## EUROPEAN RESPIRATORY journal

FLAGSHIP SCIENTIFIC JOURNAL OF ERS

### **Early View**

Original research article

# The burden of asthma, hay fever and eczema in adults in 17 countries: GAN Phase I study

Kevin Mortimer, Maia Lesosky, Luis García-Marcos, M. Innes Asher, Neil Pearce, Eamon Ellwood, Karen Bissell, Asma El Sony, Philippa Ellwood, Guy B. Marks, Antonela Martínez-Torres, Eva Morales, Virginia Perez-Fernandez, Steven Robertson, Charlotte E. Rutter, Richard J. Silverwood, David P. Strachan, Chen-Yuan Chiang

Please cite this article as: Mortimer K, Lesosky M, García-Marcos L, *et al*. The burden of asthma, hay fever and eczema in adults in 17 countries: GAN Phase I study. *Eur Respir J* 2022; in press (https://doi.org/10.1183/13993003.02865-2021).

This manuscript has recently been accepted for publication in the *European Respiratory Journal*. It is published here in its accepted form prior to copyediting and typesetting by our production team. After these production processes are complete and the authors have approved the resulting proofs, the article will move to the latest issue of the ERJ online.

Copyright ©The authors 2022. For reproduction rights and permissions contact permissions@ersnet.org

Kevin Mortimer PhD<sup>1,2\*</sup>, Maia Lesosky PhD<sup>3,4\*</sup>, Luis García-Marcos PhD<sup>5\*</sup>, M. Innes Asher MBChB<sup>6\*</sup>, Neil Pearce PhD<sup>7\*</sup>, Eamon Ellwood DipTch<sup>6</sup>, Karen Bissell DrPH<sup>8</sup>, Asma El Sony PhD<sup>9\*</sup>, Philippa Ellwood MPH<sup>6</sup>, Guy B. Marks PhD<sup>10\*</sup>, Antonela Martínez-Torres NP<sup>11</sup>, Eva Morales PhD<sup>12</sup>, Virginia Perez-Fernandez PhD<sup>13\*</sup>, Steven Robertson BA<sup>7</sup>, Charlotte E. Rutter MSc<sup>7</sup>, Richard J. Silverwood PhD<sup>7,14</sup>, David P. Strachan MD<sup>15\*</sup>, Chen-Yuan Chiang DrPhilos<sup>16\*</sup> and the Global Asthma Network Phase I Study Group<sup>17</sup>

<sup>1</sup>Department of Medicine, University of Cambridge, Cambridge, United Kingdom
<sup>2</sup>Liverpool University Hospitals NHS Foundation Trust, Liverpool, United Kingdom
<sup>3</sup>Department of Clinical Sciences, Liverpool School of Tropical Medicine, Pembroke Place, Liverpool, L3 5QA, United Kingdom

<sup>4</sup>School of Public Health and Family Medicine, University of Cape Town, South Africa <sup>5</sup>Paediatric Allergy and Pulmonology Units, Virgen de la Arrixaca University Children's Hospital, University of Murcia and IMIB Bio-health Research Institute, Murcia; and ARADyAL Allergy Network, Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, Murcia, Spain.

<sup>6</sup>Department of Paediatrics: Child and Youth Health, Faculty of Medical and Health Sciences, University of Auckland, Private Bag 92019, Auckland, New Zealand

<sup>7</sup>Department of Medical Statistics, London School of Hygiene & Tropical Medicine, Keppel Street, London WC1E 7HT, United Kingdom

<sup>8</sup>School of Population Health, Faculty of Medical and Health Sciences, University of Auckland, Private Bag 92019, Auckland, New Zealand

<sup>9</sup>Epidemiological Laboratory (Epi-Lab) for Public Health, Research and Development, Khartoum 3, Block 3- Building 11, Khartoum, Sudan

<sup>10</sup>Respiratory & Environmental Epidemiology, University of New South Wales, Goulburn St, Sydney 2085, Sydney, Australia

<sup>11</sup>Paediatric Allergy and Pulmonology Units and Nurse Research Group, Virgen de la Arrixaca University Children's Hospital; and IMIB Bio-health Research Institute, Murcia, Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, 30394 Murcia, Spain <sup>12</sup>Department of Public Health Sciences, University of Murcia, and IMIB Bio-health Research Institute, Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, Murcia, Spain

Department of Biostatistics, University of Murcia, and IMIB Bio-health Research Institute,
 Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, Murcia, Spain
 Centre for Longitudinal Studies, UCL Social Research Institute, University College London,
 Bedford Way, London WC1H OAL, United Kingdom

<sup>15</sup>Population Health Research Institute, St George's, University of London, Cranmer Terrace, London SW17 ORE, United Kingdom

<sup>16</sup>International Union Against Tuberculosis and Lung Disease, Paris, France; and Division of Pulmonary Medicine, Department of Internal Medicine, Wan Fang Hospital, Taipei Medical University; and Division of Pulmonary Medicine, Department of Internal Medicine, School of Medicine, College of Medicine, Taipei Medical University, 111 Hsin-Long Road, Section 3, Taipei, 116, Taiwan

<sup>17</sup>Global Asthma Network Phase I Study Group listed at the end of the report.

<sup>\*</sup> Full professor

**Corresponding author** 

Chen-Yuan Chiang

Division of Pulmonary Medicine, Department of Internal Medicine, Wan Fang Hospital,

Taipei Medical University; and Division of Pulmonary Medicine, Department of Internal

Medicine, School of Medicine, College of Medicine, Taipei Medical University, 111 Hsin-Long

Road, Section 3, Taipei, 116, Taiwan

Email: cychiang@tmu.edu.tw

Take home message (a 256-character (including spaces) summary)

There is a substantial global burden of asthma, hay fever and eczema in adults representing

a major global public health problem. Accessible, affordable, equitable and effective

strategies are needed to reduce this burden across the life-course.

#### **ABSTRACT**

Asthma, hay fever and eczema are three common chronic conditions. There are no recent multi-country data on the burden of these three conditions in adults; the aims of this study are to fill this evidence gap.

The Global Asthma Network (GAN) Phase I is a multi-country cross-sectional population-based study using the same core methodology as the International Study of Asthma and Allergies in Childhood (ISAAC) Phase III. It provides data on the burden of asthma, hay fever, and eczema not only in children and adolescents but also for the first time in their parents/guardians.

Data were available from 193,912 adults (104,061 female; mean age 38 (SD 7·5)) in 43 centres in 17 countries. The overall prevalences (range) of symptoms of current wheeze, asthma ever, hay fever ever and eczema ever were 6.6% (0.9% -32.7%), 4.4%(0.9% -29.0%), 14.4%(2.8%-45.7%), and 9.9%(1.6%-29.5%), respectively. Centre prevalence varied considerably both between countries and within countries. There was a moderate correlation between hay fever ever and asthma ever, and between eczema ever and hay fever ever at the centre level. There were moderate to strong correlations between indicators of the burden of disease reported in adults and the two younger age groups.

We found evidence for a substantial burden of asthma, hay fever ever and eczema ever in countries examined highlighting the major public health importance of these diseases.

Prevention strategies and equitable access to effective and affordable treatments for these three conditions would help mitigate the avoidable morbidity they cause.

#### **Funding**

International Union Against Tuberculosis and Lung Disease, Boehringer Ingelheim New Zealand, Astra Zeneca Educational Grant, National Institute for Health Research, UK, Medical Research Council, UK, European Research Council, Instituto de Salud Carlos III, Spain.

#### **INTRODUCTION**

Asthma, hay fever and eczema are three common chronic conditions that typically start in childhood and often continue across the life-course [1]. All three conditions cause considerable morbidity globally especially when basic effective treatments are unavailable [2]. Asthma is an important cause of avoidable mortality [3].

The International Study of Asthma and Allergies in Childhood (ISAAC) investigated the symptom prevalence and determinants of asthma, rhinoconjunctivitis and eczema in school children on two previous occasions (ISAAC Phase I in 1993-5 and ISAAC Phase III in 2001-3) [4-15]. The Global Asthma Network (GAN) subsequently continued the work of ISAAC through the centres of ISAAC and new centres that are interested in GAN Phase I, which is a multi-country population-based cross-sectional study designed to assess the three conditions, as well as severity, management and risk factors in 13-14-year-old adolescents, 6-7-year-old children, and their parents/guardians using the same methods of ISAAC Phase III [16].

There has been no large survey on the prevalence of asthma in adults since WHO implemented the World Health Survey (WHS) in 2002 and 2003 [17], and no such surveys for hay fever ever or eczema ever. In this paper we report data on the prevalence of asthma symptoms, hay fever ever and eczema ever in adults in GAN Phase I. We compare their global patterns, and contrast with those observed in children in the same populations.

#### **METHODS**

The GAN methodology has previously been published [16,18], and will only be briefly summarized here.

#### **Participants**

The adult participants were the parents (or guardians) of children and adolescents in GAN Phase I. Cluster sampling was applied to randomly select at least 10 schools from a geographically defined sampling frame. All schools were included if there were < 10 schools in the sampling frame. The compulsory age group was adolescents, who self-completed written questionnaires at school. Additional inclusion of 6-7-year-olds was optional.

Optionally parents/guardians were also asked to complete similar questionnaires on their own health (the adult group), and the linkage between adults and children and adolescents was documented.

#### Questionnaires

Questionnaires for the adults were developed building on questionnaires used in ISAAC and the European Community Respiratory Health Survey [19,20]. The original questionnaire was in English; with translation to local languages and back-translation to English completed using a specific methodology common to ISAAC and GAN [16].

#### **Definitions**

#### Asthma:

"Current wheeze" was defined by a positive answer to the question "Have you had wheezing or whistling in the chest in the past 12 months?". "Severe asthma symptoms" were defined as those with current wheeze who, in the past 12 months, reported having had ≥4 attacks of wheeze, or >1 night per week sleep disturbance from wheeze, or wheeze affecting speech.

