Risk and Management of Blood-Borne Infections in Health Care Workers

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INTRODUCTION

Exposure to blood-borne pathogens poses a serious risk to health care workers (HCWs). Transmission of at least 20 different pathogens by needlestick and sharps injuries has been reported (79). Despite improved methods of preventing exposure, occupational exposures will continue to occur.

Assessment of the risk of blood-borne pathogen transmission in the health care setting requires information derived from various sources, including surveillance data, studies of the frequency and preventability of blood contacts, seroprevalence studies among patients and HCWs, and prospective studies that assess the risk of seroconversion after an exposure to infected blood. Factors influencing the risk to an individual HCW over a lifetime career include the number and types of blood contact experienced by the worker, the prevalence of blood-borne pathogen infection among patients treated by the worker, and the risk of transmission of infection after a single blood contact.

In this article, we review the risk and management of the three blood-borne viruses most commonly involved in occupational transmission: human immunodeficiency virus (HIV), hepatitis B virus (HBV), and hepatitis C virus (HCV). We also will discuss current methods of preventing exposure, including standard precautions and the use of safety devices in the health care setting, as well as recommendations for postexposure prophylaxis.

TRANSMISSION OF BLOOD-BORNE PATHOGENS IN THE HEALTH CARE SETTING

Modes of Blood-Borne Pathogen Transmission

In the health care setting, blood-borne pathogen transmission occurs predominantly by percutaneous or mucosal exposure of workers to the blood or body fluids of infected patients. Occupational exposures that may result in HIV, HBV, or HCV transmission include needlestick and other sharps injuries; direct inoculation of virus into cutaneous scratches, skin lesions, abrasions, or burns; and inoculation of virus onto mucosal surfaces of the eyes, nose, or mouth through accidental splashes. HIV, HBV, and HCV do not spontaneously penetrate intact skin, and airborne transmission of these viruses does not occur.

Epidemiology of Blood Contact

To understand the nature, frequency, and prevention of percutaneous injuries and mucocutaneous blood contacts among HCWs, prospective observational studies have been performed in different patient care settings (Table 1). The percentage of procedures with at least one blood contact of any type ranged from 3% of procedures performed by invasive radiology personnel in a study in Dallas, Tex. (130), to 50% of procedures performed by surgeons in a study in Milwaukee, Wisc. (224). The percentage of procedures with at least one injury caused by a sharp instrument also varied widely, from 0.1 to 15%. These differences may be related to variations in study methods, procedures observed, and precautions used by the workers performing the procedures.

Several of these studies assessed specific risk factors for injury or exposure. For example, of the 99 percutaneous injuries observed by Tokars et al. during 1,382 operations in five different surgical specialties (general, orthopedic, gynecologic, trauma, and cardiac), most (73%) were related to suturing (256). Rates were highest (10%) during gynecologic surgeries (256). Panlilio et al. found in their study of blood contacts during surgery that risk factors for blood contacts by surgeons included performing an emergency procedure, patient blood loss greater than 250 ml, and surgery duration greater than 1 h.
Distant (6 months; resolved). Among patients with known exposure dates, the estimated median time from initial infection to the development of detectable antibody is 2.4 months; 95% of individuals develop antibodies within 6 months of infection (34). In persons who are seroconverted to HIV, 5% tested negative for HIV antibodies at > 6 months after their occupational exposure but were seropositive within 12 months (73). The two antibody tests commonly used to detect HIV are the enzyme immunoassay (EIA) and the Western blot. An HIV test result is reported as negative when the EIA result is negative. The result is reported as positive when the EIA result is repeatedly reactive and when the result of a more specific, supplemental confirmatory test, such as the Western blot, is also positive. Once an individual develops an antibody response, it usually remains detectable for life. HIV infection for longer than 6 months without detectable antibody is uncommon (73, 226).

Direct virus assays (e.g., PCR for HIV RNA) are sensitive methods for the detection of HIV infection. However, problems with laboratory contamination, false-positive rates, and increased costs limit their routine use. While PCR for HIV RNA is approved for use in established HIV infection, its reliability in detecting very early infection has not been determined (34). At present, the false-positive and false-negative rates of PCR are too high to warrant a broader role for it in routine postexposure management (207).

Detection and Diagnosis of HBV Infection

The incubation period for acute hepatitis B ranges from 45 to 160 days, with an average of 120 days. Exposure to HBV can lead to an acute infection which may result in a chronic infection. Acute hepatitis B resembles other forms of viral hepatitis and cannot be distinguished based on history, physical examination, or serum biochemical tests.

