AN OUTBREAK OF HEPATITIS A IN SCHOOL CHILDREN AT NAKHON SI THAMMARAT, SOUTHERN THAILAND

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Abstract. Hepatitis A antibody prevalence in Southeast Asia has markedly declined among children and adolescents. Therefore increasing a number of susceptible populations could result in an outbreak or epidemic. This paper reports an intensive study of an outbreak of hepatitis A in a primary school children during an endemic at Nakhon Si Thammarat, Southern Thailand. Eighty-nine children were inspected randomly from the total of 269 students, age ranged from 7-12 years old. The school children and parents were interviewed for their illness. Serological tests for antiHAV IgM and antiHAV (total antibody) were performed by ELISA. There were 36 cases of clinical acute hepatitis with school absence ranging from 1-21 days (average 7.3 days). Not all cases with acute hepatitis were positive for antiHAV IgM because serological tests were performed nearly 4 months later. Seventy of 89 children tested were positive for antiHAV and 16 of them were positive for IgM class. Seven of 16 children with antiHAV IgM positive were asymptomatic. The significant risk factors for children with positive antiHAV were occurrence of hepatitis patients in the family and no latrine (p < 0.01). Endemic transmission in this outbreak occurred rapidly. Therefore preventive measures are essential in reducing the infection rate. In addition to personal hygiene, immunoprophylaxis with either immunoglobulin or HAV vaccine is recommended.

INTRODUCTION

Hepatitis A virus (HAV) infection when symptomatic is characterized by jaundice, nausea, vomiting, abdominal discomfort, enlarged and tender liver, fever and other systemic manifestations. HAV predominantly causes acute infections, specifically inflammation of the liver. It is an important public health problem throughout the world (Chitkara and Sachdev, 1988). Now it continues to be a source of morbidity and mortality in both developed and rapidly developing countries. A decline in disease incidence in developed and developing countries, including Japan (Takahashi et al, 1990), Australia (Gust, 1992), USA (Frace and Jahre, 1991; Desenctos et al, 1991), Europe (Stroffolini et al, 1990) and Thailand (Poovorawan et al, 1993) has been attributed to general improvement in standard of living. This has resulted in a large proportion of the population that is now susceptible to HAV. Because fecal contamination of water or food sources by the virus has not been completely eliminated, subsequently periodic endemics continue to occur. In 1988 an outbreak was observed in Shanghai, China with more than 300,000 cases of hepatitis detected (Xu et al, 1991). Other incidences were also reported eg in

Jefferson County, Kentucky (Rosenblum et al, 1990); in Indian Reservations of South Dakota (Shaw et al, 1990); in Panama city, Florida (Desenclos et al, 1991) and in Eastern Pennsylvania (Frace et al, 1991). The outbreaks of HAV usually occur in particular communities such as children's institutions (Gudkov et al, 1990), day-care centers (Reves et al, 1990), hospital employees (Takahashi et al, 1990) and army units (Yan et al, 1990).

In Thailand, HAV infection is one of the major public health problems. The prevalence in 1990 was 31 cases per 100,000 population (Ann Epidemiol Thai Surv Report, 1991). The infection that occurred in Thai children was mostly asymptomatic (Poovorawan et al, 1988). Detectable clinical cases are fewer than asymptomatic infected cases, allowing this the disease to spread rapidly.

The unfortunate endemic of HAV that occurred in Nakhon Si Thammarat, Southern Thailand, during May-June 1992, was 5 times the normal prevalence rate causing a serious health problem.

The objectives of this study were to study the clinical manifestations, risk factors of HAV infection and post serological survey in an outbreak at Nakhon Si Thammarat.

MATERIALS AND METHODS

Studied population

Primary school children from a school with hepatitis A outbreak were systematic randomly selected. One third of 269 (No. = 89) of the school students were included in the study. Demographic data of school children are shown in Table 1. The students were verbally interviewed on standards of hygiene, sanitation, history of personal and family illness during March to September 1992. The risk factors which included religion, family income, family size, presence of hepatitis illness and latrines in the family were compared between children with and without positive antiHAV.