"Asthma ever" was defined as a positive answer to the question "Have you ever had asthma?"

#### Hay fever:

"Hay fever ever" was defined by a positive answer to the question: "Have you ever had hay fever?"

#### Eczema:

"Eczema ever" was defined as a positive answer to the question: "Have you ever had eczema?"

#### Sample size and study power

Sample sizes of at least 1000 and preferably 3000 were sought for the adolescents and children within each centre [17]. A participation rate of at least 80% for the adolescents and 70% for the children was aimed for [21]. The actual response rate was 90% for adolescents and 79% for children [22]. We were unable to calculate a conventional response rate for the adults because some schoolchildren have only one parent or guardian and the number of adults that received the questionnaire was not known. The estimated median participation rate of adults, by using a "per child" approach [21], was 82·9%, range 30·2% - 100%, with 4 centres unable to be calculated due to insufficient information.

#### Data handling and analysis

All centres submitted their datasets and a Centre Report documenting the methodology used to the GAN Global Centre in Auckland (New Zealand) [16]. A first quality control check was performed together with a careful review of the Centre Report for adherence to protocol. Depending on the language used locally, the dataset was then sent to one of the two GAN data centres in either Murcia (Spain) (Spanish and Portuguese speaking centres) or London (United Kingdom) (all other languages), for a standardised and coordinated data

check. Centres reported in this analysis are any centres that were included in the analysis of data from children and adolescents [18] that also collected data from adults. For prevalence estimations, positive answers to a specific symptom in the centre was divided by the number of completed questionnaires.

Global national income (GNI) category for each country was calculated using cut-off points provided by the World Bank in June 2020 [23], and countries were classified into high income country (HIC), upper middle income country (U-MIC) and, lower middle/low income country (L-MIC/LIC). The Spearman correlation coefficient was used to estimate the correlation between symptoms of different conditions and between age groups using centre level data. The correlation was defined as strong if correlation coefficient ≥0.7, moderate if ≥0.4 but <0.7, and weak if <0.4. Multilevel log-binomial regression was used to estimate how much of the variability of each symptom's prevalence was dependent on centre-level variation, additional to within-centre binomial sampling error. As the intraclass correlation coefficient was higher than 5% (ICC>0.05) in the null model in all instances, multilevel models fitting centre as a random effect were used to estimate the effect of sex and GNI in the prevalence of symptoms of the three conditions. A uniform approach to data processing, checking and analysis was used, using Stata versions 13−15 (Stata Corp LLC, College Station, Texas, USA).

#### **Centre funding and ethics**

All centres in GAN Phase I obtained their own funding and applied for ethics approval from their local ethics committee before starting the study.

#### Role of the funding source

The funding sources had no role in study design; collection, analysis, and interpretation of data; writing of the report; or the decision to submit the paper for publication.

#### **RESULTS**

Data were collected from 193,912 adults (104,061 female; mean (SD) age 38 (7·5); 5·0% >50 year-old; 16·8% current smokers; 47.2% parents of adolescents) in 43 centres (including 12 ISAAC Phase I and 19 ISAAC Phase III centres) in 17 countries between 2015 and 2020 (Web Tables 1-2). The prevalence of current wheeze was highest (10·6%, 95% Confidence Interval (CI): [10·2%, 10·9%]) among participants from HICs, followed by 8·4% [8·2%, 8·6%] among participants from U-MICs, and 3·6% [3·5%, 3·8%] among participants from L-MICs/LICs (Table 1). Similar trends across GNI categories were noted for asthma ever, severe asthma symptoms and hay fever ever with the exception of eczema ever which was lowest in U-MICs.

#### **Asthma**

The prevalence of asthma ever was 4·4%, ranging from 0·9% in Gjilan and Ferizaj, Kosovo to 29·0% in Costa Rica. Heterogeneity was high both between countries and between centres within countries (Figure 1, Web Table 1, and Web Figures 1-2). Centre level variation explained 21·8% of the variability of the prevalence in the multilevel analysis. Adult females had a higher prevalence of asthma ever than males (4·8% vs 3·9%; aRR for males 0·85, 95% CI: [0·82, 0·89]).

The overall prevalence of current wheeze was 6.6%, ranging from 0.9% in New Delhi, India to 32.7% in Tegucigalpa, Honduras. Heterogeneity of the prevalence of current wheeze was high (Figure 2, Web Table 1, and Web Figures 1-2). Centre level variation explained 13.1% of the variability of the prevalence of current wheeze in the multilevel analysis. Adult females had a higher prevalence of current wheeze than males (6.8% vs 6.2%; aRR for males 0.97, 95% CI: [0.94, 1.00]).

The prevalence of severe asthma symptoms was 2.6%, ranging from 0.2% in Bikaner, India to 20.9% in Tegucigalpa, Honduras. The prevalence of severe asthma symptoms among those who reported current wheeze was 39.0%, ranging from 15.0% in Bikaner, India to 63.9% in Tegucigalpa, Honduras (Web Table 1, and Web Figures 1-2). Centre level variation accounted for 17.0% of the variability of the prevalence of severe asthma symptoms. Adult females had a higher prevalence of severe asthma symptoms than males (2.9% vs 2.2%; aRR for males 0.83, 95% CI: [0.82, 0.90]).

#### Hay fever

The prevalence of hay fever ever was 14·4%, ranging from 2·8% in Ibadan, Nigeria to 45·7% in Bangkok, Thailand (Figure 3, Web Table 1, and Web Figures 1-2). Centre level variation explained 21·8% of the variability of the prevalence of hay fever ever. The prevalence of hay fever ever was higher in females than males (14·7% vs 14·0%; aRR for males 0·92, 95% CI: [0·90, 0·93]).

#### Eczema

The prevalence of eczema ever was 9.9%, ranging from 1.6% in Tijuana, Mexico to 29.5% in Bangkok, Thailand (Figure 4, Web Table 1, and Web Figures 1-2). Centre level variation explained 19.6% of the variability of the prevalence of eczema ever. The prevalence of eczema ever was higher in females than males (10.0% vs 9.9%; aRR for males 0.90, 95% CI: [0.88, 0.93]).

#### Correlations of prevalence between the three conditions

There was moderate correlation between the prevalence of hay fever ever and asthma ever (Rho: 0.54, 95% CI: [0.32, 0.76]), and between eczema ever and hay fever ever (0.66 [0.48,

0.83]), but no significant correlation between asthma ever and eczema ever (0.13 [-0.19, 0.45]) at the centre level (Figure 5). The correlation between the prevalence of hay fever ever and asthma ever and between the prevalence of eczema ever and hay fever ever remained after stratification by sex (Web Figure 3).

#### Relationship between age groups

There was strong correlation between the prevalence of asthma ever in adults vs adolescents (Rho: 0·87, 95%CI: [0·79, 0·95]), between asthma ever in adults vs children (0·83 [0·66, 1.00]), between current wheeze in adults vs adolescents (0·81 [0·68,0.94]), between severe asthma symptoms in adults vs adolescents (0·79 [0·67, 0·92]), between severe asthma symptoms in adults vs children (0·82 [0·65, 0·98]) between hay fever ever in adults vs adolescents (0·75 [0·57, 0·92]), between eczema ever in adults vs adolescents (0·87 [0·78, 0·95]), and between eczema ever in adults vs children (0·71 [0·51, 0.91]). There was moderate correlation between current wheeze in adults vs children, and between hay fever ever in adults vs children (Web Figure 4).

#### DISCUSSION

The major findings of GAN Phase I were: 1) the overall prevalence of symptoms of current wheeze, asthma ever, hay fever ever and eczema ever was 6·6%, 4·4%, 14·4%, and 9·9%, respectively; 2) centre prevalence varied considerably both between countries and within countries; 3) the burden of all three conditions was higher in females and in higher income countries; 4) there was a moderate correlation between hay fever ever and asthma ever, and between eczema ever and hay fever ever at the centre level; 5) there were moderate to strong correlations between the prevalence of asthma symptoms, hay fever ever and eczema ever reported in adults and the two younger age groups.

A multi-country survey on the prevalence of asthma in adults has been conducted previously by the European Community Respiratory Health Survey (ECRHS) in the 1990s [24]. The questions used in the ECRHS were "Have you had wheezing or whistling in your chest at any time in the last 12 months?" and "Have you had an attack of asthma in the last 12 months?" [25] The ECRHS reported large geographical variations in the prevalence of asthma [25]. The median prevalence of current asthma was 4·5% (range 2·0% -11·9%) in ECRHS stage one and 5·2% (range 1·2% − 13·0%) in ECRHS stage two [26]. Females had a higher prevalence of current asthma than males and the prevalence of wheeze was negatively associated with age. The ECRHS concluded that the geographical variations in the prevalence of asthma were most likely due to environmental factors [26]. The WHS enrolled 308,218 adults aged ≥18 years from 64 countries [17]. The WHS defined current wheeze symptoms as a positive response to any of the symptom questions: "during the last 12 months, have you experienced any of the following: 1) attacks of wheezing or whistling breathing? (yes/no); or 2) attacks of wheezing that came on after you stopped exercising or some other physical activity? (yes/no)". The WHS reported that global prevalence of current wheeze symptoms

was 9.2%, ranging from 2.4% in Vietnam to 24.0% in Brazil. The prevalence of current wheeze symptoms increased with age, was higher among male than female, more common in smokers than nonsmokers, and was relatively high in HICs and LICs, and relatively low in MICs [17], In our survey, the prevalence of asthma ever varied markedly between centres as did the prevalence of current wheeze symptoms with less variability within countries than seen in children and adolescents. Current severe asthma symptoms were commonly reported (range: 15.0% - 63.9%) amongst participants reporting wheeze in the past 12 months across all centres suggesting a concerning level of poor asthma control [27]. There was a clear association between GNI category and the prevalence of current wheeze was highest in HICs and lowest in L-MICs/LICs, similar to the pattern seen of a lower prevalence of current wheeze symptoms in children and adolescents in L-MICs/LICs [18]. Asthma symptoms were more common in females as seen in the adolescent group and other studies of older children [4,28] and the ECRHS. The WHS reported that males were more likely to report current wheeze symptoms, perhaps because its study population was older (34% aged >50 years), and 30% were smokers, thus may have wheeze due to chronic obstructive pulmonary disease. Whether the difference between our findings and that of WHS with regards to GNI was attributable to differences in definitions (e.g. of symptoms), study population, different prevalence of environmental risk factors and genetic backgrounds requires further investigation.