The diagnosis of acute HBV infection is confirmed by the demonstration in serum of hepatitis B surface antigen (HBsAg), which appears well before onset of symptoms and before development of antibody to hepatitis B core antigen (anti-HBc), and immunoglobulin M (IgM) antibody to HBc, which appear at approximately the same time as symptoms (143). The presence of IgM anti-HBc indicates recent HBV infection, usually within the preceding 4 to 6 months. The presence of hepatitis B e antigen (HBeAg) in serum correlates with HBV replication, high titers of HBV, and infectivity. Persons who are positive for HBeAg typically have 10^6 to 10^9 HBV particles per ml of blood (243). In persons who resolve acute HBV infection, antibody to HBsAg (anti-HBs) develops and indicates immunity. The persistence of HBsAg for 6 months after the diagnosis of acute HBV is indicative of progression to chronic HBV infection.

HBV serologic markers in different stages of infection and convalescence are summarized in Table 2. Anti-HBc indicates prior infection and lasts indefinitely. In persons who

**TABLE 2. HBV serologic markers in different stages of infection and convalescence (201a)**

<table>
<thead>
<tr>
<th>Stage of infection</th>
<th>HBsAg</th>
<th>Anti-HBs</th>
<th>Anti-HBc</th>
<th>HBeAg</th>
<th>Anti-HBc</th>
</tr>
</thead>
<tbody>
<tr>
<td>Late incubation period</td>
<td>+</td>
<td>-</td>
<td>-</td>
<td>+</td>
<td>-</td>
</tr>
<tr>
<td>Acute hepatitis B</td>
<td>+</td>
<td>-</td>
<td>+</td>
<td>+</td>
<td>-</td>
</tr>
<tr>
<td>HBsAg carrier</td>
<td>+</td>
<td>- (+ rarely)</td>
<td>+</td>
<td>+</td>
<td>-</td>
</tr>
<tr>
<td>Recent (&lt;6 months; resolved infection)</td>
<td>-</td>
<td>++</td>
<td>+</td>
<td>-</td>
<td>+ or -</td>
</tr>
<tr>
<td>Distant (&gt;6 months; resolved infection)</td>
<td>-</td>
<td>++</td>
<td>+</td>
<td>-</td>
<td>+ or -</td>
</tr>
<tr>
<td>Vaccinated</td>
<td>-</td>
<td>++</td>
<td>-</td>
<td>-</td>
<td>-</td>
</tr>
</tbody>
</table>

* a. positive; ++, strongly positive; ++++, very strongly positive; + or - , variable reaction; -, negative.
* b. The total anti-HBc assay detects both IgM and IgG antibody.
* c. Resolved, the patient no longer has the disease.

In their study of dental procedures, Cleveland et al. found that most percutaneous injuries sustained by dental residents occurred extraorally and were associated with denture impression procedures (77).

Retrospective studies and surveys have also shown high rates of blood contact among HCWs in different patient care settings. Tokars et al. found that among 3,420 participants at the American Academy of Orthopaedic Surgeons annual meeting, 87.4% of surgeons surveyed reported a blood-skin contact and 39.2% reported a percutaneous blood contact in the previous month (258). In a retrospective survey by O’Briain in 1991 (202), 56% of 36 resident and staff pathologists reported that they had sustained a cut or needlestick injury in the preceding year. In this study, pathologists reported 72 injuries, corresponding to a rate of one injury for every 37 autopsies performed and one injury for every 2,629 surgical specimens handled (202). An anonymous national survey of certified nurse-midwives by Willy et al. found that 74% had soiled their hands with blood, 51% had splashed blood or amniotic fluid in their faces, and 24% had sustained one or more needlestick injuries in the preceding 6 months (281). Among 550 medical students and residents in Los Angeles, Calif., who were surveyed anonymously by O’Neill et al., 71% reported exposures to patients’ blood and body fluids during the preceding year (204). In a recent study of third- and fourth-year medical students in San Francisco, Calif., by Osborn et al., 12% reported an exposure to infectious body substances over the 7-year study period, from 1990 to 1996 (205). There is evidence among some groups of HCWs, such as dentists, that rates of exposure are decreasing over time, temporally associated with increased awareness and compliance with the practice of standard precautions (76).

**DETECTION AND DIAGNOSIS OF BLOOD-BORNE PATHOGEN INFECTIONS**

An understanding of the detection and diagnosis of HIV, HBV, and HCV infection is vital for the appropriate management and care of HCWs exposed to or infected with blood-borne viruses.

**Detection and Diagnosis of HIV Infection**

After initial primary infection with HIV, there is a window period prior to the development of detectable antibody. In persons with known exposure dates, the estimated median time from initial infection to the development of detectable antibody is 2.4 months; 95% of individuals develop antibodies within 6 months of infection (34). Among HCWs with a documented seroconversion to HIV, 5% tested negative for HIV antibodies at >6 months after their occupational exposure but were seropositive within 12 months (73). The two antibody tests commonly used to detect HIV are the enzyme immunoassay (EIA) and the Western blot. An HIV test result is reported as negative when the EIA result is negative. The result is reported as positive when the EIA result is repeatedly reactive and when the result of a more specific, supplemental confirmatory test, such as the Western blot, is also positive. Once an individual develops an antibody response, it usually remains detectable for life. HIV infection for longer than 6 months without detectable antibody is uncommon (73, 226).

