

Symptoms of asthma, bronchial responsiveness and atopy in immigrants and emigrants in Europe

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Symptoms of asthma, bronchial responsiveness and atopy in immigrants and emigrants in Europe. A. Tobias, J.B. Soriano, S. Chinn, J.M. Anto, J. Sunyer, P. Burney, for the European Community Respiratory Health Survey. ©ERS Journals Ltd 2001.

ABSTRACT: Migration studies on asthma may provide information on its environmental causes. The European Community Respiratory Health Survey has potential advantages due to the number of countries involved, standardized collection of information, assessment of directionality of migration, and availability of physiological data on bronchial responsiveness and atopy.

Prevalence rates of symptoms associated with asthma were compared for immigrants, emigrants and nonmigrants living in centres mostly in western Europe. Similar analyses were carried out for bronchial responsiveness (provocative concentration causing a 20% fall in forced expiratory volume in one second and slope) and atopy. Medication and use of health services were also explored.

Overall, 1,678 (8.6%) of 19,516 participants were immigrants in the 18 countries participating in the study, of whom 581 were emigrants from one of the participating countries. Rates of asthma symptoms were higher in immigrants (odds ratio (OR): 1.21, 95% confidence interval (CI): 1.00–1.51) and emigrants (OR: 1.31, 95% CI: 0.96–1.51) compared to nonmigrants after controlling for area, sex, age and smoking status. However, bronchial responsiveness and atopy were equally distributed between immigrants, emigrants and nonmigrants. Use of health services was observed to be similar in migrants and nonmigrants with asthma.

In the European Community Respiratory Health Survey, migrants reported more asthma symptoms, but had similar bronchial responsiveness, atopy, and use of health services when compared with the nonmigrant population.

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Migrants may have a greater susceptibility to asthma [1]. Variations in the prevalence of disease and increased rates associated with time, migration and changing lifestyle highlight the importance of environmental agents in the induction of the disease [2]. Compared with studies of cancer and cardiovascular disease, international migrant studies of asthma are scarce and very localized, tending to compare immigrants only with the host populations. Immigrants may suffer from psychological changes including somatization due to post-traumatic stress disorder and other anxiety disorders [3]. As with the unemployed [4], migrants might lack access to medical care, as well as being more susceptible to respiratory and other conditions.

The European Community Respiratory Health Survey (ECRHS) has previously provided information on the distribution of asthma and asthma-related phenotypes in the 20–44-yr-old population within

Europe and elsewhere [5–7]. To study the effects of migration on asthma, the ECRHS has potential advantages: data from 14 European and four other countries, standardized information including direction of migration, and physiological data on atopy and bronchial responsiveness.

By means of a cross-sectional analysis of the ECRHS, the authors have compared prevalence rates of asthma symptoms, bronchial responsiveness, atopy and use of health services by those with asthma in first-generation immigrants and emigrants, and nonmigrants.

Methods

Study design and sample

The study design, methodology and objectives of the ECRHS have been described elsewhere [5]. Briefly,

the ECRHS was conducted in 38 areas of 18 countries from 1991–1993. A random sample of 3,000 males and females aged 20–44 yrs from each of the participating areas was contacted and requested to complete a short screening questionnaire on respiratory symptoms. Subsequently, a 20% random sample ($n=19,516$) of the source population was asked to complete a long questionnaire, provide blood samples, undergo skin and forced spirometric tests, and have bronchial responsiveness measured through a methacholine challenge test [7].

Migration and social data

Information on country of birth and country of residence was self-reported by participants. Immigrants were defined as those subjects who were not born in their current country of residence. A subsample of them were also considered to be emigrants, as they were born in one of the participating countries. Others were considered to be nonmigrants. Social class was defined as manual, nonmanual and other (including unemployed and housewives) as presented elsewhere [8].

Countries were stratified into zones according to whether they had a high, medium or low prevalence of asthma among nonmigrants. The groups were as follows: high: Australia, France, New Zealand, UK, USA; Medium: Denmark, Ireland, Italy, Sweden, Switzerland; low: Belgium, Germany, Iceland, Netherlands, Norway, Spain. India and Estonia had virtually no migrants and were excluded from these analyses.

