itabira	2,714 (77%):	7-8: P	14.3 ^{IQ}	13.0, 15.6
				2,468 (77%):	13-14: P	9.3 ^{IQ}	8.2, 10.4
	14	'94-'95:	Porto Alegre, Recife, Sao Paulo:	7,261 [†] (>80%):	6-7: P	23.3 ^{IS}	22.3, 24.3*
			'95-'96:	Curitiba, Porto Alegre, Recife, Salvador, Sao Paulo:	14,697 (95.1%):	13-14: S	22.7 ^{IS}
	55	'95-'96	Sao Paulo	1,132 (96%)	0.5-5 P	12.5 ^{IQ}	10.6, 14.4*
26	'99	Sao Paulo	3,003:	6-7: P	24.3 ^{IS}	22.8, 25.8*	
			3,487:	13-14: S	22.1 ^{IS}	20.7, 23.5*	
Canada	14	'94:	Hamilton:	5,755 (75.1%):	6-7: P	20.1 ^{IS}	18.8, 21.5
					13-14: S	30.6 ^{IS}	29.0, 32.2
	'93-'94:	Saskatoon:	4,952 (68.6%):	6-7: P	14.1 ^{IS}	12.7, 15.4	
				13-14: S	24.0 ^{IS}	22.1, 26.0	

(P=Parental-report
S=Self-report)

	23	Published	Montreal	1,111 (83.9%)	6-12	P	5.1	IQ	3.8, 6.4 ^{21*}
		'95							
Caribbean	56	'02	Trinidad & Tobago	4,988 (87.8%)	11-19	S	13.2	IQ	12.3, 14.1*
Chile	11	'90	La Serena	1,730 (71%):	7:	P	26.2	A	25.0, 27.5
				1,820 (71%):	12:	P	20.6	A	18.5, 22.7
				1,851 (71%):	15:	P	17.2	A	15.7, 18.8
	14	'94-'95	Central Santiago, Punta Arenas, South	10,838 [†] (>80%):	6-7:	P	17.9	IS	17.2, 18.6*
			Santiago, Valdivia	11,780(92.7%):	13-14:	S	10.2	IS	9.7, 10.7*
Costa Rica	14	'94-'95	Nationwide	2,942 [†] (>80%):	6-7:	P	32.1	IS	30.4, 33.8*
				2,925 (>80%):	13-14:	S	23.7	IS	22.2, 25.2*
	57	'98	Nationwide	1,105 (74%)	10	P	27.1	IQ	24.5, 29.7*
Ecuador	58	Published	Pichincha & Esmeraldas	4,433 (96.3%)	5-18	P	10.3	IQ	8.8, 11.9
		'03							
Mexico	14	'94-'95	Cuernavaca	3,097 [†] (>80%):	6-7:	P	8.6	IS	7.6, 9.6*
				2,863 (92.3%):	13-14:	S	6.6	IS	5.7, 7.5*
	25	Published	Ciudad Juarez, Chihuahua	3,390 (92%):	6-8:	P	9.7	IQ	8.7, 11.0
		'03		2,784 (92%):	11-14:	P	5.8	IQ	5.0, 6.8

Panama	14	'94-'95	David-Panama	3,043 [†] (>80%):	6-7:	P	23.5	IS	²² 22.0, 25.0*
				2,775 (96.2):	13-14:	S	17.6	IS	16.2, 19.0*
Paraguay	14	'94-'95	Asuncion	2,764 (93.2%)	13-14	P	19.4	IS	18.0, 20.8*
Peru	14	'94-'95	Lima	3,051 (96.6%)	13-14	S	26.0	IS	24.5, 27.5*
Uruguay	14	94-'95	Montevideo	3,071 [†] (>80%)	6-7:	P	18.0	IS	16.6, 19.4*
				2,860 (93.1%)	13-14:	S	19.0	IS	17.6, 20.4*
USA	21	'88-'94	Nationwide	2,318:	2-3:	P	26.4	IQ	24.6, 28.2*
				2,070:	4-5:	P	19.4	IQ	17.7, 21.1*
				1,545:	6-8:	P	19.2	IQ	17.2, 21.1*
				1,605:	9-11:	P	13.4	IQ	11.7, 15.0*
	59	'94	Illinois	2,693 (90%)	13-18	S	25.1	IQ	23.5, 26.7*
	60	'94-'95	Seattle	1,865 (37%)	5-9	P	18.8	B	17.0, 20.6*
	14	'94-'95	Chicago, Seattle	6,660 (88.7%)	13-14	S	21.7	IS	20.8, 22.6*
61	'97	New York City – East Harlem	1,139 (82%)	5-12	P	24.5	IQ	22.0, 27.0*	
62	Published	North Carolina	1,596 (92%)	13-14	S	21.0	IQ	19.0, 23.0*	
	'00								
22	'99-'00	North Carolina	128,568 (88%)	13-14	S	26.1	IQ	25.9, 26.3*	
63	'00	New Orleans	1,535	5-18	P/S	25.7	IQ	23.5, 27.9*	

64	'01	South Plains & Panhandle, West Texas	1,500 (64%)	<16	P	17.5	IQ	15.6, 19.4 ²³ *
65	'00-'02	Iowa	3,090 (86.6%)	6-14	P	19.1	IQ	17.7, 20.5*

Key:

'Prevalence'

IS: ISAAC study, with question "Have you had wheezing and whistling in the chest in the last 12 months?" (Yes/No)

IQ: ISAAC question, but not an ISAAC study

A: In the past 12 months has your child had a wheezing or asthma attack? (Yes/No)

B: Current wheezing without a diagnosis of asthma & Physician diagnosed asthma

C: In the last 12 months, has a wheeze (that is, a whistling noise, high or low pitched) ever been heard from your child's chest?

D: Has your child (ever) wheezed in the past 12 months?

E: Wheeze in the previous year (interview questionnaire)

F: Have you had wheezing attacks in the past year?

G: Has your child had wheezing in the chest (but not from the throat or nose)

H: Wheezy or whistling sound in the chest when having a cold or occasionally apart from colds or for most days or nights, in the past 12 months

* CI not given in the publication and calculated by author

† N is the number of questionnaires given out & response rate obtained from ISAAC study^{14 66}

Table 2. Studies of wheeze prevalence in Europe

Country	Reference	Survey Year	Area	N (Response rate)	Age (years) / ascertainment	Prevalence %	95% CI
Albania	14	'94-'95	Tirane	2,981 [†] (>80%):	6-7: P	7.6	IS 6.6, 8.6*
				2,862 (96.8%):	13-14: S	2.6	IS 2.0, 3.2*
Austria	14	'94-'95	Salzburg, Urfahr-Umgebung	5,787 [†] (>80%)	6-7: P	8.9	IS 8.2, 9.6*
				4,270 (87.40%)	13-14: S	11.6	IS 10.7, 12.5*
Belgium	67	'97	Salzburg	2,283 (95.4%)	8-10 P	7.1	IQ 6.0, 8.2*
				5,782 (88.5%):	6-7: P	7.3	IS 6.6, 8.0*
Bulgaria	14	'94-'95:	Antwerp	1,477 (97.5%):	13-14: S	12.0	IS 10.3, 13.7*
		'95-'96:		3,631 (92%)	7-11 P	14.5	IQ 13.4, 15.6*
Czech Republic	28	'96	Four urban areas	3,479 (55%)	7-11 P	14.7	IQ 13.5, 15.9*
Cyprus	68	'97	Northern Cyprus	2,529 (89.6%)	6-14 P	4.8	IQ 4.0, 5.6
Estonia	69	'92-3	Tallin & Tartu	1,519 (96.1%)	10-12 P	7.0	C 5.7, 8.3*
	70	'93-'94:	Tallinn	3,070 (90.8%):	6-7: P	9.3	IS 8.3, 10.3*
				3,506 (88.7%):	13-14: S	8.5	IS 7.6, 9.4*

(P=Parental-report
S=Self-report)

		'01-'02:		2,388 (85.7%):	6-7:	P	9.7	IS	8.5, 10.9 ^{25*}
				3,605 (93.4%):	13-14:	S	9.2	IS	8.3, 10.1*
Finland	71	'93	Kuopio	2,564 (86%)	7-12	P	5.4	D	4.5, 6.3*
	14	'94-'95	Lapland, Kuopio, Turku & Pori:	8,836:	13-14	S	14.7	IS	14.0, 15.4*
			Helsinki:	2,771:			19.8	IS	18.3, 21.3*
France	14	'94-'95	Pessac:	3,202 [†] (>80%):	6-7:	P	8.1	IS	7.2, 9.0*
			Marseilles, Montpellier, Pessac, Strasbourg, West Marne:	15,410 (83.1%):	13-14:	S	13.5	IS	13.0, 14.0*
	72	'99-'00	Bordeaux, Clermont-Ferrand, Creteil, Marseille, Strasbourg, Reims	4,901 (80.9%)	9-11	P	8.0	IQ	7.2, 8.8*
Georgia	14	'94-'95	Kutaisi, Tbilisi	6,770 [†] (>80%):	6-7:	P	7.6	IS	7.0, 8.2*
				6,098 (90.4%):	13-14:	S	3.6	IS	3.2, 4.0*
Germany	8	'91	Bochum	1,928 (93%)	12-15	S	20.0	IQ	18.2, 21.8*
	14	'94-'95	Munster (W Germany)	3,036 (81.2%):	6-7:	P	9.6	IS	8.6, 10.6*
				3,763 (94.0%):	13-14:	S	14.1	IS	13.6, 15.8*
			Greifswald (E Germany)	2,451 (85.9%):	6-7:	P	7.2	IS	6.2, 8.2*
				2,776 (87.6%):	13-14:	S	13.3	IS	12.0, 14.6*
	73	'95-'96	E Germany (Leipzig, Dresden) :	9,524 (84.3%):	5-11:	P	7.7	IQ	7.2, 8.2*

