SERO-PREVALENCE AND PREDICTIVE STATISTICAL MODEL FOR THE HEPATITIS B VIRUS EXPOSURE AMONG PEOPLE IN COMMUNITY, YASOTHON, THAILAND

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ABSTRACT

This study investigated the prevalence of Hepatitis B virus (HBV) and identified a predictive statistical model for the HBV exposure among people in the community, Yasothon, Thailand. A cross-sectional study was performed on participants over 26 years old and living in Muang district, Yasothon province, Thailand. The research was conducted from July to August 2019. All 1,258 participants were verbally screened. Four hundred and fifty nine people were the risk group and tested for HBsAg, and 18 cases were positive for HBsAg (3.9%, [95%CI 3.5-4.4]). For the predictive model, the HBV exposure connected with sex, marital status, alcohol, smoking, and knowledge. The area under the receiver operating characteristics (ROC) curve was 61.8% ([95%CI, 58.6 to 65.0]). At cut-off point -0.66, the sensitivity, specificity and accuracy were 72.6%, 42.4% and 53.4%, respectively. HBV infection was a serious health problem, it can cause cirrhosis and liver cancer in the future. The predictive model of five variables can predict risk exposure of HBV which may had other relevant factors. Verbal screening by questionnaire to classify HBsAg risk group can lower the implementation cost.

Keywords: Hepatitis B virus, Seroprevalence, Predictive model

INTRODUCTION

Hepatitis B virus (HBV) caused high morbidity and mortality worldwide. World Health Organization (WHO) estimates that 257 million people were living with chronic hepatitis B infection. HBV was the main causes of liver cancer. And more than 50% of liver cancer patients had HBV. An estimated 887,000 deaths, mostly from cirrhosis and hepatocellular carcinoma⁴.

Thailand was an epidemic area of HBV. Approximately 10 million of the population were the carriers of HBV and those born under 1992 had an HBV infection of 4.5-5.2%. It is estimated that may be the population infected with HBV of approximately 2.2 – 3 million⁵. High prevalence of HBV infection was those over 25 years³. In 2013, Yasothon province found that the mortality rate of liver cancer was the 3rd highest of Northeastern Thailand, 50.3 per ten thousand people or 270 cases were liver cancer. The HBV situation in Yasothon in 2016 has been reported 45 cases, 8.3 per ten thousand population⁴. It is estimated that there was more HBV infection in Yasothon.

HBV vaccine has been used in Thailand for the past 26 years. Thus, people born before 1991 didn’t get the HBV vaccine which affected the high incidence of HBV infection⁶. In 2016, the Ministry of Public Health (MOPH) of Thailand had determined the preventive strategy of HBV, 2017-2021 which related with Framework for the global health sector strategy on viral hepatitis, 2016-2021⁷. Strategy framework included: 1) Surveillances and data development; 2) Prevention; 3) Risk communication; 4) Case finding and care; 5) Research; and 6) Resource management which the goal decreased the HBV rate. Activities focused on expanded program for immunization (EPI) promote, chronic HBV is able to access to health care more than 50% by 2021⁸. Thailand’s Hepatitis B virus prevention strategy will be supported by the implementation of prevention and patient care to be clearer and more concrete. However, current conduction still do not have the format for drive strategies to solve the HBV problem.

In the past, HBV problem solving only conducted on health workers, men who had sex with men (MSM), sex worker, injection drug user (IDU); nevertheless, this was never implemented in the community⁵, especially a prevalence survey of people with HBV infection and risk behavior survey people of HBV⁴. Affect to health workers lacked the important data for planning to solve HBV problem. While with the risk group most was screening test and access to treatment.

This study investigated the prevalence of HBV and identifies the predictive statistical model for the...
HBV exposure among people in community, Yasothon, Thailand.

MATERIALS AND METHODS

Settings and Design
A cross-sectional study was conducted in a community, Muang District, Yasothon Province, Thailand. It aimed to achieve the primary objective of estimating the prevalence of HBs seropositivity and to identify the predictive statistical model for the HBV exposure. The study was run during July to August 2019.

Participants
People living in the community, Muang district, Yasothon province in the present was a cross section of men and women aged 26 year or older, who born after the HBV vaccine had been used in Thailand were recruited in the study. The exclusions criteria were as follows: there were hearing problems and answering the questionnaires were not complete.

