Aeroallergens, atopy and allergic rhinitis in the Middle East

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**Introduction**
The prevalence of allergic rhinitis (AR) and other allergic respiratory conditions is increasing worldwide and may be as high as 40% in children in some developed countries (1,2,3). Prevalences in developing countries are generally (but not always) lower, but are tending to rise (3). The disease burden in AR is considerable (4-7) and is associated with significant direct and indirect health and socio-economic costs (8). Furthermore, the link between AR and the subsequent development of asthma is well established (9,10). Although AR and allergic asthma are global and increasing serious health problems (3,11-14), most of the detailed epidemiological studies and clinical trials in this field have been performed in Western Europe and North America (15). In comparison, data on the Middle East region are scarce (16). The present work reviews literature data on (i) the nature and prevalence of indoor and/or outdoor aeroallergens, (ii) atopy (according to skin prick test (SPTs) and/or serum specific IgE [ssIgE] assays) in the general population or patient populations, (iii) the prevalence of AR (or self-reported symptoms compatible with a diagnosis of AR) and (iv) the management of AR in the Middle East region. For the purposes of the present work, the Middle East was defined as comprising Bahrain, Egypt, Iran, Iraq, Israel, Jordan, Kuwait, Lebanon, Oman, Palestine, Qatar, the Kingdom of Saudi Arabia (KSA), Syria, United Arab Emirates and Yemen.

**Materials and Methods**
The MEDLINE database was searched from January 1976 until May 2014 using logical combinations of the following terms (in English only): allergen*; allerg*; prevalence*; dust; mite*; pollen*; grass; airborne; indoor; outdoor; asthma*; allerg*; rhinit*; epide-
Results

Our search initially identified a total of 168 potentially relevant publications, of which 110 were analyzed in detail. The number of analyzed publications varied markedly from country to country. There were relatively high levels of documentation for Iran and KSA but relatively low levels of documentation for Bahrain, Iraq, Jordan, Oman, Qatar, Yemen and even the highly populous country of Egypt.

Direct measurements of indoor and outdoor aeroallergens in the Middle East

Direct measurements of pollen levels

Despite the stereotypical, desert image of the Middle East, levels of hundreds of different grass, weed and tree pollens have been characterized across the region. In the KSA, levels of pollen from the ornamental mesquite shrub Proopis juliflora peak in April and September, and a putative link with “urban greening” has been suggested (17,18). Rahal et al. (19) used a volumetric trap sampler to observe Cupressus, Pinus, Quercus, Paritatoria, Fraxinus, Urtica, Buxus, Brassica, Syringa, and Chenopodium pollen in Beirut (Lebanon) from March 2004 to August 2004, with peaks in March and April. Al-Qura’n (20) conducted the trap sampled in the city of Tafileh in the central highlands of Jordan. Most pollens (61.60% of the total) were from trees (mainly Pinus sp., Quercus sp., Cupressaceae, and Platanus sp) and herbaceous plants (Gramineae, Chenopodiaceae/Amaranthaceae, Helianthus sp. and Xanthium sp), with an overall peak in March to May. Al-Qura’n (21) confirmed the Pinus pollen peak was in March in northern Jordan.

Direct measurements of fungal spores

A wide range of fungal genera have been detected across the Middle East region. In the Kuwait city of Khalidiah, Moustafa and Kamel (1976) (22) identified fungal spores in the outdoor air, from 55 genera and 116 species (88 Hyphomycetes, 11 Ascomycetes, 9 Coccilomyces and 8 Zygomyces). The most prevalent genus was Alternaria (18.3% of all colonies), followed by Aspergillus (17.1%), Penicillium (14%) and Cladosporium (13.6%). Moustafa and Kamel suggested that many of the spores came from neighboring agricultural areas in Iran and Iraq. In an early study in the KSA, Sorenson et al. (23) detected 24 genera of fungi (mainly Aspergillus) in dust collected from ten homes of patients with perennial rhinitis. The researchers also performed SPTs on the 10 patients; 4 had positive SPTs for moulds. Similarly, Alternaria and Cladosporium spores have been measured in Jeddah (west coast), in Al-Khobar (east coast) and at two sites in Riyadh (central region) in the KSA (24,25). Cladosporium, Penicillium and Alternaria were also predominant in KSA and have been observed in central Iran (27), along with Candida spp., Geotrichum spp. and Trichosporon spp.

Direct measurements of house dust mites (HDMs) and animal dander

Frankland and El-Hefny (28) were the first to report on the prevalence of Dermatophagoides farinae (D far) in Egypt. Since then, many studies have confirmed the widespread presence of HDMs and other mites in domestic settings across Egypt and in the Middle East more widely. Even in a desert climate, domestic mites can grow and reproduce in homes and other buildings. For example, in a study of military hospitals in Egypt and in the homes of some of the nursing staff, Saleh et al. (29) reported the following prevalences: D far 73%, Dermatophagoides pteronyssinus (D pte) 55%, Ornithonyssus bacoti 18%, Acarus siro 36%, Tyrophagus putrecenter 55% and Laelaps nuttalli 15%. In Israel, Teplitsky et al. (30) found that whole D pte and D far were prevalent on the skin of 19 Israeli patients with atopic dermatitis and in dust from bedding and clothes in the patients’ homes. Sixteen (84.2%) of the patients had at least one positive skin sample. The presence of HDMs and/or high levels (> 2 µg/g house dust) of HDM allergens has also been confirmed in the great majority of dwellings and civic buildings in Saudi Arabia (23). Palestinian villages, cities and refugee camps (31,32), Kuwait (33), Iran (34) and Iraq (35).