The ECRHS reported that the median prevalence of nasal allergy and hay fever was 20.9% (range 9.5% - 40.9%) [26]. Subjects with perennial rhinitis were more likely to have current asthma. In our study, the prevalence of hay fever ever was 14.4% but was variable (from 2.8% in Ibadan, Nigeria, to 45.7% in Bangkok, Thailand) with less variability within countries. This was also seen in children and adolescents as was an association between hay fever and GNI

categories with the greatest burden seen in HICs [18]. Consistent with the ECRHS, our study shows moderate correlation between the prevalence of hay fever ever and asthma ever.

The overall prevalence of eczema ever was 9.9% but varied from 1.6% in Tijuana, Mexico, to 29.5% in Bangkok, Thailand, with less variability within countries. We did find an association between eczema ever and GNI categories with the greatest burden seen in HICs, which was also seen in children and adolescents in GAN Phase I [18]. In previous ISAAC surveys, the adolescents in GAN Phase I, as well as in other cohort studies, a difference between sexes (more prevalent in females) was found [7,18,29,30] and we found the same in this study of adults. No significant correlation between the prevalence of asthma ever and eczema ever in adults was identified. As eczema tends to occur in early stage of life and decreases with age [31], whether this was in part due to recall bias was unknown.

There was considerable variation in prevalence of all three conditions in adults, which is partly accounted for by centre level variation. We speculate that the difference in the prevalence of risk factors may contribute to the difference in observed prevalence between centres and countries; risk factors associated with the three conditions collected as part of GAN Phase I will be analysed and reported separately, which should provide more insight into this issue. We found moderate to strong correlations between the prevalence of asthma symptoms, hay fever ever and eczema ever reported in adults and the two younger age groups, likely indicating that parents/guardians and the two younger age groups have similar environmental and genetic risk factors of the three conditions.

The strengths and weaknesses of the ISAAC and GAN Phase I methodology have been discussed in depth [4] and have been previously summarised elsewhere [16,18]. We acknowledge limitations of the small number of GAN Phase I centres vs ISAAC Phase I and III,

the self-selection of centres potentially limiting representativeness, challenges of inferring clinical diagnoses from self-reporting via questionnaires (e.g. risk of recall bias and lack of direct physician diagnoses), difficulties around the translation of concepts such as "wheezing" into different languages. Furthermore, we did not collect information on current symptoms of hay fever and eczema, but only information of hay fever ever and eczema ever, which may not represent current prevalences of hay fever and eczema because both conditions may remit during adolescence. Moreover, it is possible that parents of children with hay fever or eczema may be more aware of these two conditions and more likely to report having hay fever ever or eczema ever than parents of children with no hay fever or eczema; such potential recall bias may affect the correlation of the prevalence of hay fever ever and eczema ever between children and adults. There was difficulty in obtaining a high response rate for some centres [22]. The correlation analysis is an ecologic analysis (at centre level) and these correlations may not hold at the individual level. Key additional strengths are the linkages between the child, adolescent and adult participants that have enabled additional analyses including exploring the relationship between symptoms reported by the different age groups. However, recruiting the parents of the child and adolescent participants will have led to a degree of selection bias and the included adult population may not be fully representative of the local population in terms of factors including age and socioeconomic status.

In conclusion, the present study offers a unique picture of current symptoms related to asthma, and lifetime history of asthma, hay fever and eczema. Our findings in adults were largely consistent with our findings in children and adolescents (particularly) [18] and the burden of the three conditions seems to correlate across the three age groups. Further studies are needed to confirm whether findings from one group may be cautiously extrapolated to the others.

#### **ILLUSTRATIONS**

#### Figure 1

Map of the prevalence of asthma ever. The symbols indicate prevalence values of <5% (blue squares), 5 to <10% (green circle), 10 to <20% (yellow diamonds) and  $\geq$ 20% (red stars).

#### Figure 2

Map of the prevalence of current wheeze. The symbols indicate prevalence values of <5% (blue squares), 5 to <10% (green circle), 10 to <20% (yellow diamonds) and ≥20% (red stars).

#### Figure 3

Map of the prevalence of hay fever ever. The symbols indicate prevalence values of <5% (blue squares), 5 to <10% (green circle), 10 to <20% (yellow diamonds) and ≥20% (red stars).

#### Figure 4

Map of the prevalence of eczema ever. The symbols indicate prevalence values of <5% (blue squares), 5 to <10% (green circle), 10 to <20% (yellow diamonds) and ≥20% (red stars).

#### Figure 5

Rank correlation values and scatter plots of prevalence of the three conditions at the centre level. The dashed line is the identity line. Rank correlation coefficient and 95%CI is shown in each graph.

#### Web Figure 1

Ranking of centres for the symptom prevalences of current wheeze, asthma ever, hay fever ever and eczema ever.

#### Web Figure 2

Ranking of centres for the symptom prevalences of current wheeze, asthma ever, hay fever ever and eczema ever by sex (males on left).

#### Web Figure 3

Rank correlation values and scatter plots of prevalence of the three conditions at the centre level by sex (males on left). The dashed line is the identity line. Rank correlation coefficient and 95%CI is shown in each graph.

#### Web Figure 4

Rank correlation comparing centre prevalence (%) of reporting current wheeze, asthma ever, severe asthma symptoms, hay fever ever and eczema ever between the three age groups (children, adolescents, adults) included in GAN Phase I. The dashed line is the identity line.

Rank correlation coefficient and 95%CI is shown in each graph.

#### **TABLES**

#### Table 1

Symptom prevalence of asthma, hay fever and eczema in centres grouped by Gross National Income (GNI).

#### Web Table 1

Demographic summary by centre.

#### Web Table 2

Symptom prevalence of asthma, hay fever and eczema by centre.

#### **Authors individual contributions**

The following individual contributions were made: conceptualisation IA, KB, C-YC, AES, PE, LG-M, GM, NP, DS; data curation EE, PE, LG-M, EM, VP-F, CR, SR, RS, ML; verification of the underlying data CR, NP, VP-F, DS; formal analysis ML, NP, CR, DS; investigation IA; methodology IA, C-YC, PE, LG-M, NP, CR, DS, RS; project administration, IA, EE; PE; resources IA; supervision LG-M, NP, DS, RS; validation PE; visualisation EE, PE, CR; writing — original draft KM, C-YC; writing — review/editing IA, GM, AM-T, SR, CR, KB, AES, EE, PE, LG-M, EM, VP-F, NP, DS, RS, ML and the Global Asthma Network Phase I Study Group; the latter contributed original data to the analyses.. All authors shared responsibility for the decision to submit the manuscript.

#### **Declaration of interests**

The authors declare that they have no conflict of interest.

#### Acknowledgements

We are grateful to the children, adolescents and adults who willingly participated with the help of schools and field workers in GAN Phase I.

We thank the children, adolescents and parents who participated in GAN Phase I; the school staff for their assistance and help with coordination; the principal investigators and their colleagues; the many funding bodies throughout the world that supported the individual GAN centres.

The GAN Global Centre in Auckland was funded by The University of Auckland with additional funding from the International Union Against Tuberculosis and Lung Disease, Boehringer Ingelheim NZ, Astra Zeneca Educational Grant. The London Data Centre was supported by a PhD studentship [to CR] from the UK Medical Research Council (grant number MR/N013638/1) and funding from the European Research Council under the

European Union's Seventh Framework Programme (FP7/2007-2013, ERC grant agreement number 668954). The Murcia Data Centre was supported by the University of Murcia and by Instituto de Salud Carlos III, fund PI17/0170. ML was supported in part by the Academy of Medical Sciences Newton Advanced Fellowship (NAF\R2\180681).We thank the NIHR Global Health Research Unit on Lung Health and TB in Africa at Liverpool School of Tropical Medicine - "IMPALA" for helping to make this work possible (grant number 16/136/35); IMPALA was commissioned by the National Institute for Health Research (NIHR) Global Health Research (GHR) using UK aid from the UK Government. The views expressed in this publication are those of the authors and not necessarily those of any of the funders. Individual centres involved in GAN Phase I data collection were funded by the following organisations:

Brazil, Uruguaiana, funded by Dr Marilyn Urrutia Pereira; Cameroon, Yaounde, funded by Elvis Ndikum (95%) and from family, friends (5%); Costa Rica, partially funded by an unrestricted grant from Astra Zeneca for logistic purposes; India; [Bikaner, Chandigarh, Jaipur, Kolkata, Kottayam, Lucknow, Mysuru, New Delhi, Pune], GAN Phase I was undertaken by Asthma Bhawan in India which was supported by Cipla Foundation; Iran, Karaj, Alborz University of Medical Sciences; Kosovo, Gjakova, Municipality of Gjakova and the Directorate for Health and Education; Mexico, Puerto Vallarta, Centro Universitario de la Costa, Universidad de Guadalajara; New Zealand, Auckland, Auckland Asthma Charitable Trust; Nigeria, Ibadan, funded by National Institute for Health Research (NIHR) (IMPALA, grant reference 16/136/35) using UK aid from the UK Government to support global health research; Poland, Katowice, funded by the Medical University of Silesia.

#### **Data sharing**

The study protocol including a recommended informed consent form and statistical analysis plan are in the public domain (http://globalasthmanetwork.org/surveillance/

surveillance.php). The GAN Phase I data, including de-identified individual participant data, will be password protected and made available on the Global Asthma Network website <a href="http://www.globalasthmanetwork.org/">http://www.globalasthmanetwork.org/</a> within 12 months of all GAN Phase I analyses being published. Access for non GAN researchers will require a formal request to the GAN Steering Group for consideration, by submission of a written proposal and on acceptance, a signed data access agreement.