Symptoms questionnaire, respiratory function and immunological measurements

Any individual was considered symptomatic of asthma if he/she answered yes to any of the following questions of the long questionnaire: "Have you been woken by an attack of shortness of breath in the last 12 months?", "Have you had an attack of asthma during the last 12 months?", or "Are you currently taking medicines for asthma?". For the assessment of smoking status, participants were asked whether they had smoked at least one cigarette a day, one cigar a week for 1 yr or 360 g tobacco in a lifetime [9]. The subjects were categorized into three groups: non-smokers, exsmokers and current smokers.

Bronchial responsiveness was defined as a provocative concentration causing a 20% fall in forced expiratory volume in one second (FEV₁) (PD₂₀) from the largest postdiluent volume during methacholine challenge with an estimated cumulative dose of 5.1 μmol methacholine [7]. Because many participants will have censored data on bronchial responsiveness as measured by PD₂₀, analysis was also carried out by calculating the slope, or regression coefficient of percentage decline in FEV₁ with log dose [7].

Atopy was assessed by means of serum-specific immunoglobulin-E (IgE) [6]. A participant was considered as having a measurable specific serum IgE when he/she had a specific IgE $>0.35 \text{ kU}\cdot\text{L}^{-1}$ by the

Pharmacia CAP method (Pharmacia Diagnostics, Uppsala, Sweden) to any of the following allergens: *Dermatophagoides pteronyssinus*, timothy grass, cat, Cladosporium, or a local allergen. The local allergen was birch for northern Europe, Parietaria for southern Europe and ragweed for the USA, New Zealand and Australia.

Medication and use of health services

Use of respiratory medications during the last year was estimated for inhaled, oral and other medicines. Inhaled medicines included β_2 -agonists, nonspecific β -agonists, antimuscarinics, steroids, compound bronchodilators and other inhalers. Oral medicines included β_2 -agonists, nonspecific β -agonists, antimuscarinic drugs, methylxanthines, steroids, antihistamines, and bronchodilators. Finally, other respiratory medications included vaccines, injections, suppositories, and other remedies. Participants were also asked if medication was prescribed by a doctor, and the frequency of taking medication, namely every day or only for attacks of breathlessness. The use of health services for any respiratory problem, frequency of medication and the type of medical practitioner last contacted were recorded. All analyses on the use of health care services focused only on subjects reporting symptoms associated with asthma.

Statistical analysis

The statistical analysis was done using Stata (StataCorp., College Station, TX, USA) [10]. The Chi-squared test was used in the univariate analysis. Multinomial logistic regression [11] was applied to obtain adjusted odds ratios (OR) and their 95% confidence intervals (CI) by country of residence. Regression models were adjusted for sex, age (categorized into five groups, 20–24, 25–29, 30–34, 35–39 and 40–45 yrs), social class and smoking status. Random effects meta-analysis was used to pool the results over countries and test for heterogeneity [12].

Results

Migration rates

There were 1,678 immigrants in the present study, constituting 8.6% of the ECRHS study population. Overall, 581 (3.0%) were born in one of the participating countries, therefore being considered emigrants from these countries.

Migration rates varied widely between countries, the centres of the USA (27.9%), Switzerland (22.4%) and Australia (20.5%) had the highest rates of immigrants observed, while those of Italy (1.3%), Spain (2.0%) and Iceland (2.1%) had the lowest. India and Estonia had practically no immigrants, and therefore were excluded in the analyses. There were statistically significant differences in the rates of migration by age groups ($p<0.01$), the oldest group

(40–45 yrs) having the highest rate of migration (10.7%) and social class, with a higher percentage of manual class workers among immigrants than nonmigrants. There were no statistically significant differences in the rates of migration by sex or smoking status (table 1).

Symptoms in migrants

Higher prevalence rates of asthma symptoms were reported by immigrants (11.2%) and emigrants (11.0%) than nonmigrants (8.6%), both differences being statistically significant ($p < 0.01$). The corresponding ORs adjusted by sex, age, and smoking were 1.21 (95% CI: 1.00–1.51) in immigrants and 1.31 (95% CI: 0.96–1.78) in emigrants compared to nonmigrants (table 2). Risk estimates could be considered homogeneous between countries for immigrants (Chi-squared test for heterogeneity=12.22, degrees of

freedom (df)=14, $p=0.59$) and for emigrants (Chi-squared=11.79, df=10, $p=0.30$), being significantly different only for immigrants to Australia (OR: 2.18, 95% CI: 1.25–3.80) and for emigrants from Germany (OR: 2.08, 95% CI: 1.07–4.05) and New Zealand (OR: 6.10, 95% CI: 1.79–20.69).