Atopy rates in the Middle East

Data on atopy in patients with AR or in the general population (i.e. sensitization to allergens, as assessed by SPTs or ssIgE assays) were available for almost all the Middle East countries studied here. Large bodies of data were reported for Iran and the KSA, which are reviewed first. For the other countries, the level of detail varied greatly. Lastly, no publications on atopy in Oman, Syria and Yemen were found.

Atopy in Iran

In 2003, Kashef et al. (36) reported on the results of SPTs in 212 patients with chronic rhinitis (mean age: 18.2 years) attending a clinic in the city of Shirazin (southwest Iran). The
most frequent sensitizations were to weed pollens (69.6% of SPT-positive patients), followed by grass pollens (59%), tree pollen (51.5%), D. pte (15.5%), D. far (12.8%), mixed fungi (9.8%), Alternaria (3.7%), Aspergillus (2.2%) and Candida (0.7%). The low prevalence of sensitizations to mite and fungal allergens was ascribed to the dry climate. In the same year, Khazaee et al. (37) reported briefly on the results of SPTs in 1286 patients in the Gulf province of Sistan-Balouchestan; the highest sensitization rates were for HDMs (89.7%), feathers (70.3%), Aspergillus (65.1%), Alternaria (57.4%), Cladosporium (47.1%), grass pollen (43.4%), tree pollen (41.3%), Penicillium (39.2%), fruits (38.4%) and weeds (32.5%). In Farhoudi et al.’s (38) study of 226 patients at private clinic close to Tehran, 68% of the patients had at least one positive SPT. The most common aeroallergens were “herbaceae II” pollen mix (positive for 62% of the patients with at least one SPT), sycamore pollen (57%), Chenopodium pollen (53%), tree pollen mix (50%), with D. pte (19%) and D. far (18%) well behind. This profile was confirmed by Mohammadi et al. (39) for Tehran and Shiraz. Even higher rates were reported by Arshi et al. (40) in Tehran, where 97.1% of the studied patients had at least one positive SPT (pollens: 92.7%; mites: 64.9%; moulds: 50.2%). Arshi et al. (40) had suggested that the low prevalence of HDM sensitization in these cities was due to the dry climate. Surprisingly, low mite sensitization rates (D. far: 24.8%; D. pte 25.3%) were reported by Ghaffari et al. (41) for the humid city of Sari. The city of Mashhad in north-eastern Iran is characterized by a high polysensitization rate (81%) among patients, high sensitization rates for grass, weed and tree pollens (e.g. Cynodon dactylon: 56.8%; Poa pratensis: 59.5%; Amanthus palmeri: 83.3%; Fraxinus americana: 61.6%), moderate rates for mites (D. pte 25.2% and D. far 21.8%) (42,43) and high rate for Alternaria (53.4%) (44). Assarehzeadegan et al. (43) speculated that dust storms caused by climate change may influence aeroallergen exposure and sensitization. High sensitization rates for pollen and low to moderate rates for HDMs (despite the strong presence of mite allergens in dwellings (45)) were confirmed in the cities of Ahvaz (43), Yazd (46) and Qazvin (47).

Atopy in the KSA

A large body of data was recorded in the KSA, evidencing the importance of shrub pollen sensitization (attributed to irrigation and the increased local plantation of trees and flowers) and moderate levels of HDM sensitization. In a preliminary study, Sørensen et al. (23) performed SPTs on 10 patients with persistent AR, with four positive for mould extracts, three for HDMs; two for cat allergens, and three for cow dander, four for Bermuda grass pollen. A year later, the same researchers (48) prospectively screened 100 patients with perennial rhinitis (mean age: 28.5; age range: 16-57; females: 34%) in Al-Khobar. Overall, 54% of the patients had at least one positive SPT. The most frequent sensitizations were to Bermuda grass pollen (61% of the sensitized patients), goat hair/dander (39%), cat hair/dander (30%), D pte (28%) and Phragmites communis reed pollen (24%). In 1987, Al Nahdi et al. (49) performed SPTs on 210 allergy patients in the city of Dammam in eastern KSA. Fifty-five patients (26.2%) had at least one positive SPT and most were polysensitized. Surprisingly, the grass pollen SPT was positive in less than 5% of the participants. Al-Qorain and Al-Nahdi (50) reported on 129 AR patients (mostly aged between 20 and 39) in Al-Khobar. Skin prick tests revealed sensitizations to house dust extract (19.3% of patients), cat dander (16.3%), D pte (13.2%), Chenopodium album pollen (13.2%), flower pollen (13.2%) and dog dander (11.6%), amongst others. Zakzouk and Gad-El-Rab (51) studied 80 patients (age range: 10-50; females: 43.7%) with nasal symptoms consulting in the desert city of Riyadh: 66.2% had at least one positive SPT, with 50.9% for Bermuda grass, 41.5% for HDM mix and 39.6% for cat dander (despite the fact that pet-keeping was uncommon). Interestingly, 13.2% of the patients were sensitized to Iris germanica (orris root), used locally in toothpaste. Al-Annazy and Zakzouk (52) commented on the changing rural and urban environments during the 1980s/1990s (irrigation, street greening, imported flora, etc.) when studying SPTs in 68 children over the age of 4 years with a history of AR-compatible symptoms; 72% tested positive for at least one allergen. The most common positive SPTs were for cat fur (24 of the 49 positive patients: 49%), Prosopis juliflora pollen (51%), Bermuda grass (41%), and house dust mix (33%). Hasnain et al. (24) also screened 616 patients in Qassim, Abha, Gizan, Hofuf, Jeddah and Makkah with Alternaria extract SPTs: 21.6% of the patients had a positive SPT. In 2004, Hasnain et al. (25) reported on a similar study of Cladosporium levels at the same centres. Al-Frayh et al. (18) also reported that high proportions of allergic asthma AA sufferers were sensitized to pollen from Prosopis juliflora. In a specific radioallergosorbent test (RAST), 11.3% of 284 serum samples were positive. In the Qassim agricultural region, positive SPT rates were also high for Chenopodium album (79%), Sasola ten- nifolia (78.2%), Cynodon dactylon (69.5%) and Atriplex poly- carpa (71%). In a report from 2004, Hasnain et al. (25) also described the results of SPTs with a Cladosporium herbarum extract in 605 patients; 19.67% had a positive test. Koshak et al. (53) studied 41 patients with AR being treated in Jeddah. Thirty patients (73%) had one or more positive SPTs (D pte: 70% of positive patients; D far: 66.7%; cat dander: 33.3%; cockroach: 33.3%, Salsola pestifer pollen: 23.3%; Aspergillus: 20%). In a group of 151 asthma patients in Jeddah, Koshak (54) later found that 131 had at least one positive SPT to in-
door aeroallergens. Similar results were obtained by Al Saeed (55) in Riyadh.

