Abstract

Background: In 2010, Qatar adopted the target of reducing hepatitis B prevalence to < 1% in children by 2015. The World Health Organization Region for the Eastern Mediterranean is identified with intermediate hepatitis B virus (HBV) endemicity, ranging from 2% to 7%. It is estimated that 4.3 million individuals are living with HBV infection in the Region.

Aims: This study was conducted to assess hepatitis B seroprevalence in children, hepatitis B vaccination coverage, potential exposure to risk factors, and knowledge among parents/guardians about hepatitis B infection.

Methods: We carried out this cross-sectional study in Qatar during the academic year 2015/16. Multistage cluster sampling was used to select a nationally representative sample of 2735 grade 1 school students aged ≥ 5 years. Blood was collected by finger prick and tested using the point-of-care test/rapid test. A self-administered, precoded questionnaire was used to assess parent/guardian knowledge about HBV and collect information on the child’s HBV vaccination coverage.
Results: All blood samples were HBsAg negative. Qataris had a vaccination card and were totally vaccinated but 17.7% of non-Qataris did not hold a vaccination card and most parents/guardians were not aware of the vaccination status of their children. Children were exposed to various hepatitis B risk practices. Knowledge about hepatitis B among parents/guardians was low.

Conclusions: Qatar has averted the hepatitis B threat and maintained high vaccination coverage for children.

Keywords: Qatar, hepatitis B, seroprevalence, children, vaccination


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Introduction

The hepatitis B virus (HBV) infection takes a heavy toll on lives worldwide. In 2014, the World Health Organization (WHO) estimated that 30% of the world’s population, around 2 billion people, have been infected with HBV and around 360 million have developed chronic liver disease (1,2). Furthermore, HBV has been identified as the cause of up to 80% of all cases of hepatocellular carcinoma worldwide (3–5). Although HBV-related acute hepatitis may be asymptomatic and may resolve spontaneously, it may also lead to chronic lifelong infection. The main danger of HBV is in acquiring the infection early in life as progress of the condition towards severe pathological consequences, e.g. liver cirrhosis, liver failure and hepatocellular
carcinoma, in adulthood may reach up to 90% of infected cases (2,6–10). Additionally, mother-to-child transmission results in chronic infection later in life in 90% of infants infected during the perinatal period and in 80–90% of infants infected in the first year of life. The risk of chronic HBV infection declines with age, reaching 30–50% for children infected between 1 and 4 years of age and only 2–5% for adolescents and adults (6).

The WHO Eastern Mediterranean Region, which includes Qatar, is identified with intermediate HBV endemicity ranging from 2 to 7%. It is estimated that 4.3 million persons are living with HBV infection in the countries of the Region (11,12). Numerous studies have documented HBV infection as an important cause of liver cirrhosis and hepatocellular carcinoma in the countries of the Region (13–19). In addition to the burden on people’s lives, HBV infection drains the resources of the health system because of the cost of treatment for liver cirrhosis and hepatocellular carcinoma.

The limited number of studies on HBV infection in Qatar confirm that the country has not been free of the virus. A study in 1985 showed that 23% of adult patients with acute viral hepatitis admitted to Hamad General Hospital had HBV infection (20). During 2000–2005, the pathology reports at Qatar Principal Reference Laboratory indicated hepatitis B incidence was 2.5% among liver disease patients (21). In addition, the incidence of hepatitis B was reported in 2002–2006 as 4.7% of screened and diagnosed viral hepatitis patients in Hamad General Hospital (22). A study in 2010 reported HBV prevalence of 2.2% and lack of HBV immunity of 53.3% among professional male athletes who attended the Qatar Orthopaedic and Sports Medicine Hospital in Doha (23). As reported in these studies, HBV infection was more prevalent among non-Qataris than Qataris and among males than females.

The HBV vaccine has been part of the routine infant immunization programme in Qatar since August 1989, when a 3-dose schedule of monovalent vaccine was introduced. In 2000, a birth dose was added, aimed at preventing 70–95% of HBV infections. Thereafter in July 2003, a new 4-dose vaccine was introduced in the Expanded Programme on Immunization schedule to accommodate the use of a pentavalent vaccine (DTwP-HepB-Hib) and hexavalent vaccine (DtaP, Hep B, Hib and IPV) in addition to the monovalent vaccine at birth. Hepatitis B vaccination coverage has attained maturity over the past 5 years at 93–99% of all infants.