Definition of clinical hepatitis A

A child with clinical hepatitis A was defined as an illness (during the period of an outbreak) with symptoms severe enough to cause school absence. The clinical symptoms included fever, malaise, fatigue, nausea, vomiting, abdominal discomfort enlarged and tender liver, dark urine and jaundice. The collected data were validated by counter checking with school records.

Serologic study

Blood samples were taken 3-4 months after an outbreak and collected at -20°C until testing. AntiHAV IgG was determined by using the ELISA technique (Abbott Laboratories, North Chicago, III). This technique was based on the principle of binding antiHAV in a patient's specimen (10 μ l) or antiHAV

Table 1
Demographic data on primary school children.

| Characteristics | | No. of children |
|-----------------|----------|-----------------|
| Sex | male | 44 |
| | female | 45 |
| Religion | Buddhism | 48 |
| | Muslim | 41 |

: horseradish peroxidase (antiHAV: HRPO) conjugate (200 μ l) to an HAV-coated bead. An HAV-coated bead was incubated (40°C, 3 hours, 15-30°C, 18-24 hours) in a reaction well with a mixture of serum or plasma and the antiHAV: HRPO. If antiHAV was present in the specimen, less antiHAV: HRPO would be bound to the bead. Within limits, the greater the amount of antiHAV in the specimen, the lower the absorbance (492 mm) value for that specimen. In case of results near the cut off point, the test were repeated again. AntiHAV IgM was also determined by ELISA test (Abbott Laboratories, North Chicago, III).

Statistical analysis

Percentage of immunological responses and the presence antiHAV for age-specific prevalence, ratio of symptomatic and asymptomatic patients with positive antiHAV IgM and range and average of liver enzyme levels (SGOT and SGPT) were calculated. The risk factors were analyzed by applying chi-square test.

RESULTS

During the two months of an outbreak, there were 36 cases of acute symptomatic hepatitis A. The prevalence was high among the age group between 8-11 years old. Clinical evidence included fever, malaise, fatigue, nausea, vomiting, abdominal discomfort, dark urine and jaundice. Fever was the principal symptom and jaundice was the second one of HAV infection (Table 2).

Table 2
Clinical manifestations of symptomatic children with hepatitis a virus infection.

| Signs/symptoms | No. of cases n = 36 cases | <u>%</u> |
|----------------------|------------------------------|----------|
| Fever | 33 | 97.22 |
| Malaise | 16 | 50 |
| Fatigue | 20 | 55.55 |
| Nausea/vomiting | 19 | 52.77 |
| Abdominal discomfort | 15 | 41.66 |
| Jaundice | 29 | 80.55 |
| icteric : anicteric | 4:1 | |

Among the patients with clinical hepatitis, the icteric and anicteric ratio was 4 to 1. Age specific serological prevalence of antiHAV in the school children studied after an endemic was shown in the Fig 1. Seventy of 89 children (79%) were positive for antiHAV post endemic. Sixteen were still antiHAV positive. Of the children with antiHAV IgM, 9 children had symptomatic hepatitis while 7 children had not. The symptomatic to asymtomatic ratio in a group with positive antiHAV IgM was 1.3:1. Liver enzyme levels (SGOT and SGPT) decreased to normal within four months. No complications or mortality occurred during the study period.

Interestingly, having no latrine, and occurrence of symptomatic hepatitis in the family were the significant risk factors in children with positive antiHAV (p < 0.01), while religion, family income, family size showed no significant difference (Table 3).

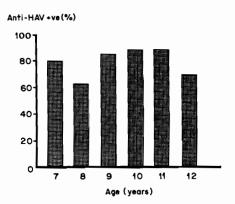


Fig 1- The scroprevalence of antiHAV in primary school children after an outbreak at Nakhon Si Thammarat.

Table 3

Risk factors for virus hepatitis A infections during an outbreak.

| Risk factors | X² | p-value |
|---------------------------------------|--------|----------|
| Religion | 0.022 | NS |
| Family income (3,390 + 1,537.72 baht) | 5.854 | NS |
| Family size $(5.42 + 1.25)$ | 3.215 | NS |
| Occurrence of patients in the family | 26.218 | p < 0.01 |
| No latrine | 9.968 | p < 0.01 |

DISCUSSION

Nakhon Si Thamrat, a large province in the center of the Southern part of Thailand, is an agricultural and fishing area. The total population is approximately 1.5 million. There were 3 endemic hepatitis outbreaks in the last decade. The first and second outbreaks arose in the year 1982 and 1989, the third one in 1992.