In a rare multinational study (9 centres in the KSA, 1 centre in the UAE and 1 in Sudan), Hasnain et al. (56) investigated sensitization to indigenous pollens and moulds in a total of 492 consecutive patients (females: 56%; mean age: 30). The highest sensitization rates for pollens were observed for Chenopodium murale (52% of the patients in Khartoum, Sudan) and Salsola imbricata (30% in Riyadh, KSA) and Prosopis juliflora (24% in Riyadh). Sensitization to mites was also very prevalent in Khartoum: 72%; Abu Dhabi (United Arab Emirates): 46%; and Jeddah (KSA): 30%. This is surprising for Khartoum, which has a continental climate with low general humidity and dry, hot summers. However, Hasnain et al. suggested that cultural differences between the Sudan on one hand and Saudi Arabia and the UAE on the other may explain differences in sensitization.

Atopy in Egypt

Only one non-detailed publication dealt with atopy in Egypt. Attia and Mohamed (57) used in-house SPTs and two different IgE assays to screen 150 allergic patients for sensitization to a range of food allergens and aeroallergens between 2009 and 2012. The comparative diagnostic performances of the various tests were reported but sensitization rates for each allergen were not reported.

Atopy in Jordan

Only one publication on atopy in Jordan was identified. Aburuz et al. (58) evaluated SPT reactivity to aeroallergens in 538 over-18 AR patients attending hospital outpatient clinics (mean age: 34.4; females: 61.2%; mean ± SD time since the onset of AR: 6 ± 5 years). In all, 9.6% of the patients were not sensitized to any of the 18 tested allergens, 9.0% were monosensitized and 81.4% were polysensitized. The most frequent sensitizations >30% were for grass pollen mix: (51.4%), "thistleweed": (46.9%), olive tree pollen (45.3%), cat dander (41.6%), cereal pollen mix (40.6%), D pte (36.4%) and Betulaceae pollen (30.2%).

Atopy in Iraq

Data on atopy in Iraq were scarce, with just three publications. Al-Dulaimy and Alwan (35) tested 391 patients with AR and/or AA (age range: 4-63; AR only: 49.1%; females: 62%) attending an allergy clinic in the city of Baquba in 2009. The most common sensitizations were HDMs (25.3% of all patients tested), Bermuda grass (23%) and mixed grasses (19.6%). Saleh et al. (65) screened 62 patients with AR attending the allergic clinic at Tikrit Teaching Hospital (Tikrit, northern Iraq) with 18 allergen extracts. Three patients (5%) were non-sensitized, 8 (13%) were monosensitized and 51 (82%) were sensitized to two or more allergens. The most notable sensitizations were for Bermuda grass (66% of all patients), grass mix (28%), moulds (at least 34%), D pte (18%) and D far (9%). The city of Mosul (northern Iraq) was the setting for a study of allergy patients by Zakaria and Basima (66); the overall sensitization rate in intradermal tests was 44% for HDMs and 38% for grass pollen.

Atopy in Israel

There were only a few reports on atopy in populations in Israel. In 2002, Graif et al. (67) reported SPT data for 175
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asthmatics, 100 controls and 150 subjects with suspected asthma (all aged between 18 and 24) attending a defence force clinic. The proportions of participants with at least one positive SPT were 95.4% in asthmatics, 54% in controls and 69% in suspected asthmatics. Data on the other individual extracts were not provided. In Zeldin et al.’s report from 2008 (68), the SPT results showed that 98% of AR patients were sensitized to HDMs, with 43% for tree pollens, 34% for grass pollens, 23% for weed pollens, 25% for cat dander and 8% to dog dander. Sade et al. (69) performed SPTs with mite extracts on 117 patients (mean age: 21.3; age range: 16-44; females, 43.6%) attending an outpatient allergy clinic in Tel Aviv. Ninety-five (81%) of the patients had at least one positive mite extract SPT, with 57 (49%) sensitized to at least one tree, grass, or weed pollen. Sade et al. considered that the humid climate of much of Israel favoured the proliferation of domestic mites (69).

Atopy in Bahrain

In the only identified study in Bahrain, Tabbara et al. (70) used the ImmunoCAP assay to study atopy to aeroallergens and food allergens in 95 children with a clinical diagnosis of asthma (females: 35.8%; Bahraini nationals: 82.1%; mean ± SD age: 6.6 ± 3.8). Overall, 31.6% were not atopic, 28.4% were sensitized to common aeroallergens and food allergens, and 28.4% were sensitized to common aeroallergens alone, and 10.6% were sensitized to food allergens alone. Sensitization to aeroallergens was significantly associated with clinical AR (p = 0.004).