In 2010, Qatar adopted the WHO target of reducing HBV prevalence among children below 5 years of age to
Thus, this study was conducted to monitor the impact of the HBV vaccination programme in achieving the national and regional target among children aged at least 5 years who received the total vaccination doses, including the birth dose. In other words, the objectives of the study were to measure the HBV seroprevalence in children who benefitted from the National vaccination programme; assess the HBV vaccination coverage for these children and the potential exposure to HBV-related risk factors; and assess parent/guardian knowledge about HBV infection. Thus, the study was conducted to fill the existing gap in knowledge on hepatitis B prevalence and to add depth to the vaccination programme in Qatar. The significance of the study goes beyond its contribution to knowledge and regional discourse on the effectiveness of the HBV vaccination programme to draw the attention of the population to the national effort to protect children in Qatar from the HBV threat.

### Methods

#### Study design and target population

As it was hypothesized that the new 4-dose HBV vaccination programme had succeeded in reducing hepatitis B prevalence in children of at least 5 years of age to < 1% and these children are mostly enrolled in grade 1, a school-based survey was conducted during the academic year 2015–2016 targeting children and their parents/guardians.

#### Sample size

The sample size was calculated using the Epi-Info Statcalc package for prevalence of 1%, α-level 0.05 and design effect 1.5. For the 95% confidence level, the sample size was estimated to be 2279 children. As it was expected that up to 20% of the parents/guardians and/or the children themselves may refuse to participate in the survey, the sample was inflated by 20%, thus a total sample of 2735 children was targeted.

#### Sampling method

A multistage cluster sampling technique was used to select a nationally representative sample of grade 1 school students attending public and private schools and their parents/guardians. There were 19 195 children enrolled in grade 1 in 200 public and private schools in Qatar. A total of 52 schools were selected at random from the list of schools. As the number of students varied markedly by nationality, type of school and sex, sampling was done proportionately. Sampling frames and measure of size were prepared for each municipality by type of school (independent, private international, private approved by education council and private foreign community) and sex (mixed, boys and girls). A total of 51 primary sampling units from 50 schools were selected by systematic random sampling process proportional to school type and sex. In very few cases, notably when only 1 cluster was needed per school type and sex, a simple random process was used.
In the selected schools, registries were used to provide the list of classes and students in grade 1. If the number of students within a school exceeded the sample size, classes were selected at random and all students within those classes were enrolled in the study. Parents/guardians of selected students were also invited to participate in the study.

Data collection and analysis

Data collection was carried out during November 2015–February 2016 and was consistent in all schools. There were 2 components to the data. The first was the serologic component which included HBsAg testing of students using the point-of-care test/rapid test. This test was used as it is easy to administer and highly flexible. It requires minimal training, no equipment and can be relied upon for fast, accurate diagnosis in the most basic settings. Detecting HBsAg was conducted using the Alere Determine HBsAg rapid test (Abbott Laboratories, Santa Clara, California). The results were produced within 15 minutes with sensitivity 95.16% and specificity 99.95%. Blood was collected by finger prick to reduce fears among parents/guardians and children regarding venepuncture as this may affect the participation rate. Chronic HBV infection was assessed by measuring the proportion of HBsAg-positive by rapid test.

The second data collection tool was a standardized, self administered, precoded parent / guardian questionnaire. This was developed in English, translated into Arabic and adapted in order to ensure that it took into account the specific sociocultural realities of the country. The questionnaire was used to assess parent/guardian knowledge about HBV and collect information on child HBV vaccination coverage and the potential exposure of children to HBV-related risk factors. Vaccination coverage data were obtained from the child’s vaccination card. Where the child did not hold a vaccination card, the parent/guardian was asked to provide the information. The survey was pilot tested in 1 school on 152 children and the questionnaire was pretested on the parents/guardians prior to the actual fieldwork. The results of the pilot study were included in the final data analysis.

The vaccination status for hepatitis B vaccination at birth (HepB-birth) was defined as:

- fully vaccinated: HepB-birth administered < 24h from birth;
- partial vaccinated: HepB-birth administered > 24h to < 72h from birth;
not vaccinated: no full or partial HepB-birth vaccination.

