A knowledge survey of obstetrics and gynecology staff on the prevention of mother-to-child transmission of hepatitis B virus

Yali Hu1,2, Xinwei Dai1, Yi-Hua Zhou3, Huixia Yang4

1Department of Obstetrics and Gynecology, Nanjing Drum Tower Hospital, Nanjing Medical University, Jiangsu, China
2Department of Obstetrics and Gynecology, Nanjing Drum Tower Hospital, Nanjing University Medical School, Jiangsu, China
3Departments of Experimental Medicine and Infectious Diseases, Nanjing Drum Tower Hospital, Nanjing University Medical School, Jiangsu, China
4Department of Obstetrics and Gynecology, Peking University First Hospital, Peking University, Beijing, China

Abstract
Introduction: This survey was designed to investigate the knowledge awareness of obstetrics and gynaecology staff (Obs/Gyn staff) on the prevention of mother-to-child transmission (PMTCT) of hepatitis B virus (HBV).

Methodology: Obs/Gyn staff from 21 of the 31 Chinese mainland provinces, who attended medical meetings or training classes from July to October 2011, were invited to complete a questionnaire regarding PMTCT of HBV. The questionnaire included the clinical implications of HBV serologic markers and PMTCT preventive measures for both pregnant women and infants.

Results: A total of 828 questionnaires were distributed, 617 (74.5%) Obs/Gyn staff participated in the survey, and 559 (90.6%) questionnaires met the inclusion criteria. Overall, 90% of participants correctly determined the positive hepatitis B surface antigen (HBsAg) as infectious, but up to 27.5% mistakenly considered the presence of anti-HBe and/or anti-HBc with negative HBsAg as infectious. In total, 96.3% respondents knew that pregnant women should be screened for HBV infection, and 95.3% realized that infants of HBsAg-positive mothers should be injected with hepatitis B immunoglobulin and vaccine. On the other hand, with the available immunoprophylaxis, 13.8% participants mistakenly believed caesarean section may prevent HBV mother-to-child transmission, and only 13% correctly answered that newborns of HBsAg positive mothers may be breastfed.

Conclusion: Obs/Gyn staff in China have mastered the strategies of HBV PMTCT, but there is obvious insufficiency in details of the application. Intensified efforts to train the Obs/Gyn staff are required to improve the current suboptimal medical service in HBV-exposed infants and to control mother-to-infant transmission of HBV.

Key words: hepatitis B virus; mother-to-child transmission; immunoprophylaxis; knowledge awareness


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Introduction
Chronic hepatitis B virus (HBV) infection is a severe global health threat causing chronic hepatitis, cirrhosis, and hepatocellular carcinoma (HCC) [1]. It is estimated that 350 to 400 million people are chronically infected [2]. In China, the overall positive rate of hepatitis B surface antigen (HBsAg) is 7.18% [3], and annually approximately 300,000 individuals die from HBV-related diseases including HCC (40% to 50% of that of the entire world) [4]. Mother-to-child transmission of HBV leads to 80% to 90% chronic carriers [5]. Thus prevention of mother-to-child transmission (PMTCT) of HBV is of great importance.

The standard immunoprophylaxis against HBV is to give three doses of hepatitis B vaccine to all newborns as follows: one at birth, and then again at one month and six months after birth. In addition, infants born to HBsAg carrier mothers should receive hepatitis B immunoglobulin (HBIG) within 12 hours after birth [6]. The strategy is highly effective in the prevention of mother-to-infant transmission of HBV, with 99.71% to 100% and 85% to 95% protective rates in children of HBeAg-negative and HBeAg-positive carrier mothers respectively [7-12]. However, the strategy is not very well implemented in developing countries, including China [13,14]. For instance, many high-risk infants born to HBsAg positive mothers failed to receive HBIG and the timely first dose vaccine, in both urban and rural areas [15,16], and some unnecessary tests, such as testing for breast milk
HBV DNA [17], and non-evidence-based interventions, such as Caesarean section for PMTCT [18], are still in practice. Obstetrics and gynecology medical staff (Obs/Gyn staff) play critical roles in PMTCT of HBV since they administer HBIG and/or first dose hepatitis B vaccine to newborn infants. We conducted this survey to determine whether these inappropriate measures are due to the lack of knowledge in Obs/Gyn staff, so that education programs can be developed to reinforce the ability of Obs/Gyn staff to conduct PMTCT.