Atopy in Lebanon

In the only identified publication on this topic (Irani et al.’s (71) study of 200 patients with AR and/or AA), 24.2% of the participants were sensitized to HDMs, with values of 17.75% for grass pollen, 24.2% for Parietaria, 16.2% for olive pollen and 15.32% for Cupressaceae pollen. Around three-quarters of the participants were polysensitized.

Atopy in Palestine

In a study reported in 2003, El-Sharif et al. (72) screened children for atopy with SPTs and serum IgE assays. 19.5% of children with recent wheezing had at least one positive SPT. The most frequent SPT sensitizations in children with physician-diagnosed asthma were 12% for cockroach, 9.5% for a D far/D pte mix, 3.6% for dog dander, 2.4% for cat dander, 2.4% for olive tree pollen, and 1.2% for Alternaria tenuis. The sensitization rates in children with physician-diagnosed asthma were higher in sIgE assays, with 25.6% for mixed grass pollen, 23.5% for D pte, 23.2% for olive tree pollen, 18.4% for D far, 14.6% for dog dander, 9.4% for cockroach, 8.2% for cat dander and 5.3% for Alternaria tenuis.

Atopy in Qatar

Two publications were found. Sattar et al. (73) studied 1106 patients (mean age: 30; age range: 12-48; females: 54.9%) with clinical signs of allergy attending a clinic at a general hospital in the city of Doha. 51.4% had at least one positive SPT. The most common sensitizations were D pte (41.6% of the patients), D far (36.9%), cockroach (32.2%), Chenopodium (13.6%) and 12-grass mix (12.4%). Janahi et al. (74) studied 134 children (mean ± SD age: 4.5 ± 0.3; age range: 1 month to 10 years; females: 70.9%) attending an allergy clinic in Doha. In all, 19.4% of the children were sensitized to at least one aeroallergen, and the prevalence increased with age (7.7% in the youngest age group and 69.2% in the oldest age group). The most frequent sensitizations were to allergens in cat dander (16.4%), dust (14.9%), dog dander (11.9%) and ray grass (2.2%).

Atopy in the UAE

In the mid-1990s, Lestringant et al. (75) performed a retrospective study of SPT (n = 244 patients, all UAE nationals) and RAST (n = variable) results. 46% and 16.7% of the patients were diagnosed with AR and asthma, respectively. A battery of 15 local and panregional allergens were used for SPTs but RASTs for the same allergens were not performed for all patients. 71.8% of the patients were sensitized to at least one allergen (according to SPTs and RASTs). The most frequent sensitizations (according to the SPTs) were for Salsola kali pollen, (33.2%), Kochia scoparia pollen (25.8%), Suaedi moquini pollen (19.7%) and other pollens, whereas rates to HDMs and dander were below 15%. In a subsequent study in the same setting, Benet et al. (76) again identified Prosopis juliflora as the most common sensitizer (45.5%) in allergic patients, followed by grass mix (40.7%), cottonwood (33.1%), Bermuda grass (31.3%), Kochi (25.8%) and other pollens. The prevalence for HDM mix was 11.9%. Benet et al. attributed these rates to the ongoing “westernization” of the UAE.

The prevalence of allergic rhinitis in the Middle East

Much of the data on the prevalence of rhinitis and/or asthma in the Middle East has come from the International Study of Asthma and Allergies in Childhood (ISAAC) (3) or ISAAC-inspired studies of schoolchildren and (more rarely) university students. Although most studies had focused on asthma, the self-reported or parent-reported prevalence of current (12-month) rhinitis symptoms across the region ranged from 9% to 38% (table I, figure 1). The most extensive data again concerned the country of Iran.
The prevalence of allergic rhinitis in Egypt

Only one publication was found. Georgy et al. (77) used an Arabic translation of the ISAAC questionnaire to screen 2,645 11- to 15-year-old schoolchildren (females: 49.8%) in the capital city of Cairo. The “overall” prevalences of AR symptoms, allergic rhinoconjunctivitis symptoms and physician-diagnosed AR were 40.0%, 15.3% and 10.8%, respectively. The prevalences of 12-month wheeze and physician-diagnosed asthma were 14.7% and 9.4%, respectively. Georgy et al. pointed out that there is no everyday word in Arabic for “wheeze”, which may have biased the questionnaire results (77).

The prevalence of allergic rhinitis in Iran

Masjedi et al. (78) reported on the results of the ISAAC phase III study in Tehran (n = 3015 questionnaires). A 12-month history of wheezing concerned 8.6% of the 6-7 age group and 10.9% of the 13-14 age group (i.e. little change since the ISAAC phase I). In the cold-arid city of Urmia, Rad et al. (79) determined the
Figure 1 - Maps of the Middle East with per-country data on the 12-month prevalence of rhinitis and asthma symptoms in 6- to 7-year-olds and 13- to 14-year-olds in the ISAAC or in ISAAC-like studies (for values and references, see table I). a: the 12-month prevalence of rhinitis symptoms in 6- to 7-year-olds. b: the 12-month prevalence of rhinitis symptoms in 13- to 14-year-olds. c: the 12-month prevalence of asthma symptoms in 6- to 7-year-olds. d: the 12-month prevalence of asthma symptoms in 13- to 14-year-olds.
Figure 1 - Maps of the Middle East with per-country data on the 12-month prevalence of rhinitis and asthma symptoms in 6- to 7-year-olds and 13- to 14-year-olds in the ISAAC or in ISAAC-like studies (for values and references, see table I). a: the 12-month prevalence of rhinitis symptoms in 6- to 7-year-olds. b: the 12-month prevalence of rhinitis symptoms in 13- to 14-year-olds. c: the 12-month prevalence of asthma symptoms in 6- to 7-year-olds. d: the 12-month prevalence of asthma symptoms in 13- to 14-year-olds.
12-month prevalence of self-reported rhinitis (using the ISAAC written questionnaire) to be 23.6% (28.6% in boys and 18.7% in girls) and the 12-month prevalence of self-reported wheezing was 14.5% (20.7% in boys and 8.2% in girls). Similar values were reported by Morsi and Ghazi et al. (80) for 10- to 14-year-olds in Tehran. Rad and Hamzadeh (81) published the results of a study from 2999 six-to-seven-year-old children in the city of Urmia; the 12-month prevalence of self-reported rhinitis was 9.3% (12.0% in boys and 6.8% in girls) and the 12-month prevalence of self-reported wheezing was 9.4% (11.3% in boys and 7.6% in girls). The prevalence of self-reported AR was higher (22%) in the south-western city of Shiraz (82). Very few studies in the Middle East have applied the Allergic Rhinitis and its Impact on Asthma classification to local populations. Alyasin and Amin (83) studied 96 patients in Shiraz and found mild intermittent: 2%; moderate-severe intermittent: 4%; mild persistent: 24%; and moderate-severe persistent: 58%. In the desert city of Yazd, prevalence of ever AR in children exposed to pets (27.7%) was significantly higher than in non-exposed children (OR = 1.74; 95% CI = 1.36-2.23; p = 0.001) (84).