Country of birth and country of residence

In table 3, an analysis is presented by the country of residence and the country of birth of the migrants, stratifying the countries by whether they had a high, medium or low prevalence of asthma. Despite small figures in some cells, it can be seen that no pattern is observed, and that migrants moving from a high to low country or a low to high country show no consistent pattern. Similarly, emigrants from South America, Asia or Africa yielded no pattern.

Table 1. – Description of the study population by migration status

	Nonmigrants	Immigrants	Emigrants
Country of residence*			
Australia	532	137	20
Belgium	1080	42	14
Denmark	372	22	9
Estonia	431	0	1
France	1866	259	29
Germany	1939	44	93
Iceland	550	12	3
India	592	1	17
Ireland	410	44	11
Italy	882	12	69
Netherlands	1193	54	24
New Zealand	1011	243	11
Norway	784	51	18
Spain	1902	40	31
Sweden	1650	206	14
Switzerland	662	191	11
UK	1461	118	173
USA	521	202	33
Sex			
M	8483 (48)	781 (47)	292 (50)
F	9355 (52)	897 (53)	289 (50)
Age yrs*			
20–25	3064 (17)	180 (11)	58 (10)
25–30	3517 (20)	247 (15)	85 (15)
30–35	3513 (20)	361 (22)	115 (20)
35–40	3487 (20)	379 (23)	125 (22)
40–45	4257 (24)	511 (30)	198 (34)
Smoking status [#]			
Non	7853 (44)	706 (45)	255 (44)
Current	6349 (36)	535 (34)	202 (35)
Ex	3571 (20)	326 (21)	121 (21)
Social class* [#]			
Nonmanual	8476 (61)	591 (61)	316 (66)
Manual	4195 (30)	317 (33)	145 (30)
Others	1174 (8)	54 (6)	17 (4)
Total	17838 (100)	1678 (100)	581 (100)

Data are presented as n and n (%). M: male; F: female.
[#]: Some figures do not add to totals due to nonresponse.
 *: $p < 0.05$.

Bronchial responsiveness and atopy

Similar prevalence rates of bronchial responsiveness (14.2% for immigrants, 15.0% for emigrants and 14.4% for nonmigrants) and atopy (40.6% for immigrants, 37.6% for emigrants and 38.8% for nonmigrants) were observed. Overall, migration did not modify the risk of either bronchial responsiveness or atopy (table 4). The OR of having bronchial responsiveness adjusted by sex, age and smoking was 0.90 (95% CI: 0.74–1.19) in immigrants and 1.03 (95% CI: 0.74–1.43) in emigrants compared to nonmigrants. Analysis of slope, instead of PD20, gave similar results. Similarly, the OR of having atopy adjusted by sex, age and smoking was 0.93 (95% CI: 0.79–1.11) in immigrants and 0.88 (95% CI: 0.67–1.14) in emigrants compared to nonmigrants.

Use of health services

Finally, the association between migration and use of medication in the last year, and use of health services, was explored in the individuals who reported symptoms associated with asthma (table 5). An approximately similar proportion of immigrants, emigrants and nonmigrants with asthma symptoms had taken inhaled and other medications, although symptomatic immigrants had an increased prevalence of oral medication use, compared to nonmigrants (OR: 1.41, 95% CI: 1.001–1.99). Immigrants and emigrants, considered separated or grouped, showed no consistent differences in the frequency of medication, the type of practitioner last contacted for breathing problems, or visits to hospital. Compared to nonmigrants, migrants were more likely to be taking oral medication, medicines every day, and medicines only for attacks of breathlessness. Access to health care, as indicated by prescriptions, visits to the doctor or admissions to hospitals yielded ORs close to unity.