			W Germany (Munich):	4,777:	5-11:	P	8.8	IQ	8.0, 9.6 ²⁶ *
	74	'95-'96	Zerbst, Bitterfeld, Hettstedt	2,814 (74.7%)	5-14	P	24.8	IQ	23.1, 26.5*
	75	'99-'00	Munster (W Germany)	3,529 (82%):	6-7:	P	13.2	IS	2.1, 14.3*
				3,816 (94%):	13-14:	S	17.5	IS	16.3, 18.7*
Greece	14	'94-'95	Athens	1,654 [†] (>80%)	6-7:	P	7.6	IS	6.3, 8.9*
				2,228 (87.0%)	13-14:	S	3.7	IS	3.0, 4.4*
Hungary	28	'96	Five urban areas	3,721 (66%)	7-11	P	6.5	IQ	5.7, 7.3*
Ireland	76	'92-'93	Dublin, Wicklow, Kildare	1,416:	4-10:	P	16.5	C	14.6, 18.4*
				1,317:	11-19:	P	14.9	C	13.0, 16.8*
	14	'95	Nationwide	2,898 (92.1%)	13-14	S	29.1	IS	27.4, 30.8
	29	'00	Counties Louth & Mead	1,899 (61.4%)	6-7	P	17.4	IQ	15.7, 19.1*
Italy	14	'93-'95	Northern & Central Italy	19,982 (96%):	6-7:	P	7.3	IS	6.9, 7.7*
				25,497 (96.3%):	13-14:	S	8.9	IS	8.6, 9.2*
	77	'02	North, Central & South Italy	20,016 (89%):	6-8:	P	7.9	IQ	7.5, 8.3*
				13,266 (81%):	13-14:	P+S	8.3	IQ	7.8, 8.8*
Latvia	14	'94-'95	Riga:	3,003 [†] (>80%):	6-7:	P	7.3	IS	6.4, 8.2*
			Riga, Rural Latvia:	5,755 (93.6%):	13-14:	S	8.4	IS	7.7, 9.1*
Malta	78	'95	Nationwide	4,184 (88.7%)	13-15	S	16.0	IS	14.9, 17.1*

	79	Published '02	Malta and Gozo	3,506 (78.5%)	5-8	P	8.8	IS	7.9, 9.7 ^{*27}
Norway	30	Published '97	Oslo, Hallingdal, Odde	924 (88%):	6-8:	P	13.6	IQ	11.4, 15.8
				1,877 (88%):	9-12:	P	11.4	IQ	10.0, 12.8
				1,213 (88%):	13-16:	P	12.2	IQ	10.4, 14.0
Poland	80	'95-'97	Nord-Trondelag County	8,571 (86%)	13-19	S	26.1	IQ	25.2, 27.0 [*]
	14	'93-'95	Krakow, Poznan	4,974 [†] (>80%):	6-7:	P	10.9	IS	10.0, 11.8 [*]
				9,282 (91.3%):	13-14:	S	8.1	IS	7.6, 8.6 [*]
	28	'96	Four urban areas	2,932 (66%)	7-11	P	9.6.	IQ	8.5, 10.7 [*]
Portugal	81	'96-'98	Swietochlowice, Kedzierzyn-Kozle, Pszczyna & Kielce	1,561 (80%)	9-11	P	8.6	D	7.2, 10.0 [*]
	14	'96	Funchal, Lisbon, Portimao:	5,129 [†] (>80%):	6-7:	P	13.2	IS	12.4, 14.2 [*]
			Funchal, Lisbon, Portimao, Porto:	9,977 (92.8%):	13-14:	S	9.5	IS	8.9, 10.1 [*]
Romania	14	'94-'95	Cluj	3,362 (99.0%)	13-14	S	3.0	IS	2.4, 3.6 [*]
	28	'96	Four urban areas	3,470 (66%)	7-11	P	6.4	IQ	5.6, 7.2 [*]
Russia	14	'94-'95	Moscow	2,910 (85.3%)	13-14	S	4.4	IS	3.7, 5.1 [*]
Slovak	82	Published '04	9 cities	5,951 (98%)	8-12	P	13.4	IQ	12.5, 14.3 [*]
	28	'96	Four urban areas	3,038 (66%)	7-11	P	4.5	IQ	3.8, 5.2 [*]

Republic									
Spain	14	'94-'95	5 cities:	16,884 [†] (>80%):	6-7:	P	6.2	IS	5.8, 6.6*
			8 cities:	23,770 (95.0%):	13-14:	S	10.3	IS	9.9, 10.7*
	83	'02-'03	11 cities	18,599 (71.5%):	6-7:	P	9.4	IS	9.0, 9.8*
				24,214 (87.3%):	13-14:	S	9.2	IS	8.8, 9.6*
	84	Published '05	3 cities	2,720 (33.8%)	9-12	P ^{IQ}	13.1	IQ	11.8, 14.4*
Sweden	10	'90	Norrkoping	1,276 (90.4%)	12	P	9.2	D	7.6, 10.8*
	14	'94-'95	Stockholm, Uppsala:	3,029 [†] (>80%)	6-7:	P	10.4	IS	9.3, 11.5*
			Linkoping, Stockholm, Uppsala:	6,039 (93.6%)	13-14:	S	12.9	IS	12.1, 13.7*
	85	'91: (cohort)	Goteborg, Karuna	2,481 (87.6%):	7-8:	P	7.6	E	6.6, 8.6*
	86	'96-'98:	Karuna, Lulea, Pitea	3,247 (92%)	7-10:	P	10.9	IQ	9.8, 12.0*
	87	'97	Linkoping, Ostersund	2,108 (84.2%)	10-11	P	9.2	IQ	8.0, 10.4*
	88	'00	Varmland	10,851 (79%)	1-6	P	19.3	IQ	18.6, 20.0*
	89	'02	Lulea, Pitea, Karuna	3,345 (95.2%)	13-14	S	10.5	IQ	9.5, 11.5*
Switzerland	90	'90	Canton of St Galon	5,338 (97.8%)	7:	P	7.4	A	5.9, 8.8
					12:	P	6.0	A	4.7, 7.3
					15:	P	4.5	A	3.5, 5.6
	91	'92-'93:	4 urban & 4 rural communities	1,324 (74.9%):	13-14	S	9.7	IQ	8.1, 11.3*

			'95-'97:	1,668 (80.6%):			9.2	IQ	7.8, 10.6 ^{29*}	
			'99-'00:	1,250 (73.9%):			10.9	IQ	9.2, 12.6*	
	92		'92-'93:	Urban, suburban, rural and alpine areas	4,026:	6-14	P	8.2	IQ	5.6, 11.8*
			'98-'01:	areas	9,591:			6.1	IQ	4.2, 8.9*
The Netherlands	93		Published '92	Helmond	3,344 (73%)	6-12	P	10.1	C	9.1, 11.1*
	94		'91-'92	Nationwide (Child Health Monitoring System)	2,677 (92%):	4-9:	P	8.9	D	7.8, 10.0*
					2,509 (92%):	10-15:		5.5	D	4.6, 6.4*
						(P<12 / S)				
	95		'93	Melick-Herkenbosch, Leek	1,190 (97%)	6-12	P	14.4	D	12.4, 16.4*
	96		'89:	Westelijke Mijnstreek	1,794 (>95%):	8-9	P	13.4	D	11.8, 15.0*
			'93:		1,526 (>95%):			13.3	D	11.6, 15.0*
			'97:		1,670 (>95%):			11.9	D	10.3, 13.5*
			'01:		1,102 (95.5%):			9.1	D	7.4, 10.8*
Uzbekistan	14		'94-'95	Samarkand, Tashkent	4,354 (93.4%)	13-14	S	9.2	IS	8.4, 10.0*
UK	31		'75:	Nationwide	12,402:	5:	P	9.9	D	9.4, 10.4*
			'80:		13,575:	10:	P	7.6	D	7.2, 8.0*
			'86:		9,006:	16:	S	6.6	D	6.1, 7.1*

97	'87:	Southampton	3,187 (90%):	6-8:	P	14.6 ^D	13.4, 15.8 ^{30*}
	'95:		2,289 (75.5%):	14-16:	P	18.2 ^D	16.6, 19.8 [*]
98	'89	Walsall	1,334 (81.8%)	6-7	P	11.1 ^{IQ}	9.4, 12.8 [*]
40	'89-'90	Southampton	1,033 (76.7%)	7-12	P	17.2 ^D	14.9, 19.5 [*]
99	'90	Leicestershire	1,422 (86.2%)	0-5	P	13.0 ^F	11.3, 14.8
100	'78:	Croydon	4,147 (87%):	7.5-8.5	P	11.1 ^F	10.1, 12.1 [*]
	'91:		3,070 (81%):			12.9 ^F	11.7, 14.1 [*]
8	'91	West Sussex	2,097 (92%)	12-15	S	29.1 ^{IQ}	27.2, 31.0 [*]
52	'91:	Sheffield	4,580 (85.3%):	8-9:	P	17.0 ^{IQ}	15.9, 18.1
	'99:		5,011 (83.2%):	8-9:	P	19.4 ^{IQ}	18.3, 20.5
101	'92	National	2,866:	5-10:	P	16.3 ^{IQ}	14.9, 17.7 [*]
			2,606:	11-17:	P	13.6 ^{IQ}	12.3, 14.9 [*]
102	'93	East Surrey	1,936 (97%)	13-14	S	24.3 ^{IQ}	22.4, 26.2 [*]
103	'88:	Nottingham	13,579 (78%):	4-11	P	12.5 ^F	11.9, 13.1 [*]
	'95:		22,968 (83%):			15.1 ^F	14.6, 15.6
104	'92:	Scottish Highlands	1,825:	12,14:	P	19.2 ^C	17.4, 21.0 [*]
	'94:		1,537 (85%):	12,14:	P	17.4 ^C	15.5, 19.3 [*]
14	'94-'95	Sunderland:	1,864 [†] (>80%):	6-7:	P	18.4 ^{IS}	16.6, 20.2 [*]

		Nationwide:	35,485 (86.9%)	13-14:	S	32.2	IS	31.8, 32.8 ^{31*}
105	'96	Nottingham	27,826	11-16:	S	19.0	F	18.5, 19.5*
106	'90:	Leicestershire	1,422 (86%):	1-5:	P	12.3	F	10.6, 14.0*
	'98:		2,127 (84%):	1-5:	P	25.4	F	23.6, 27.2*
107	Published '99	North-East England	3,000 (80.0%)	6-7	P	18.0	IQ	16.6, 19.4
32	'99	Sheffield	4,311 (83.2%)	8-9	P	20.2	IQ	19.0, 21.4*
108	'99-'00	Isle of Wight	1,373 (94.3%)	10	P	18.9	IQ	16.8, 21.0*
109	'93:	Manchester	2,478:	<16	P	27	IQ	25, 29
	'95:		2,228:			24	IQ	22, 26
	'99:		1,985:			25	IQ	23, 27
	'01:		1,781:			22	IQ	20, 24
110	'00-'01	Aberdeen	1,374 (71.4%)	2	P	15.3	G	13.4, 17.2*
111	Published '02	Swindon, Basingstoke, Hook, Fleet	1,732 (56.2%)	7-9	P	18.9	IQ	17.1, 20.7*
112	'02	Nationwide	15,755	12-14	S	27.5	IQ	26.8, 28.2*
113	'03	East Midlands, Eastern Region	11,562	4-6	P	14.1	IQ	13.5, 14.7*

'Prevalence' key, see Table 1.