The vaccination status for the 3 hepatitis B doses (HepB-3 dose) was defined as:

fully vaccinated: HBV first dose (HepB1) at 4 months, second dose (HepB2) at 6 months and third dose (HepB3) at 15 months;

partially vaccinated: invalid HepB2 and/or HepB3 vaccination;

not vaccinated: no full or partial HepB-3 dose vaccination.

The vaccination status for HepB total was defined as:

fully vaccinated: HepB-birth, HepB1, HepB2 and HepB3;

partial vaccination: partial Hep-birth, HepB1 and/or HepB2 and/or HepB3;

not vaccinated: no full or partial HepB-birth and HepB-3 dose.

Data entry and analysis were done using the SPSS computer package. The chi-squared test was used to assess significant differences between proportions, and Fisher’s exact test was used for 2 × 2 tables when the chi-squared test was not valid. Independent t-test was used to detect significant difference between 2 independent means. The significance level was P < 0.05.

Ethical considerations
The study proposal was reviewed and approved by the ethical review board of Hamed Medical Cooperation. All selected children and their parents/guardians were invited to participate in the study. Written informed consent from parents/guardians and assent from children were taken prior to data collection. All data were anonymous and confidential. Children and parents/guardians were told that they could refuse to answer any question and withdraw from the study any time.

**Results**

**Background characteristics of the study sample**

A total of 2803 grade 1 school students were enrolled in the study, 745 (26.6%) Qatari and 2058 non-Qatari. Boys accounted for 52.8% of the sample and ages ranged from 5 to 8 years [mean = 6.2, standard deviation (SD) = 0.6]. Over half of the children (58.7%) were born in Qatar and 51.7% were in public schools.

The majority of the non-Qatari children were from The Eastern Mediterranean Region (66.8%), and South East Asia Region (27.8%). The rest were from Europe (3.7%), Africa (1.1%), the United States of America and Canada (0.3%) and Western Pacific Region (0.1%).

**Table 1** provides the background characteristics of grade 1 school students in Qatar. The proportion of boys was 55.4% among Qatari students and 51.8% among the non-Qatari. The difference was not statistically significant. The age range for both Qatari and non-Qatari was around 5–8 years, and Qatari students were significantly slightly older than non-Qatari (Qatari: mean = 6.4, SD = 0.9 years; non-Qatari mean = 6.2, SD = 0.7 years). Almost all the Qatari (98.5%) were born in Qatar compared to only 44.3% of non-Qatari (P < 0.001).

**Hepatitis B seroprevalence**

As shown in **Table 2**, 2562 of our students (91.4% of the study sample) agreed to have the HBsAg testing using the point-of-care test, and 2538 tests (99.1%) were valid. All blood samples tested negative for HBsAg. These results clearly indicate that none of the tested students had HBV infection.

**Hepatitis B vaccination coverage**

All Qatari students and 82.3% of non-Qatari had a vaccination card (**Table 3**). All Qatari students in the sample, whether born in Qatar or outside, were fully vaccinated with HepB-birth and HepB-3dose. For the non-Qataris holding a vaccination card, 96.3% were full vaccinated with HepB-birth, 99.0% with HepB-3 dose, with a total of 96.2% fully vaccinated. There were 64
students (3.8% of non-Qatars holding a vaccination card) who were partially vaccinated as they were either partially or not vaccinated with HepB-birth and/or HepB-3dose. They were all from India, born in their home country and 74.2% of them were in private schools.

All of the 364 non-Qatari grade 1 school students who had no vaccination card were born outside Qatar, mainly in countries of the WHO Eastern Mediterranean Region (69.8% of non-vaccination card holders) and South East Asia (22.5%). They were from Egypt (33.8%), Syrian Arab Republic (15.4%) and India (17.3%), and 70.9% were private school students. Two-thirds of the parents/guardians did not know whether their children had received HepB-birth and 86.8% were not able to declare the vaccination status of their children. However, 28.8% of these children were known to have received HepB-birth, 12.4% were known to have received HepB-3-dose and 7.4% were known to be totally vaccinated.