In one of the few studies to assess disease-specific QoL in the Middle East, Shariat et al. (85) used the English version of the RQLQ (completed by the physician) to assess 110 adult AR patients consulting in Tehran. 55% of the patients reported a major degradation of their QoL by AR. Furthermore, there was a significant relationship between high nasal congestion on one hand and low overall QoL and sleep quality on the other. Lastly, Amizadeh et al. (86) measured the Score for Allergic Rhinitis and general QoL (with the SF-36) in 1511 high school students in the city of Kerman (south-east Iran). The prevalence of self-reported AR (19.3%) was correlated with the “physical function” and “bodily pain” domains of the SF-36. The authors commented that differences with the literature data might have been due to cultural factors and differences in the patient profiles.

The prevalence of allergic rhinitis in Iraq

Alsamarai et al. (87) analyzed survey data for 16,736 heterogeneous subjects in five pooled studies. The overall prevalence of physician-diagnosed asthma was 10.2%. On the basis of total IgE levels and a rather low threshold of 100 IU/ml, Alsaimary (88) reported that 88.8% of the population in Basrah was likely to have an allergy.

The prevalence of allergic rhinitis in Israel

In 2000, Shohat et al. (89) used a modified ISAAC questionnaire to study the prevalence of asthma in 13-14-year-old children across Israel (n = 10057). The prevalence of wheezing in the last 12 months was 17.9%. Jewish population group was associated with wheezing in the last 12 months (odds ratio [95% CI] = 2.11 [1.79-2.48], p = 0.0001). In a subsequent report on the same dataset, Shohat et al. (90) suggested that genetic, nutritional and other environmental factors may account for the higher prevalence of asthma in Jewish schoolchildren. In addition to Jewish ethnicity, smoking among mothers was a major contributing factor to atopic diseases in children and adolescents included in a study by Kivity et al. (91). They surveyed a Jewish town (Zichron Yaakov) and a neighbouring Arab town (Paradis). Symptoms of asthma, AR and atopic dermatitis were reported in 11.2%, 19.9% and 5.5% of study population. Chronic rhinitis was reported in 9.7% of Arab children and 19.7% of Jewish children (p = 0.0043). Kivity et al. suggested that this difference was due in part to a lower proportion of smokers among the Arab children’s mothers (91). Using a computer-assisted telephone interview, Shahar and Lorber (92) questioned 1,141 Israeli Jews, 233 recent Jewish immigrants from the former USSR, and 263 Israeli Arabs). The overall self-reported prevalences were as follows: asthma: 14%; AR: 14%; “hay fever”: 11%; perennial AR and values were significantly higher in the Israeli Arabs. Romano-Zelekha et al. (93) reported that the prevalence of current AR symptoms was 9.4% in 1997 and 10.5% in 2003 (probably as a result of an increase in prevalence of Israeli Arabs in the population. The “ever” and current prevalences of AR symptoms were 41.6% and 9.4% in Graif et al.’s study (94) of 10,057 13- to 14-year-olds, and current asthma, a parental history of asthma and Jewish population group were found to be risk factors for AR.

The prevalence of allergic rhinitis in Jordan

Only one publication for Jordan (on asthma but not AR) was identified. In the ISAAC part II, Abu-Ekteish et al. (95) measured prevalences of physician-diagnosed asthma in 6- to 7-year-old schoolchildren (n = 4791) and 13- to 14-year-old schoolchildren (n = 4317) in Amman (the capital city) and Al-Mafraq (a rural Bedouin area of northern Jordan). The prevalence of physician-diagnosed asthma was 10.0% in Amman and 8.7% in Al-Mafraq in the 6- to 7-year-old age group and 8.8% in Amman and 9.5% in Al-Mafraq in the 13- to 14-year-old age group. Abu-Ekteish et al. concluded that asthma was not uncommon in Jordan (95).