Methodology

Study subjects and questionnaires

This survey targeted the Obs/Gyn medical personnel from hospitals in 21 out of the 31 provinces in mainland China, who attended medical meetings or training classes from July to October 2011, excluding non-Obs/Gyn staff, undergraduate medical students, and those who had practiced in Obs/Gyn for fewer than two months. A questionnaire regarding PMTCT of HBV, containing the clinical significance of HBV serologic markers and relevant PMTCT measures, including those for both pregnant women and infants, was distributed among the subjects. The questionnaire was anonymous. The personal data and employers’ information were kept confidential. We collected the questionnaires on the spot when they were completed. The questionnaire was in Chinese, but an English version can be provided upon request.

Data analysis

The data were statistically analyzed using SPSS Package version 13.0 (IBM SPSS Inc., Chicago, IL, USA). We used analysis of variance to compare the mean numbers, and used chi square test as well as Fisher exact probability test to compare proportions. P < 0.05 was considered to be statistically significant.

Results

Demographics

Of the total 828 Obs/Gyn staff who were asked to complete the questionnaires, 617 (74.5%) participated in the survey. In the end, 559 (90.6%) questionnaires were included in this study, since 35 questionnaires were unfinished and the 23 others who did not meet the inclusion criteria were excluded. Of the 559 respondents, 194 (34.7%), 193 (34.5%), and 172 (30.8%) had worked for over 20, 11 to 20, and 1 to 10 years, respectively; 240 (42.9%), 124 (22.2%), and 111 (19.9%) were chief, attending, and resident doctors, respectively, and 84 (15.0%) were nurses. Overall, 389 (69.6%) respondents were employed by general hospitals and 279 (49.9%) were employed by Level 3 hospitals, which is the top level in China.

Basic knowledge/awareness of hepatitis B

The knowledge regarding HBV PMTCT of the participants was obtained from medical school (71.6%), academic lectures (69.9%), self-consulting relevant data (64.0%), and colleagues (32.0%), respectively. The proportion sum exceeding 100% is due to the multiple selections. No statistically significant differences were found among the accuracy rates of the basic knowledge of hepatitis B of respondents with the above knowledge sources (data not shown).

We designed a 12-question test about HBV serologic markers determination, consisting of 12 combination of the five HBV serologic markers (HBsAg, anti-HBs, HBeAg, anti-HBe, and anti-HBc), with positive or negative status for each. Respondents were asked to clarify the clinical significance of each combination. Overall, 90% of the participants correctly determined the positive HBsAg as infectious, but up to 27.5% mistakenly considered the presence of anti-HBe and/or anti-HBc with negative HBsAg as infectious; 81.4% of the respondents correctly determined the positive anti-HBs as immune to HBV, but 35.6% erroneously considered having the presence of anti-HBe and/or anti-HBc in the absence of anti-HBs as immune to HBV.

The correct rates with regard to HBV serologic markers determination were related to respondents’ education levels, titles, work seniorities, as well as levels and properties of the hospitals where they were working. Generally, respondents with higher education backgrounds had higher accuracy rates, and those working longer, with higher titles, or working in higher level hospitals were inexplicably less correct (P < 0.05).

PMTCT measures for pregnant women

As shown in Table 1, most (96.3%) of the participants knew that pregnant women should be routinely screened for HBV infection, while 3.3% still believed it should be determined based on specific situations (e.g., with an HBsAg-positive family history, or with abnormal liver function). With respect to whether HBsAg-positive pregnant women can breastfeed, only 13.0% respondents correctly answered that “newborns could be breastfed after administration of HBIG and full-dose vaccine”; while
<table>
<thead>
<tr>
<th>Measures (correct answer)</th>
<th>Correct/Total (%)</th>
<th>Hospital level†</th>
<th>Hospital Property</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td>1</td>
<td>2</td>
</tr>
<tr>
<td>Need for HBV screening during pregnancy? (yes)</td>
<td>519/539 (96.3)</td>
<td>68/72 (94.4)</td>
<td>201/203 (99.0)</td>
</tr>
<tr>
<td>Need caesarean section for PMTCT? (no)</td>
<td>462/536 (86.2)</td>
<td>64/72 (88.9)</td>
<td>177/202 (87.6)</td>
</tr>
<tr>
<td>Need HBV DNA test for breast milk? (no)</td>
<td>103/534 (19.3)</td>
<td>14/72 (19.4)</td>
<td>46/202 (22.8)</td>
</tr>
<tr>
<td>Can HBV carrier mothers breastfeed? (yes)</td>
<td>69/532 (13.0)</td>
<td>9/72 (12.5)</td>
<td>23/201 (11.4)</td>
</tr>
</tbody>
</table>

