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Migration and allergic diseases in a rural area of a developing country

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28 Capsule summary

- 29 Migration processes as the absence of the mother at home through temporary or permanent
- 30 migration could be an important determinant of the increase of allergic diseases in rural
- 31 areas of developing regions.
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- 33 Key words: allergic diseases, developing country, migration, rural area
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- 35 Word count: 975
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38 To the Editor:

Studies in Developing Countries (DCs) have frequently reported a lower prevalence of 39 allergic diseases (AllDis) in rural areas compared with urban settings, and this has been 40 attributed to the protective effects of environmental exposures such as rural lifestyle.[1] 41 42 Recent evidence from studies conducted in Africa and Asia showed that AllDis are increasing in urban and even in rural settings, reducing the urban-rural prevalence gap.[2,3] 43 It has been hypothesized that temporal increases in AllDis prevalence might be associated 44 45 with urbanization processes, especially with the change from rural to more modern urban 46 lifestyles.[1]

Migration is an important component of the urbanization process and involves 47 socioeconomic, environmental and lifestyle changes in rural and urban populations. 48 However, the effects of migration on AllDis in urban and rural settings of DCs have not been 49 50 explored.[4] The impact of migration on AllDis has been largely investigated by comparing populations that have migrated from DCs (presumed low-risk for AllDis) to developed 51 countries (presumed high-risk).[5] These studies have shown that being born in a country of 52 low risk provides protection against asthma, but this protection may decline with the length of 53 54 residence in the new environment.[5] Others studies have shown that age of migration and time since migration are associated with the risk of asthma and other AllDis, often leading to 55 a higher risk of atopy and allergy among migrants than the local population.[6] 56

The SCAALA (Social Changes, Asthma and Allergy in Latin America) study has been investigating the effects of migration on the prevalence of AllDis in schoolchildren living in rural and urban areas.[4] We studied 4295 rural and 2510 urban children aged 5-16 years attending a convenience sample of schools in Esmeraldas province, Ecuador. Data on potential risk factors, migration (direction and distance of migration, age at migration, and time since migration), and wheeze, rhinitis, eczema symptoms within the previous 12 months were collected using an investigator-administered questionnaire that included the core

allergy questions of ISAAC phase II.[4] Atopy was measured by skin prick testing to 7
 aeroallergens.

Results from the rural area showed that children who migrated during the first year of life had 66 a greater risk of wheeze and rhinitis compared to non-migrant children, and children with 67 history of international migration (children from rural areas of Colombia) had a higher 68 prevalence of rhinitis than non-migrant children (Table 1). The study also evaluated the 69 effects of maternal migration on allergic outcomes in children using the variables, maternal 70 history of migration and children living with one or no parent. These analyses suggested that 71 72 children whose mothers had a history of migration had a greater risk of eczema than children whose mother did not and children who did not live with any parent had more wheeze than 73 children living with both parents (Table 1). The magnitude of the latter association was 74 greater for all allergic symptoms among children of migrant mothers (Table 2). No 75 76 associations were observed for atopy (at least one positive allergen skin test).

77 The present study is unique in investigating migrants within a rural area of a DC, where migrants come from urban and rural settings. In this setting, age at migration and 78 international migration were important factors associated with a higher risk of AllDis in rural 79 populations. A novel observation was the effect on the prevalence of AllDis of migrant status 80 81 of the mother: children of migrant mothers not living with either parent had a two-fold greater risk of all 3 AllDis compared to children living with both parents. These data raise a question: 82 Could it be that social effects of migration, such as absence of parents at home, are 83 important determinants of the increase in AllDis in rural populations of DCs? In order to 84 85 answer this question, we need to consider some demographic patterns in these regions. It is well known that people in rural villages move to urban areas, temporally or permanently, in 86 search of work to improve their quality of life. A high proportion of these rural migrants are 87 single women who provide economic support for their families. Most of these women leave 88 their children in the community of origin to be cared for by relatives. Some of these 89 90 immigrants are able to settle in the city while others return to their rural communities.[7] In

91 the SCAALA rural population 31% of the children and 23% of the mothers had history of
92 migration, and 15% of the children lived with no parent.