The prevalence of allergic rhinitis in Kuwait

Behbehani et al. (96) studied 13- to 14-year-old children (n = 3,110) throughout Kuwait as part of the ISAAC. The prevalence rates (95% CI) for symptoms of AR ever, current symptoms of AR, and physician-diagnosed AR were 43.9%, 30.7% and 17.1%, respectively (i.e. higher than the values in the neighbouring KSA). The corresponding rates for ever, current and physician-diagnosed asthma were 25.9%, 16.4% and 16.8%,
respectively. In 2008, Owayed et al. (97) compared the ISAAC part 1 data (collected in 1995-1996) for 13- to 14-year-old children with those of a very similar study in 2001-2002. The prevalence rates for current symptoms of AR fell from 30.7% in ISAAC phase I to 27.6% in the 2001-2002 study (p = 0.008). However, the prevalence of physician-diagnosed AR increased from 17.1% in ISAAC phase I to 22.2% in the 2001-2002 study (p = 0.001). Behbehani et al. (98) observed two peaks per year (April-May and September-October) in the number of AR patients referred to an allergy clinic in Sulaiykh, these were strongly correlated with the pollen count. Abal et al. (99) reported on an ISAAC-like study of 2,117 5- to 7-year-old children from throughout Kuwait. Overall, 15% of the children had reportedly displayed symptoms of AR in the previous 12 months and 11% had been diagnosed with AR by a physician. Although Der p 1 was found in around 20% of dust samples, levels were always low (< 0.10 μg/g). Indeed, Abal et al. considered that the levels of Der p 1 were too low to induce allergic disease (99).

The prevalence of allergic rhinitis in Lebanon

Our literature search identified two studies. Waked and Salameh (100) carried out an ISAAC-like study of 5522 schoolchildren aged 5 to 14. They found that 24.5% and 19.5% of the children had symptoms suggestive of AR and asthma, respectively. Musharrafieh et al.'s 2009 study (101) of 3115 schoolchildren aged 13 or 14 revealed the prevalences of ever rhinitis and current rhinitis as 45.2% and 38.6%, respectively. The prevalence of wheezing in the previous 12 months was 24.1% and a value of 8.3% was reported for the prevalence of asthma (presumably physician-diagnosed asthma, since the prevalence of wheezing ever was 32.9%). Musharrafieh et al. concluded that the prevalences of allergic diseases are high in Lebanon and are approaching the rates recorded in developed countries (101).

The prevalence of allergic rhinitis in Oman

Only two studies were identified. Al-Riyami et al. (102) based their work on the ISAAC Phase I questionnaire (completed by 3174 children aged 13-14 and the parents of 3893 children aged 6-7). The 12-month prevalence of rhinitis symptoms was 15.0% in the 6-7 age group and 23.8% in the 13-14 age group. The “ever” prevalences of AR for the age groups were 22.0% and 34.6%, respectively (p = 0.03). The 12-month prevalence of wheeze was 7.1% in the 6-7 age group and 8.9% in the 13-14 age group. Al-Riyami et al. concluded that allergic conditions were common in Omani schoolchildren and were associated with significant morbidity. Al-Rawas et al. (103) then compared the ISAAC Phase I and Phase III results. There were few significant changes, although the prevalence of current (12-month) wheeze in the 6-7 age group rose from 7.1% in 1995 to 8.4% in 2001.

The prevalence of allergic rhinitis in Palestine

Our literature search identified several studies but most focused on asthma and not AR. Relatively low prevalences were reported. In 2000, Hasan et al. (104) surveyed 894 schoolchildren aged 6-7 and 13-14 in the city of Ramallah (n = 599) and the northern village of Anin (n = 295). The prevalences of “ever wheeze”, current wheeze and physician-diagnosed asthma in the 6-7 age group were 16.8%, 9.1% and 4.1%, respectively. The prevalences of “ever wheeze”, current wheeze and physician-diagnosed asthma in the 13-14 age group were 12.7%, 8.6% and 3.3%, respectively. El-Sharif et al. have carried out a series of studies of allergic disease in children in Palestinian villages, cities and refugee camps. The prevalences of “ever wheeze”, current wheeze and physician-diagnosed asthma were 17.1%, 8.8% and 9.4%, respectively (105). Children from refugee camps appeared to be at greater higher risk of asthma than children from neighbouring villages or cities. A subsequent study by the same group (106) confirmed that a family history of atopic disease and the indoor presence of cats and moulds are significant predictors of childhood asthma in Palestinian children. Prevalence rates for asthma and asthma symptoms were higher for younger children living in northern Gaza (vs. Ramallah) and higher for older children in Ramallah district (108). The only data on AR were reported by Ghazal Musmar et al. (108) for 1000 university students (females: 52%; age range 18-27) in the city of Nablus: “the percentage of patients who reported to have allergic rhinitis was 38.1%”.

The prevalence of allergic rhinitis in Qatar

A single study was identified. Bener and Janahi (109) used an ISAAC questionnaire to survey 3,283 Qatari schoolchildren (females: 47.7%; mean ± SD age: 9.02 ± 1.99; age range: 6-14) in 2003. The overall prevalence rate of AR (as reported by the parents) was 30.5%. The same results were reported by Janahi et al. (110).

The prevalence of allergic rhinitis in the KSA

Several large ISAAC and ISAAC-like studies have been performed in the KSA. Hijazi et al. (111) assess AR and asthma symptoms in 1020 children in the city of Jeddah and in 424 rural children 180-250 km from there. The 12-month prevalence of “itchy eyes/nose” was significantly higher in the urban area (13.9%) than in the rural area (8.0%) (p = 0.002), as was the prevalence of “diagnosed hay fever” (2.7% vs. 0.2%, respectively; p < 0.006). In the same study population, Hijazi et al. later
reported (112) that dietary factors during childhood (notably low vitamin E intake) were associated with the diseases measured in the ISAAC. Sobki and Zakzouk (113) used a modified ISAAC questionnaire to estimate the point prevalence of AR in 9540 children; 26.51% were considered to have rhinitis and of these, 25.66% of these had physician-diagnosed asthma. 61.8% of a subsample of 304 children were found to be atopic. In a study in Riyadh, Harfi et al. (114) concluded that the prevalence of AR was falling (relative to previous studies). The prevalence of AR and asthma were 12.7% and 11.4%, respectively, in 11000 children (815 boys and 285 girls) aged 6 to 14 (separate data for 6-7 and 13-14 age groups were not published). Furthermore, 2343 mothers and fathers also reported on their allergies, and the prevalences of self-reported AR and asthma were 12.6% and 3.9%. Other researchers have suggested that the low observed rates were due to a high proportion of males in the study population (116). Sabry (116) used a novel, Arabic-language questionnaire to estimate the prevalence of allergic diseases in Saudi students, employees and staff from the Taif University of Ta'if (Mecca Province): 52.81% had AR-like symptoms, 51.66% had mild AR and 48.34% had moderate/severe AR. 45.89% of the AR group reported evidence of pollen allergy. Nahhas et al. (117) observed quite high rates of rhinitis in 5188 children aged 6 to 8 children in the desert city of Madinah (also known as Medina). The prevalences of ever rhinitis, 12-month rhinitis and physician-diagnosed rhinitis were 24.2%, 18.2% and 4.2%, respectively. The study's results prompted Nahhas et al. to write that "children in Madinah have amongst the highest prevalence of allergic problems in the world" (117). Al-Ghobain et al. (115) also used an ISAAC questionnaire to evaluate 3073 high-school students in Riyadh (females: 51%; age range: 16 to 18). The "ever rhinitis" and "12-month" rhinitis prevalences were 43.8% and 38.6%, respectively.