Table 2. – Adjusted odds ratios (ORs), by area, sex, age and smoking status for asthma symptoms among immigrants and emigrants compared to nonmigrants (reference group), by country

Country	Asthma no:yes			OR (95% CI)	
	Nonmigrants	Immigrants	Emigrants	Immigrants	Emigrants
Australia	478:54	114:23	17:3	2.18 (1.25–3.80)*	1.59 (0.44–5.70)
Belgium	1004:73	37:5	14:0	1.86 (0.70–4.94)	nc
Denmark	338:34	21:1	9:0	0.54 (0.07–4.27)	nc
France	1660:206	226:33	24:5	1.33 (0.89–2.00)	1.77 (0.66–4.73)
Germany	1828:111	41:3	82:11	1.27 (0.38–4.17)	2.08 (1.07–4.05)*
Iceland	531:17	11:0	3:0	nc	nc
Ireland	371:39	26:2	10:1	0.71 (0.16–3.18)	1.10 (0.13–9.51)
Italy	800:82	10:2	63:6	2.14 (0.45–10.23)	0.85 (0.35–2.07)
Netherlands	1123:70	49:3	22:2	1.16 (0.35–3.87)	1.39 (0.31–6.20)
New Zealand	868:143	211:31	6:5	1.03 (0.63–1.69)	6.10 (1.79–20.69)*
Norway	736:48	46:5	16:2	1.71 (0.64–4.58)	1.99 (0.44–9.04)
Spain	1774:128	34:4	28:3	1.69 (0.59–4.87)	1.32 (0.39–4.45)
Sweden	1515:135	188:18	14:0	1.09 (0.65–1.83)	nc
Switzerland	606:56	168:23	11:0	1.56 (0.93–2.63)#	nc
UK	1278:183	79:7	159:19	0.63 (0.28–1.38)	0.87 (0.51–1.46)
USA	461:60	32:3	28:5	0.67 (0.20–2.28)	1.38 (0.50–3.77)
Total	15371:1439	1293:163	501:62	1.21 (1.00–1.51)*	1.31 (0.96–1.78)#

CI: confidence interval; nc: model did not achieve convergence due to lack of migrants. *: $p < 0.05$; #: $p < 0.1$.

Discussion

In this general population study of adults aged 20–45 yrs in Europe, USA, Australia and New Zealand, migrants (both immigrants and emigrants) had a slightly higher prevalence of symptoms associated with asthma than those subjects defined as nonmigrants. The higher prevalence rates in immigrants could not be attributed to other variables such as sex, smoking or social class. There was a wide variation in the use of medication and health services in the previous year, as previously reported in subjects with asthma-related symptoms [13, 14]. However, symptomatic individuals were treated and used health services evenly, irrespective of migration status. When country of birth and current country of residence were

considered together, there was a lower prevalence of asthma in migrants in those areas of medium prevalence than in those areas of high prevalence, however, there was no clear pattern overall.

Although this is compatible with the observations of STRACHAN *et al.* [15] and LEUNG *et al.* [16] that subjects who migrate tend to adopt the prevalence of the area to which they migrate, the fact that no pattern was observed on the rates of migration to/from countries where the asthma burden is very high (*i.e.* English speaking countries) or to/from countries where the asthma burden is lower, and that migration was not associated with higher rates of bronchial responsiveness or atopy, leads the authors to consider the present study as less conclusive.

A number of limitations must be considered, namely nonresponse, sampling, the low number of immigrants and the lack of extended questionnaire data. There was a substantial nonresponse rate in the ECRHS, and despite intense efforts, the participation rate was about 60% for the long questionnaire and 40–50% for respiratory and blood tests. However, this lack of response has not necessarily affected the validity of the results. It is unlikely that immigrants were selectively persuaded to participate in the study. Finally, because sampling strategy in each country was from official sources, only legal immigrants may have been included, although there is universal coverage of health benefits in most European countries for legal and illegal immigrants.

One major limiting factor was the low number of immigrants reported in most of the participating countries, leading to a lack of power in analyses particularly when adjusting by confounding variables. There were limited data on migration in the questionnaire, and more specific analyses regarding to when asthma symptoms developed in relation to migration could not be performed.