#### **Global Asthma Network Phase I Study Group:**

Global Asthma Network Steering Group: MI Asher, Department of Paediatrics: Child and Youth Health, Faculty of Medical and Health Sciences, University of Auckland, Private Bag 92019, Auckland, New Zealand; K Bissell, School of Population Health, Faculty of Medical and Health Sciences, University of Auckland, Auckland, New Zealand; C-Y Chiang, International Union Against Tuberculosis and Lung Disease, Paris, France; and Division of Pulmonary Medicine, Department of Internal Medicine, Wan Fang Hospital, Taipei Medical University; and Division of Pulmonary Medicine, Department of Internal Medicine, School of Medicine, College of Medicine, Taipei Medical University, 111 Hsin-Long Road, Section 3, Taipei, 116, Taiwan; A El Sony, Epidemiological Laboratory for Public Health and Research, Khartoum 3 Block3-Building 11, Khartoum, Sudan; E Ellwood, P Ellwood, Department of Paediatrics: Child and Youth Health, Faculty of Medical and Health Sciences, Private Bag 92019, University of Auckland, Auckland, New Zealand; L García-Marcos, Pediatric Allergy and Pulmonology Units, Virgen de la Arrixaca University Children's Hospital, University of Murcia and IMIB Bioresearch Institute, Murcia; and ARADyAL Allergy Network, Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, 30394 Murcia Spain; GB Marks, Respiratory & Environmental Epidemiology, University of New South Wales, Goulburn St, Sydney 2085, Sydney, Australia; R Masekela, Department of Paediatrics and Child Health, Nelson R Mandela School of Clinical Medicine, College of Health Sciences,

University of KwaZulu Natal, Durban, South Africa; E Morales, Department of Public Health Sciences, University of Murcia, and IMIB Bio-health Research Institute, Edificio

Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, Murcia, Spain; K Mortimer,
Liverpool School of Tropical Medicine, Pembroke Place, Liverpool L3 5QA, United Kingdom;
N Pearce, Department of Medical Statistics, London School of Hygiene & Tropical Medicine,
Keppel Street, London WC1E 7HT, United Kingdom; DP Strachan, Population Health
Research Institute, St George's, University of London, Cranmer Terrace, London SW17 ORE,
United Kingdom.

#### **Global Asthma Network International Data Centres:**

**GAN Global Centre:** P Ellwood, E Ellwood, MI Asher, Department of Paediatrics: Child and Youth Health, Faculty of Medical and Health Sciences, Private Bag 92019, University of Auckland, Auckland, New Zealand.

Murcia, Spain: L García-Marcos, Pediatric Allergy and Pulmonology Units, Virgen de la Arrixaca University Children's Hospital, University of Murcia and IMIB Bioresearch Institute, Murcia; and ARADyAL Allergy Network, Edificio Departamental-Laib, Murcia, Spain; V Perez-Fernández, Department of Paediatrics, University of Murcia; and IMIB Bio-health Research Institute, Murcia, Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, 30394 Murcia Spain; E Morales, Department of Public Health Sciences, University of Murcia, and IMIB Bio-health Research Institute, Murcia, Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, 30394 Murcia, Spain; A Martinez-Torres, Paediatric Allergy and Pulmonology Units and Nurse Research Group, Virgen de la Arrixaca University Children's Hospital, University of Murcia and IMIB Bio-health Research Institute, Murcia, Edificio Departamental-Laib, Avenida Buenavista s/n, 30120 El Palmar, 30394 Murcia, Spain. London, United Kingdom: DP Strachan, Population Health Research Institute, St George's, University of London, Cranmer Terrace, London SW17 ORE, United Kingdom; N Pearce, S

Robertson, CE Rutter, Department of Medical Statistics, London School of Hygiene & Tropical Medicine, Keppel Street, London WC1E 7HT, United Kingdom; RJ Silverwood, Department of Medical Statistics, London School of Hygiene & Tropical Medicine, Keppel Street, London WC1E 7HT, United Kingdom and Centre for Longitudinal Studies, UCL Social Research Institute, University College London, 20 Bedford Way, London WC1H 0AL, United Kingdom. Global Asthma Network Adult Principal Investigators: Brazil: M Urrutia-Pereira, Federal University of Pampa, UNIPAMPA (Uruguaiana); Cameroon: GA Ajeagah, The University of Yaounde 1 (Yaounde); Costa Rica: ME Soto-Martinez, Hospital Nacional de Niños "Dr. Carlos Saénz Herrera, Caja Costarricense Seguro Social - Universidad de Costa Rica, San José, Costa Rica (Costa Rica); Greece: K Priftis, National and Kapodistrian University of Athens (Athens); Honduras: J Sanchez, Instituto Nacional Cardiopulmonar (Tegucigalpa); India: SK Kochar, Sardar Patel Medical College (Bikaner); M Singh, Postgraduate Institute of Medical Education and Research (Chandigarh); N Singh, Asthma Bhawan (Jaipur); N Sit, National Allergy Asthma Bronchitis Institute (Kolkata (19)); TU Sukumaran, Pushpagiri Institute of Medical Sciences and Research, Thiruvalla, Kottayam (Kottayam); S Awasthi, King George's Medical University (Lucknow); PA Mahesh, JSS Medical College, JSSAHER (Mysuru); S Sinha, All India Institute of Medical Sciences (New Delhi); M Barne, Chest Research Foundation (Pune); Iran: M Tavakol, Alborz University of Medical Sciences (Karaj); N Behniafard, Shahid Sadoughi University of Medical Sciences (Yazd); Kingdom of Saudi Arabia: SA Alomary, Ministry of Health (Kingdom of Saudi Arabia); Kosovo: I Bucaliu-Ismajli, The Principal Center of Family Care (Ferizaj); L Hana-Lleshi, General Hospital "Isa Grezda" Gjakova, Kosovo (Gjakova); V Gashi, American Hospital in Kosovo (Gjilan); X Kurhasani, UBT College Kosovo (Peja); B Gacaferri-Lumezi, University of Prishtina Hasan Prishtina (Peja 6-7); LN Ahmetaj\*, University Hospital (Prishtina); V Lokaj-Berisha, University of Prishtina (Prizren); México: MG Sanchez Coronel, COMPEDIA (Colegio Mexicano de Pediatras Especialistas en Inmunología y Alergia) (Aguascalientes); G Ochoa-Lopez, Department of Pediatric Allergology (Ciudad Juárez); R

García-Almaráz, Hospital Infantil de Tamaulipas (Ciudad Victoria); JA Sacre Hazouri, Instituto Privado de Alergia, (Córdoba); MdJ Ambriz-Moreno, Hospital General de Matamoros Tamaulipas Mexico "Dr. Alfredo Pumarejo Lafaurie" (Matamoros); JV Mérida-Palacio, Centro de Investigacion de Enfermedades Alergicas y Respiratorias (Mexicali); OJ Saucedo-Ramirez, Hospital Angeles Pedregal (Mexico City North); LO Hernández-Mondragón, CRIT de Michoacán (Michoacán); A Arias-Cruz, Hospital Universitario (Monterrey); CA Jiménez González, Universidad Autonoma of San Luis Potosí (San Luis Potosí); AJ Escalante-Dominguez, Hospital General Tijuana [Isesalud] (Tijuana); FJ Linares-Zapién, Centro De Enfermedades Alergicas Y Asma de Toluca (Toluca Rural); EM Navarrete-Rodriguez, Hospital Infantil de México Federico Gómez (Toluca Urban); New Zealand: I Asher, University of Auckland (Auckland); Nigeria: AG Falade, University of Ibadan and University College Hospital (Ibadan); Poland: G Brożek, Medical University of Silesia (Katowice); Russia: K Kyzmicheva, Tyumen State Medical University (Tyumen); Spain: L García-Marcos\*, Pediatric Allergy and Pulmonology Units, Virgen de la Arrixaca University Children's Hospital, University of Murcia and IMIB Bioresearch Institute, Murcia; (Cartagena); Taiwan: K Yeh, Chang Gung Memorial Hospital (Taipei); Thailand: S Chinratanapisit, Department of Pediatrics, Bhumibol Adulyadej Hospital, Royal Thai Air Force (Bangkok).

\* National Coordinators

Global Asthma Network National Co-ordinators not named above: Brazil: D Solé, Escola

Paulista de Medicina, Federal University of São Paulo; Costa Rica: ME Soto-Quirós, University

of Costa Rica; India: V Singh, Asthma Bhawan; Kingdom of Saudi Arabia: WA Althagafi,

Ministry of Health; México: BE Del Río Navarro, Service of Allergy and Clinical Immunology,

Hospital Infantil de México; Thailand: P Vichyanond, Mahidol University.

#### **REFERENCES**

- 1. Meghji J, Mortimer K, Agusti A, et al. Improving lung health in low-income and middle-income countries: from challenges to solutions. *Lancet* 2021; **397**(10277): 928-40.
- 2. Vos T, Lim SS, Abbafati C, et al. Global burden of 369 diseases and injuries in 204 countries and territories, 1990-2019: a systematic analysis for the Global Burden of Disease Study 2019. *Lancet* 2020; **396**(10258): 1204-22.
- 3. Asher I, Bissell K, Chiang C-Y, et al. Calling time on asthma deaths in tropical regions: how much longer must people wait for essential medicines? *Lancet Respir Med* 2019; **7**(1): 13-5.
- 4. The International Study of Asthma and Allergies in Childhood (ISAAC) Steering Committee. Worldwide variation in prevalence of symptoms of asthma, allergic rhinoconjunctivitis, and atopic eczema: ISAAC. *Lancet* 1998; **351**(9111): 1225-32.
- 5. The International Study of Asthma and Allergies in Childhood (ISAAC) Steering Committee. Worldwide variations in the prevalence of asthma symptoms: the International Study of Asthma and Allergies in Childhood (ISAAC). *Eur Respir J* 1998; **12**(2): 315-35.
- 6. Strachan D, Sibbald B, Weiland S, et al. Worldwide variations in prevalence of symptoms of allergic rhinoconjunctivitis in children: the International Study of Asthma and Allergies in Childhood (ISAAC). *Pediatr Allergy Immunol* 1997; **8**(4): 161-8.
- 7. Williams H, Robertson C, Stewart A, et al. Worldwide variations in the prevalence of symptoms of atopic eczema in the international study of asthma and allergies in childhood. *J Allergy Clin Immunol* 1999; **103**(1): 125-38.
- 8. Ellwood P, Asher MI, Björkstén B, et al. Diet and asthma, allergic rhinoconjunctivitis and atopic eczema symptom prevalence: an ecological analysis of the International Study of Asthma and Allergies in Childhood (ISAAC) data. ISAAC Phase One Study Group. *Eur Respir J* 2001; **17**(3): 436-43.