93 If the absence of parents at home (especially the mother) is an important determinant of the increase of AllDis in DCs, then two migration trends that have occurred over recent decades 94 might help us understand temporal trends in AllDis. In the past, most economic migrants 95 were young men, but now "feminization of migration" is a growing trend worldwide because 96 of a greater demand for female labour.[8] Second, "circular migration" is a common 97 phenomenon in regions that are undergoing high levels of urbanization, and it refers to 98 99 repeated migrations between rural and urban areas due to improvements in transport and modern forms of communication.[9] 100

Migration affects not only the individual who migrates but also their family. Migration impacts 101 on roles, support structures, and responsibilities of family members resulting in changes in 102 103 social and psychological factors. In the case of maternal migration, children who remain in 104 their community may experience heightened levels of stress and depression due to separation from their primary carer. Psychological mechanisms have been proposed to 105 explain how emotional factors, in the context of family, might affect the development of 106 allergic diseases.[10] For this reason, we propose that the absence of the parents at home, 107 108 through temporary or permanent migration, may contribute to the increase of AllDis in rural and urban populations of DCs. 109

Finally, further analyses in different populations living in rural and urban areas evaluating the effects on migration on AllDis are required. A better understanding of the social, psychological and environmental effects of migration on AllDis in DCs is required.

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118 **References:**

von Hertzen L, Haahtela T. Disconnection of man and the soil: reason for the asthma and
 atopy epidemic? J Allergy Clin Immunol. 2006;117(2):334-344.
 doi:10.1016/j.jaci.2005.11.013.

Addo-Yobo EOD, Woodcock A, Allotey A, Baffoe-Bonnie B, Strachan D, Custovic A.
 Exercise-induced bronchospasm and atopy in Ghana: Two surveys ten years apart. PLoS
 Med. 2007;4(2):0355-0360. doi:10.1371/journal.pmed.0040070.

Selcuk ZT, Demir AU, Tabakoglu E, Caglar T. Prevalence of asthma and allergic
 diseases in primary school children in Edirne, Turkey, two surveys 10 years apart. Pediatr
 Allergy Immunol. 2010;21(4 Pt 2):e711-e717. doi:10.1111/j.1399-3038.2010.01008.x.

 4. Cooper PJ, Chico ME, Vaca MG, et al. Risk factors for asthma and allergy associated with urban migration: background and methodology of a cross-sectional study in Afro-Ecuadorian school children in Northeastern Ecuador (Esmeraldas-SCAALA Study). BMC Pulm Med. 2006;6:24. doi:10.1186/1471-2466-6-24.

5. Cabieses B, Uphoff E, Pinart M, Antó JM, Wright J. A Systematic Review on the
Development of Asthma and Allergic Diseases in Relation to International Immigration: The
Leading Role of the Environment Confirmed. PLoS One. 2014;9(8):e105347.
doi:10.1371/journal.pone.0105347.

- 6. Rottem M, Szyper-Kravitz M, Shoenfeld Y. Atopy and asthma in migrants. Int Arch Allergy
 Immunol. 2005;136:198-204. doi:10.1159/000083894.
- 138

7. Mujeres migrantes de América Latina y el Caribe: derechos humanos, mitos y duras realidades. Santiago de Chile: Comisión Económica para América Latina y el Caribe; 2005.
Available at: http://www.cepal.org/es/publicaciones/7200-mujeres-migrantes-de-americalatina-y-el-caribe-derechos-humanos-mitos-y-duras. Accessed February 22, 2015

143

8. Chammartin G. The feminization of international migration. Int Migr Program Int Labour
 Organ. 2002:37-40. <u>http://library.fes.de/pdf-files/gurn/00072.pdf</u>.

146 147 9 Beguly D Bocgi

Beguy D, Bocquier P, Zulu EM. Circular migration patterns and determinants in Nairobi
 slum settlements. Demogr Res. 2010;23:549-586. doi:10.4054/DemRes.2010.23.20.

10. Kaugars AS, Klinnert MD, Bender BG. Family influences on pediatric asthma. J Pediatr
 Psychol. 2004;29:475-491. doi:10.1093/jpepsy/jsh051

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Table 1. Odds ratios (OR) and 95% confidence intervals (95% Cl) for associations between migration variables and allergic symptoms adjusted for sex, age and socioeconomic status.