Table 3. Studies of wheeze prevalence in the Eastern Mediterranean and Africa

Country	Reference	Survey Year	Area	N (Response rate)	Age (years) / ascertainment	Prevalence %	95% CI
Algeria	¹⁴	'94-'95	Algiers	1,147 (97.8%)	13-14 S	7.8 ^{IS}	7.3, 8.3*
Ethiopia	¹⁴	'94-'95	Addis Ababa, Jima	5,906 (98.8%)	13-14 S	6.2 ^{IS}	5.6, 6.8*
	²⁰	'96	Jimma and 3 rural communities	2,929:	0-9: P	2.0 ^D	1.5, 2.5*
				3,957:	10-19: S	1.7 ^D	1.3, 2.1*
	¹¹⁴	'97	Gondar Zuria, Dembia, Wegera, Lai Armachiho	3,365 (98.4%)	13-14 S	16.2 ^{IQ}	15.0, 17.4*
	¹¹⁵	'00-'01	Jimma & 3 rural communities	7,155	1-4 P	3.4 ^{IQ}	3.0, 3.8*
Iran	¹⁴	'94-'95	Rasht, Tehran	5,469 [†] (>80%):	6-7: P	5.4 ^{IS}	4.8, 6.0*
				5,474 (93.2%):	13-14: S	10.9 ^{IS}	10.1, 11.7*
Israel	³⁴	'97	National	10,057 (86.4%):	13-14: S	17.9 ^{IQ}	17.2, 18.6*
Kenya	¹¹⁶	'93	Muranga, Nairobi	1,172 (91.8%)	10 P	5.9 ^E	4.6, 7.2*
	¹⁴	'94-'95	Eldoret, Nairobi	6,236 (99.5%)	13-14 S	13.9 ^{IS}	13.0, 14.8*
	¹¹⁷	'95:	Uasin Gishu district	3,018:	13-14: S	10.2 ^{IQ}	9.1, 11.3*

(P=Parental-report)

S=Self-report)

		'01:		3,258:	13-14:	S	13.8	^{IQ}	12.6, 15.0*
Kuwait	³⁵	'95 & '96	National	3,110	13-14	S	16.1	^{IS}	14.8, 17.4
Lebanon	¹⁴	'94-'95	Beirut	2,993 (100%)	13-14	S	14.4	^{IS}	13.1, 15.7*
Malta	¹⁴	'94-'95	Nationwide	3,493 [†] (>80%):	6-7:	P	8.8	^{IS}	7.9, 9.7*
				3,711 (88.7%):	13-14:	S	16.0	^{IS}	14.9, 17.1*
Morocco	¹⁴	'94-'95	Casablanca, Marrakech, Rabat	8,900 (95.1%)	13-14	S	7.5	^{IS}	7.8, 8.0*
Nigeria	¹⁴	'95	Ibadan	3,057 (76.5%)	13-14	S	10.7	^{IS}	9.6, 11.8*
Oman	¹⁴	'94-'95	Al-Khod	3,891 [†] (>80%):	6-7:	P	7.1	^{IS}	6.3, 7.9*
				2,984 (94.0%):	13-14:	S	8.9	^{IS}	7.9, 9.9*
Palestine	¹¹⁸	'00	Ramallah	1,048 (86%)	6-7:	P	10.2	^{IQ}	8.4, 12.0*
-				2,334 (86%)	8-12:	P	8.4	^{IQ}	7.3, 9.5*
West									
Bank									
	¹¹⁹	'00-'01	Palestine, Ramallah, North	14,650 (84%):	5-8:	P	9.5	^{IQ}	9.0, 10.0*
			Gaza	14,060 (90%):	12-15:	S	7.3	^{IQ}	6.9, 7.7*
South	¹⁰	'90	Cape Town	1,239 (98.9%)	12	P	17.8	^D	15.7, 19.9*

Africa

33	'93	Mitchell's Plain area	1,955 (90.0%)	7-8	P	26.8 ^D	24.8, 28.8
14	'94-'95	Cape Town	4,283 (82.8%)	13-14	S	16.1 ^{IS}	15.1, 17.1*
120	Published '02	Cape Town	4,706 (82.8%)	13-14	S	16.0 ^{IQ}	15.0, 17.0*
Turkey							
121	'92:	Ankara	1,036:	6-13	P	11.9 ^D	10.0, 14.0
122	'94	Edirne	5,412 (85.8%):	7-12	P	5.8 ^D	5.2, 6.4
123	'95	Istanbul	2,216 (94.9%)	6-12	P	8.2 ^{IQ}	7.1, 9.3
124	'96	Ankara	2,784 (88.3%)	7-14	P	4.7 ^A	3.9, 5.5
125	'96-'97	Istanbul	2276 (87.5%)	6-15	P	7.2 ^D	6.1, 8.3*
126	'96	Nationwide	14,492 (93.6%):	0-4:	P	5.1 ^A	4.7, 5.5*
			17,873 (93.6%):	5-10:	P	3.2 ^A	2.9, 3.5*
			14,412 (93.6%):	11-17:	P	1.8 ^A	1.6, 2.0*
127	'99-'00	Ankara	3,041 (88.7%)	8-11	P	11.5 ^{IQ}	10.4, 12.6*
128	'00-'01	Afyon	1,366 (94.5%)	13-18	S	12.2 ^F	10.5, 13.9*

'Prevalence' key, see Table 1.

Table 4. Studies of wheeze prevalence in Asia

Country	Reference	Survey Year	Area	N (Response rate)	Age (years) / ascertainment	Prevalence %	95% CI
China	¹⁴	'94-'95	Beijing, Chongqing, Guangzhou, Shanghai, Wulumuqi	18,704 (98.4%)	13-14 S	4.2 ^{IS}	3.9, 4.5*
	¹²⁹	'95-'96	Beijing:	7,668 (98.9%)	6-7 P	6.0 ^{IS}	5.1, 6.9
			Urumqi:			2.9 ^{IS}	2.3, 3.5
	¹³⁰	'97-'98	Hong Kong:	10,902 (94%)	9-11 P	5.8 ^{IQ}	5.0, 6.7
			Guangzhou:			3.4 ^{IQ}	2.8, 4.1
			Beijing:			3.8 ^{IQ}	3.3, 4.4
	¹³¹	'99	Tou-Cheng City	8,754 (97.2%)	13-15 S	8.2 ^{IQ}	7.6, 8.8*
¹³²	'01	Wuhun	4,185	14-15 S	4.2 ^{IQ}	3.6, 4.8*	
Hong Kong	⁹	'92	Hong Kong	1,062 (89.2%)	11-20 P	3.7 ^D	2.6, 4.8
	¹⁴	'95:	Hong Kong	3,509 (97%):	6-7: P	9.1 ^{IS}	8.1, 10.1*
		'94-'95:	Hong Kong	4,526 (97%):	13-14: S	12.4 ^{IS}	11.4, 13.4*
	¹³³	'95	Hong Kong	2,292 (87%)	8-12 P	10.8 ^H	9.5, 12.1*

(P=Parental-report

S=Self-report)

	134	'97-'98	Hong Kong	3,110 (97%)	10	P	6.6	IQ	5.7, 7.5*
	135	'01	Hong Kong	4,448 (95%)	6-7	P	9.5	IQ	8.5, 10.3*
	136	'02	Hong Kong	3,321 (99%)	13-14	S	8.7	IS	7.8 – 9.7
India	14	'94-'95	11 cities	31,697 [†] (>80%):	6-7:	P	5.6	IS	5.3, 5.9*
				35,461(95.4%):	13-14:	S	6.0	IS	5.8, 6.2*
Indonesia	14	'94-'95	Bandung	1,390 [†] (>80%):	6-7:	P	4.1	IS	3.1, 5.1*
				2,152 (95.7%):	13-14:	S	2.1	IS	1.5, 2.7*
Japan	14	'94-'95	Fukuoka	2,900 [†] (>80%):	6-7:	P	17.3	IS	15.9, 18.7*
				2,667 (94.2%):	13-14:	S	13.4	IS	12.1, 14.7*
	137	'01	Suita City	5,614 (76%)	12-15	P	6.7	IQ	6.0, 7.4*
Korea	36	'95	Seoul and Provincial cities	5,494 (92.5%):	6-7:	P	13.6	IQ	12.7, 14.6
				10,015(97.3%):	13-14:	S	7.9	IQ	7.4, 8.5
	138	'98	Rural areas	2,087:	7-9:	P	11.5	IQ	10.1, 12.9*
				2,137:	10-12:	P	11.2	IQ	9.9, 12.5*
				3,030:	13-15:	P	9.4	IQ	8.4, 10.4*
				(91.2% overall)					
	139	'00	Seoul & Provincial cities	15,894	12-15	S	9.3	IQ	8.9, 9.8
				(96.4%)					