- 9. Foliaki S, Nielsen SK, Björkstén B, et al. Antibiotic sales and the prevalence of symptoms of asthma, rhinitis, and eczema: The International Study of Asthma and Allergies in Childhood (ISAAC). *Int J Epidemiol* 2004; **33**(3): 558-63.
- 10. Weiland SK, Hüsing A, Strachan DP, et al. Climate and the prevalence of symptoms of asthma, allergic rhinitis, and atopic eczema in children. *Occup Environ Med* 2004; **61**(7): 609-15.
- 11. Burr ML, Emberlin JC, Treu R, et al. Pollen counts in relation to the prevalence of allergic rhinoconjunctivitis, asthma and atopic eczema in the International Study of Asthma and Allergies in Childhood (ISAAC). *Clin Exp Allergy* 2003; **33**(12): 1675-80.
- 12. Asher MI, Montefort S, Björkstén B, 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**(9537): 733-43.
- 13. Pearce N, Aït-Khaled N, Beasley R, et al. Worldwide trends in the prevalence of asthma symptoms: phase III of the International Study of Asthma and Allergies in Childhood (ISAAC). *Thorax* 2007; **62**(9): 758-66.
- 14. Lai CKW, Beasley R, Crane J, et al. Global variation in the prevalence and severity of asthma symptoms: Phase Three of the International Study of Asthma and Allergies in Childhood (ISAAC). *Thorax* 2009; **64**(6): 476-83.
- 15. Ellwood P, Williams H, Aït-Khaled N, et al. Translation of questions: The International Study of Asthma and Allergies in Childhood (ISAAC) experience. Int J Tuberc Lung Dis. 2009; 13: 1174-1182.
- 16. Ellwood P, Asher MI, Billo NE, et al. The Global Asthma Network rationale and methods for Phase I global surveillance: prevalence, severity, management and risk factors. *Eur Respir J* 2017; **49**(1): 1601605.

- 17. Sembajwe G, Cifuentes M, Tak SW, et al. National income, self-reported wheezing and asthma diagnosis from the World Health Survey. *European Respiratory Journal* 2010; **35**(2): 279-86.
- 18. Garcia-Marcos L, Asher MI, Pearce N, et al. The burden of asthma, hay fever and eczema in children in 25 countries: GAN Phase I. Eur Respir J 2022 (in press).
- 19. The European Community Respiratory Health Survey II Steering Committee. The European Community Respiratory Health Survey II. *Eur Respir J* 2002; **20**(5): 1071-9.
- 20. Ellwood P, Asher MI, Ellwood E, et al. The Global Asthma Network Manual for Global Surveillance: Prevalence, Severity and Risk Factors; August 2015. Available from: http://www.globalasthmanetwork.org/surveillance/manual/manual.php.
- 21. Ellwood P, Ellwood E, Rutter C, et al. Global Asthma Network Phase I Surveillance: Geographical Coverage and Response Rates. *J Clin Med* 2020; 9:3688.
- 22. Asher MI, Rutter CE, Bissell K, et al. Worldwide trends in the burden of asthma symptoms in school children:Global Asthma Network Phase I cross-sectional studies. *Lancet* 2021;398:1569-80.
- 23. World Bank. New World Bank country classifications by income level: 2020-2021. https://blogsworldbankorg/opendata/new-world-bank-country-classifications-income-level-2020-2021 (accessed 10/05/2021).
- 24. Burney P, Luczynska C, Chinn S, et al. The European Community Respiratory Health Survey. *Eur Respir J* 1994; **7**(5): 954-60.
- 25. European Community Respiratory Health Survey. Variations in the prevalence of respiratory symptoms, self-reported asthma attacks, and use of asthma medication in the European Community Respiratory Health Survey (ECRHS). *Eur Respir J* 1996; **9**(4): 687-95.
- 26. Janson C, Anto J, Burney P, et al. The European Community Respiratory Health Survey: what are the main results so far? *Eur Respir J* 2001; **18**(3): 598-611.

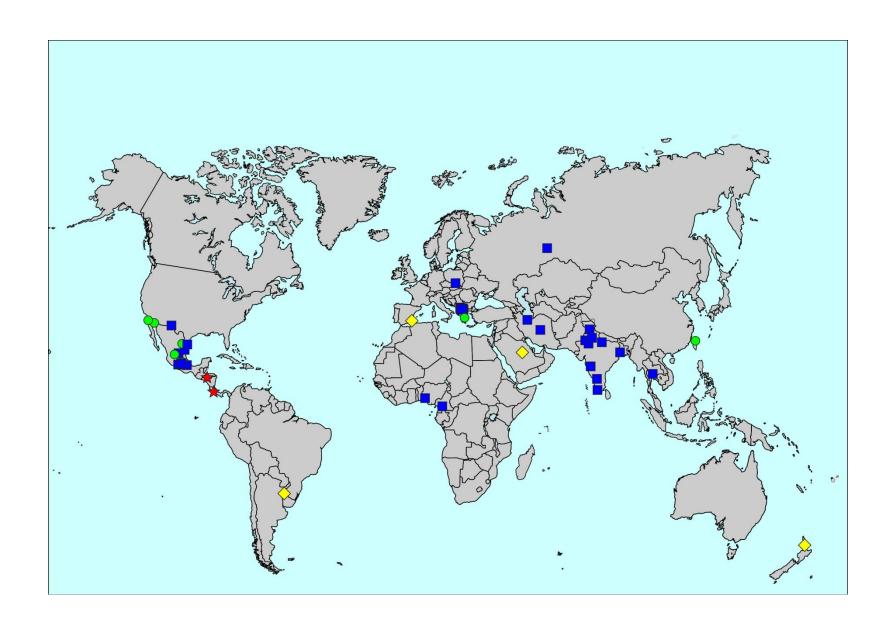
- 27. Chiang C-Y, Bissell K, Macé C, et al. Revisiting Asthma Drug Facility and management of asthma in resource limited settings: the way forward. Int J Tuberc Lung Dis 2022 (in press).
- 28. Almqvist C, Worm M, Leynaert B. Impact of gender on asthma in childhood and adolescence: a GA2LEN review. *Allergy* 2008; **63**(1): 47-57.
- 29. Odhiambo JA, Williams HC, Clayton TO, et al. Global variations in prevalence of eczema symptoms in children from ISAAC Phase Three. *J Allergy Clin Immunol* 2009; **124**(6): 1251-8.e23.
- 30. Ballardini N, Kull I, Söderhäll C, et al. Eczema severity in preadolescent children and its relation to sex, filaggrin mutations, asthma, rhinitis, aggravating factors and topical treatment: a report from the BAMSE birth cohort. *Br J Dermatol* 2013; **168**(3): 588-94.
- 31. Hill DA, Spergel JM. The atopic march: Critical evidence and clinical relevance. *Ann Allergy Asthma Immunol* 2018; **120**(2): 131-7.

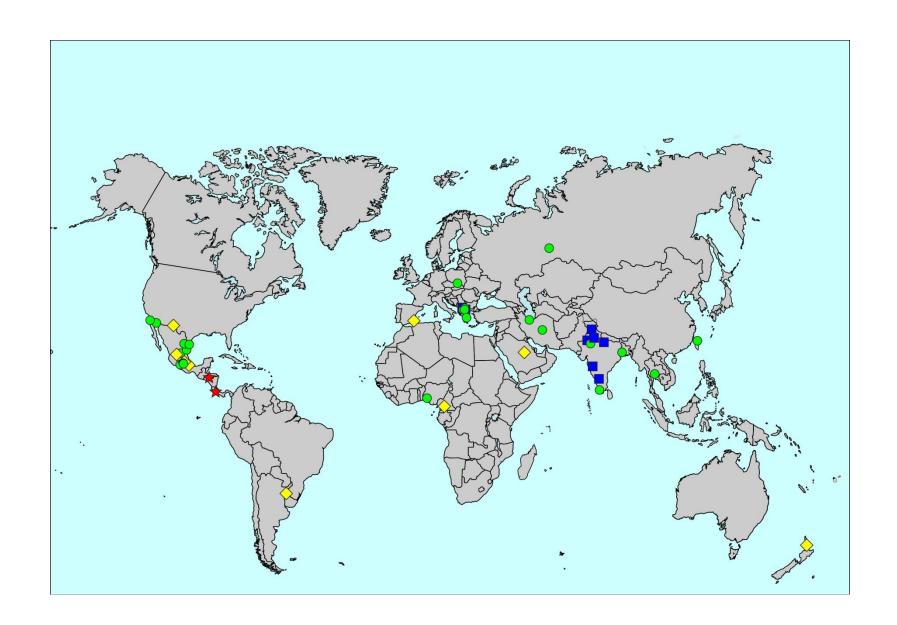
Table 1: Prevalence of asthma, hay fever and eczema in centres grouped by Gross National Income (GNI)

