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Vol. 45/No. 33, August 17, 1985 TEXAS STATE DOCUMEN

Texas Preventable Disease

contents:

- BUREAU OF EPIDEMIOLOGY

1100 West 49th Street, Austin, Texas 78756 (512-458-7207) -

Non-B Hepatitis

ACIP: Recommendations for Protection Against Viral Hepatitis A and Non-A,

ACIP:

RECOMMENDATIONS FOR PROTECTION AGAINST VIRAL HEPATITIS A AND NON-A, NON-B HEPATITIS

The following recommendations from the Immunization Practices Advisory Committee are excerpted from MMWR, Vol.34/No.22, June 7, 1985, and update all previous recommendations on use of immune globulins for protection against viral hepatitis A. Please refer to the above issue of MMWR for recommendations regarding the use of hepatitis B vaccine and hepatitis B immune globulin for prophylaxis of hepatitis B.

#### INTRODUCTION

The term "viral hepatitis" is commonly used for several clinically similar diseases that are etiologically and epidemiologically distinct. Two of these, hepatitis A (formally called infectious hepatitis) and hepatitis B (formally called serum hepatitis) have been recognized as separate entities since the early 1940s and can be diagnosed with specific serologic tests. The third, currently known as non-A, non-B hepatitis, is probably caused by at least two different agents, and lacking specific diagnostic tests, remains a disease diagnosed by exclusion. It is an important form of acute viral hepatitis in adults and currently accounts for most post-transfusion hepatitis in the United States. An epidemic type of non-A, non-B hepatitis, which is probably spread by the fecal-oral route and is different from the types seen in the United States, has been described in parts of Asia and North Africa.

A fourth type of hepatitis, delta hepatitis, has recently been characterized as an infection dependent on hepatitis B virus. It may occur as a coinfection with acute hepatitis B infection or as a superinfection of a hepatitis B carrier.

## HEPATITIS SURVEILLANCE

Approximately 21,500 cases of hepatitis A, 24,300 cases of hepatitis B, 3,500 cases of non-A, non-B hepatitis, and 7,100 cases of hepatitis type unspecified were reported in the United States in 1983. Most cases of each type occur among young adults. Since reporting from many localities is incomplete, the actual number of hepatitis cases occurring annually is thought to be several times the reported number.

### IMMUNE GLOBULINS

Immune globulins used in medical practice are sterile solutions of antibodies (immuno-globulins) from human plasma. They are prepared by cold ethanol fractionation of large plasma pools and contain 10% to 18% protein. In the United States, plasma is primarily obtained from professional donors. Only plasma shown to be free of hepatitis B surface antigen (HBsAg) is used to prepare immune globulins.

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Immune globulin (IG) produced in the United States contains antibodies against the hepatitis A virus (anti-HAV) and the hepatitis B surface antigen (anti-HBs). Tests of IG lots prepared since 1977 indicate that both types of antibody have been present uniformly. Hepatitis B immune globulin (HBIG) is an IG prepared from plasma containing high titers of anti-HBs.

Neither IG nor HBIG commercially available in the United States transmits hepatitis or other viral infections. There is no evidence that the causative agent of AIDS (human T-lymphotropic virus type III/lymphadenopathy-associated virus [HTLV-III/LAV]) has been transmitted by IG or HBIG.

Serious adverse effects from immune globulins administered as recommended have been exceedingly rare. Standard immune globulins are prepared for intramuscular use and should not be given intravenously. Two preparations for intravenous use in immunodeficient and other selected patients have recently become available in the United States but are not recommended for hepatitis prophylaxis. Immune globulins are not contraindicated for pregnant women.

HEPATITIS A

Hepatitis A is caused by the hepatitis A virus (HAV), a 27-nm ribonucleic acid (RNA) agent that is a member of the picornavirus family. The illness caused by HAV characteristically has an abrupt onset with fever, malaise, anorexia, nausea, abdominal discomfort, and jaundice. Severity is related to age. In children, most infections are asymptomatic, and illness is usually not accompanied by jaundice. Most infected adults become symptomatically ill with jaundice. Fatality among reported cases is infrequent (about 0.6%).

Hepatitis A is primarily transmitted by person-to-person contact, generally through fecal contamination. Transmission is facilitated by poor personal hygiene, poor sanitation, and intimate (intrahousehold or sexual) contact. Common-source epidemics from contaminated food and water also occur. Sharing utensils or cigarettes or kissing are not believed to transmit the infection.

The incubation period of hepatitis A is 15 to 50 days (average 28 to 30). High concentrations of HAV ( $10^8$  particles/g) are found in stools of infected persons. Fecal virus excretion reaches its highest concentration late in the incubation period and early in the prodromal phase of illness, and diminishes rapidly once jaundice appears. Greatest infectivity is during the two-week period immediately before the onset of jaundice. Viremia is of short duration; virus has not been found in urine or other body fluids. A chronic carrier state with HAV in blood or feces has not been demonstrated. Transmission of HAV by blood transfusion has occurred but is rare.