The sample size was 1,258 participants assuming a prevalence of HBsAg of 5%, from a study that reported among the general population category[4]. Sample size was derived using Cochran WG's formula, which estimate population proportion when the finite population the following formula: (n^2 = (Z^2a / 2Q) / rP, n = n^2 / 1 + n^2/ N), where n^2 = initial sample size; n = sample size; Z^2a / 2 = standard normal distribution absissa corresponding to 95% confidence interval (1.96); P = proportion of HBsAg reported in similar study noted above (5%); Q = (1-P); r = relative standard error (5%) and N = populations aged 26 years or older that live in a community (4,048). Simple random sampling without replacement.

Collecting data and instrumentation
The questionnaires were designed by a research team to consist of five sections: (1) demographic data, (2) other health data, (3) HBV history, (4) risk behavior of HBV, and (5) HBV knowledge. Development and psychometric testing of the questionnaire were performed. The content validity index (CVI) was found of 0.83 – 1.00 from six content experts, and the content validity of the overall scale (S-CVI) was discovered of 0.99. After that, the bright questionnaire was tried out in 30 samples for reliability testing which found Cronbach’s alpha of 0.83.

Serology
All participants were verbally screened by the questionnaire. They were interviewed by health officers. Then, those who had the risk exposure up to 2 items, were referred to blood testing by finger test for Hepatitis B surface antigen (HBsAg). The samples who had HBsAg positive were consequently confirmed by Real-Time Polymerase Chain Reaction (RT-PCR) technic of Yasothon hospital.

Data analyzing and statistical
Data was analyzed using Statistical Package for the Social Sciences (SPSS) software, version 18.0 (SPSS®, Chicago, USA). Descriptive statistic was used to summarize demographic data. Risk behavior or HBV exposure score was classified into two levels, two or more scores were “high risk group”, less than two scores were “none or low risk group”. Knowledge score was classified into three levels which included low, medium and high7. Unconditional multiple logistic regression was used to identify predictive statistical model for the HBV exposure. Independent (x) variables were demographic data, knowledge, alcohol, smoking and underlying. Dependent variable (y) was risk behavior from the high risk group and none or low risk group. Chi-square test was used to evaluate the bivariate relationship between the dependent variables and independent variables. The independent variables were selected with p value set at < 0.25 that suggested by Hosmer and Lemeshow10 into initial regression model of multivariate analysis. Before conducted multiple logistic regression, which was an assumption test to include multi-collinearity by Variance Inflation Factor (VIF) and interaction effect that founded all independent not violated assumption. Fitted the model by backward elimination technique, the research only kept the significant variables in the final model, after that a test model was discharged by Hosmer-Lemeshow goodness-of-fit test. Identified Cut-off-point for precision of the model for predict HBV exposure was used area under the receiver operating characteristics (ROC) curve. P value <0.05 was considered significant.

RESULTS

Of the 1,258 participants, mostly were female (60.2%), more than 45 years old (75.2%), Single/Divorced/Widowed (75.4%), agriculture (68.1%) and lower secondary school (76.2%) in table 1.

Risk behaviors toward hepatitis B were high risk group (36.5). HBV knowledge mostly was at good levels (57.6%) but was of low levels at 42.4% in table 2.

459 participants in high risk group were HBsAg testing and found 18 positives (3.9% [95%CI 3.5-4.4]). Mostly of HBsAg positive which had HBV, cirrhosis and liver cancer family history as 61.1% (11/18) in table 3.

Bivariate analysis for select independent variable into initial regression model of multivariate analysis found six variables that p-value < 0.25 and all variable had VIF value < 10 in table 4.
Table 1  Demographic characteristics of the study participants (N = 1,258)

| Variables            | n   | %    |
|----------------------|-----|------|
| Sex                  |     |      |
| Male                 | 501 | 39.8 |
| Female               | 757 | 60.2 |
| Age (year)           |     |      |
| 26-44                | 312 | 24.8 |
| ≥45                  | 946 | 75.2 |
| Mean (SD), Median (min: max) | 53.1 (12.8), 53 (26: 96) |
| Marital status       |     |      |
| Married/Partnered    | 309 | 24.6 |
| Single/Divorced/Widowed | 949 | 75.4 |
| Occupation           |     |      |
| Agriculture          | 856 | 68.1 |
| Employed             | 223 | 17.7 |
| Unemployed           | 179 | 14.2 |
| Education            |     |      |
| Lower secondary school | 958 | 76.2 |
| Secondary school and upper | 300 | 23.8 |