A potential bias may come from language problems

Table 3. – Prevalence of asthma symptoms in migrants grouping by prevalence of asthma zone in nonmigrants

	Migrant's zone of residence		
	High	Medium	Low
Overall			
Migrants	700 (12.8)	563 (11.0)	223 (7.3)
Nonmigrants	3525 (12.5)	5885 (9.1)	6748 (6.1)
Migrant's zone of birth			
High	199 (13.6)	102 (7.8)	28 (10.7)
Medium	48 (14.6)	39 (5.1)	43 (4.7)
Low	26 (15.4)	62 (6.5)	24 (12.5)
West Europe	16 (7.1)	52 (12.5)	31 (15.0)
South America	28 (6.2)	33 (11.5)	26 (9.7)
Asia	61 (17.9)	52 (12.1)	35 (0.0)
Africa	28 (16.4)	176 (13.5)	20 (5.7)
Not known	294 (8.5)	47 (22.6)	16 (0.0)

Data are presented as n (%). Prevalences of asthma; High: Australia, France, New Zealand, UK, USA; Medium: Denmark, Ireland, Italy, Sweden, Switzerland; Low: Belgium, Germany, Iceland, Netherlands, Norway, Spain.

Table 4. – Adjusted odds ratios (OR), by area, sex, age and smoking status for bronchial responsiveness and atopy in immigrants and emigrants compared to nonmigrants (reference group) by country

Country	Bronchial responsiveness		Atopy (serum IgE)	
	Immigrants	Emigrants	Immigrants	Emigrants
Australia	1.22 (0.72–2.06)	1.70 (0.50–5.83)	0.79 (0.51–1.20)	0.70 (0.23–2.15)
Belgium	3.83 (1.41–10.44)*	nc	0.22 (0.05–0.99)*	0.65 (0.13–3.31)
Denmark [#]	1.00 (0.19–5.16)	nc		
France	0.91 (0.60–1.37)	0.43 (0.10–1.92)	1.21 (0.95–1.75) [¶]	1.01 (0.35–2.89)
Germany	0.74 (0.25–2.13)	0.65 (0.30–1.40)	1.10 (0.54–2.24)	0.73 (0.42–1.27)
Iceland	nc	nc	0.34 (0.04–2.77)	6.08 (0.32–116.1)
Ireland	1.17 (0.31–4.40)	1.00 (0.11–8.24)	3.01 (1.04–8.70)*	1.63 (0.68–2.09)
Italy	2.20 (0.45–10.85)	2.56 (1.10–5.96)*	0.92 (0.19–4.55)	1.07 (0.54–2.10)
Netherlands	nc	3.12 (0.79–12.38)	0.62 (0.28–1.34)	0.73 (0.22–2.36)
New Zealand	0.90 (0.55–1.47)	1.14 (0.22–6.03)	0.82 (0.51–1.34)	1.19 (0.33–4.31)
Norway	0.53 (0.07–4.09)	1.02 (0.13–8.28)	0.94 (0.46–1.93)	1.93 (0.60–6.23)
Spain	0.61 (0.14–2.61)	0.86 (0.20–3.80)	1.69 (0.74–3.84)	0.65 (0.24–1.76)
Sweden	0.62 (0.32–1.22)	0.97 (0.12–8.05)	0.95 (0.66–1.36)	0.29 (0.04–2.36)
Switzerland	1.16 (0.61–2.20)	nc	0.95 (0.63–1.44)	0.88 (0.20–3.59)
UK	0.50 (0.19–1.29)	1.22 (0.76–1.98)	0.68 (0.38–1.23)	0.86 (0.57–1.29)
USA	1.19 (0.31–4.58)	0.34 (0.08–1.54)	4.05 (1.39–11.80)*	0.37 (0.12–1.14) [¶]
Total	0.90 (0.74–1.19)	1.03 (0.74–1.43)	0.93 (0.79–1.11)	0.88 (0.67–1.14)

Data are presented as OR (95% confidence interval (CI)). IgE: immunoglobulin-E; nc: model did not achieve convergence due to lack of migrants. [#]: no data for IgE. *: p<0.05; [¶]: p<0.1.

among immigrants, who might systematically report higher morbidity due to respiratory symptoms, similar to observations in some psychiatric surveys [17]. Indeed, misclassification of asthma for shortness of breath *via* anxiety and hyperventilation cannot be ruled out in any migration study of asthma. Within the present study, a report of shortness of breath was more frequent in migrants than nonmigrants both in a crude analysis (OR: 1.35, 95% CI: 1.10–1.65) and in an analysis adjusted by country of residence, sex, age and smoking status (OR: 1.32, 95% CI: 1.07–1.64) (data not shown). However, adjusted risks were not significantly increased in migrants for reported attack of asthma (OR: 1.09, 95% CI: 0.85–1.42) and for currently taking medicines for asthma (OR: 1.08,

95% CI: 0.81–1.44). The increased report of asthma in migrants is, therefore, unlikely to be due to a labelling or diagnostic bias. Language difficulties could lead to migrants having poor asthma education, and therefore poor compliance [18] and more symptoms. However, there is no evidence that the migrants in this study had less access to services (table 4).

