Epidemiological survey on pandemic influenza A (H1N1) virus infection in Kurdistan province, Islamic Republic of Iran, 2009

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مسح لوبائيات العدوى بفيروس الإنفلونزا الجائحة A (HINI) في ولاية كردستان، جمهورية إيران الإسلامية 2009 شهلا افراسيابيان، بهزاد محسن بور، كتايون حاجي باقري، محمد براري، إبراهيم قادري، روجين هاشمي، فردين غريبي

الخلاصة: تهدف هذه الدراسة لتقييم وبائيات الحالات المستبهة بالعدوى بفيروس الإنفلونزا الجائحة A (HINI) في عامي 2009 – 2010 في ولاية كردستان، وهي إحدى الولايات الحدودية في جمهورية إيران الإسلامية. وقد استوفى المرض الذين يزورون المراكز الصحية والمستشفيات في تلك الولاية استبياناً يغطي السيات الديموغرافية والمشهد السريري والنتائج وتاريخ التعرض والسفر. وتـم تحليل والمستشفيات في تلك الولاية استبياناً يغطي السيات الديموغرافية والمشهد السريري والنتائج وتاريخ التعرض والسفر. وتـم تحليل مسحات أنفية وبلعومية بطريقة النفاعل السلساي للبوليميراز – الوقت الحقيقي، وتـم تقييم 2059 حالة مشتبه بها، تأكد من بينها مسحات أنفية وبلعومية وبلعومية والنه من الحالات المؤكدة (20%) بين الأطفال دون عمر السنة، وأظهر التحليل المتعدد المتغيرات أن الخاطة المسبقة مع مرضى ظهرت لديمم أعراض الإنفلوانزا (نسبة الأرجحية 217) والإدخال في المعردات أن أخذي والازان المعامي والمائلة، وقامي والمعند المنعيرات أن المعام المائلة من بينها مسحات أنفية وماعومية بطريقة التفاعل السلساي للبوليميراز – الوقت الحقيقي، وتـم تقييم 2059 حالة مشتبه بها، تأكد من بينها معدحات أنفية ومائلة ورائدة (20%) بين الأطفال دون عمر السنة، وأظهر التحليل المتعدد المتغيرات أن المخالطة المسبقة مع مرضى ظهرت لديم أعراض الإنفلوانزا (نسبة الأرجحية 217) والإدخال في المستشفى (نسبة الأرجحية 217) والإدخال في المستشفى (نسبة الأرجحية 318) المحاطة المحاطة المسبقة مع مرضى ظهرت لديم أعراض الإنفلوانزا (نسبة الأرجحية 217) والإدخال في المائلة وأطبون المالاتان يُعْتَدُ بها إحصائياً للعدوى المؤكدة بفيروس الإنفلونزا A (HINI)، أما العوامل الأحرى مثل هما عامل الاختطار الوحيدان اللدان يُعْتَدُ بها إحصائياً للعدوى المؤكدة بفيروس الإنفلونزا A (HINI)، أما العوامل الأحرى مثل العمر والجنس والأعراض السروالا المائلة، علم أحمي والانفيزا A والالا) ولدي مالا ولاحرى مثل معمو والجنس والأعراض اللدان يُعْتَدُ من يكون قد المولي فلا يُعتَدُ من خلال المسائي، علم أرالا المال والذا A (HINI) قد التشر في جمهورية إيران الإسلامية ومن المحمل أن يكون قد انتقل من خلال المافرين إلى ولاية كردستان.