Table 2  Risk behaviors toward hepatitis B and hepatitis B knowledge (N = 1,258)

| Variables                        | n   | %    |
|----------------------------------|-----|------|
| Risk behaviors of HBV            |     |      |
| Low or None                      | 799 | 63.5 |
| High                             | 459 | 36.5 |
| HBV knowledge                    |     |      |
| Good (≥60%)                      | 533 | 42.4 |
| Low (<60%)                       | 725 | 57.6 |

Table 3  Prevalence of Hepatitis B virus (HBV) stratified by member of family was HBV/Cirrhosis/Liver cancer

| Family history of HBV/Cirrhosis/Liver cancer | Verbal screening | High risk group and blood testing | HBV positive |
|---------------------------------------------|------------------|-----------------------------------|--------------|
| n                                           | n (%)            | n (%)                             | n (%)        |
| Yes                                         | 224              | 148 (66.1)                        | 11 (7.4)     |
| No                                          | 1,034            | 311 (30.1)                        | 7 (2.3)      |
| Total                                       | 1,258            | 459 (36.5)                        | 18 (3.9)     |

Multivariate analysis by Unconditional multiple logistic regression, the final model found five factors associated with HBV exposure include sex (Adjusted OR 2.0, 95%CI 1.4 to 2.8), marital status (Adjusted OR 1.5, 95%CI 1.1 to 1.9), alcohol (Adjusted OR 1.6, 95%CI 1.2 to 2.0), smoking (Adjusted OR 1.5, 95%CI 1.1 to 2.2), and knowledge (Adjusted OR 1.7, 95%CI 1.3 to 2.1 in table 5).

DISCUSSION
HBV infection was a serious health problem in Yasothon province. Despite the recent data demonstrated a prevalence rate less than the prevalence estimated from Thailand, a prevalence rate higher than some studies. The result of this study showed that people in the community were a risk group to HBV infection similar prevalence of general population. Also, HBV infection to have a higher prevalence in those who have family history of HBV infection. Many studies mentioned that people who had the family history of HBV, cirrhosis and liver cancer are high risk populations for HBV infection. These can affect to increase for HBV chronic, cirrhosis and liver cancer in the future. Suh et al. reported that HBV was the main causes of liver cancer. And more than 50% of liver cancer patients had HBV. Liver cancer is the sixth most common cancer worldwide. About 83% occurred in less developed countries, and the highest incidence was observed in Asia and Africa.
Tables 4  Factors associated with HBV exposure (Bivariate analysis) (N = 1,258)

| Factors                        | N   | High risk group n (%) | Crude OR | 95% CI | P-value |
|--------------------------------|-----|-----------------------|----------|--------|---------|
| Sex                            |     |                       |          |        |         |
| Female                         | 757 | 298 (39.4)            |          |        | 0.009*  |
| Male                           | 501 | 161 (32.1)            | 1.4      | 1.1, 1.7 |         |
| Age                            |     |                       |          |        |         |
| 26-44 year                     | 312 | 119 (38.1)            |          |        | 0.484   |
| ≥45 year                       | 946 | 340 (35.9)            | 0.9      | 0.9, 1.9 |         |
| Status                         |     |                       |          |        |         |
| Married/Partnered              | 949 | 325 (34.2)            |          |        | 0.004*  |
| Single/Divorced/Widowed        | 309 | 134 (43.4)            | 1.5      | 1.1, 1.9 | 0.533   |
| Education                      |     |                       |          |        |         |
| Less than middle school        | 958 | 345 (36.0)            | 1.1      | 0.8, 1.4 | 0.065*  |
| Since middle school            | 300 | 114 (38.0)            |          |        |         |
| Occupation                     |     |                       |          |        |         |
| Agriculture                    | 859 | 302 (35.2)            |          |        |         |
| Employed                       | 220 | 81 (36.8)             | 1.1      | 0.8, 1.5 |         |
| Unemployed                     | 179 | 76 (42.5)             | 1.4      | 0.9, 1.9 |         |
| Underlying factors             |     |                       |          |        |         |
| (DM/HT/Heart/kidney)           |     |                       |          |        |         |
| No                             | 836 | 297 (35.5)            |          |        |         |
| Yes                            | 422 | 162 (38.4)            | 1.1      | 0.9, 1.4 |         |
| Smoking                        |     |                       |          |        |         |
| No                             | 901 | 315 (35.0)            |          |        | 0.074*  |
| Yes                            | 357 | 144 (40.3)            | 1.3      | 0.9, 1.6 |         |
| Alcohol drinking               |     |                       |          |        |         |
| No                             | 587 | 194 (33.0)            |          |        | 0.018*  |
| Yes                            | 671 | 265 (39.5)            | 1.3      | 1.0, 1.7 |         |
| HBV knowledge                  |     |                       |          |        | <0.001* |
| Good (≥60%)                    | 533 | 160 (30.0)            |          |        |         |
| Low (<60%)                     | 725 | 299 (41.2)            | 1.6      | 1.3, 2.1 |         |