*The data represents “correct answers/total answers”, followed by the correct rates (%) in brackets. HBV, hepatitis B virus; PMTCT, prevention of mother-to-child transmission; HBIG, hepatitis B immunoglobulin; FPC, family planning centre. †Level 3 refers to the highest level hospital in China.
69.0% respondents answered that “it depends on HBV serologic markers status or HBV DNA in breast milk”. Similarly, 81.7% incorrectly considered that it is necessary to test HBV DNA in milk. In addition, 13.8% of the respondents believed that Caesarean section can reduce mother-to-child transmission risk of HBV, although this is not the case.

With regard to the likelihood of antiviral therapy reducing mother-to-infant transmission, 11.7% of the participants considered it necessary to implement antiviral treatment with nucleoside drugs (e.g., lamivudine) for HBsAg carrier pregnant women, whereas 42.1% considered it not necessary and 21.6% considered it dependent upon HBV serologic markers; and 13% others considered that further study is needed. There was a significant difference among the correct rates of respondents from different types of hospitals (P = 0.008).

**PMTCT measures for newborns**

Over 95% of the participants realized that infants of mothers with positive HBsAg should be injected with HBIG and vaccine (Table 2); nevertheless, only 61.6% and 80.4% knew the right injection method of HBIG and proper immunization program of the vaccine, respectively. In spite of that, just 84.5% of the respondents answered that HBIG was “almost always” injected into infants of HBsAg carrier mothers in practice. In addition, 61.8% of the respondents mistakenly believed that those infants should be tested for the presence of HBV markers in the cord blood.

In practice, when a pregnant woman’s HBsAg status remained unknown at delivery, 71.9% respondents would have an emergency test, while others would treat it as negative (19.3%) or positive (6.6%). Over 51% of the respondents said they would “almost always” take the initiative to follow up the infants of HBsAg-positive mothers, and 27% would “sometimes” do that.

**Discussion**

This survey shows that the vast majority of the Obs/Gyn staff in China are aware of routine screening for HBV serologic markers in pregnant women, and know the immunoprophylaxis of combination HBIG and vaccine to the high-risk infants for PMTCT. However, the Obs/Gyn staff in China does not fully understand the significance of HBV serologic markers. Additionally, many Obs/Gyn staff are confused on the issues of whether HBsAg-positive pregnant women need detection of milk HBV DNA, whether they can breastfeed their infants, and whether cord blood needs measurement of HBV markers to determine if the infants are infected.

Antenatal screening for HBV serologic markers during pregnancy is of great significance in the prevention of HBV mother-to-infant transmission [19]. Given the possibility that some pregnant women may not tell the true histories of HBV infection because of their worries about the potential discrimination against HBV carriers, and that ad-hoc infection may occur during the pregnancy, Obs/Gyn physicians cannot exclude HBV infection in pregnant women only based on the history or pre-pregnancy testing results. On the other hand, if anti-HBs were once positive before pregnancy, there is little possibility of re-infection with HBV since immunity has been established. Hwang et al. [20] found financial barriers and the health provider’s knowledge mainly influence HBV screening among Asian Americans. Considering Obs/Gyn physicians in our survey have high awareness of routine HBV screening for pregnant women, financial barriers may be the leading cause that affects the completeness of universal screening for pregnant women in developing countries.

Historically, some measures were implemented for PMTCT of HBV, such as performing Caesarean section [21] and avoiding breastfeeding [22]. However, many studies have confirmed that these two measures cannot reduce the rate of mother-to-child transmission after standard immunoprophylaxis [23-27]. Although most respondents (86.2%) considered it unnecessary to perform Caesarean section for PMTCT, the awareness rate still needs to be further improved. As for the breastfeeding issue, the majority of respondents believed it depends on HBV serologic markers of milk to determine whether HBsAg carrier mothers can breastfeed. The lack of this knowledge may result from delayed updates of relevant textbooks and reference books in Chinese.