The diagnosis of acute hepatitis A is confirmed by finding IgM-class anti-HAV in serum collected during the acute or early convalescent phase of disease. IgG-class anti-HAV, which appears in the convalescent phase of disease and remains detectable in serum thereafter, apparently confers enduring protection against disease. Commercial tests are available to detect IgM anti-HAV and total anti-HAV in serum.

Although the incidence of hepatitis A in the United States has decreased over the last 15 years, it is still a common infection in older children and young adults. About 38% of reported hepatitis cases in this country are attributable to hepatitis A.

Recommendations for IG prophylaxis of hepatitis A. Numerous field studies conducted in the past four decades confirm that IG given before exposure or during the incubation period of hepatitis A is protective against clinical illness. Its 5

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prophylactic value is greatest (80% to 90%) when given early in the incubation period and declines thereafter.

Preexposure prophylaxis. The major group for whom preexposure prophylaxis is recommended is international travelers. The risk of hepatitis A for US citizens traveling abroad varies with living conditions, incidence of hepatitis A infection in areas visited, and length of stay. In general, travelers to developed areas of western Europe, Japan, and Australia are at no greater risk of infection than in the United States. In contrast, travelers to developing countries may be at significant risk of infection. In such areas, the best way to prevent hepatitis A and other enteric diseases is to avoid potentially contaminated water or food. Drinking water (or beverages with ice) of unknown purity and eating uncooked shellfish or uncooked fruits or vegetables that are not peeled (or prepared) by the traveler should be avoided.

IG is recommended for travelers to developing countries if they will be eating in settings of poor or uncertain sanitation (some restaurants or homes) or will be visiting extensively with local persons, especially young children, in settings with poor sanitary conditions. Persons who plan to reside in developing areas for long periods should receive IG regularly if they anticipate exposure as described above or will be living in rural areas with poor sanitation.

- For such travelers, a single dose of IG (0.02 ml/kg) is recommended if travel is for less than 2 months. For prolonged travel, 0.06 ml/kg should be given every 5 months. For persons who require repeated IG prophylaxis, screening for total anti-HAV antibodies before travel may be useful to define susceptibility and eliminate unnecessary doses of IG in those who are immune.
- Postexposure prophylaxis. A serologic test for the diagnosis of acute hepatitis A is now widely available. Since only 38% of acute hepatitis cases in the United States result from hepatitis A, serologic confirmation of hepatitis A in the index case is recommended before treatment of contacts. Serologic screening of contacts for anti HAV before giving IG is not recommended because screening is more costly than IG and would delay its administration.
- IG should be given as soon as possible after exposure; giving IG more than 2 weeks after exposure is not indicated.
- Specific recommendations for IG prophylaxis of hepatitis A depend on the nature of the HAV exposure:
  - 1. Close personal contact. IG is recommended for all household and sexual contacts of persons with hepatitis A.
  - 2. Day-care centers. Day-care facilities with children in diapers can be important settings for HAV transmission. IG should be administered to all staff and attendees of day-care centers or homes if: a) one or more hepatitis A cases are recognized among children or employees, or b) cases are recognized in two or more households of center attendees. When an outbreak (hepatitis cases in three or more families) occurs, IG should also be considered for members of households whose diapered children attend. In centers not enrolling children in diapers, IG need only be given to classroom contacts of an index case.
  - 3. Schools. Contact at elementary and secondary schools is usually not an important means of transmitting hepatitis A. Routine administration of IG is not indicated for pupils and teachers in contact with a patient. However, when epidemiologic study clearly shows the existence of a

school- or classroom-centered outbreak, IG may be given to those who have close personal contact with patients.

- Institutions for custodial care. Living conditions in some 4. institutions, such as prisons and facilities for the developmentally disabled, favor transmission of hepatitis A. When outbreaks occur, giving IG to residents and staff who have close contact with patients with hepatitis A may reduce the spread of disease. Depending on the epidemiologic circumstances, prophylaxis can be limited in extent or can involve the entire institution.
- Routine IG prophylaxis for hospital personnel is not 5. Hospitals. indicated. Rather, sound hygienic practices should be emphasized. Staff education should point out the risk of exposure to hepatitis A and emphasize precautions regarding direct contact with potentially infective materials.
- Offices and factories. Routine IG administration is not indicated 6. under the usual office or factory conditions for persons exposed to a fellow worker with hepatitis A. Experience shows that casual contact in the work setting does not result in viral transmission.
- 7. Common-source exposure. IG might be effective in preventing foodborne or waterborne hepatitis A if exposure is recognized in time. However, IG is not recommended for persons exposed to a common source of hepatitis infection after cases have begun to occur in those exposed, since the two-week period during which IG is effective will have been exceeded.

If a foodhandler is diagnosed as having hepatitis A, common-source transmission is possible but uncommon. IG should be administered to other foodhandlers but is usually not recommended for patrons. However, IG administration to patrons may be considered if: a) the infected person is directly involved in handling, without gloves, foods that will not be cooked before they are eaten; b) the hygienic practices of the foodhandler are deficient; and c) patrons can be identified and treated within two weeks of exposure. Situations where repeated exposures may have occurred, such as in institutional cafeterias, may warrant stronger consideration of IG use.