In the past endemic study, 79% of children were antiHAV detectable which included positive antiHAV IgM 18%. Only 21% of children were still susceptible to the disease. Because the serological study was done 3-4 months after an outbreak of the disease, therefore few patients had antiHAV IgM. In cases with antiHAV IgG or IgM, the patients had the same clinical features during this period, which suggested that they were infected by HAV during this outbreak. Without an outbreak, the seroprevalence of antiHAV among primary school children in Thailand was 25-50% (Poovorawan et al, 1991; Poovorawan et al, 1993). It clearly indicated that, prior to the outbreaks, the primary school children under 12 years old were susceptible to HAV infection.

Hepatitis A has been a major problem in many countries. Improvement of living standards have postponed the childhood infection into adulthood. HAV infection in early childhood is usually subclinical, while symptomatic hepatitis A infections occurs with increasing age (Hadler et al, 1980; Benenson et al, 1980). Our study showed that the ratio of symptomatic to asymptomatic hepatitis cases with positive IgM was 1.3 to 1.

The antiHAV prevalence in primary school Thai children with low socioeconomic status was seen in higher age group and markedly decreased in younger children from 1988-1993 as compared to the previous reports. Therefore endemicity is more likely because the majority of the population is susceptible to hepatitis A virus (Poovorawan et al, 1993).

In asymptomatic hepatitis A infection, the patients can spread the disease rapidly, so when there is an outbreak, people in the endemic area must be alerted to improve their personal hygiene and possibly undertake immunomodulations. In contrast, previous studies done on clinical manifestations reported that in younger children the ratio of symptomatic to asymptomatic was 1:4 (Hadler et al, 1980).

Clinical manifestations were similar to those reported by Rayman *et al* (1992). However, serological test is mandatory for confirming the HAV infection.

Among the risk factors of HAV infection, the occurrence of symptomatic hepatitis patients and no latrine in the family were the important ones which were similar to previous reports (Papaevangelou, 1984) because large amounts of HAV were in feces of the infected subjects (Castkoya et al, 1988). Thus members of the family or others, especially of those without latrine and with general poor standard hygiene might be infected by the contaminated virus in the feces. Additionally, the spread of the disease through everyday contact had been confirmed (Gudkov et al, 1990), particularly in the family with persons already infected by HAV.

Future incidents of this kind are likely to occur in Nakhon Si Thammarat. We would strongly recommend that latrine installation should be increased rapidly in time. Moreover, improvement of personal hygiene will assist the decline in disease transmission. Food consumption habits of people should be carefully considered, since other risk factors have been reported such as consumption of fresh products (Rosenblum et al, 1990; Ramsay et al, 1989) and half-cooked or raw seafood (Desenclos et al, 1991; Wang et al, 1990; Tang et al, 1991).

The changing incidence of age specific anti HAV prevalence refected the better living standard among those people, probably resulting in an increasing number of susceptible children. Therefore, preventive measures are essential in reducing the infection rate. Other than improvement of personal hygiene, mass vaccination is another promising preventive measure, particularly during an epidemic. Hepatitis A vaccine is widely used in several countries in Europe and Asia, particularly among travelers and soldiers entering hyperendemic countries. The vaccine is highly effective during an outbreak and when given as mass vaccination. However the cost effectiveness of the vaccine should be considered.