To date, there have been no definite conclusions on the issue of whether HBV infected pregnant women during the third trimester should take the antiviral drugs for PMTCT [28-30]. Further research remains to be conducted to clarify this issue. However, 11.7% of the respondents in this survey thought antiviral therapy is necessary. Similarly, a survey conducted in the United States [31] also found physicians’ attitudes on this issue to be undefined. In that survey, 51.8% physicians would initiate antiviral treatment during pregnancy. Absence of appropriate guidelines (60.4%) and lack of evidence to support either safety (32.1%) or efficacy (7.5%) are responsible for the caution of...
<table>
<thead>
<tr>
<th>Measures (correct answer)</th>
<th>Correct/Total (%)</th>
<th>Correct rates</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>$\chi^2/P$</th>
<th>General</th>
<th>Maternity</th>
<th>FPC</th>
<th>$\chi^2/P$</th>
</tr>
</thead>
<tbody>
<tr>
<td>Need for HBIG injection at birth? (yes)</td>
<td>Correct rates</td>
<td>511/536 (95.3)</td>
<td>69/72</td>
<td>196/203</td>
<td>246/261</td>
<td>1.403/0.496</td>
<td>359/373</td>
<td>122/131</td>
<td>30/32</td>
<td>2.311/0.315</td>
</tr>
<tr>
<td>Is HBIG administered to newborns in your hospital in practical work?</td>
<td>Almost always</td>
<td>453/536 (84.5)</td>
<td>63/72</td>
<td>175/203</td>
<td>215/261</td>
<td>1.403/0.496</td>
<td>359/373</td>
<td>122/131</td>
<td>30/32</td>
<td>2.311/0.315</td>
</tr>
<tr>
<td></td>
<td>Sometimes</td>
<td>42/536 (7.8)</td>
<td>5/72</td>
<td>15/203</td>
<td>22/261</td>
<td>2.168/0.705</td>
<td>359/373</td>
<td>122/131</td>
<td>30/32</td>
<td>6.844/0.126</td>
</tr>
<tr>
<td></td>
<td>Others§</td>
<td>41/536 (7.6)</td>
<td>4/72</td>
<td>13/203</td>
<td>24/261</td>
<td>2.168/0.705</td>
<td>359/373</td>
<td>122/131</td>
<td>30/32</td>
<td>6.844/0.126</td>
</tr>
<tr>
<td>Need detection of HBV markers in cord blood? (no)</td>
<td>Correct rates</td>
<td>203/531 (38.2)</td>
<td>28/72</td>
<td>61/201</td>
<td>114/258</td>
<td>9.177/0.010</td>
<td>132/368</td>
<td>58/131</td>
<td>13/32</td>
<td>2.973/0.226</td>
</tr>
<tr>
<td>How do you manage the pregnant women with unknown HBsAg status at delivery?</td>
<td>Determine as negative</td>
<td>103/534 (19.3)</td>
<td>9/72</td>
<td>42/203</td>
<td>52/259</td>
<td>7.737/0.006</td>
<td>77/372</td>
<td>23/130</td>
<td>3/32</td>
<td>6.237/0.357</td>
</tr>
<tr>
<td></td>
<td>Determine as positive</td>
<td>35/534 (6.6)</td>
<td>2/72</td>
<td>13/203</td>
<td>20/259</td>
<td>2.168/0.705</td>
<td>21/372</td>
<td>13/130</td>
<td>1/32</td>
<td>6.237/0.357</td>
</tr>
<tr>
<td></td>
<td>Emergency test</td>
<td>384/534 (71.9)</td>
<td>56/72</td>
<td>146/203</td>
<td>182/259</td>
<td>11.097/0.074</td>
<td>265/372</td>
<td>91/130</td>
<td>28/32</td>
<td>6.237/0.357</td>
</tr>
<tr>
<td></td>
<td>Nothing special</td>
<td>12/534 (2.2)</td>
<td>5/72</td>
<td>2/203</td>
<td>5/259</td>
<td>2.168/0.705</td>
<td>9/372</td>
<td>3/130</td>
<td>0/32</td>
<td>6.237/0.357</td>
</tr>
<tr>
<td>Do you follow up the infants born to HBsAg carrier mothers?</td>
<td>Almost always</td>
<td>274/534 (51.3)</td>
<td>45/72</td>
<td>89/202</td>
<td>140/260</td>
<td>11.455/0.022</td>
<td>183/371</td>
<td>67/131</td>
<td>24/32</td>
<td>9.450/0.051</td>
</tr>
<tr>
<td></td>
<td>Sometimes</td>
<td>144/534 (27.0)</td>
<td>19/72</td>
<td>64/202</td>
<td>61/260</td>
<td>11.455/0.022</td>
<td>108/371</td>
<td>31/131</td>
<td>5/32</td>
<td>9.450/0.051</td>
</tr>
<tr>
<td></td>
<td>Others§</td>
<td>116/534 (21.7)</td>
<td>8/72</td>
<td>49/202</td>
<td>59/260</td>
<td>11.455/0.022</td>
<td>80/371</td>
<td>33/131</td>
<td>3/32</td>
<td>9.450/0.051</td>
</tr>
</tbody>
</table>