The prevalence of allergic rhinitis in Syria

Syria participated in the ISAAC Phase III (study period: 2001-2003). The written and video ISAAC questionnaires and an additional questionnaire on environmental factors were used by Mohammad et al. (118) to survey children aged 6-7 (n = 5107) or aged 13-14 (n = 9068) in Aleppo, Lattakia and Tartous (see table I for the prevalences).

The prevalence of allergic rhinitis in the UAE

Al-Maskari et al. (119) used an ISAAC questionnaire to survey 3002 UAE schoolchildren (females: 41.3%) aged between 6 and 13. The prevalence of physician-diagnosed asthma was 13%. The reported prevalence of "hay fever" was 14.9% (13.5% in the 6-9 age group and 13.8% in the 9-13 age group). These values were similar to literature values for other Gulf countries. Al-sowaidi et al. (120) performed a cross-sectional survey of 6543 adolescent schoolchildren and their parents/carers (median age: 30; age range: 8-93) using an Arabic ISAAC questionnaire. The 12-month prevalences of current wheeze and current symptoms of AR were 4.8% and 27.4%. The prevalences of AR and asthma were much higher in UAE nationals than in non-nationals. John et al. (121) studied 255 undergraduate students (66.7% females; mean age 20 ±2.6; UAE nationals: 12.2%) at a university in Ajman, UAE: 45% self-reported current (12-month) AR and 27% self-reported current bronchial asthma. A family history of allergies (but not nationality) was associated with the presence of allergic conditions.

The management of allergic rhinitis in the Middle East

There are few reports on the diagnosis and management of AR in the Middle East. In the detailed, multinational Allergies in Middle East Survey (AIMES) survey by Abdulrahman et al. (16) (covering five countries: Egypt, Iran, Lebanon, the KSA and the UAE), 92% of the 501 participants reported that they had taken a medication to treat their AR in the previous 12 months. Overall, 84% had taken nasal steroids. Nasal steroid use was most common in Iran (98%) and least common in the UAE (51%). Despite these measures, only 40% of the respondents stated that their symptoms were under control. Eighty percent of the respondents had consulted a physician in the previous 12 months because of their AR. In terms of diagnosis, ENT specialists were the most frequently consulted physician (54% of respondents), followed by internal medicine specialists (15%), allergists and family physicians (both 12%). In a study of children over the age of 4 years with a history of AR-compatible symptoms in the KSA, Al-Anazy and Zakzouk (52) reported that 60.3% of their study cohort used topical corticosteroids. Many of the publications reviewed here reported moderate-to-high rates of physician-diagnosed AR and/or asthma. However, IgE sensitization was not always confirmed by an SPT or an ssIgE assay. This is exemplified by the results of the AIMES survey by Abdulrahman et al. (16), in which only 46% of the respondents (all of whom had physician-diagnosed AR) had undergone an SPT and/or an ssIgE assay. Similarly, Tabbara et al’s study in Bahrain (70) found that 31.6% of 95 patients with a clinical diagnosis of asthma were not atopic (as judged by the results of an ImmunoCAP assay).