	140	Published '01	Urban and rural areas	2,055	7-16	P	8.2	IQ	7.0, 9.4
	141	Published '01	Industrial factories area & less polluted area	6,886 (91.7%)	7-12	P	10.3	IQ	9.6, 11.0
	142	Published '02	Urban and rural areas (86.6%)	16,624	7-18	P<10/S	9.3	IQ	8.9, 9.7*
	143	Published '02	Rural areas	1,727 (86.1%)	16-18	S	13.0	IQ	11.4, 14.6*
Malaysia	14	'94-'95	Alor Setar, Ipoh, Klang Valley, Kota Bharu, Muar	15,285 [†] (>80%): 17,313(92.9%):	6-7: 13-14:	P S	6.1 9.6	IS IS	5.7, 6.5* 9.2, 10.0*
	144	'01:	Kota Bharu	3,157: 3,004:	6-7: 13-14:	P S	4.3 5.7	IQ IQ	3.6, 5.0* 4.9, 6.5*
Pakistan	14	'94-'95	Karachi	1,829 (100%)	13-14	S	8.5	IS	7.2, 9.8*
Philippines	14	'94-'95	Metro Manilla	3,558 [†] (>80%): 3,063 (95.5%):	6-7: 13-14:	P S	11.3 12.3	IS IS	10.3, 12.3* 11.2, 13.4*
Singapore	14	'94	Nationwide	2,118 (90%): 3,785 (90%):	6-7: 13-14:	P S	15.7 9.7	IS IS	14.2, 17.2* 8.8, 10.6*
	145	'01	Nationwide	5,305: 4,058:	6-7: 12-15:	P S	10.2 11.9	IQ IQ	9.4, 11.0* 10.9, 12.9*
Taiwan	14	'94-'95	Taipei	4,806 [†] (>80%):	6-7:	P	9.6	IS	8.8, 10.4*

				10,636(93.3%):	13-14:	S	5.2	IS	4.8, 5.6*
	¹⁴⁶	'02	Taoyuan	3,079:	6-8:	P	7.5	IQ	6.6, 8.5
				3,111:	13-15:	S	4.2	IQ	3.5, 5.3
Thailand	¹⁴	'95	Bankok	3,629 [†] (>80%):	6-7:	P	11.0	IS	10.0, 12.0*
				2,777 (74.8%):	13-14:	S	13.5	IS	12.2, 14.8*
	¹⁴	'95	Chiang Mai	3,522 (92%):	6-7:	P	5.5	IS	4.7, 6.3*
				3,731 (95%):	13-14:	S	12.6	IS	11.5, 13.7*

'Prevalence' key, see Table 1.

Table 5. Studies of wheeze prevalence in Australasia

Country	Reference	Survey Year	Area	N (Response rate)	Age (years) / ascertainment	Prevalence %	95% CI
Australia	11	'90:	Melbourne	3,325 (89%):	7: P	23.1 ^A	21.7, 24.5
				2,899 (89%):	12: P	20.9 ^A	19.4, 22.4
				2,968 (89%):	15: P	18.6 ^A	17.0, 19.8
	147	'90	Sidney, Melbourne, Brisbane, Hunter Valley	8,753 (84%)	8.7(Mean), 2.1(SD) P	19.5 ^D	18.7, 20.3*
	148	'90-'91	Campbelltown	4,550 (82%)	12-18 S	18.1 ^D	17.0, 19.2*
	8	'91	Adelaide & Sydney	2,947 (87%)	12-15 S	29.7 ^{IQ}	28.1, 31.1*
	149	'92	Burra, Gladstone, Kingston	1,032 (80.0%)	5-18 P	25.0 ^D	22.4, 27.6*
	150	'93	Sydney, Brisbane, Melborne, & Hunter Valley NSW	10,106 (84.4%)	5-12 P	21.2 ^D	20.4, 22.0*
	151	'91-'93	State of NSW	6,388 (78.4%)	8-11 P	24.1 ^D	23.1, 25.1*
	152	'93	South Australian state	14,124 (72.8%)	4 P	25.2 ^D	24.8, 25.6

(P=Parental-report

S=Self-report)

	14	'93 &'94	Melbourne, Sydney, Adelaide, Perth	9,155 (84%): 11,541 (94%):	6-7: 13-14:	P S	24.6 ^{IS} 29.4 ^{IS}	23.7, 25.5* 28.6, 30.2*
	153	'95	Tazmania	6,378 (92%)	7	P	22.0 ^{IQ}	21.0, 23.0*
	154	'97	Moree, Wagga Wagga	1,499 (67.3%)	7-12	P	27.7 ^D	25.4, 30.0*
	155	Published '99	Melbourne, Government schools	9,794 (75%)	13-19 mainly 14-15	P	18.9 ^A	18.0, 19.9
	156	'99	Torres Strait & Northern Peninsula area	1,650 (98%)	0-17	P	12.4 ^{IQ}	10.8, 14.0
	157	'99-'01	Aboriginal & Torres Strait Islander and Australian Capital Territory	10,452 (80%)	4-6	P	15.5 ^{IQ}	14.8, 16.2*
Fiji	38	'90 Sept	Suva city	2,117 (97.4%)	9-10	P	21.0 ^{IQ}	19.2, 22.8
New	158	'85:	Auckland:	1,084:	7-10:	P	14.8 ^D	12.7, 16.9
Zealand		'91:	Auckland:	1,901 (80.4%):	7-10:	P	18.7 ^D	16.9, 20.5
	8	'91	Wellington	1,863 (87%)	12-15	S	28.3 ^{IS}	26.3, 30.3*
	14	'92-'93	Auckland, Bay of Plenty, Christchurch, Hawke's Bay, Nelson, Wellington	16,898 (91%): 19,023 (93%):	6-7: 13-14:	P S	24.5 ^{IS} 30.2 ^{IS}	23.9, 25.1* 29.5, 30.9*

159	'00	City of Hastings & Borough of Havelock North	1,287 (84%)	10-12	P	22.0 ^{IQ}	19.7, 24.3*	41
160	'02	Wellington	2,539 (47%)	6-7	P	24.3 ^{IQ}	22.6, 26.0*	

'Prevalence' key, see Table 1.

Table 6. Multiple logistic regression analysis using all studies in the UK and Australia of country, year, area, age and type of study on the prevalence of wheeze

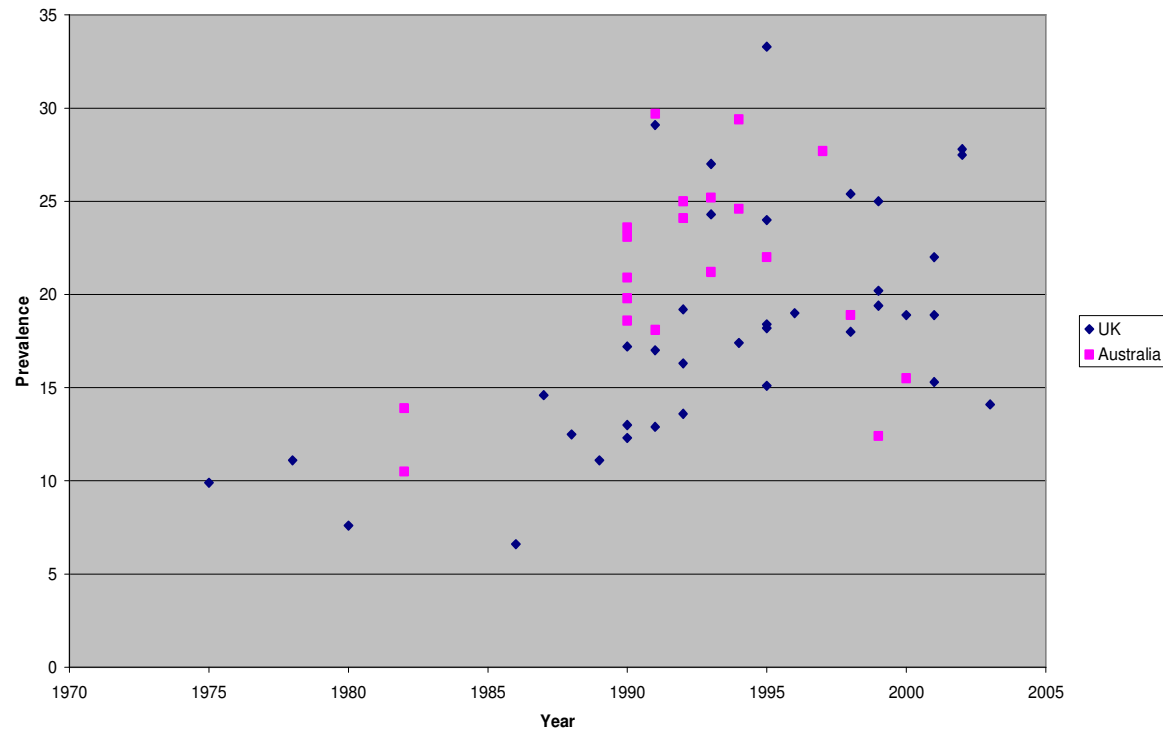
Variable	All studies	ISAAC studies only	Non-ISAAC studies only	ISAAC question only
	Odds Ratio (+95% CI)(+p-value)	Odds Ratio (+95% CI)(+p-value)	Odds Ratio (+95% CI)(+p-value)	Odds Ratio (+95% CI)(+p-value)
Country: UK (reference) vs Australia	1.343 [1.318, 1.369] p<0.0005	0.987 [0.958, 1.016] p=0.373	1.470 [1.438, 1.502] p<0.0005	0.986 [0.957, 1.017] p=0.0005
Year of survey	1.025 [1.023, 1.026] p<0.0005	0.975 [0.971, 0.979] p<0.0005	1.026 [1.024, 1.028] p<0.0005	0.977 [0.973, 0.981] p<0.0005
Age of children surveyed	0.993 [0.989, 0.997] p<0.0005	1.007 [0.998, 1.016] p=0.124	0.989 [0.985, 0.993] p<0.0005	1.006 [0.995, 1.017] p=0.0005
Parental- (reference) vs Self-response	1.290 [1.246, 1.335] p<0.0005	§ - -	1.242 [1.197, 1.287] p<0.0005	1.739 [1.624, 1.862] p<0.0005
ISAAC study: No (reference) vs Yes	1.638 [1.598, 1.678] p<0.0005	- -	- -	- -

§ OR not given as age of child (7-8 & 13-14 yrs) in ISAAC is parent-report & self-report respectively

Table 7. Multiple logistic regression analysis using all studies in UK and Australia, as for Table 6, with interaction term (country x year)