*variables are p value < 0.25 to initial model of multiple logistic regression

Table 5 Predictive statistical model for the risk behaviors toward hepatitis B (Multivariable analysis by multiple logistic regression) (N = 1,258)

| Variables                        | Co-efficient | SE  | Crude OR | Adj. OR | 95% CI for OR | P-value |
|----------------------------------|--------------|-----|----------|---------|---------------|---------|
| Sex (male)                       | 0.69         | 0.17| 1.4      | 2.0     | 1.4, 2.8      | <0.001  |
| Marital status                   | 0.37         | 0.14| 1.5      | 1.5     | 1.1, 1.9      | 0.007   |
| (Single/Divorced/Widowed)        |              |     |          |         |               |         |
| Alcohol                          | 0.44         | 0.13| 1.3      | 1.6     | 1.2, 2.0      | 0.001   |
| Smoking                          | 0.42         | 0.19| 1.1      | 1.5     | 1.1, 2.2      | 0.025   |
| HBV knowledge (low)              | 0.52         | 0.12| 1.6      | 1.7     | 1.3, 2.1      | 0.001   |
| Constants                        | -1.72        | 0.19| -        | -       | -             | -       |

Verbal screening by questionnaire for classified risk group into HBsAg testing in this study can decrease a cost for the implementation. HBV vaccination for persons who did not have the immune and HBsAg testing in the risk group to be cost-effective. Therefore, these referred to treatment or health-promoting 14,15. The predictive model of five variables can predict the risk exposure of HBV significantly including sex, marital status, alcohol, smoking and knowledge.

Males had the risk exposure more than females because males had a chance to show off sex behavior such as they had a tattoo and sex with multiple partners. In a study carried out in blood donors by Torre et al14 and Xaydalasouk et al17

Figure 1. Area under the receiver operating characteristics (ROC)
show that people who have multiple partners or history of sexual risk behaviors are associated with HBV infection.

The subjects who had marital status single, divorced and widowed seemed to have the risk exposure more than those who had married, may be those who did not have a regular partner, could be having sex with another partner. The study carried out by Pereira et al\textsuperscript{10} showed significant association with married status.

Moreover, those who used smoking, alcohol drinking and low HBV knowledge will have an increased chance to expose the risk of HBV similar findings were reported from Ataef et al\textsuperscript{9} and Frew et al\textsuperscript{17}. On the other hand, those who don’t smoke or drink alcohol and have a high knowledge will have a decreased chance to expose the risk of HBV\textsuperscript{18,19}.

Hosmer-Lemeshow goodness-of-fit test (GOF) and the area under the receiver operating characteristics (ROC) showed that if the model was fit, predicted the risk exposure of HBV were at a poor level but 95% CI of ROC that the narrow showed the model was precise. Sensitivity value was fair, even specificity and accuracy value to be fail\textsuperscript{20}. However, the model can be used to plan for HBV prevention, especially using screening for health promoting and HBsAg testing which were cost-effective\textsuperscript{14}.

A highly effective program for universal immunization was implemented in Thailand more than two decades ago. However, a large number of individuals infected with HBV were born earlier and people live in community are one of important risk group. The high burden of HBV in Yasothon province still is a major challenge for the incorporation of programs to prevent HBV complications within health care systems such as the development the surveillance system in order for early detection and refer people with HBV infection to be treated. While educational programs for health must focus in particular on the prevention of people who had the family history of HBV, cirrhosis and liver cancer.

LIMITATION

This study conducted among people in community who lived in the Northeast maybe have a difference about culture. Therefore, they should be careful when using the finding from this study for implantation in other risk groups.

SUGGESTIONS

Health officer can use the finding from this study for application into HBV prevention. And they can also bring the questionnaire to verbal screening the risk group of HBV. Next study, should be a study of other factors that associates with HBV. And follow people that had HBsAg positive that became cirrhosis and liver cancer.

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Conflict of interest

The authors have no conflicts of interest to declare.

Ethical approval

The present study was approved by Human Research Ethics Committee of the Mahasarakham University (protocol number 058/2019). Patients were enrolled after written informed consent.

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