ABSTRACT This study evaluated the epidemiology of suspected cases of pandemic influenza A (H1N1) virus infection in 2009–2010 in Kurdistan province, a frontier province of the Islamic Republic of Iran. A questionnaire covering demographic characteristics, clinical presentation and outcome, and history of exposure and travel was completed by patients attending health centres and hospitals in the province. Nasal and throat swabs were analysed by RT-PCR. A total of 1059 suspected cases were assessed; H1N1 influenza A was confirmed in 157 (14.8%). The highest proportion of confirmed cases was 30.0%, among children aged <1 year. In multivariate analysis, previous contact with symptomatic influenza patients (OR = 2.17) and hospitalization (OR = 3.88) were the only significant risk factors for confirmed H1N1 infection. Age, sex, residency, presenting symptoms and history of national or international travel were not significant. Influenza A (H1N1) virus has spread in Islamic Republic of Iran; probably transmitted by travellers to Kurdistan.

Enquête épidémiologique sur l'infection par le virus de la grippe pandémique A(H1N1) dans la province du Kurdistan (République islamique d'Iran) en 2009

RÉSUMÉ La présente étude visait à évaluer l'épidémiologie des cas suspectés d'infection par le virus de la grippe pandémique A(H1N1) en 2009–2010 dans la province du Kurdistan, une province frontalière de la République islamique d'Iran. Un questionnaire couvrant les caractéristiques démographiques, la présentation clinique et les résultats ainsi que les antécédents d'exposition et de déplacements a été rempli par les patients consultant dans les centres de santé et les hôpitaux de la province. Des prélèvements de nez et de gorge ont été analysés par PCR en temps réel. Au total, 1059 cas suspects ont été évalués ; le virus de la grippe A(H1N1) a été confirmé dans 157 cas (14,8 %). Le pourcentage le plus élevé de cas confirmés était 30 % chez les enfants de moins d'un an. À l'analyse multivariée, un contact antérieur avec des patients porteurs de symptômes de la grippe (OR = 2,17) et une hospitalisation (OR = 3,88) étaient les seuls facteurs de risque importants pour une infection à H1N1 confirmée. L'âge, le sexe, le lieu de résidence, les symptômes initiaux ainsi que des antécédents de voyages dans le pays ou à l'étranger n'étaient pas des facteurs significatifs. Le virus de la grippe A(H1N1) s'est propagé en République islamique d'Iran, probablement transmis par des voyageurs qui se rendaient au Kurdistan.

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Introduction

The H1N1 influenza A virus is a relatively new virus of swine origin that can spread quickly and has the potential to cause a worldwide pandemic [1-3]. The virus was first detected in humans in Islamic Republic of Iran in 2009. During a short period the disease spread to several other countries [4] including neighbouring countries of the Islamic Republic of Iran [5].

The symptoms of pandemic influenza A (H1N1) are similar to seasonal influenza and include fever, sore throat, body pain, cough, rhinitis, headache, shivering and gastrointestinal complications. Although some symptoms are specific to the severity and regions of the infection [6], most patients show slight or self-limiting disease [4,7,8]. As most patients have mild symptoms, it is likely that just a small proportion of infected patients are registered [8] and therefore incidence estimates are not exact. About half of H1N1-infected patients have previous history of contact with a symptomatic influenza patient and one-third of them have a history of foreign travel [7]. In most cases of H1N1 infection, patients are younger than 30 years [7,9]. Unfortunately, routine seasonal vaccination is not effective against pandemic influenza A (H1N1) virus [10].

Two peaks of disease incidence occurred in the Islamic Republic of Iran during the period 1 June to 11 November 2009. Travelling abroad and the start of the educational year were the main explanations for the first and second peaks respectively [11]. Kurdistan province with a population of about 1.4 million, is located in western part of the Islamic Republic of Iran. This is a frontier province with 230 km common border with its neighbouring country of Iraq. There are some cross-border shopping markets in this province attracting many Iranian travellers. Therefore, monitoring the incidence of diseases is important in Kurdistan province. Furthermore, there is no efficient health

surveillance system in the Kurdistan region of Iraq. Therefore, evaluation of the characteristics of confirmed cases of H1N1 is of prime importance to adopt effective prevention and controlling strategies regarding H1N1. The present study aimed to evaluate the prevalence and time trend of pandemic influenza A (H1N1) virus in Kurdistan province of Islamic Republic of Iran to define the main factors influencing the spread of this pandemic virus. The confirmed and suspected cases of H1N1 virus infection and their characteristics were evaluated in this frontier province.