Variable		Odds Ratio (+95% CI)	p-value
Country	[UK (reference) versus Australia]	1.406 [1.379, 1.433]	p<0.0005
Year of survey		1.032 [1.030, 1.034]	p<0.0005
Age of children surveyed		0.993 [0.989, 0.997]	p=0.016
Parental-response (reference) vs Self-complete		1.239 [1.197, 1.283]	p<0.0005
ISAAC study	[No (reference) versus Yes]	1.649 [1.609, 1.690]	p<0.0005
Country by Year interaction		0.938 [0.933, 0.943]	p<0.0005

Figure 1. Prevalence of “wheezing in the past year” by calendar year of survey in all children (aged 0-16), reported in published studies in Australia and UK



References

1. Lee DA, Winslow NR, Speight ANP, Hey EN. Prevalence and spectrum of asthma in childhood. *British Medical Journal* 1983;286:1256-8.
2. Gergen PJ, Mullally DI, Evans R. National survey of prevalence of asthma among children in the United States, 1976 to 1980. *Pediatrics* 1988;81(1):1-7.
3. Burney PGJ, Laitinen LA, Perdrizet S, et al. Validity and repeatability of the IUALTD (1984) bronchial symptoms questionnaire: An international comparison. *European Respiratory Journal* 1989;2:940-5.
4. Shaw RA, Crane J, Pearce NE, Bremner P, Burgess C, Woodman K, et al. Validation of a video questionnaire for assessing asthma prevalence. *Clinical and Experimental Allergy* 1992;22:562-9.
5. Shaw RA, Woodman K, Ayson M, et al. Measuring the prevalence of bronchial hyper-responsiveness in children. *International Journal of Epidemiology* 1995;24:597-602.
6. Baumann A, Hunt J, Young L, Larkin P, Peat JK. Asthma under-recognition and under-treatment in Australian community. *Australian and New Zealand Journal of Medicine* 1992;22:36-40.
7. Dales RE, Raizenne M, El-Saadanny S, Brook J, Burnett R. Prevalence of childhood asthma across Canada. *International Journal of Epidemiology* 1994;23:775-81.
8. Pearce NE, Weiland S, Keil U, et al. Self-reported prevalence of asthma symptoms in children in Australia, England, Germany and New Zealand: An international comparison using the ISAAC written and video questionnaires. *European Respiratory Journal* 1993;6:1455-61.
9. Leung R, Ho P. Asthma, allergy and atopy in three Southeast Asian populations. *Thorax* 1994;49:1205-10.
10. Burr ML, Limb ES, Andrae S, Barry DMJ, Nagel F. Childhood asthma in four countries: a comparative survey. *International Journal of Epidemiology* 1994;23:341-7.
11. Robertson CF, Bishop J, Sennhauser FH, Mallol J. International comparison of asthma prevalence in children: Australia, Switzerland, Chile. *Pediatric Pulmonology* 1993;16:219-26.
12. Sunyer J, Anto JM, Tobias A, et al. Generational increase of self-reported first attack of asthma in fifteen industrialised countries. *European Respiratory Journal* 1999;14:885-91.
13. Asher MI, Keil U, Anderson HR, et al. International Study of Asthma and Allergies in Childhood (ISAAC): Rationale and methods. *European Respiratory Journal* 1995;8:483-91.
14. ISAAC Steering Committee. Worldwide variations in the prevalence of asthma symptoms: the International Study of Asthma and Allergies in Childhood (ISAAC). *European Respiratory Journal* 1998;12:315-35.
15. Asher MI, Montefort S, Bjorksten B, Lai CKW, Strachan DP, Weiland SK, et al. Worldwide time trends in the prevalence of symptoms of asthma, allergic rhinoconjunctivitis, and eczema in childhood: ISAAC Phases One and Three repeat multicountry cross-sectional surveys. *Lancet* 2006;368:733-43.
16. ISAAC Co-ordinating Committee. *Manual for the International Study of Asthma and Allergies in Childhood (ISAAC)*. 2nd Edn ed. Auckland, New Zealand, Munster, Germany, 1993.
17. Kuehni CE, Brooke AM, Silverman M. Prevalence of wheeze during childhood: retrospective and prospective assessment. *European Respiratory Journal* 2000;16:81-5.
18. Pezzichini MM, Rennie D, Senthilselvan A, et al. Limited agreement between written and video asthma symptom questionnaires. *Pediatric Pulmonology* 2000;30:307-12.
19. McCullagh P, Nelder JA. *Generalized linear models*. 2nd ed. London: Chapman and Hall, 1989.

20. Yemaneberhan H, Bekele Z, Venn A, Lewis S, Parry E, Britton J. Prevalence of wheeze and asthma and relation to atopy in urban and rural Ethiopia. *Lancet* 1997;350:85-90.
21. Eldeirawi K, Persky VW. History of ear infections and prevalence of asthma in a national sample of children aged 2 to 11 years: the Third National Health and Nutrition Examination Survey, 1988 to 1994. *Chest* 2004;125(5):1685-92.
22. Sotir M, Yeatts K, Shy C. Presence of asthma risk factors and environmental exposures related to upper respiratory infection-triggered wheezing in middle school-age children. *Environmental Health Perspectives* 2003;111:657-62.
23. Ernst P, Demissie K, Joseph L, Locher U, Becklake MR. Socioeconomic status and indicators of asthma in children. *American Journal of Respiratory and Critical Care Medicine* 1995;152:570-5.
24. Osterman JW, Armstrong BG, Ledoux E, Sloan M, Ernst P. Comparison of French and English versions of the American Thoracic Society respiratory questionnaire in a bilingual working population. *International Journal of Epidemiology* 1991;20:138-43.
25. Villarreal AB, Aguirre LHS, Rojo MMT, Navarro ML, Romieu I. Risk factors for asthma in school children from Ciudad Juarez, Chihuahua. *Journal of Asthma* 2003;40(4):413-23.
26. Sole D, Camelo-Nunes IC, Wandalsen GF, Melo KC, Naspitz CK. Is rhinitis alone or associated with atopic eczema a risk factor for severe asthma in children? *Pediatric Allergy and Immunology* 2005;16(2):121-125.
27. Jeffs D, Grainger R, Powell P. Is childhood allergy more common amongst an island population? *Journal of the Royal Society for the Promotion of Health* 2000;120:236-41.
28. Leonardi GS, Houthuijs D, B. N, J. V, Rudnai P, Zejda J, et al. Respiratory symptoms, bronchitis and asthma in children of Central and Eastern Europe. *European Respiratory Journal* 2002;20:890-8.
29. Harty SB, Sheridan A, Howell F, Nicholson A. Wheeze, eczema and rhinitis in 6-7 year old Irish schoolchildren. *Irish Medical Journal* 2003;96:102-4.
30. Nystad W, Magnus P, Roksund O, Svidal B, Hetlevik O. The prevalence of respiratory symptoms and asthma among schoolchildren in three different areas of Norway. *Pediatric Allergy and Immunology* 1997;8:35-40.
31. Lewis S, Richards D, Bynner J, Butler N, Britton J. Prospective study of risk factors for early and persistent wheezing in childhood. *European Respiratory Journal* 1995;8:349-56.
32. Ng-Man-Kwong G, Das C, Proctor A, Whyte MKB, Primhak RA. Diagnostic and treatment behaviour in children with chronic respiratory symptoms: relationship with socioeconomic factors. *Thorax* 2002;57:701-4.
33. Ehrlich RI, Du-Toit D, Jordaan E, Volmink JA, Weinberg EG, Zwarenstein M. Prevalence and reliability of asthma symptoms in primary schoolchildren in Cape Town. *International Journal of Epidemiology* 1995;24:1138-45.
34. Shohat T, Golan G, Tamir R, Green MS, Livne I, Davidson Y, et al. Prevalence of asthma in 13-14 yr-old schoolchildren across Israel. *European Respiratory Journal* 2000;15:725-9.
35. Behbehani NA, Abal A, Syabbalo NC, Abd-Azeem A, Shareef E, Al-Momen J. Prevalence of asthma, allergic rhinitis, and eczema in 13-14-year-old children in Kuwait: an ISAAC study. *Annals of Allergy, Asthma and Immunology* 2000;85:58-63.
36. Lee SI, Shin MH, Lee HB, Lee JS, Son BK, Koh YY, et al. Prevalences of symptoms of asthma and other allergic diseases in Korean children: a nationwide questionnaire survey. *Journal of Korean Medical Science* 2001;16:155-64.
37. Wang XS, Tan TN, Shek LPC, Chng SY, Hia CPP, Ong NBH, et al. The prevalence of asthma and allergies in Singapore; data from two ISAAC surveys seven years apart. *Archives of Disease in Childhood* 2004;89:423-6.