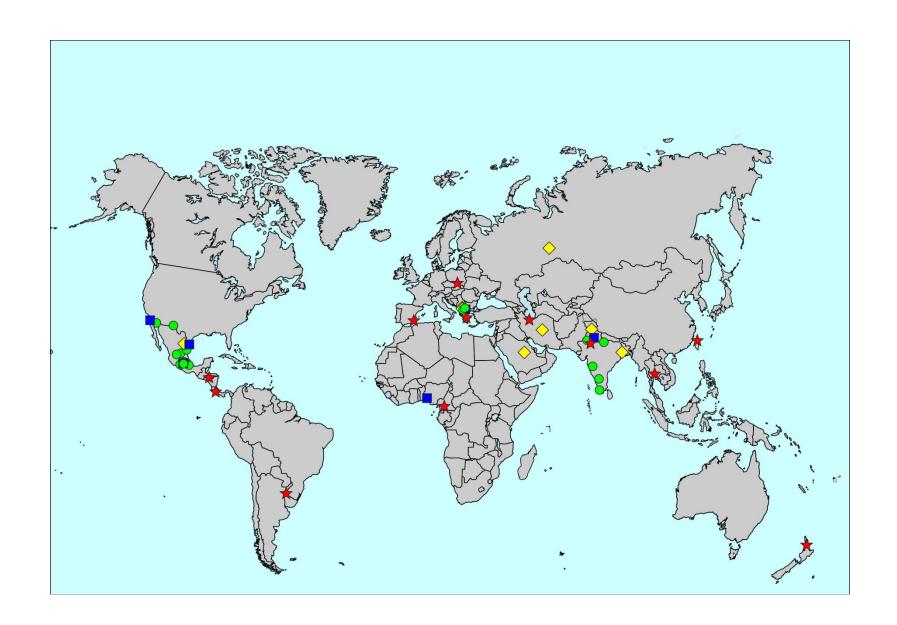
GNI	Years	Centres	No.	Current wheeze	Asthma ever	Severe asthma symptoms*	Severe asthma symptoms* (population denominator)	Eczema ever	Hay fever ever
High	2015-20	6	30556	3231 (10.6%)	3106 (10.2%)	1179 (36.5%)	1179 (3.9%)	5081 (16.6%)	9453 (30.9%)
Upper middle	2015-20	15	74897	6299 (8.4%)	3502 (4.7%)	2669 (42.4%)	2669 (3.6%)	5377 (7.2%)	9736 (13.0%)
Lower middle/Low	2017-19	12	88459	3208 (3.6%)	1926 (2.2%)	1161 (36.2%) 1161 (1.3%)		8791 (9.9%)	8695 (9.8%)
Total		33	193912	12738 (6.6%)	8534 (4.4%)	5009 (39.3%)	5009 (2.6%)	19249 (9.9%)	27884 (14.4%)

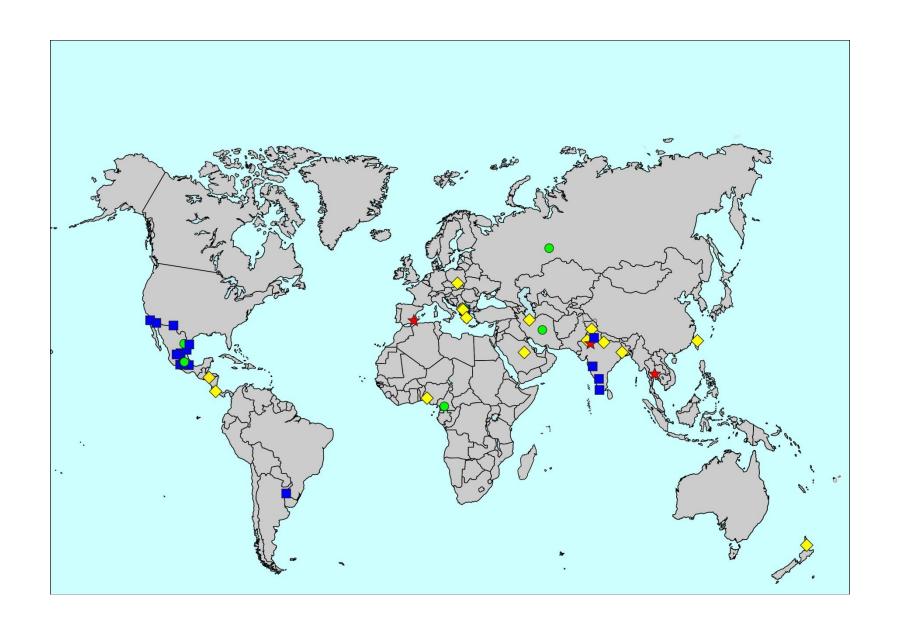
All values as number and (percentage), P-values for comparison between GNI categories: Current wheeze <0.0001, Asthma ever <0.0001, Severe asthma symptoms (population denominator) <0.0001, eczema ever <0.0001, hay fever ever <0.0001. Test by Chi-square.

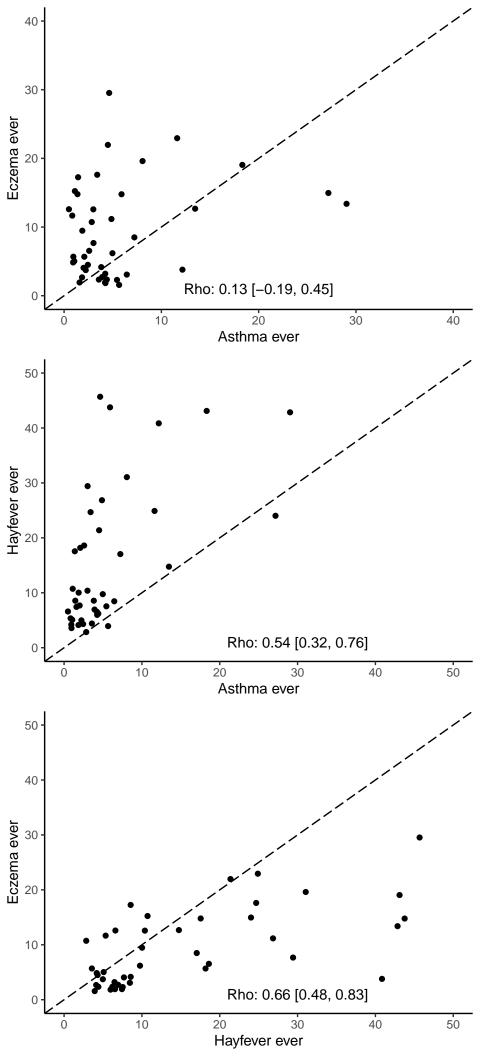
<sup>\*</sup> Current wheeze denominator; †Total participants denominator