Methods

This was a follow-up survey conducted during April 2009 to March 2010 in Kurdistan province.

Sample

The research population consisted of all suspected cases of pandemic influenza A (H1N1) virus infection who had attended health-care centres and hospitals in Kurdistan province over the study period. A suspected case was defined as follows: fever above 38 °C, or having at least 2 respiratory symptoms including rhinorrhoea, sore throat or cough in addition to at least 1 of the following: (1) history of travel to an H1N1 epidemic area during the last 7 days; (2) history of contact with an H1N1 infected patient; (3) severe pulmonary disease or any unknown pulmonary disease causing death.

Data collection

According to the Iranian national H1N1 influenza surveillance, the suspected patients were reported to the national centre for disease control. Trained staff took nasal and throat swabs from all suspected patients and sent samples to the national reference laboratory at Tehran University of Medical Sciences, Islamic Republic of Iran. The samples were examined by real-time polymerase chain reaction (RT-PCR) as developed by the United States Centers for Disease Control and Prevention for the detection of pandemic H1N1 virus, as recommended by World Health Organization (WHO) [12]. A researcherdesigned questionnaire containing items related to demographic data (age, sex and residency), clinical symptoms, contact history and national and international travel history was completed by all subjects. Outcomes, including hospitalization and death, were noted for presenting patients.

Data analysis

The data were entered into SPSS software, version 11.5. The percentages, means, standard deviation (SD) and frequency distributions of variables were calculated using descriptive statistics. Univariate analyses were performed using the chi-squared and *t*-test to assess the difference between the confirmed and unconfirmed cases of H1N1. In the next step, odds ratio (OR) and its confidence intervals (CI) were calculated. Variables with P-values < 0.25 in univariate analysis were entered into a multivariate analysis using logistic regression to find predictable characteristics related to H1N-infected patients after adjustment for other variables.

Results

A total of 1059 people, registered as suspected cases of pandemic influenza A (H1N1) virus in Kurdistan province, were included in the study. Of the total group 558 people were female (52.7%) and 911 were urban residents (86.0%). Most cases of suspected H1N1 infection were detected in October (Figure 1).

The disease was confirmed in 157 cases (14.8%). The mean age of the confirmed and unconfirmed H1N1-infected patients was 21.8 (SD 15.2) years and 20.7 (SD 19.3) years respectively (P = 0.44). The ages of confirmed





Figure 1 Frequency of confirmed and unconfirmed cases of pandemic influenza A (H1N1) based on month of occurrence (bold numbers = H1N1 patients)

H1N1-infected patients ranged from < 1 to 80 years and of the unconfirmed group from < 1 year to 86 years. The highest percentage of H1N1-confirmed patients (30.0%) was observed among children aged < 1 year (Figure 2).

The mean time interval between the onset of symptoms and first health care attendance of the confirmed cases was 2.7 (SD 2.6) days and the median was 2 (ranged 0 to 15) days. A total of 158 people (14.9%) reported previous contact with an influenza symptomatic patient: 29.3% of the confirmed cases and 12.4% of the unconfined cases (P < 0.001). Three H1N1-infected patients had a history of travel abroad: 1 of them had visited Saudi Arabia and the other 2 did not mention the place. In the unconfirmed group, 11 persons had visited Saudi Arabia, 4 Dubai and 1 Iraq and 3 had visited other countries. Overall, 788 suspected cases (74.4%) were hospitalized: 91.7% of the confirmed cases and 71.4% of the unconfined cases (P < 0.001). No deaths were reported among the



Figure 2 Comparison of the percentage of confirmed and unconfirmed cases of pandemic influenza A (H1N1) by age group