38. Flynn MGL. Respiratory symptoms, bronchial responsiveness, and atopy in Fijian and Indian children. *American Journal of Respiratory and Critical Care Medicine* 1994;150:415-20.
39. Kaur B, Anderson HR, Austin J, Burr M, Harkins LS, Strachan DP, et al. Prevalence of asthma symptoms, diagnosis and treatment in 12-14yr old children across Great Britain (ISAAC UK). *British Medical Journal* 1998;316:118-24.
40. Pararajasingam CD, Sittampalam L, Damani P, Pattemore PK, Holgate ST. Comparison of the prevalence of asthma among Asian and European children in Southampton. *Thorax* 1992;47:529-32.
41. Peat JK, Van Den Berg RH, Green WF, et al. Changing prevalence of asthma in Australian children. *British Medical Journal* 1994;308:1591-1596.
42. Kemp T, Pearce N, Crane J, Beasley R. Problems of measuring asthma prevalence. *Respirology* 1996;3:183-8.
43. Strachan DP. Family size, infection and atopy: the first decade of the "hygiene hypothesis". *Thorax* 2000;55(S1):2-10.
44. Renzoni E, Forastiere F, Biggeri A, Viegi G, Bisanti L, Chellini E, et al. Differences in parental- and self-report of asthma, rhinitis and eczema among Italian adolescents. *European Respiratory Journal* 1999;14:597-604.
45. Sherman CB, Tosteson TD, Tager IB, et al. Early childhood predictors of asthma. *American Journal of Epidemiology* 1990;132:83-95.
46. Peat JK, Tovey E, Tpoelle BG, et al. House dust mite allergens: A major risk factor for childhood asthma in Australia. *American Journal of Respiratory and Critical Care Medicine* 1996;153:141-6.
47. Strachan DP, Cook DG. Parental smoking and childhood asthma: longitudinal and case-control studies. *Thorax* 1998;53:204-12.
48. Dold S, Wjst M, von Mutius E, et al. Genetic risk for asthma, allergic rhinitis and atopic dermatitis. *Archives of Disease in Childhood* 1992;67:1018-22.
49. Hodge L, Salome C, Peat J, Haby M, Xuan W, Woolcock A. Consumption of oily fish and childhood asthma risk. *Medical Journal of Australia* 1996;164:137-40.
50. Brooks AM, Byrd RS, Weitzman M, Auinger P, McBride JT. Impact of low birth weight on early childhood asthma in the United States. *Archives of Pediatrics and Adolescent Medicine* 2001;155:401-6.
51. Anderson HR, Gupta R. *Trends in asthma*. London: Aesculapius, 2002.
52. Ng-Man-Kwong G, Proctor A, Billings C, Duggan R, Das C, Whyte MK, et al. Increasing prevalence of asthma diagnosis and symptoms in children is confined to mild symptoms. *Thorax* 2001;56:312-4.
53. Anderson HR, Bland JM, Patel S, Peckham C. The natural history of asthma in childhood. *Journal of Epidemiology and Community Health* 1986;40:121-9.
54. Werneck G, Ruiz S, Hart R, White M, Romieu I. Prevalence of asthma and other childhood allergies in Brazilian schoolchildren. *Journal of Asthma* 1999;36:677-90.
55. Benicio MHD, Ferreira MU, Cardoso MRA, Konno SC, Monteiro CA. Wheezing conditions in early childhood: prevalence and risk factors in the city of Sao Paulo, Brazil. *Bulletin of The World Health Organization* 2004;82(7):516-22.
56. Monteil MA, Joseph G, Changkit C, Wheeler G, Antoine RM. Comparison of prevalence and severity of asthma among adolescents in the Caribbean islands of Trinidad and Tobago: results of a nationwide cross-sectional survey. *BMC Public Health* 2005;5:96.
57. Soto-Quiros ME, Soto-Martinez M, Hanson LA. Epidemiological studies of the very high prevalence of asthma and related symptoms among school children in Costa Rica from 1989 to 1998. *Pediatric Allergy and Immunology* 2002;13:342-9.
58. Cooper PJ, Chico ME, Bland M, Griffin GE, Nutman TB. Allergic symptoms, atopy and Geohelminth infections in a rural area of equador. *American Journal of Respiratory and Critical Care Medicine* 2003;168:313-7.

59. Fagan JK, Scheff PA, Hryhorczuk D, Ramakrishnan V, Ross M, Persky V. Prevalence of asthma and other allergic diseases in an adolescent population: association with gender and race. *Annals of Allergy, Asthma and Immunology* 2001;86:177-84.
60. Maier WC, Arrighi HM, Morray B, Llewellyn C, Redding GJ. The impact of asthma and asthma-like illness in Seattle school children. *Journal of Clinical Epidemiology* 1998;51:557-68.
61. Findley S, Lawler K, Bindra M, Maggio L, Penachio MM, Maylahn C. Elevated asthma and indoor environmental exposures among Puerto Rican children of East Harlem. *Journal of Asthma* 2003;40(5):557-69.
62. Yeatts K, Shy C, Wiley J, Music S. Statewide adolescent asthma surveillance. *Journal of Asthma* 2000;37:425-34.
63. Mvula M, Larzelere M, Kraus M, Moisiwicz K, Morgan C, Pierce S, et al. Prevalence of asthma and asthma-like symptoms in inner-city schoolchildren. *Journal of Asthma* 2005;42(1):9-16.
64. Arif AA, Borders TF, Patterson PJ, Rohrer JE, Xu KT. Prevalence and correlates of paediatric asthma and wheezing in a largely rural USA population. *Journal of Paediatrics and Child Health* 2004;40(4):189-94.
65. Chrischilles E, Ahrens R, Kuehl A, Kelly K, Thorne P, Burmeister L, et al. Asthma prevalence and morbidity among rural Iowa schoolchildren. *Journal of Allergy and Clinical Immunology* 2004;113(1):66-71.
66. ISAAC Steering Committee. Worldwide variation in prevalence symptoms of asthma, allergic rhinoconjunctivitis and atopic eczema: ISAAC. *Lancet* 1998;351:1225-32.
67. Riedler J, Eder W, Oberfeld G, Schreuer M. Austrian children living on a farm have less hay fever, asthma and allergic sensitization. *Clinical and Experimental Allergy* 2000;30:194-200.
68. Kalayci O, Saraclar Y, Sekerel BE, Adalioglu G, Kuyucu S, Egor G, et al. Prevalence of asthma symptoms among Turkish Cypriot schoolchildren. *Turkish Journal of Pediatrics* 1999;41:413-20.
69. Riikjarv MA, Julge K, Vasar M, Braback L, Knutsson A, Bjorksten B. The prevalence of atopic sensitization and respiratory symptoms among Estonian schoolchildren. *Clinical and Experimental Allergy* 1995;25:1198-204.
70. Annus T, Riikjarv M-A, Rahu K, Bjorksten B. Modest increase in seasonal allergic rhinitis and eczema over 8 years among Estonian schoolchildren. *Pediatric Allergy and Immunology* 2005;16(4):315-20.
71. Timonen KL, Pekkanen J, Korppi M, Vahteristo M, Salonen RO. Prevalence and characteristics of children with chronic respiratory symptoms in eastern Finland. *European Respiratory Journal* 1995;8:1155-60.
72. Penard-Morand C, Charpin D, Raheison C, Kopferschmitt C, Caillaud D, Lavaud F, et al. Long-term exposure to background air pollution related to respiratory and allergic health in schoolchildren. *Clinical and Experimental Allergy* 2005;35(10):1279-87.
73. Weiland SK, von Mutius E, Hirsch T, Duhme H, Fritzsche C, Werner B, et al. Prevalence of respiratory and atopic disorders among children in the East and West of Germany five years after unification. *European Respiratory Journal* 1999;14:862-70.
74. Wjst M, Hoelscher B, Frye C, Wichmann HE, Dold S, Heinrich J. Early antibiotic treatment and later asthma. *European Journal of Medical Research* 2001;6:263-71.
75. Maziak W, Behrens T, Brasky TM, Duhme H, Rzehak P, Weiland SK, et al. Are asthma and allergies in children and adolescents increasing? Results from ISAAC phase I and phase III surveys in Munster, Germany. *Allergy* 2003;58:572-9.
76. Taylor MR, Holland CV, O'Lorcain P. Asthma and wheeze in schoolchildren. *Irish Medical Journal* 1996;89(1):34-5.
77. Simoni M, Lombardi E, Berti G, Rusconi F, La Grutta S, Piffer S, et al. Mould/dampness exposure at home is associated with respiratory disorders in Italian children and adolescents: the SIDRIA-2 Study. *Occupational and Environmental Medicine* 2005;62(9):616-22.

78. Montefort S, Lenicker HM, Caruna S, Agius-Muscat H. Asthma, rhinitis and eczema in Maltese 13-15 year-old schoolchildren - prevalence, severity and associated factors [ISAAC]. *Clinical and Experimental Allergy* 1998;28:1089-99.
79. Montefort S, Muscat HA, Caruna S, Lenicker H. Allergic conditions in 5-8 year-old Maltese schoolchildren: Prevalence, severity and associated risk factors [ISAAC]. *Pediatric Allergy and Immunology* 2002;13:98-104.
80. Henriksen AH, Holmen TL, Bjermer L. Gender differences in asthma prevalence may depend on how asthma is defined. *Respiratory Medicine* 2003;97:491-7.
81. Zlotkowska R, Zejda JE. Fetal and postnatal exposure to tobacco smoke and respiratory health in children. *European Journal of Epidemiology* 2005;20:719-27.
82. Spengler JD, Jaakkola JJK, Parise H, Katsnelson BA, Privalova LI, Kosheleva AA. Housing characteristics and children's respiratory health in the Russian Federation. *American Journal of Public Health* 2004;94(4):657-62.
83. Garcia-Marcos L, Quiros AB, Hernandez GG, Guillen-Grima F, Diaz CG, Urena IC, et al. Stabilization of asthma prevalence among adolescents and increase among schoolchildren (ISAAC phases I and III) in Spain. *Allergy* 2004;59(12):1301-7.
84. Garcia-Marcos L, Castro-Rodriguez JA, Suarez-Varela MM, Garrido JB, Hernandez GG, Gimeno AM, et al. A different pattern of risk factors for atopic and non-atopic wheezing in 9-12-year-old children. *Pediatric Allergy and Immunology* 2005;16(6):471-7.
85. Hesselmar B, Aberg B, Eriksson B, Aberg N. Asthma in children: prevalence, treatment and sensitization. *Pediatric Allergy and Immunology* 2000;11:74-9.
86. Ronmark E, Perzanowski M, Platts-Mills T, Lundback B. Incidence rates and risk factors for asthma among school children: A 2-year follow-up report from the Obstructive Lung Disease in Northern Sweden (OLIN) studies. *Respiratory Medicine* 2002;96:1006-13.
87. Braback L, Kjellman NI, Sandin A, Bjorksten B. Atopy among schoolchildren in northern and southern Sweden in relation to pet ownership and early life events. *Pediatric Allergy and Immunology* 2001;12:4-10.
88. Bornehag CG, Sundell J, Hagerhed L, Janson S. Pet-keeping in early childhood and airway, nose and skin symptoms later in life. *Allergy* 2003;58:939-44.
89. Hedman L, Lindgren B, Perzanowski M, Ronmark E. Agreement between parental and self-completed questionnaires about asthma in teenagers. *Pediatric Allergy and Immunology* 2005;16(2):176-81.
90. Sennhauser FH, Kuhni CE. Prevalence of respiratory symptoms in Swiss children: is bronchial asthma really more prevalent in boys? *Pediatric Pulmonology* 1995;19:161-6.
91. Braun-Fahrlander C, Gassner M, Grize L, Takken-Sahli K, Neu U, Stricker T, et al. No further increase in asthma, hay fever and atopic sensitisation in adolescents living in Switzerland. *European Respiratory Journal* 2004;23(3):407-13.
92. Bayer-Oglesby L, Grize L, Gassner M, Takken-Sahli K, Sennhauser FH, Neu U, et al. Decline of ambient air pollution levels and improved respiratory health in Swiss children. *Environmental Health Perspectives* 2005;113(11):1632-7.
93. Brunekreef B, Groot B, Hoek G. Pets, allergy and respiratory symptoms in children. *International Journal of Epidemiology* 1992;21:338-42.
94. Spee-van-der-Wekke J, Meulmeester JF, Radder JJ, Verloove-Vanhorick SP. School absence and treatment in schoolchildren with respiratory symptoms in The Netherlands: data from the Child Health Monitoring System. *Journal of Epidemiology and Community Health* 1998;52:359-63.
95. de-Kok ME, Mertens PL, Cuijpers CE, Swaen GM, Wesseling GJ, Broer J, et al. The rate of respiratory symptoms among primary schoolchildren in two Dutch regions. *European Journal of Pediatrics* 1996;155(6):506-11.

