

**Table 1.** Amino acid sequences of the V3 region of HIV and antibody titers of HIV-infected subjects in Bangkok.

Patient no.	Amino acid sequence	Antibody titers by (ELISA)			Route of infection
		Thai A	Thai B	HGP-30	
1	CTRPSNNTRTSITIGPGQVFYRTGDIIGDIRKAYC	>6400	<200	200	Heterosexual intercourse
2	-----	1600	800	800	Heterosexual intercourse
3	-----	>6400	<200	200	Drug injection
4	-----	6400	400	<200	Heterosexual intercourse
5	-----P-----	>6400	400	1600	Heterosexual intercourse
6	-----A-----	6400	800	1600	Heterosexual intercourse
7	-----P-A-----	>6400	>6400	800	Heterosexual intercourse
8	-----P-A-----	6400	1600	200	Heterosexual intercourse
9	-----A-E-----	6400	3200	3200	Heterosexual intercourse
10	-----L-A-T-----	6400	3200	800	Heterosexual intercourse
11	-----V-A-T-N-----	6400	200	200	Drug injection
12	-----M-A-V-T-N-----	200	400	<200	Homosexual intercourse
13	-----LP-A-T-----	6400	400	400	Drug injection
14	-----M-V-N-----	6400	800	200	Heterosexual intercourse
15	-----Q-M-V-T-----	3200	200	1600	Heterosexual intercourse
16	CTRPNNTRKSIHLGPGQAWYTTGQIIGDIRQAHC	800	3200	800	Heterosexual intercourse
17	-----	3200	6400	400	Heterosexual intercourse
18	-----	1600	3200	800	Heterosexual intercourse
19	-----	3200	>6400	1600	Heterosexual intercourse
20	-----V-F-D-----	1600	3200	800	Heterosexual intercourse
21	-----G-V-R-----	800	3200	800	Heterosexual intercourse

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#### Control of a Hepatitis A Outbreak by Active Immunization of High-Risk Susceptible Subjects

**Colleagues**—Hepatitis A is an enterically transmitted disease whose prevalence is related to economic development. In Thailand, which is rapidly developing into an industrialized nation,

All subjects gave informed consent, and the study was approved by the ethical committee of Chulalongkorn University, Bangkok.

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there has been a rapid decline in anti-hepatitis A virus (HAV) seroprevalence among persons <20 years old, rendering a substantial percentage susceptible to disease [1, 2]. This has resulted in numerous recent outbreaks of hepatitis A among young Thai adults. Control of such outbreaks has previously depended on the use of immune globulin, which provides only short-term protection and no public health benefit. Therefore, we investigated the feasibility of actively vaccinating seronegative susceptible subjects as a means to control hepatitis A. The vaccine selected for use was a novel "virosome" formulation that has previously been found to elicit rapid seroconversion (97%-100% by 14 days after immunization) after a single dose [3].

An outbreak of hepatitis A occurred among a class of 61 radiology students at Chulalongkorn Hospital. The index case became infected after exposure as part of her clinical duties to a child with acute disease. About 2 weeks later, two classmates presented with fever, anorexia, and jaundice. Both were diagnosed as having hepatitis A by a positive anti-HAV IgM antibody test (Abbott Laboratories, Abbott Park, IL). The remaining students were immediately tested for anti-HAV antibodies with 26 (13 male, 13 female; mean age, 19.5 years) found to be seronegative and therefore susceptible.

All 26 subjects received a single intramuscular dose of vaccine. Mild, transient local reactions, characterized by tenderness at the injection site, were reported by 42% of the vaccinees. The immune response was measured by an IgG-specific ELISA [3]. All subjects were seropositive ( $\geq 20$  mIU/mL) at 1 and 6 months after vaccination, with geometric mean titers of 432 and 463 mIU/mL, respectively. There was no clinical evidence of active hepatitis A in this group. Liver function test results remained within normal ranges over the 6-month observation period. Therefore, the use of a hepatitis A vaccine in high-risk subjects appeared to be effective at preventing disease spread.

Hepatitis A vaccines have recently become commercially

available in several countries. On the basis of these promising results, their use, either alone or in combination with immune globulin, should be considered as a means to control outbreaks or epidemics of hepatitis A.

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### Hepatitis E Virus and Posttransfusion Hepatitis

**Colleagues**—Since the cloning of hepatitis E virus (HEV) and the development of an ELISA [1], many seroepidemiologic studies have been reported [2-4]. Recent studies of human volunteer and animal experiments have revealed viremic status [5, 6]. Therefore, the possibility of transfusion-related HEV infection has been considered [5]. In a prospective study of posttransfusion hepatitis (PTH) in Taiwan, we have encountered 6 patients with non-A, non-B, non-C hepatitis [7]. Accordingly, we used the HEV nested polymerase chain reaction (PCR) and an ELISA to study the role of HEV in these patients.

Six patients developed PTH in a prospective study [7]. They received blood or blood components from donors negative for hepatitis B virus (HBV) surface antigen and with serum alanine aminotransferase (ALT)  $< 45$  IU/L [7]. They were negative by serology for IgM antibodies to the following: hepatitis A virus, HBV core antigen, cytomegalovirus, Epstein-Barr virus, hepatitis C virus (HCV), and hepatitis D virus. Sera were also negative for HBV DNA and HCV RNA by PCR [7]. Subjects were natives of Taiwan and had no history of traveling abroad prior to disease onset. There were 4 men and 2 women with mean age of 42 years (range, 20-68), mean peak ALT of 286 IU/L (range, 102-537), and mean incubation period of 63 days (range, 15-126). All patients recovered within 3 months, and none had evidence of chronicity in 3-5 years of follow-up. Serial serum

samples of these 6 patients were tested in duplicate for HEV antibodies by ELISA (Diagnostic Biotechnology, Singapore) [3] and for HEV RNA by PCR. RNA from 100  $\mu$ L of serum was extracted [7], reverse-transcribed into cDNA, and amplified by nested PCR using primers as previously described [8].

All 6 patients were negative for HEV RNA by PCR. By ELISA, all the pretransfusion samples were negative. However, positive results were found in 2 sera of a patient (6 and 9 months after transfusion) and in 3 sera of another patient (3, 4, and 8 months after transfusion). The cutoff values of these 5 samples were between 1.8 and 3.4. The other serial specimens were negative.

Transmission of HEV has been documented by the fecal-oral route [8]. Although the possibility of HEV infection resulting from transfusion of HEV-containing blood has been raised, screening of ALT should reduce the possibility, as HEV viremia appears only at the acute stage of infection [5, 6]. Therefore, it was reasonable that none of our patients were infected with HEV. This might be another reason for ALT screening after antibodies to HCV have appeared. It is now well known that anti-HEV IgG is long-lasting once it appears [2, 5, 6]. In the 5 seropositive samples, because of the negative PCR results and because the appearance of HEV antibody was not consistent, these results were considered to be falsely positive. Therefore, the interpretation of current HEV ELISA results should be cautious as to clinical and seroepidemiologic diagnosis until confirmatory tests are available.

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