According to clinical practice guidelines, AIT (available as subcutaneous and sublingual formulations, SLIT and SCIT) is treatment option for patients with IgE-linked AR (122). There are few publications on the use of AIT in the Middle East. In the multinational AIMES survey by Abdulrahman et al. (16), fully 42% of the respondents had received AIT of some sort (ranging from 57% in Lebanon to 19% in the KSA). Abdulrahman et al. commented that this relatively high proportion probably re-
lected that fact that physician-diagnosed AR was an inclusion criterion for the detailed survey (16). Separate data on SLIT and SCIT were not reported. Other publications on AIT came from Iraq and Israel (for SCIT) and Iran (for SCIT and SLIT). Farid et al. (123) reported on their experience of six years of SCIT in 156 patients in Iran (mean age: 36.5; females: 69.2%) with AR (n = 120) or AA (n = 29) or both (n = 7), diagnosed on the basis of a positive SPT and corresponding symptoms. The AIT regimen was based on weekly injections of “common aeroallergens and house dust mite extract” for ten weeks, fortnightly injections for the next ten weeks and then monthly injections. The patients were questioned on their symptoms before and after the course of AIT. The outcome measure was a “good” response to treatment (70% of AR patients), a “moderate” response (18.3%) and a lack of response (11.4%). In a study in Iraq, Alsamarai et al. (124) reported on the safety of SCIT in 693 patients with AR and/or AA (mean age: 33 ± 15; age range: 6 to 75; polysensitized: 90%). Over a nine-year period, a total of 39,281 injections were administered, including a range of allergen extracts. Overall, 11.8% of the patients experienced a systemic reaction (2.1 events per 1000 injections) although none were life-threatening. However, subcutaneous epinephrine was used at least once. Systemic reactions were most common in subjects receiving mould SCIT (82.9% of systemic reactions) and HDM SCIT (75.6%). The frequency of systemic reactions was correlated with the number of sensitizations. Efficacy was not reported on. The only double-blind, placebo-controlled, randomized, parallel-group study was that by Ahmadihashar et al. (125). Twenty-four young patients with AR were randomized to pre- and co-seasonal rye grass pollen SLIT (900 IR, three times a week) or placebo. Twenty patients completed the study. Although detailed numerical data and statistical analysis were not presented, the researchers reported significantly lower symptom scores in the SLIT group (vs. placebo) after 21 weeks of treatment and significantly lower medication scores after 15 weeks of treatment. Lastly, Zeldin et al. (68) described their experience of at least one year of SCIT with one or more aeroallergens (weeds, trees and grass pollen and/or HDMs) in 133 patients (mean age: 22.7) in Israel. The mean disease activity score (on a 10 cm visual analogue scale) decreased from 8.1 to 3.3 (for AR) and from 4.8 to 2.4 (for asthma) (p < 0.001 for both). In the 44 patients with both AR and AA, the mean asthma disease activity score fell from 4.8 to 2.4 (p < 0.001). The 14 immediate-onset systemic reactions (in 8 different patients) were all grade 1 or 2.

Discussion

A wide range of “global” and “local” aeroallergens is present in the Middle East. Although direct measurements were not available in all Middle East countries, a very wide range of indoor and outdoor aeroallergens have been observed across the region: (i) pollen from hundreds of different grasses, weeds, shrubs and trees, (ii) spores from hundreds of species of moulds, (iii) insect allergens, such as cockroach and mosquito, (iv) dander from domestic animals, rodents and farm animals, and (v) dozens of types of house dust mite and storage mites. “Local” allergens include pollen from the date palm and pollen from the initially foreign but now well-established shrub Prosopis juliflora. Although less work has been performed on absolute domestic levels of allergens, studies such as that by El-Sharif et al. (32) have reported geometric mean levels of HDM major allergens in dust that are as high as measured in many Western European countries. Many Middle East countries have regions with humid climates that favour the persistence of HDMs (69). Indeed, the broad variety of aeroallergens in the Middle East is related to the differing climates and indoor environments encountered across the region (126). The Middle East is stereotypically considered to have a dry, arid, desert-like, with hot summers (with mean maximum temperatures of up to 45°C), mild dry winters (with mean maximum temperatures of around 15-20°C) and almost no rainfall. While this is true of much of the land masses of Saudi Arabia, Egypt, Oman, Yemen, Syria and Iraq, these countries also have coastal regions that experience a subtropical or Mediterranean climate and/or mountainous regions that experience lower summer temperatures and colder, wetter winters. Summer humidity levels are also way above desert levels in the Gulf States (especially as Bahrain, an archipelago of 33 islands in the western Persian Gulf). Syria, Iran, Jordan and Yemen all have mountainous regions in which the climate is colder and wetter, with up between 750 and 1000 mm of rain per year. A country such as Oman has all three of the above-mentioned climates: a Mediterranean-like and quite humid climate in the north and east, a humid subtropical climate in the south (with a monsoon between June and September) and a desert climate in the central region. Some authors (43) have speculated that specific weather features (such as sandstorms) may influence aeroallergen exposure and sensitization. In the Middle East, levels of atopy in patients with known respiratory allergies (and, to a lesser extent, in the general population) appear to be just as high as in many European countries. This concerns “global” aeroallergens (such as HDMs) but also now-local plants (such as the shrub Prosopis juliflora (17,43,59,61,63) and the amaranth weed Salsola (42,43,64)). However, the very high polysensitization rates observed in some studies (42, 43) suggest abnormally low thresholds for wheal size or IgE levels. Researchers from the Middle East have commented on a number of factors that may have contributed to the rise in the prevalence of AR over recent decades: (i) irrigation for agriculture of previously desert zones; (ii) the ornamental "green-
ing” of desert cities with often imported plants (17,50,52), (iii) the increased use of air-conditioning in dwellings (61), meaning that mites are present in dwellings and public places even in hot, arid desert climates, and (iv), more generally, a progressive shift to many aspects of a western lifestyle (52,76). While rates of diagnosis and treatment appear to be high, there are few reports on patient satisfaction and potential unmet needs in AR sufferers. There is also a need for more data on diagnosis of respiratory allergies, the use of symptomatic medications and the potential role of a guidelines-recommended treatment option such as AIT (16,122,128-130).

Conclusions

A wide range of indoor and outdoor aeroallergens have been observed in the Middle East region: grass, weed, shrub and tree pollen, moulds, house dust and storage mites, insects and dander from domestic animals, rodents and farm animals. Levels of atopy and the prevalence of current, self-reported allergy-like symptoms in much of the Middle East are as just as high in some European countries and many developing countries worldwide. Although atopy in patients with AR is relatively well documented, there are few data on the management of AR and unmet treatment needs in the general population. Across the region, the prevalence of current 12-month self-reported or parent-reported symptoms of rhinitis ranged from 9% to 38%. Allergic rhinitis appears to be treated with conventional symptomatic medications and AIT. There is a need for research on the feasibility, safety and efficacy of AIT with “global” or local aeroallergens in the Middle East. Physician and patient education must play a central role in treatment decision making, particularly in the Middle East, to achieve higher patient satisfaction.

Conflict of interest

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References


Aeroallergens, atopy and allergic rhinitis in the Middle East


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