96. Mommers M, Gielkens-Sijstermans C, Swaen GMH, van Schayck CP. Trends in the prevalence of respiratory symptoms and treatment in Dutch children over a 12 year period: results of the fourth consecutive survey. *Thorax* 2005;60(2):97-9.
97. Withers NJ, Low L, Holgate ST, Clough JB. The natural history of respiratory symptoms in a cohort of adolescents. *American Journal of Respiratory and Critical Care Medicine* 1998;158:352-7.
98. Symington P, Coggon D, Holgate S. Respiratory symptoms in children at schools near a foundry. *British Journal of Industrial Medicine* 1991;48:588-91.
99. Luyt DK, Burton PR, Simpson H. Epidemiological study of wheeze, doctor diagnosed asthma and cough in preschool children in Leicestershire. *British Medical Journal* 1993;306:1386-90.
100. Anderson HR, Butland BK, Strachan DP. Trends in prevalence and severity of childhood asthma. *British Medical Journal* 1994;308:1600-4.
101. Strachan DP, Anderson HR, Limb ES, O'Neill A, Wells N. A national survey of asthma prevalence, severity and treatment in Great Britain. *Archives of Disease in Childhood* 1994;70:174-8.
102. Waldron G, Pottle B, Dod J. Asthma and the motorways - one District's experience. *Journal of Public Health Medicine* 1995;17:85-9.
103. Venn A, Lewis S, Cooper M, Hill J, Britton J. Increasing prevalence of wheeze and asthma in Nottingham primary schoolchildren 1988-1995. *European Respiratory Journal* 1998;11:1324-8.
104. Austin JB, Russell G. Wheeze, cough, atopy and indoor environment in the Scottish Highlands. *Archives of Disease in Childhood* 1997;76:22-6.
105. Venn A, Lewis S, Cooper M, Hill J, Britton J. Questionnaire study of effect of sex and age on the prevalence of wheeze and asthma in adolescence. *British Medical Journal* 1998;316:1945-6.
106. Kuehni CE, Davis A, Brooke AM, Silverman M. Are all wheezing disorders in very young (preschool) children increasing in prevalence? *Lancet* 2001;357:1821-5.
107. Shamssain MH, Shamsian N. Prevalence and severity of asthma, rhinitis and atopic eczema: the north east study. *Archives of Disease in Childhood* 1999;81:313-7.
108. Kurukulaaratchy RJ, Fenn M, Twiselton R, Matthews S, Arshad SH. The prevalence of asthma and wheezing illnesses amongst 10-year-old schoolchildren. *Respiratory Medicine* 2002;96:163-9.
109. Frank PI, Wicks PD, Hazell ML, Linehan MF, Hirsch S, Hannaford PC, et al. Temporal change in the prevalence of respiratory symptoms and obstructive airways disease 1993-2001. *British Journal of General Practice* 2005;55(517):596-602.
110. Martindale S, McNeill G, Devereux G, Campbell D, Russell G, Seaton A. Antioxidant intake in pregnancy in relation to wheeze and eczema in the first two years of life. *American Journal of Respiratory and Critical Care Medicine* 2005;171(2):121-8.
111. McCann D, McWhirter J, Coleman H, Devall I, M. C, Weare K, et al. The prevalence and management of asthma in primary-aged schoolchildren in the south of England. *Health Education Research* 2002;17:181-94.
112. Anderson HR, Ruggles R, Strachan DP, Austin JB, Burr M, Jeffs D, et al. Trends in prevalence of symptoms of asthma, hay fever, and eczema in 12-14 year olds in the British Isles, 1995-2002: questionnaire survey. *British Medical Journal* 2004;328:1052-3.
113. Lewis SA, Antoniak M, Venn AJ, Davies L, Goodwin A, Salfeld N, et al. Secondhand smoke, dietary fruit intake, road traffic exposures, and the prevalence of asthma: a cross-sectional study in young children. *American Journal of Epidemiology* 2005;161(5):406-11.
114. Hailu S, Tessema T, Silverman M. Prevalence of symptoms of asthma and allergies in schoolchildren in Gondar town and its vicinity, Northwest Ethiopia. *Pediatric Pulmonology* 2003;35:427-32.

115. Dagoye D, Bekele Z, Woldemichael K, Nida H, Yimam M, Hahh A, et al. Wheezing, allergy, and parasite infection in children in urban and rural Ethiopia. *American Journal of Respiratory and Critical Care Medicine* 2003;167:1369-73.
116. Odhiambo JA, Ng'ang'a LW, Mungai MW, Gicheha CM, Nyamwaya JK, Karimi F, et al. Urban-rural differences in questionnaire-derived markers of asthma in Kenyan school children. *European Respiratory Journal* 1998;12:1105-12.
117. Esamai F, Ayaya S, Nyandiko W. Prevalence of asthma, allergic rhinitis and dermatitis in primary school children in Uasin Gishu district, Kenya. *East African Medical Journal* 2002;79:514-8.
118. El-Sharif N, Abdeen Z, Qasrawi R, Moens G, Nemery B. Asthma prevalence in children living in villages, cities and refugee camps in Palestine. *European Respiratory Journal* 2002;19:1026-34.
119. El-Sharif NA, Nemery B, Barghuthy F, Mortaja S, Qasrawi R, Abdeen Z. Geographical variations of asthma and asthma symptoms among schoolchildren aged 5 to 8 years and 12-15 years in Palestine: the ISAAC. *Annals of Allergy, Asthma and Immunology* 2003;90:63-71.
120. Poyser MA, Nelson H, Ehrlich RI, Bateman ED, Parnell S, Puternam A, et al. Socioeconomic deprivation and asthma prevalence and severity in young adolescents. *European Respiratory Journal* 2002;19:892-8.
121. Kalyoncu AF, Selcuk ZT, Karakoca Y, Emri AS, Coplu L, Sahin AA, et al. Prevalence of childhood asthma and allergic diseases in Ankara, Turkey. *Allergy* 1994;49:485-8.
122. Selcuk ZT, Caglar T, Enunlu T, Topal T. The prevalence of allergic diseases in primary school children in Edirne, Turkey. *Clinical and Experimental Allergy* 1997;27:262-9.
123. Ones U, Sapan N, Somer A, Disci R, Salman N, Guler N, et al. Prevalence of childhood asthma in Istanbul, Turkey. *Allergy* 1997;52:570-5.
124. Saraclar Y, Sekerel BE, Kalayci O, Cetinkaya F, Adalioglu G, Tuncer A, et al. Prevalence of asthma symptoms in school children in Ankara, Turkey. *Respiratory Medicine* 1998;92:203-7.
125. Akcakaya N, Kulak K, Hassanzadeh A, Camcioglu Y, Cokugras H. Prevalence of bronchial asthma and allergic rhinitis in Istanbul school children. *European Journal of Epidemiology* 2000;16:693-9.
126. Turktas I, Selcuk ZT, Kalyoncu AF. Prevalence of asthma-associated symptoms in Turkish children. *Turkish Journal of Pediatrics* 2001;43:1-11.
127. Saraclar Y, Kuyucu S, Tuncer A, Sekerel B, Sackesen C, Kocabas C. Prevalence of asthmatic phenotypes and bronchial hyperresponsiveness in Turkish schoolchildren: an ISAAC phase 2 study. *Annals of Allergy, Asthma and Immunology* 2003;91:477-84.
128. Unlu M, Orman A, N. D. The prevalence of asthma among secondary school students in Afyon, Turkey. *Asian Pacific Journal of Allergy Immunology* 2002;20:1-6.
129. Zhao TB, Wang A, Chen Y, Xiao M, Duo L, Lui G, et al. Prevalence of childhood asthma, allergic rhinitis and eczema in Urumqi and Beijing. *Journal of Paediatrics and Child Health* 2000;36:128-33.
130. Wong GW, Hui DS, Chan HH, Fok TF, Leung R, Zhong NS, et al. Prevalence of respiratory and atopic disorders in Chinese schoolchildren. *Clinical and Experimental Allergy* 2001;31:1225-31.
131. Chen CF, Wu KG, Hsu MC, Tang RB. Prevalence and relationship between allergic diseases and infectious diseases. *Journal of Microbiology, Immunology and Infection* 2001;34:57-62.
132. Salo PM, Xia J, Johnson CA, Li Y, Avol EL, Gong J, et al. Indoor allergens, asthma, and asthma-related symptoms among adolescents in Wuhan, China. *Annals of Epidemiology* 2004;14(8):543-50.
133. Yu ITS, Wong TW, Li W. Using child reported respiratory symptoms to diagnose asthma in the community. *Archives of Disease in Childhood* 2004;89(6):544-8.