Variable	Confirmed H1N1 casesUnconfirmed(n = 157)(n = 902)		med cases 902)	Unadjusted OR (95% CI)	P-value	
	No.	%	No.	%		
Sex						
Male	65	41.4	436	48.3	0.76 (0.54–1.07)	0.10.9
Female	92	58.6	466	51.7		0.100
Residency						
Urban	141	89.8	770	85.4	1.51 (0.87-2.62)	0.129
Rural	16	10.2	132	14.6		0.130
Domestic travel history						
Yes	1	0.6	24	2.7	0.24 (0.03-1.75)	0 122
No	156	99.4	878	97.3		0.125
International travel history						
Yes	3	1.9	27	3.0	0.63 (0.19–2.11)	0.451
No	154	98.1	875	97.0		0.451
Contact history						
Yes	46	29.3	112	12.4	2.92 (1.97-4.35)	< 0.001
No	111	70.7	790	87.6		< 0.001
Hospitalization						
Yes	144	91.7	644	71.4	4.44 (2.47-7.97)	< 0.001
No	13	8.3	258	28.6		< 0.001
Fever						
Yes	151	96.2	819	90.8	2.55 (1.09-5.95)	0.025
No	6	3.8	83	9.2		0.025
Body pain						
Yes	118	75.2	588	65.2	1.62 (1.10–2.38)	0.014
No	39	24.8	314	34.8		0.014
Cough						
Yes	140	89.2	754	83.6	1.62 (0.95-2.76)	0.075
No	17	10.8	148	16.4		0.075

Table 1 Comparison of confirmed and unconfirmed cases of pandemic influenza A (H1N1) virus infection with respect to the study variables: univariate analysis

OR = odds ratio; CI = confidence interval.

unconfirmed and confirmed H1N1infected cases (Table 1).

In univariate analysis, there was no significant difference between confirmed and unconfirmed H1N1 patients (P > 0.05) with respect to sex, urban/rural residency, Domestic and international travel history but we found significant differences between confirmed and unconfirmed H1N1 patients according to contact history, hospitalization, fever and body pain (P< 0.05). However, in multivariate analysis, previous contact with symptomatic influenza patients (OR = 2.17; 95% CI: 1.41–3.32) and getting hospitalized (OR = 3.88; 95% CI: 2.12-7.11) were the only significantly risk factors for confirmed H1N1-infection (P < 0.001) (Table 2).

Discussion

Our findings showed that both the frequency of previous contact with symptomatic influenza patients and the rate of hospitalization were higher in patients confirmed with pandemic influenza A (H1N1) virus infection compared with those not confirmed. As a history of international travel was not

a significant risk factor for confirmed disease, travelling to neighbouring Iraq presumably did not have an impact on the influenza epidemic in Kurdistan province of the Islamic Republic of Iran. The peak incidence of the disease occurred in September and October and, considering the age group of samples, these peaks can be attributed to transmission of the disease in universities and schools. Only 1 of the confirmed cases had a history of a domestic trip. Therefore, it seems likely that the disease was transmitted by travellers who came to Kurdistan. The findings of Gooya et al.'s study in the Islamic Republic of

Variable	β	Adjusted OR (95% CI)	<i>P</i> -value
Age	-0.002	0.99 (0.99–1.01)	0.657
Sex (male/female)	-0.216	0.81 (0.56–1.16)	0.240
Residency (rural/urban)	0.248	1.28 (0.73–2.26)	0.393
Domestic travel history (yes/no)	-0.605	0.55 (0.07-4.31)	0.566
Contact history (yes/no)	0.773	2.17 (1.41-3.32)	< 0.001
Fever (yes/no)	0.757	2.13 (0.89-5.09)	0.089
Body pain (yes/no)	0.271	1.31 (0.85–2.03)	0.224
Cough (yes/no)	0.080	1.08 (0.62–1.90)	0.781
Hospitalization (yes/no)	1.357	3.88 (2.12–7.11)	< 0.001

Table 2 Relationship between confirmed cases of pandemic influenza A (H1N1) virus infection and the study variable	es:
multivariate logistic regression analysis	

OR = odds ratio; *CI* = confidence interval.