For postexposure IG prophylaxis, a single intramuscular dose of 0.02 ml/kg is recommended.

### NON-A. NON-B HEPATITIS

United States. Non-A, non-B hepatitis that presently occurs in the United States has epidemiologic characteristics similar to those of hepatitis B, occurring most commonly following blood transfusion and parenteral drug abuse. Multiple episodes of non-A, non-B hepatitis have been observed in the same individuals and may be due to different agents. Chronic hepatitis following acute non-A, non-B hepatitis infection varies in frequency from 20% to 70%. Experimental studies in chimpanzees have confirmed the existence of a carrier state, which may be present in up to 8% of the population.

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Table 1. Hepatitis nomenclature.

Abbreviation	Term	Comments
	Hepatit	is A
HAV	Hepatitis A virus	Etiologic agent of "infectious"
		hepatitis; a picornavirus; single
	· · · · · · · · · · · · · · · · · · ·	serotype.
Anti-HAV	Antibody to HAV	Detectable at onset of symptoms;
		lifetime persistence.
lgM anti-HAV	IgM class antibody	Indicates recent infection with
	to HAV	hepatitis A; positive up to 4-6 months
		after infection.
<u> </u>	Hepatit	is B
HBV	Hepatitis B virus	Etiologic agent of "serum" or "long-
		incubation" hepatitis; also known as
		Dane particle.
HBsAg	Hepatitis B surface	Surface antigen(s) of HBV detectable in
-	antigen	large quantity in serum: several
	-	subtypes identified.
-BeAg	Hepatitis B e antigen	Soluble antigen: correlates with HBV
		replication, high titer HRV in serum
		and infectivity of serum
HBcAa	Henatitis B core antigen	No commercial test available
Anti-HBs	Antibody to HBsAg	Indicates past infection with and
	Antibody to hoshg	immunity to UPV pageive antibody from
		BRIC of impune seconds from URV
		Hold, or immune response from Hov
Anti-HBe	Antibody to HBEAg	Presence in serum of HBsAg carrier
• • • • • •		suggests lower titer of HBV.
Anti-HBc	Antibody to HBcAg	Indicates past infection with HBV at
		some undefined time.
lgM anti-HBc	lgM class antibody to	Indicates recent infection with HBV;
	HBcAg	positive for 4-6 months after
		infection.
	Delta hen	atitis
S virus	Delta virus	Etiologic agent of delta hepatitis; may
		only cause infection in presence of
		HBV.
s —Aa	Delta antigen	Detectable in early acute delta
, ··· <b>j</b>		infection
Anti-S	Antibody to delta	Indicates past or present infection
	antigen	with delta virus
	antigen	with derta virus.
	Non-A, non-B	hepatitis
NANB	Non-A, non-B hepatitis	Diagnosis of exclusion. At least two
		candidate viruses; epidemiology
		parallels that of hepatitis B.
	Epidemic non-A, n	on-B hepatitis
Epidemic NANB	Epidemic non-A. non-B	Causes large epidemics in Asia. North
•	hepatitis	Africa: fecal-oral or waterborne.
18-18-19-19-19-19-19-1-1-1-1-1-1-1-1-1-1	·	
16	Immune globulin	builds Contains antibodies to HAV low titer
10	(proviously ISC immune	antibodies to URV
	(previously isu, immune	AILIDULIES LO HDV.
	serum giobulin, or gamma	
	globulin)	
HBIG	Hepatitis B immune	Contains high titer antibodies to HBV.
	globulin	

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Although several studies have attempted to assess the value of prophylaxis with IG against non-A, non-B hepatitis, the results have been equivocal, and no specific recommendations can be made. However, for persons with percutaneous exposure to blood from a patient with non-A, non-B hepatitis, it may be reasonable to administer IG (0.06 ml/kg) as soon as possible after exposure.

Epidemic (fecal-oral) non-A, non-B hepatitis. In recent years, epidemics of non-A, non-B hepatitis spread by water or close personal contact have been reported from several areas of Southeast Asia (Indian subcontinent, Burma) and north Africa. Such epidemics generally affect adults and cause unusually high mortality in pregnant women. The disease has been transmitted to experimental animals, and candidate viruses have been identified; however, no serologic tests have yet been developed.

Epidemic non-A, non-B hepatitis has not been recognized in the United States or western Europe, and it is unknown whether the causative agent is present in these areas.

Travelers to areas having epidemic non-A, non-B hepatitis may be at some risk of acquiring this disease by close contact or by contaminated food or water. The value of IG in preventing this infection is unknown. The best prevention of infection is to avoid potentially contaminated food or water, as with hepatitis A and other enteric infections.

TEXAS PREVENTABLE DISEASE NEWS (ISSN 8750-9474) is a free, weekly publication of the Texas Department of Health, \* 1100 West 49th Street, Austin, TX, 78756-3180. Second-class postage paid at Austin, TX. POSTMASTER: Send address changes to TEXAS PREVENTABLE DISEASE NEWS, 1100 West 49th Street, Austin, TX 78756-3180.

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