134. Wong GW, Hui DS, Tam CM, Chan HH, Fok TF, Chan-Yeung M, et al. Asthma, atopy and tuberculin responses in Chinese schoolchildren in Hong Kong. *Thorax* 2001;56:770-3.
135. Lee S-L, Wong W, Lau Y-L. Increasing prevalence of allergic rhinitis but not asthma among children in Hong Kong from 1995 to 2001 (Phase 3 International Study of Asthma and Allergies in Childhood). *Pediatric Allergy and Immunology* 2004;15(1):72-8.
136. Wong GWK, Leung TF, Ko FWS, Lee KKM, Lam P, Hui DSC, et al. Declining asthma prevalence in Hong Kong Chinese schoolchildren. *Clinical And Experimental Allergy* 2004;34(10 (Print)):1550-1555.
137. Miyake Y, Yura A, Masayuki I. Relationship between distance from major roads and adolescent health in Japan. *Journal of Epidemiology* 2002;12:418-23.
138. Kim YK, Park HS, Kim HY, Jee YK, Son JW, Bae JM, et al. Citrus red mite (*Panonychus citri*) may be an important allergen in the development of asthma among exposed children. *Clinical and Experimental Allergy* 2001;31:582-9.
139. Hong S-J, Lee M-S, Sohn MH, Shim JY, Han YS, Park KS, et al. Self-reported prevalence and risk factors of asthma among Korean adolescents: 5-year follow-up study, 1995-2000. *Clinical and Experimental Allergy* 2004;34(10):1556-62.
140. Lee MH, Kim YK, Min KU, Lee BJ, Bahn JW, Son JW, et al. Differences in sensitization rates to outdoor aeroallergens, especially citrus red mite (*Panonychus citri*), between urban and rural children. *Annals of Allergy, Asthma and Immunology* 2001;86:691-5.
141. Kim YK, Baek D, Koh YI, Cho SH, Choi IS, Min KU, et al. Outdoor air pollutants derived from industrial processes may be causally related to the development of asthma in children. *Annals of Allergy, Asthma and Immunology* 2001;86:456-60.
142. Kim YK, Chang YS, Lee MH, Hong SC, Bae JM, Jee YK, et al. Role of environmental exposure to spider mites in the sensitization and the clinical manifestation of asthma and rhinitis in children and adolescents living in rural and urban areas. *Clinical and Experimental Allergy* 2002;32:1305-9.
143. Kim SH, Kim YK, Lee MH, Hong SC, Bae JM, Min KU, et al. Relationship between sensitization to citrus red mite (*Panonychus citri*) and the prevalence of atopic diseases in adolescents living near citrus orchards. *Clinical and Experimental Allergy* 2002;32:1054-8.
144. Quah BS, Wan-Pauzi I, Ariffin N, Mazidah AR. Prevalence of asthma, eczema and allergic rhinitis: two surveys, 6 years apart, in Kota Bharu, Malaysia. *Respirology* 2005;10(2):244-9.
145. Wang XS, Tan TN, Shek LPC, Chng SY, Hia CPP, Ong NBH, et al. The prevalence of asthma and allergies in Singapore; data from two ISAAC surveys seven years apart. *Archives of Disease in Childhood* 2004;89(5):423-6.
146. Kao C-C, Huang J-L, Ou L-S, See L-C. The prevalence, severity and seasonal variations of asthma, rhinitis and eczema in Taiwanese schoolchildren. *Pediatric Allergy and Immunology* 2005;16(5):408-15.
147. Bauman A, Mitchell CA, Henry RL, Robertson CF, Abramson MJ, Comino EJ, et al. Asthma morbidity in Australia: an epidemiological study. *Medical Journal of Australia* 1992;156:827-31.
148. Forero R, Bauman A, Young L, Larkin P. Asthma prevalence and management in Australian adolescents: results from three community surveys. *Journal of Adolescent Health* 1992;13:707-12.
149. Crockett AJ, Cranston JM, Alpers JH. The changing prevalence of asthma-like respiratory symptoms in South Australian rural schoolchildren. *Journal of Paediatrics and Child Health* 1995;31:213-7.
150. Comino EJ, Mitchell CA, Bauman A, Henry RL, Robertson CF, Abramson MJ, et al. Asthma management in eastern Australia, 1990 and 1993. *Medical Journal of Australia* 1996;164:403-6.

151. Peat JK, Toelle BG, Gray EJ, Haby MM, Belousova E, Mellis CM, et al. Prevalence and severity of childhood asthma and allergic sensitisation in seven climatic regions of New South Wales. *Medical Journal of Australia* 1995;163:22-6.
152. Volkmer RE, Ruffin RE, Wigg NR, Davies N. The prevalence of respiratory symptoms in South Australian preschool children: I. Geographic location. *Journal of Paediatrics and Child Health* 1995;31:112-5.
153. Ponsonby AL, Couper D, Dwyer T, Carmichael A. Cross sectional study of the relation between sibling number and asthma, hay fever, and eczema. *Archives of Disease in Childhood* 1998;79:328-33.
154. Downs SH, Marks GB, Belosouva EG, Peat JK. Asthma and hayfever in Aboriginal and non-Aboriginal children living in non-remote rural towns. *Medical Journal of Australia* 2001;175:10-3.
155. Powell CV, Nolan TM, Carlin JB, Bennett CM, Johnson PD. Respiratory symptoms and duration of residence in immigrant teenagers living in Melbourne, Australia. *Archives of Disease in Childhood* 1999;81:159-62.
156. Valery PC, Chang AB, Shibasaki S, Gibson O, Purdie DM, Shannon C, et al. High prevalence of asthma in five remote indigenous communities in Australia. *European Respiratory Journal* 2001;17:1089-96.
157. Glasgow NJ, Goodchild EA, Yates R, Ponsonby AL. Respiratory health in Aboriginal and Torres Strait Islander children in the Australian Capital Territory. *Journal of Paediatrics and Child Health* 2003;39:534-9.
158. Mitchell EA, Asher MI. Prevalence, severity and medical management of asthma in European schoolchildren in 1985 and 1991. *Journal of Paediatrics and Child Health* 1994;30:398-402.
159. Wickens K, Barry D, Friezema A, Rhodius R, Bone N, Purdie G, et al. Obesity and asthma in 11-12 year old New Zealand children in 1989 and 2000. *Thorax* 2005;60(1):7-12.
160. Cohet C, Cheng S, MacDonald C, Baker M, Foliaki S, Huntington N, et al. Infections, medication use, and the prevalence of symptoms of asthma, rhinitis, and eczema in childhood. *Journal of Epidemiology and Community Health* 2004;58(10):852-7.
161. Weiland SK, Björkstén B, Brunekreef B, Cookson WO, von Mutius E, Strachan DP and the International Study of Asthma and Allergies in Childhood Phase II Study Group. Phase II of the International Study of Asthma and Allergies in Childhood (ISAAC II): rationale and methods. *European Respiratory Journal* 2004; 24(3): 406-12.
162. Flohr C, Weiland SK, Weinmayr G, Björkstén B, Bråbäck L, Brunekreef B, Büchele G, Clausen M, Cookson WOC, von Mutius E, Strachan DP, Williams HC, and the ISAAC Phase Two Study Group. The role of atopic sensitization in flexural eczema: Findings from the International Study of Asthma and Allergies in Childhood Phase Two. *Journal of Allergy and Clinical Immunology* 2008;121(1):141-7.
163. Long S. Respiratory syncytial virus infection and recurrent wheezing: a complex relationship (editorial). *Journal of Pediatrics* 2007; 151: 6-7.
164. Court CS, Cook DG, Strachan DP. Comparative epidemiology of atopic and non-atopic wheeze and diagnosed asthma in a national sample of English adults. *Thorax* 2002; 57: 951-7.

Figure legends

Figure 1. Prevalence of “wheezing in the last year” by calendar year of survey in all children (aged 0-16), reported in published studies in Australia and UK

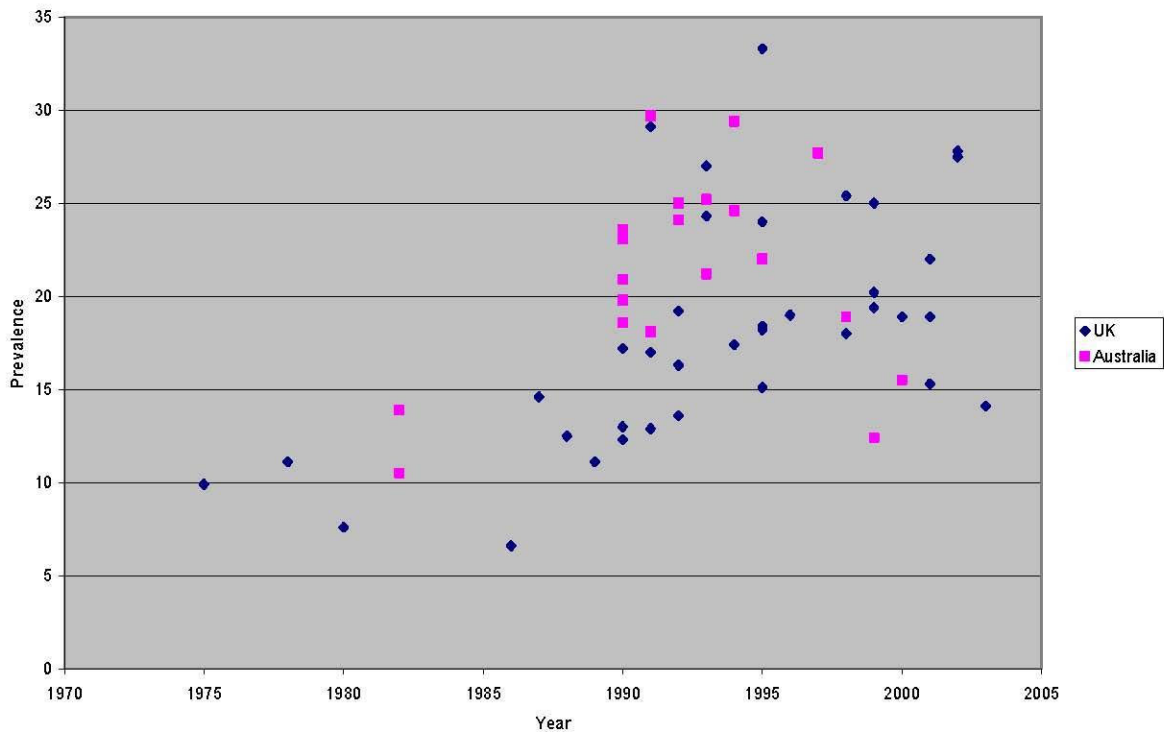


Figure 1

Additional files provided with this submission:

Additional file 1: asthma prevalence meta-analysis resubmission letter (environ
hea, 70K

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resub, 1343K

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