			Wheeze	Rhinitis	Eczema
Variables	Categories ^A	n	OR (95% CI)	OR (95% CI)	OR (95% CI)
Direction of	No Migrant	2964	1	1	1
migration	Rural to Rural	555	1.13 (0.84-1.52)	1.02 (0.7-1.49)	1.23(0.82-1.83)
	Urban to Rural	776	0.97 (0.74-1.27)	1.18 (0.86-1.61)	1.16 (0.81-1.66)
Distance	No Migrant	2964	1	1	1
of migration	National	1263	0.99 (0.79-1.25)	1.04 (0.79-1.38)	1.21(0.90-1.64)
	International	68	1.71 (0.88-3.32)	2.39(1.16-4.92)*	0.64(0.16-2.66)
Age at	No Migrant	2964	1	1	1
migration (years)	<1	269	1.47(1.02-2.12)*	1.59(1.03-2.46)*	1.25 (0.73-2.14)
	1-5	560	0.96 (0.71-1.31)	1.18 (0.83-1.69)	1.17 (0.78-1.75)
	>5	502	0.88 (0.62-1.24)	0.76 (0.48-1.19)	1.16 (0.75-1.79)
Time since	No Migrant	2964	1	1	1
migration (years)	<3 vs NM	383	0.98 (0.68-1.4)	0.94 (0.6-1.49)	0.96 (0.57-1.61)
(Jouro)	3-5 vs NM	197	0.56 (0.31-1.02)	0.9 (0.48-1.69)	1.53 (0.86-2.7)
	>5 vs NM	751	1.21 (0.94-1.58)	1.26 (0.92-1.73)	1.21 (0.85-1.73)
Maternal	No	3314	1	1	1
history of Migration	Yes	981	1.22 (0.96-1.53)	1.24 (0.93-1.65)	1.88(1.39-2.53)
Parents living	Both	2490	1	1	1
in the child's	One	1146	1.07 (0.84-1.36)	1.16 (0.87-1.54)	1.21 (0.88-1.67)
house	None	659	1.57 (1.2-2.05)*	1.29 (0.92-1.81)	1.27 (0.86-1.86

162 Outcomes were defined as: recent wheeze—reported wheezing during the previous 12 months; recent eczema—having a 163 reported itchy rash with a flexural distribution in the previous 12 months; and recent rhinitis—nasal stuffiness or sneezing 164 without a cold accompanied by itchy eyes in the previous 12 months. * p value < 0.05

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Table 2. Odds ratios (OR) and 95% confidence intervals (95% Cl) for associations
between allergic symptoms and parents living in the child's home (live with parents)
stratified by maternal history of migration. ORs adjusted for sex, age and
socioeconomic status.

Live with parents OR 95% Cl p value OR 95% Cl p value Wheeze One vs. both 1 0.76-1.34 0.976 1.2 0.77-1.87 0.429 None vs. Both 1.44 1.06-1.95 0.02 2.17 1.25-3.77 0.006 Rhinitis One vs. both 1.03 0.73-1.46 0.858 1.46 0.85-2.52 0.171 None vs. Both 1.1 0.74-1.64 0.627 2.07 1.05-4.08 0.036			Maternal history of migration NO YES						
Wheeze One vs. both 1 0.76-1.34 0.976 1.2 0.77-1.87 0.429 None vs. Both 1.44 1.06-1.95 0.02 2.17 1.25-3.77 0.006 Rhinitis One vs. both 1.03 0.73-1.46 0.858 1.46 0.85-2.52 0.171 None vs. Both 1.1 0.74-1.64 0.627 2.07 1.05-4.08 0.036 Eczema One vs. both 0.96 0.63-1.46 0.857 1.63 0.95-2.77 0.074				NU	TES	YES			
None vs. Both 1.44 1.06-1.95 0.02 2.17 1.25-3.77 0.006 Rhinitis One vs. both 1.03 0.73-1.46 0.858 1.46 0.85-2.52 0.171 None vs. Both 1.1 0.74-1.64 0.627 2.07 1.05-4.08 0.036 Eczema One vs. both 0.96 0.63-1.46 0.857 1.63 0.95-2.77 0.074		Live with parents	OR	95% CI	p value	OR	95% CI	p value	
Rhinitis One vs. both 1.03 0.73-1.46 0.858 1.46 0.85-2.52 0.171 None vs. Both 1.1 0.74-1.64 0.627 2.07 1.05-4.08 0.036 Eczema One vs. both 0.96 0.63-1.46 0.857 1.63 0.95-2.77 0.074	Wheeze	One vs. both	1	0.76-1.34	0.976	1.2	0.77-1.87	0.429	
None vs. Both 1.1 0.74-1.64 0.627 2.07 1.05-4.08 0.036 Eczema One vs. both 0.96 0.63-1.46 0.857 1.63 0.95-2.77 0.074		None vs. Both	1.44	1.06-1.95	0.02	2.17	1.25-3.77	0.006	
Eczema One vs. both 0.96 0.63-1.46 0.857 1.63 0.95-2.77 0.074	Rhinitis	One vs. both	1.03	0.73-1.46	0.858	1.46	0.85-2.52	0.171	
		None vs. Both	1.1	0.74-1.64	0.627	2.07	1.05-4.08	0.036	
None vs. Both 1.03 0.64-1.65 0.916 2.12 1.07-4.17 0.031	Eczema	One vs. both	0.96	0.63-1.46	0.857	1.63	0.95-2.77	0.074	
		None vs. Both	1.03	0.64-1.65	0.916	2.12	1.07-4.17	0.031	
				Y					