**Previous exposure to hepatitis B risk factors**

As with children in this age group, grade 1 school students are liable to falling sick and receiving health care (Table 4). Around a third of the children had had dental treatment, 21.9% were ever hospitalized, 21.4% had received injections in hospitals, 13.4% had a surgical treatment and 4.7% had received injections at home. The proportion of Qatari children receiving dental care was significantly higher than the non-Qatari children while the proportion of children receiving other medical care procedures was significantly higher in non-Qatars than Qatari (Table 4). Around 10% of the children had received health care in the form of surgery, dental care and/or blood transfusion outside Qatar with significantly higher proportions among non-Qatars (11.7%) than Qatari (4.0%).

Other risk practices are detailed on Table 4. Few children (0.5%) had tattoos, and this was the same for Qatari and non-Qatari.

**Parent/guardian knowledge about hepatitis B viral infection**

Table 5 shows the knowledge of parents/guardians of grade 1 school students about hepatitis B viral infection. We found that 70.0% of parents/guardians had heard about HBV infection, 16.8% knew of a person who had died from the infection but just 17.0% had received information about HBV infection in the 6 months prior to data collection.

The proportion of parents/guardians who knew that blood was a route of transmission ranged from 49.0–58.2%. The proportion of parents/guardians who knew of the other routes of transmission was low, 31.0% for sexual transmission, 38.4% for vertical mother-to-child
transmission and 18.9% for transmission through breast-feeding. In general, non-Qataris had significantly better knowledge about HBV infection than Qataris. However, at least one-third of parents/guardians had a number of misconceptions about HBV, and this was significantly higher among non-Qataris than Qataris.

**Discussion**

Since the introduction of the new 4-dose HBV vaccine in the national Expanded Programme on Immunization, great progress has been achieved in Qatar towards the elimination of HBV infection in children. All grade 1 school children tested for HBsAg in our study tested negative. There is very limited information on HBV prevalence in children; the latest available figures document a prevalence of 0.8% in 5–14-year-old children (21), which is an indication that Qatar was moving towards achieving the regional target of reducing the prevalence of HBV infection to < 1% by 2015. Furthermore, there were no HBV cases reported in Qatar during 2013–2015 in the 0–4-year-old children (unpublished data, Ministry of Public Health national statistics, 2016). Thus, our results confirm this progress and demonstrate the success of Qatar in eliminating HBV infection in children who benefited from the new 4-dose vaccine in the national Expanded Programme on Immunization. The reason for this success resides in the HBV control strategy in Qatar. The country offers the 4-dose Expanded Programme on Immunization services free of charge to all citizens. The primary health care centres throughout the country offer free hepatitis B vaccination to all children regardless of nationality. HepB-birth is administered to all children born in hospitals, i.e. almost all births in the country. There is high-quality vaccine management in place to address shortage and ensure availability for all children. The Department of Health Protection and Communicable Diseases in the Ministry of Public Health has a tracking system to count the number of children requiring vaccination and ensure coverage. Our findings provide evidence on the high vaccination coverage in the country which reaches 100% in Qatari children. Our results have also shown that 82.3% of non-Qatari children had a vaccination card and almost all were fully vaccinated, giving a total coverage for these children of 96.2%. These results correspond with the data provided by administrative records in 2015 showing HBV vaccination coverage of 95–99% in the same age group (unpublished data, Ministry of Public Health, 2015). Previous experience in the Region using the 4-dose schedule in the Expanded Programme on Immunization and a similar HBV control strategy (24) has shown a dramatic decrease in chronic HBV seroprevalence in children, as is the case of Qatar.

It is clear that despite the efforts exerted by Qatar public health authorities, there is still a need for further action to maintain the current success in HBV control. Qatar faces several challenges in its fight against HBV infection. Over the past decade Qatar has offered numerous job opportunities to expatriates from all over the globe to support its labour force. This strategy has attracted many who found a place to work and live with their families. Our results show that some countries do not have strict enforcement of the vaccination programme: 31.8% of the non-Qatari children born outside Qatar did not have a vaccination card and the majority of parents/guardians did not know the vaccination status of their children. This makes it difficult for the health system to spot these children and apparently some remain unreached. This is further
aggravated by the fact that around 10% of non-Qatari children are more prone to seek health care outside Qatar. This has been demonstrated before for adults, a situation that put them at risk of contracting viral hepatitis (25). It was postulated that non-Qataris prefer to seek health care outside Qatar for several reasons such as limited financial resources, work overload, belief that their home country’s healthcare system responds better to their health needs and fear of losing their job.