*The data represents “correct answers (or the selections of the relevant options)/total answers”, followed by the correct rates or the proportion (%) in brackets. HBV, hepatitis B virus; PMTCT, prevention of mother-to-child transmission; HBIG, hepatitis B immunoglobulin; FPC, family planning centre. †Level 3 refers the highest level hospital in China. §Others include “occasionally” or “scarcely”.

Table 2. Preventive measures towards newborns of HBsAg carrier mothers
the other 48.2% of physicians who did not recommend antiviral therapy.

It is recommended that infants born to HBsAg carrier mothers should be tested for HBsAg and anti-HBs at age 9 to 18 months to determine whether the prophylaxis is successful [28]. Owing to the possible maternal blood contamination and transplacental leakage of maternal HBsAg and HBeAg to cord blood [32], detection of HBV serologic markers in cord blood or neonatal peripheral blood shortly after birth cannot determine whether the infants are infected [33]. However, only 38.2% of the respondents correctly answered this question in the present investigation. Indeed, such unreliable tests are performed in clinical practice [34], which may lead to waste of medical resources and excessive worries about infant infections if any of the cord blood HBV markers is ever positive.

When a pregnant woman’s HBV status at delivery is unknown, a blood test is recommended as soon as possible, and her infant should receive HBIG and the first dose of vaccine within 12 hours of birth [28]. Although 71.9% of the respondents in the survey expressed that they would perform an emergency blood test, few hospitals can report the results in time, especially at night and holidays. Thus, in high-endemic areas of HBV infection such as China, considering the high HBsAg carrier rates of pregnant women and the extremely high chronicity rates when infants are infected with HBV, newborns of pregnant women with unknown HBsAg status should receive HBIG even though it may cause some waste.

Noticeably, even if 95.3% of the respondents knew the necessity of giving of HBIG injection to infants of HBsAg positive mothers, fewer than 85% would “almost always” do so in daily practice, indicating that some medical staff do not strictly follow the proper preventive measures. Thus increased supervisory vigilance is warranted. Additionally, we found that the senior Obs/Gyn staff in tertiary hospitals unexpectedly had a lower correct rate in answering the questions on basic knowledge of HBV infection and prevention. This finding indicates that the relevant training should be strengthened not only in the junior staff but also in senior staff.

In this survey, the study subjects were mainly attendants who participated in national and local medical meetings. Since China has a vast, geographically diverse territory with unequal financial development, health providers in remote or isolated areas, as well as rural areas, seldom have opportunities to attend medical meetings. Therefore, the data of this survey reflects only the situations in Chinese cities and towns. Based on the insufficient knowledge in the Obs/Gyn staff in this survey, we can imagine that the scenario in rural and remote areas may be worse.

Conclusion

Obs/Gyn staff in China have basically mastered the prophylaxis strategies of HBV PMTCT, but there is obvious insufficiency in details of the application. Therefore, intensified efforts to train the Obs/Gyn staff are required to improve the currently suboptimal medical service provided to HBV-exposed infants and to control mother-to-infant transmission of HBV.

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References


Corresponding author
Huixia Yang
Department of Obstetrics and Gynecology
Peking University First Hospital
Peking University
Beijing, China
Telephone/Fax: +86-10-6651-3519
Email: huixiayang2012@126.com

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