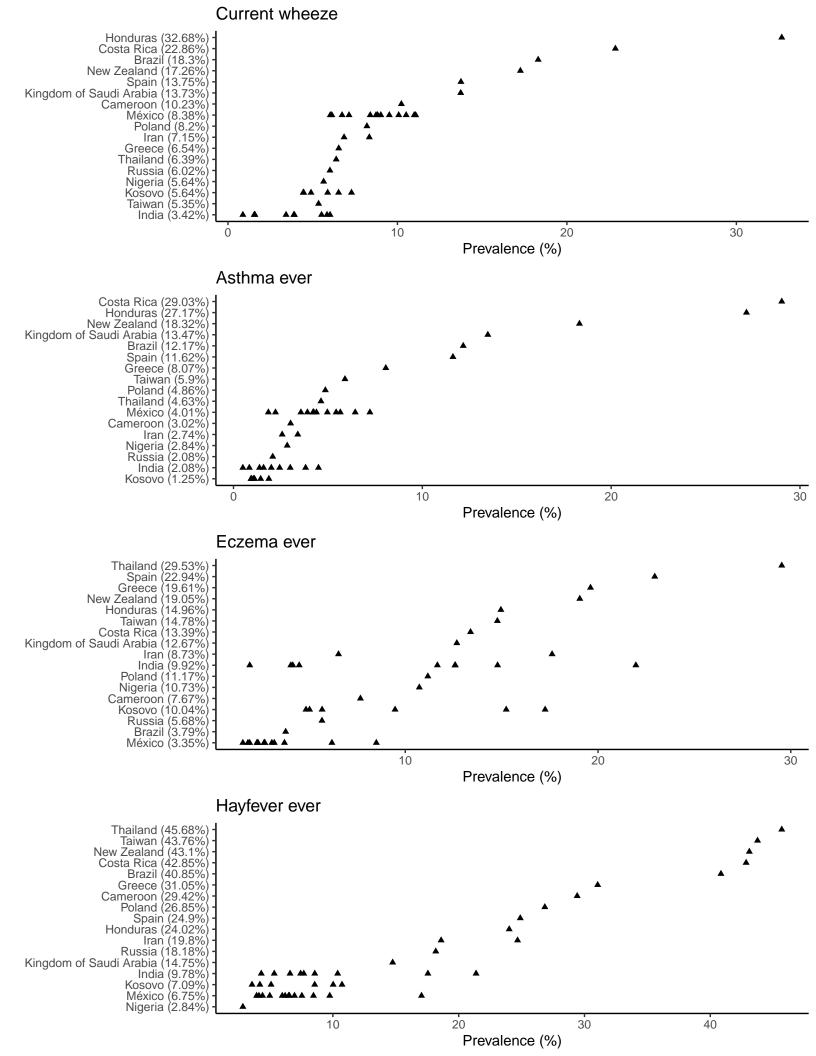


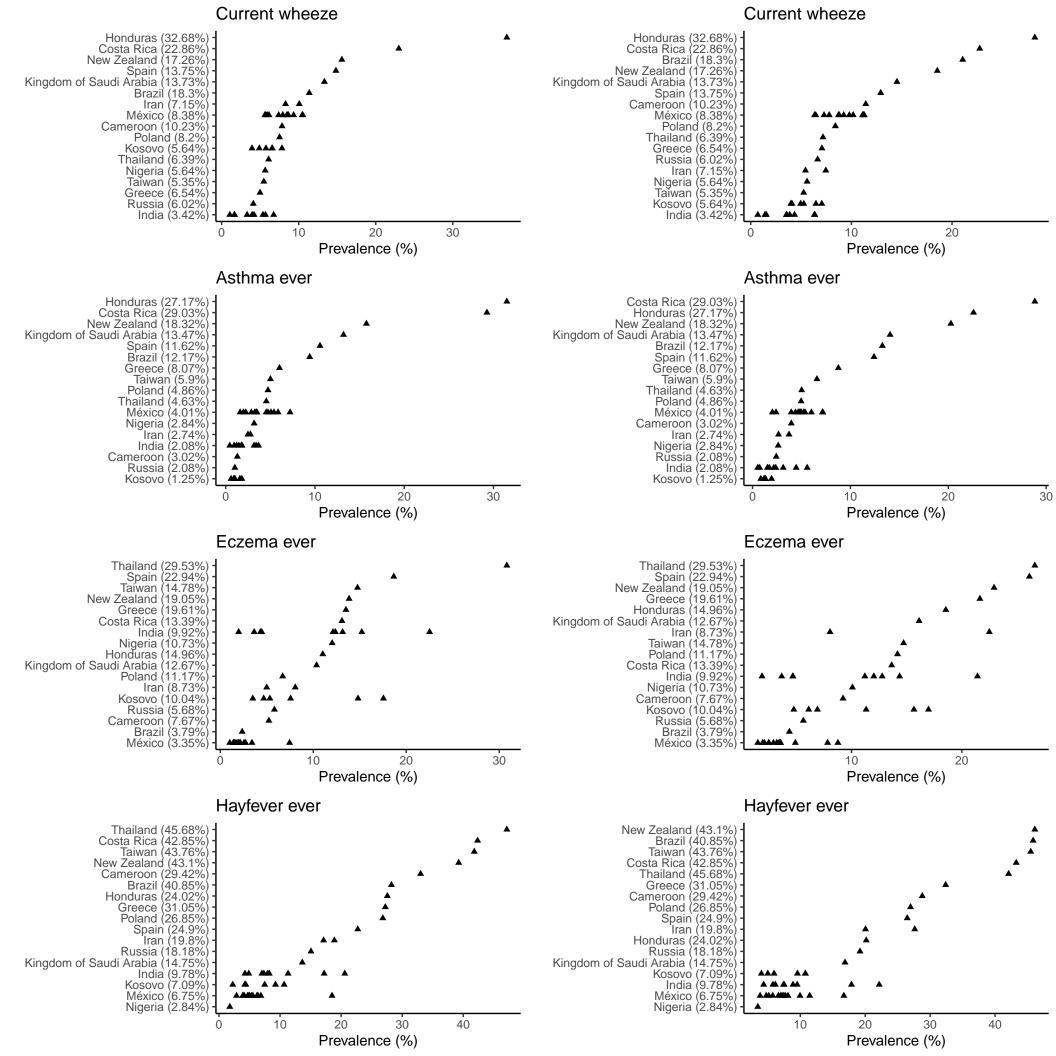


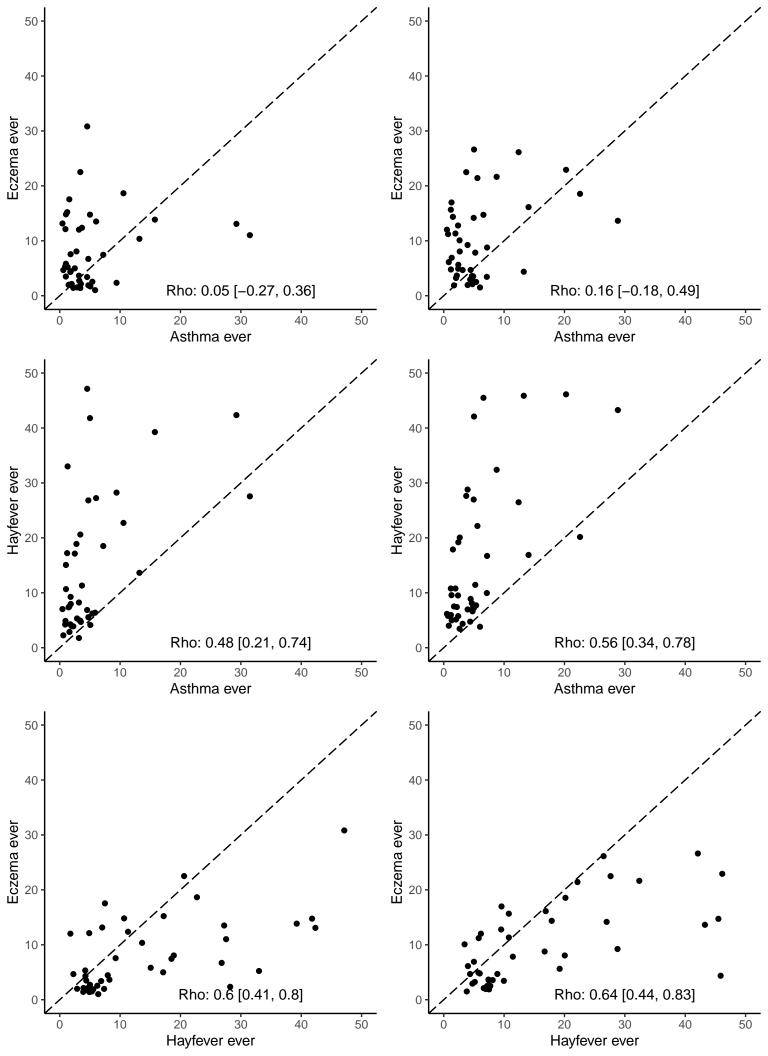


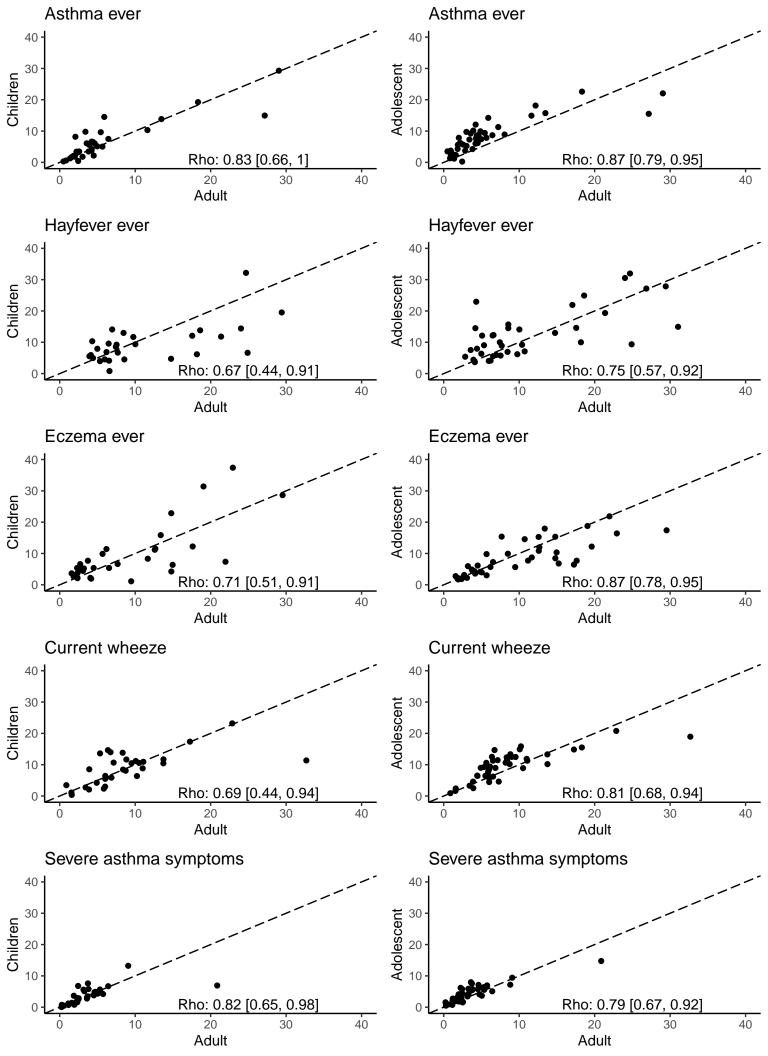












Web Table 1 : Demographic summary by centre.

Country	Centre	Proportion female	Mean age (years)	Standard deviation age (years)
Cameroon	Yaounde	0.53	36.2	8.1
Nigeria	Ibadan	0.62	43.5	9.9
Taiwan	Taipei	0.54	43.0	6.7
Thailand	Bangkok	0.26	42.4	6.8
Iran	Yazd	0.50	36.0	6.1
Iran	Karaj	0.66	39.9	6.7
Kingdom of Saudi Arabia	Kingdom of Saudi Arabia	0.40	38.5	12.2
India	Kottayam	0.50	40.9	7.1
India	New Delhi (7)	0.50	39.9	3.9
India	Chandigarh	0.50	37.0	5.5
India	Bikaner	0.50	35.5	6.7
India	Jaipur	0.50	36.5	6.3
India	Lucknow	0.50	36.5	6.3
India	Kolkata	0.52	41.3	6.7
India	Pune	0.50	35.9	6.5
India	Mysuru (Mysore)	0.50	38.2	6.4
Brazil	Uruguaiana	0.72	42.0	9.0
Costa Rica	Costa Rica	0.55	35.3	4.2
Honduras	Tegucigalpa	0.49	-	-
México	Monterrey	0.80	33.9	8.8
México	México City (North Area)	0.63	37.6	8.8
México	Mexicali	0.60	36.8	8.2
México	Ciudad Victoria	0.60	38.4	7.6
México	San Luis Potosí	0.58	38.0	8.1
México	Tijuana	0.62	34.4	8.1
México	Toluca Urban Area	0.58	34.9	7.1
México	Toluca Rural	0.55	35.8	7.6
México	Ciudad Juárez	0.62	37.2	8.5
México	Michoacan	0.67	34.2	7.5
México	Córdoba	0.60	38.2	8.5
México	Aguascalientes	0.86	37.4	7.7
México	Matamoros	0.62	34.7	7.9
Kosovo	Prishtina	0.50	42.9	6.6
Kosovo	Gjakova	0.50	43.4	6.6
Kosovo	Peja	0.51	39.7	6.9
Kosovo	Prizren	0.51	42.2	6.1
Kosovo	Gjilan	0.52	43.4	5.6
Kosovo	Ferizaj	0.50	42.2	6.3
Poland	Katowice	0.56	38.1	5.5
Russia	Tyumen	0.75	37.0	6.3
New Zealand	Auckland	0.58	39.6	8.3
Greece	Athens	0.74	46.3	5.3
Spain	Cartagena	0.57	42.7	6.4

Web Table 2: Symptom prevalence of asthma, hay fever and eczema by centre.