Health needs and practices in the childhood period cannot be neglected. Children from different countries may have different health needs; different cultural norms and practices may put them at risk of HBV transmission. Our results have shown that Qataris are more in need of dental care while non-Qataris are more in need of medical care. Furthermore, ear piercing, tattooing and sharing tooth brush are practiced by both non-Qataris and Qataris. All these practices need effective control measures to prevent the spread of blood-borne infections, including HBV.

Our study reports that there was a gap in HBV-related knowledge among parents/guardians, with misconceptions prevalent among both Qataris and non-Qataris. Non-Qataris had better HBV knowledge than Qataris, yet they had more misconceptions and showed less appreciation to the role of vaccination in controlling HBV infection. This highlights the need for rigorous awareness campaigns responding to the needs of different traditions and cultures and directed to both Qataris and non-Qataris.

It is apparent from our results that some schools, particularly in the private sector, do not request the child vaccination card upon school entry. It is, therefore, clear that with the poor knowledge base of parents/guardians, these children may remain out of the reach of the health services.

This study had some limitations. As the questionnaires were self-administered, parents/guardians may not have been accurate in responding to some information. Yet, parents/guardians were asked to copy the vaccination status of the children from the vaccination card and for non-vaccination card holders, results were analysed separately to avoid any information bias. Furthermore, the research team had close contact with the parents/guardians to ensure completeness of information and avoid non-response. Chronic HBV infection was assessed by measuring the proportion of HBsAg-positive seroprevalence by the rapid test to avoid false positive results. We planned to confirm positive results by taking a venous blood sample, however all results were negative.

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**Competing interests:** None declared.

Évaluation d’un programme de vaccination contre l’hépatite B parmi des élèves au Qatar

**Résumé**

**Contexte:** En 2010, le Qatar a adopté une cible consistant à réduire la prévalence de l’hépatite B à moins de 1 % chez l’enfant d’ici 2015. La Région OMS de la Méditerranée orientale a été identifiée comme une région d’endémicité intermédiaire pour le virus de l’hépatite B, avec un taux compris entre 2 et 7 %. On estime que 4,3 millions de personnes dans la Région vivent avec une infection par le virus de l’hépatite B.

**Objectif:** La présente étude a été menée dans le but d’évaluer la séroprévalence de l’hépatite B chez l’enfant, la couverture vaccinale associée, le risque d’exposition aux facteurs de risque, et la connaissance de l’infection par le virus de l’hépatite B parmi les parents/tuteurs.

**Méthodes:** La présente étude transversale a été réalisée au Qatar pendant l’année scolaire 2015-2016. L’échantillonnage en grappe à plusieurs niveaux a été la technique utilisée pour sélectionner un échantillon national représentatif de 2735 élèves de première année du primaire, âgés de cinq ans ou plus. Un prélèvement de sang a été réalisé par piqûre au doigt et un dépistage a été effectué à l’aide d’un test sur le lieu de soins/test rapide. Un questionnaire auto-administré précédé a été utilisé pour évaluer les connaissances des parents/tuteurs en matière d’hépatite B et recueillir des informations sur la couverture vaccinale des enfants contre l’hépatite B.
Résultats: Tous les échantillons sanguins étaient négatifs pour l'antigène de surface du virus de l'hépatite B. Les Qatari possédaient un carnet de vaccination et étaient complètement vaccinés, mais 17,7 % des non-Qatari ne possédaient pas de carnet de vaccination et la plupart des parents/tuteurs ne connaissaient pas le statut vaccinal de leurs enfants. Les enfants étaient exposés à différentes pratiques à risque pour l'hépatite B. Les connaissances des parents/tuteurs sur l'hépatite B étaient faibles.

Conclusion: Le Qatar a su écarter la menace de l'hépatite B et maintenir une couverture vaccinale élevée pour les enfants.
17.7% of the participants had a history of vaccination for hepatitis B. However, only 26% of students in Qatar had completed the vaccination, while 15% of those who completed the vaccination were vaccinated. Among the participants, a significant number of students did not possess any hepatitis B vaccination card, and among those, the majority were asymptomatic carriers of hepatitis B. This highlights the importance of awareness and education about hepatitis B.

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