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REFERENCES

- Benenson MW, Takafuji ET, Bancroft WH, Lemon SM, Callahan MC, Leach DA. A military community outbreak of hepatitis type A related to transmission in a child care facility. Am J Epidemiol 1980; 112: 471-81.
- Castkoya J, Nemecek V, Dvorakova E, Snejdarova V, Helcl J. Family contacts of patients with viral hepatitis type A as potential source of hepatitis A virus. Cesk Epidemiol Mikrobiol Immunol 1988; 37: 193-8.
- Chitkara AJ, Sachdev HPS. Hepatitis vaccine. *Indian Pediatr* 1983; 25: 453-9.
- Desenclos JC, Klontz KC, Wilder MH, Naian OV, Margolis HS, Gunn RA. A multistate outbreak of hepatitis A caused by the consumption of raw oysters. Am J Public Health 1991; 81: 1268-72.
- Frace RM, Jahre JA. Policy for managing a community infectious disease outbreak. Infect Control Hosp Epidemiol 1991; 12:364-7.
- Gudkov VG, Rytik PG, Tomal LS, et al. Analysis of seasonal epidemic of viral hepatitis A in children's institution and ways for the improvement of sanitary and epidemiological surveillance on this infection among "organized" children. Zh Mikrobiol Epidemiol Immunobiol 1990; 3: 39-44.
- Gust ID. How an outbreak of hepatitis A in Melbourne led to the development of a vaccine. Vaccine 1992; 10: 59.
- Hadler SC, Webster HM, Erben JJ, Swanson JE, Maynard JE. Hepatitis A in day-care center: a community-wild assessment. N Engl J Med 1980; 302: 1222-7.
- Papaevangelou GJ. Global epidemiology of hepatitis A. In: Gerety RJ, ed. Hepatitis A. Orlando: Academic Press, 1984: 123.
- Poovorawan Y, Theamboonlers A, Pongpunlert W, Sakulramrung R, Chumdermpadetsuk S. The antibody to hepatitis A in secondary school children in Bangkok. Siriraj Hosp Gaz 1988; 40: 219-23.
- Poovorawan Y, Paiboonkasemsuthi S, Theamboonlers A, Kamolratanakul P, Chumdermpadetsuk S. Seroepidemiology of antibody to hepatitis A in the rural eastern part of Thailand. Southeast Asian J Trop Med Public Health 1991; 22: 35-8.

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- Poovorawan Y, Theamboonlers A, Chumdermpadetsuk S. Changing seroepidemiology of hepatitis A virus infection in Thailand. Southeast Asian J Trop Med Public Health 1993; 24: 250-4.
- Ramsay CN, Upton PA. Hepatitis A and frozen rasberries. Lancet 1989; 1:43-4.
- Raymond SK. Clinical manifestations and diagnosis of hepatitis A virus infection. J Vaccine 1992; 10: 15-7
- Reves RR, Pickering LK. Infections in child day care centers as they relate to internal medicine. Ann Rev Med 1990; 41: 383-91.
- Rosenblum LS, Mirkin IR, Allen DT, Safford S, Hadler SC. A mutifocal outbreak of hepatitis A traced to commercially distributed lettuce. Am J Public Health 1990; 80: 1075-9.
- Shaw FE Jr, Shapiro CN, Welty TK, Dill W, Reddington J, Hadler SC. Hepatitis transmission among the Sioux Indians of South Dakota. Am J Public Health 1990; 80: 1091-4.
- Stroffolini T, Biagini W, Lorenzoni L, Palazzesi GP, Divizia M, Frongillo R. An outbreak of hepatitis A in young

- adults in central Italy. Eur J Epidemiol 1990; 6: 156-9.
- Takahashi M, Yamada G, Tsuji T. Sporadic acute hepatitis in hospital employees: mainly non-A, non-B type. Acta Med Okayama 1990; 44: 315-20.
- Tang YW, Wang JX, Xu ZY, Guo YF, Qian WH, Xu JX. A serologically confirmed, case-control study, of a large outbreak of hepatitis A in China, associated with consumption of calms. *Epidemiol Infect* 1991; 107: 651-7.
- Wang JY, Hu SL, Liu HY, Hong YL, Cao SZ, Wu LF. Risk factor analysis of an epidemic of hepatitis A in a factory in Shanghai. *Int J Epidemiol* 1990; 19: 435-8.
- Xu ZY, Fu TY, Margolis HS, et al. Impact and control of viral hepatitis in China. In: Hollinger, FB, Lemon SH, Margolis HS, eds. Viral hepatitis and liver disease. Baltimore: Williams and Wilkinson 1991: 700-6.
- Yan YP, Xu DZ, Wang SS, Zhou H, Chen YJ. The seroepidemiologic investigation and analysis of acute hepatitis in an army unit. J Fourth Mil Med Univ 1990; 11: 32-4.