Iran showed a high incidence of pandemic influenza A (H1N1) virus in neighbouring provinces to Kurdistan [11]. In their study, travelling abroad had a significant impact on the occurrence of the first peak of pandemic H1N1 virus infection in the Islamic Republic of Iran, and reopening the universities and schools at the middle of September caused the second peak. Letting students stay at home and closing schools and classes in regions with high numbers of suspected cases of influenza were effective strategies in controlling the disease. These strategies were recommended by the Iranian centre for disease control. This 2-peak pandemic trend was observed in the other provinces as well [11].

The mean age of the confirmed cases in our study was 21.8 years and this was similar to other studies [3,5,6,13-16]. In our study there was just 1 person aged > 65 years. The lower level of infection and susceptibility to H1N1 virus in this age group compared with younger age groups can be explained by pre-existing immunity, fewer contacts with other people and lower chances of being in crowded places. On the other hand, in Islamic countries elderly people are more likely to make religious pilgrimages, for example to Saudi Arabia, and this puts them at risk of contagious diseases. Therefore, this age group needs constant monitoring.

Cough and fever were the most common symptoms of the disease. However, the frequency of symptoms varies widely in different countries [3,5,6,13-19]. Different studies showed that most cases of infected H1N1 were mild, and severe cases of the disease were rare [4,5,8,20]. It is probable that in Kurdistan some patients with mild symptoms did not attend medical clinics and/or health staff did not consider some patients as suspected cases of H1N1 virus influenza and only assessed those with more severe symptoms. Therefore, it is necessary to perform studies assessing the level of awareness, knowledge and practice about H1N1 virus infection among health-care workers.

In our study the percentage of hospitalized patients was higher than other similar studies [3,7,14,21]. Unnecessary hospitalization may squander financial and human resources in health systems, and physicians therefore need proper training about H1N1 influenza A detection in order to avoid unnecessary referrals. As the rate of hospitalization in the confirmed H1N1 group was 91.7% compared with 71.4% for the unconfirmed group, and as no deaths occurred in the hospitalized patients, it can be concluded that hospital filing and caring systems in our hospitals are effective and satisfactory. The reported

death rate from pandemic influenza A (H1N1) virus is low in many countries, ranging from 0% to 2.2% of cases [4,5,9–11,14,22]. In the present study none of the patients died. We used the WHO guidelines for treatment of patients and this may be responsible for the low virulence level of the virus.

About 30% of the province's population live in rural areas, and an active surveillance system is implemented in these rural areas. Therefore, we expected more patients to be have been screened in these areas. In our present study, most of the H1N1infected patients were urban residents (89.8%), which can be attributed to the more frequent contacts and being in more crowded places. Furthermore, few travellers went to rural regions. Therefore, more training should be provided for urban schools and public places as well as for those villagers travelling to the cities.

There was no sex difference in the incidence of confirmed pandemic influenza A (H1N1) in our study, although other studies have reported sex differences [13,15,18,20] and this has been explained by the higher level of social involvement of men than women [14].

We did not study the incubation period of the virus, but only noted that the mean time interval between onset of symptoms and first health care attendance of all participants was 2.7 (SD 2.6) days. In other studies the mean delay ranged from 1.8–3 days [3,5,13,14,16,18,20].

In conclusion, it seems that influenza A (H1N1) virus has spread in the Islamic Republic of Iran and

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was probably transmitted by travellers who came to Kurdistan. Older people are less likely to become infected. We also conclude that caring and reporting systems related to influenza A surveillance in the province hospitals are well implemented.

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- CDC protocol of realtime RT-PCR for swine influenza A (H1N1). April 28 2009. Revision 1 (30 April 2009). Geneva, World Health Organization, 2009 (http://www.who.int/csr/resources/publications/swineflu/CDCrealtimeRTPCRprotocol_20090428.pdf, accessed 30 July 2013).
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