A small number of published papers have assessed not only questionnaire data, but also other "harder" outcomes as in this study. Positivity to birch allergen by prick test and/or radioallergosorbent test in a survey performed in immigrants living in Sweden, was 16% in the persons who had been there ~2.5 yrs and increased to 53% in those who had been living there >10.5 yrs [19, 20]. In contrast, LEUNG *et al.* [16] noted

Table 5. – Medication and use of health services of immigrants and emigrants compared with nonmigrants in subjects reporting asthma symptoms

	No:yes			OR [#] (95% CI)	
	Nonmigrants	Immigrants	Emigrants	Immigrants	Emigrants
Medication in last 12 months					
Inhaled medicines	704:730	79:81	27:34	0.98 (0.70–1.38)	1.11 (0.66–1.88)
Oral medicines	960:470	94:62	38:22	1.41 (1.00–1.99)*	1.15 (0.66–1.98)
Others	1218:218	139:20	57:4	0.83 (0.50–1.36)	0.40 (0.14–1.11) [¶]
Frequency of medication					
Every day	1198:217	123:30	46:14	1.38 (0.90–2.12)	1.60 (0.86–2.97)
Only for attacks	796:617	81:73	31:29	1.22 (0.87–1.72)	1.17 (0.69–1.98)
Health services					
Prescription by a doctor	481:834	48:91	22:39	1.15 (0.79–1.69)	0.95 (0.55–1.64)
Ever visited hospital	1137:298	124:32	47:14	0.98 (0.64–1.49)	1.14 (0.62–2.11)
Admitted to a hospital	1246:189	141:15	51:10	0.72 (0.41–1.27)	1.23 (0.61–2.49)
Ever seen a doctor	437:996	42:113	12:48	1.28 (0.87–1.89)	1.66 (0.87–3.19)
General practitioner	753:680	75:80	26:34	1.24 (0.88–1.74)	1.41 (0.83–2.39)
Specialist	1204:229	131:24	50:10	0.97 (0.61–1.54)	0.99 (0.50–2.00)
Other	1360:73	150:5	58:2	0.66 (0.26–1.68)	0.69 (0.16–2.88)

OR: odds ratio; CI: confidence interval. [#]: adjusted for sex, age and smoking. *: p<0.05; [¶]: p<0.1.

that both Asians born in Asia and Asians born in Australia had a much higher prevalence of atopy than Australians who were not of Asian origin (78.4% and 81.0% *versus* 45.6%), mainly due to sensitivity to pollen and mites. NonAsian Australians also had the lowest prevalence of hay fever, but the prevalence increased markedly with time since migration. However, non-Asian Australians had a higher prevalence of wheeze and asthma than Asian Australians who had been born in Australia, who, in turn, had higher rates than those who had been born in Asia. Among the migrants, there was a tendency for rates to increase with time since migration, though this increase was not significant after adjustment for age at arrival, sex and atopic status. Similarly, immigration to the Tucson (AZ, USA) area was a major factor in determining changes in skin reactivity prevalence to five allergens, from 39.1% at the initial survey to 50.7% after 8 yrs of follow-up [21]. Recently, ORMEROD *et al.* [22] observed that the prevalence of asthma was higher in those Asians born in the UK and in those who had lived in the UK for longest, but these associations were explained by the older age of these subjects.

To conclude, despite migrants reporting more symptoms associated with asthma than nonmigrants in the European Community Respiratory Health Survey, this study has to be considered as inconclusive. No consistency was observed when migrants moved to and from countries with high or low asthma, and no differences were observed in the level of bronchial responsiveness or atopy. In a recent editorial, it was stated that, in a civilized society, it is important that all those in need of healthcare should have equal access to it and benefit equally from that which is available [23]. Very importantly, the access to healthcare by migrants with symptoms of asthma seems no different to the nonmigrant population.

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