			Total	Current wheeze	Asthma ever	Severe asthma symptoms*	Severe asthma among population†	Eczema ever	Hay fever ever
Country	Centre	Mean date collection	N	N (%)	N (%)	N (%)	N (%)	N (%)	N (%)
Africa and E	astern Mediterranean	*							
Cameroon	Yaounde	2019-01	860	88 (10.2%)	26 (3%)	31 (35.2%)	31 (3.6%)	66 (7.7%)	253 (29.4%)
Nigeria	lbadan	2018-05	2321	131 (5.6%)	66 (2.8%)	52 (39.7%)	52 (2.2%)	249 (10.7%)	66 (2.8%)
Iran	Yazd	2020-06	4773	327 (6.9%)	123 (2.6%)	102 (31.2%)	102 (2.1%)	312 (6.5%)	888 (18.6%)
Iran	Karaj	2020-02	1175	98 (8.3%)	40 (3.4%)	37 (37.8%)	37 (3.1%)	207 (17.6%)	290 (24.7%)
Kingdom of Saudi Arabia	Kingdom of Saudi Arabia	2019-03	6786	932 (13.7%)	914 (13.5%)	364 (39.1%)	364 (5.4%)	860 (12.7%)	1001 (14.8%)
	Regional Total	2019-07	15915	1576 (9.9%)	1169 (7.3%)	586 (37.2%)	586 (3.7%)	1694 (10.6%)	2498 (15.7%)
South-East	Asia and Western Pac	ific							
India	Kottayam	2017-11	6940	418 (6%)	265 (3.8%)	174 (41.6%)	174 (2.5%)	289 (4.2%)	594 (8.6%)
India	New Delhi (7)	2017-12	9449	83 (0.9%)	231 (2.4%)	27 (32.5%)	27 (0.3%)	425 (4.5%)	407 (4.3%)
India	Chandigarh	2017-10	10386	407 (3.9%)	143 (1.4%)	118 (29%)	118 (1.1%)	1536 (14.8%)	1823 (17.6%)
India	Bikaner	2017-12	10495	167 (1.6%)	52 (0.5%)	25 (15%)	25 (0.2%)	1322 (12.6%)	692 (6.6%)
India	Jaipur	2017-11	8933	522 (5.8%)	402 (4.5%)	173 (33.1%)	173 (1.9%)	1962 (22%)	1910 (21.4%)
India	Lucknow	2017-10	11820	183 (1.5%)	99 (0.8%)	67 (36.6%)	67 (0.6%)	1379 (11.7%)	631 (5.3%)
India	Kolkata	2017-11	7823	432 (5.5%)	235 (3%)	168 (38.9%)	168 (2.1%)	984 (12.6%)	812 (10.4%)
India	Pune	2017-10	8000	311 (3.9%)	160 (2%)	93 (29.9%)	93 (1.2%)	325 (4.1%)	615 (7.7%)
India	Mysuru (Mysore)	2017-11	11178	383 (3.4%)	178 (1.6%)	180 (47%)	180 (1.6%)	216 (1.9%)	831 (7.4%)
New Zealand	Auckland	2018-08	3002	518 (17.3%)	550 (18.3%)	193 (37.3%)	193 (6.4%)	572 (19.1%)	1294 (43.1%)

Taiwan	Taipei	2017-10	9690	518 (5.3%)	572 (5.9%)	181 (34.9%)	181 (1.9%)	1432 (14.8%)	4240 (43.8%)
Thailand	Bangkok	2017-08	5418	346 (6.4%)	251 (4.6%)	132 (38.2%)	132 (2.4%)	1600 (29.5%)	2475 (45.7%)
	Regional Total	2017-11	103134	4288 (4.2%)	3138 (3%)	1531 (35.7%)	1531 (1.5%)	12042 (11.7%)	16324 (15.8%)
America									
Brazil	Uruguaiana	2017-12	896	164 (18.3%)	109 (12.2%)	79 (48.2%)	79 (8.8%)	34 (3.8%)	366 (40.8%)
Costa Rica	Costa Rica	2018-01	3272	748 (22.9%)	950 (29%)	297 (39.7%)	297 (9.1%)	438 (13.4%)	1402 (42.8%)
Honduras	Tegucigalpa	2018-06	254	83 (32.7%)	69 (27.2%)	53 (63.9%)	53 (20.9%)	38 (15%)	61 (24%)
México	Monterrey	2018-04	2118	191 (9%)	153 (7.2%)	80 (41.9%)	80 (3.8%)	180 (8.5%)	361 (17%)
México	México City (North Area)	2016-04	5231	550 (10.5%)	260 (5%)	264 (48%)	264 (5%)	324 (6.2%)	510 (9.7%)
México	Mexicali	2016-02	2436	164 (6.7%)	157 (6.4%)	91 (55.5%)	91 (3.7%)	75 (3.1%)	206 (8.5%)
México	Ciudad Victoria	2016-02	6239	551 (8.8%)	274 (4.4%)	238 (43.2%)	238 (3.8%)	147 (2.4%)	387 (6.2%)
México	San Luis Potosí	2016-04	2835	238 (8.4%)	111 (3.9%)	105 (44.1%)	105 (3.7%)	77 (2.7%)	197 (6.9%)
México	Tijuana	2015-12	1397	133 (9.5%)	79 (5.7%)	66 (49.6%)	66 (4.7%)	22 (1.6%)	55 (3.9%)
México	Toluca Urban Area	2016-02	6162	372 (6%)	137 (2.2%)	143 (38.4%)	143 (2.3%)	230 (3.7%)	307 (5%)
México	Toluca Rural	2016-01	7587	465 (6.1%)	140 (1.8%)	189 (40.6%)	189 (2.5%)	203 (2.7%)	313 (4.1%)
México	Ciudad Juárez	2016-02	2611	287 (11%)	111 (4.3%)	150 (52.3%)	150 (5.7%)	48 (1.8%)	156 (6%)
México	Michoacan	2016-08	2232	195 (8.7%)	94 (4.2%)	101 (51.8%)	101 (4.5%)	43 (1.9%)	146 (6.5%)
México	Córdoba	2016-04	2839	286 (10.1%)	120 (4.2%)	131 (45.8%)	131 (4.6%)	91 (3.2%)	184 (6.5%)
México	Aguascalientes	2016-04	2907	322 (11.1%)	158 (5.4%)	155 (48.1%)	155 (5.3%)	67 (2.3%)	219 (7.5%)
México	Matamoros	2016-03	1316	94 (7.1%)	47 (3.6%)	43 (45.7%)	43 (3.3%)	31 (2.4%)	58 (4.4%)
	Regional Total	2016-06	50332	4842 (9.6%)	2969 (5.9%)	2185 (45.1%)	2185 (4.3%)	2048 (4.1%)	4928 (9.8%)
Europe									
Kosovo	Prishtina	2017-08	2006	118 (5.9%)	21 (1%)	35 (29.7%)	35 (1.7%)	101 (5%)	102 (5.1%)
Kosovo	Gjakova	2018-06	1352	60 (4.4%)	15 (1.1%)	18 (30%)	18 (1.3%)	206 (15.2%)	145 (10.7%)
Kosovo	Peja	2017-06	1816	89 (4.9%)	34 (1.9%)	36 (40.4%)	36 (2%)	172 (9.5%)	182 (10%)

Kosovo	Prizren	2017-03	2712	177 (6.5%)	39 (1.4%)	65 (36.7%)	65 (2.4%)	468 (17.3%)	232 (8.6%)
Kosovo	Gjilan	2017-06	1835	82 (4.5%)	17 (0.9%)	22 (26.8%)	22 (1.2%)	89 (4.9%)	77 (4.2%)
Kosovo	Ferizaj	2017-09	1372	100 (7.3%)	13 (0.9%)	35 (35%)	35 (2.6%)	78 (5.7%)	49 (3.6%)
Poland	Katowice	2018-01	2220	182 (8.2%)	108 (4.9%)	73 (40.1%)	73 (3.3%)	248 (11.2%)	596 (26.8%)
Russia	Tyumen	2019-05	2360	142 (6%)	49 (2.1%)	55 (38.7%)	55 (2.3%)	134 (5.7%)	429 (18.2%)
Greece	Athens	2020-02	1897	124 (6.5%)	153 (8.1%)	38 (30.6%)	38 (2%)	372 (19.6%)	589 (31%)
Spain	Cartagena	2015-12	6961	957 (13.7%)	809 (11.6%)	330 (34.5%)	330 (4.7%)	1597 (22.9%)	1733 (24.9%)
	Regional Total	2017-07	24531	2031 (8.3%)	1258 (5.1%)	707 (34.8%)	707 (2.9%)	3465 (14.1%)	4134 (16.9%)

<sup>\*</sup> Proportion of severe asthma symptoms among those with current wheeze.

P-values comparing frequencies of symptoms within each region carried out using Chi-square test had p < 0.0001 for all comparisons with no adjustment for multiple testing.

<sup>†</sup> Proportion of severe asthma symptoms among all participants.