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HBV serologic markers in different stages of infection and convalescence are summarized in Table 2. Anti-HBc indicates prior infection and lasts indefinitely. In persons who
respond to the hepatitis B vaccine, anti-HBs is the only antibody that is elicited. Persons with chronic infection who have mutations in the precore region of the HBV genome that prevent the expression of HBeAg but allow the expression of infectious virus have been described (40, 260). High titers of HBsAg can be observed in these persons even though they are HBeAg negative. The prevalence of these precore mutations in persons in the United States is unknown. The prevalence may be relatively high in certain parts of the world (41, 124, 171, 173, 197).

Detection and Diagnosis of HCV Infection

The incubation period for acute HCV infection ranges from 2 to 24 weeks, with an average of 6 to 7 weeks (166, 179; L. B. Seef, Letter, Ann. Intern. Med. 115:411, 1991). Because different types of viral hepatitis are indistinguishable based on clinical symptoms alone, serologic testing (Table 3) is necessary to establish a specific diagnosis of hepatitis C (121). Screening EIA and supplemental immunoblot assays are licensed and commercially available to detect antibodies to HCV (anti-HCV) (283). Because the rate of false positivity for the screening EIA is high in many populations, including HCs, supplemental immunoblot assays must be used to judge the validity of repeatedly reactive EIA results. Anti-HCV may be detected within 5 to 6 weeks after the onset of infection and remains detectable long after the primary infection. In general, the interpretation of serologic tests for anti-HCV is limited by the following factors: (i) assays for anti-HCV do not distinguish between acute, chronic, or past infection; (ii) in acute infection there may be a prolonged interval between onset of illness and anti-HCV seroconversion (though most infected individuals seroconvert within 3 months of exposure); and (iii) the detection of anti-HCV does not necessarily indicate active HCV replication (8).

HCV RNA can be detected in serum or plasma within 1 to 2 weeks of exposure to the virus and several weeks before onset of alanine aminotransferase (ALT) elevations or the appearance of anti-HCV (103). In patients with chronic HCV infection, HCV RNA levels may remain relatively stable or can fluctuate over 1,000,000-fold. Fluctuations in HCV RNA may or may not correlate with elevations in transaminase levels. Rarely, the detection of HCV RNA may be the only evidence of HCV infection (14).

PCR techniques to amplify reverse-transcribed cDNA are currently the most sensitive methods for detecting HCV RNA. Both qualitative (122) and quantitative (87, 229) methods can be used to detect HCV RNA. Quantitative assays are less sensitive than qualitative assays and should not be used as a primary test to confirm or exclude the diagnosis of HCV infection (212). Currently, testing for HCV RNA is available on a research basis and no tests have been approved by the U.S. Food and Drug Administration. Because of assay variability, results of HCV RNA testing must be interpreted cautiously.

There are at least six different genotypes and more than 90 subtypes of HCV (33). About 70% of HCV-infected persons in the United States are infected with genotype 1; subtype 1a predominates over subtype 1b. Several different nucleic acid detection methods are commercially available to group isolates of HCV based on genotypes and subtypes (172).
RISK OF OCCUPATIONAL TRANSMISSION OF HIV FROM PATIENTS TO WORKERS

Risk of HIV Infection Postexposure

Prospective studies of HCWs have estimated that the average risk for HIV transmission after a percutaneous exposure to HIV-infected blood is approximately 0.3% (95% confidence interval = 0.2 to 0.5%) (25) and that after a mucous membrane exposure it is 0.09% (95% confidence interval = 0.006 to 0.5%) (147). The risk after a cutaneous exposure is less but has not been well quantified since no HCW enrolled in a prospective study has seroconverted after an isolated skin exposure. There are insufficient data to quantify the risk of transmission after occupational exposure to potentially infectious tissues or fluids other than blood. However, in a study by Fahey et al., none of 559 participants reporting cutaneous exposures to blood, sputum, urine, feces, or other body substances from patients presumed infected with HIV acquired HIV infection (102). There is also no evidence of a risk for HIV transmission by the aerosol route. Transmission of HIV by aerosol would require the generation of aerosolized particles of blood, the presence of infective HIV in these aerosolized particles, and the deposition of a sufficient number of infective particles in the respiratory tract or on the mucous membranes of a susceptible host to cause infection. Biological or epidemiologic evidence that HIV can be transmitted by aerosols via the respiratory route currently does not exist (22). Although not specifically designed to assess the possibility of aerosol transmission of HIV, the 1991 seroprevalence survey of attendees of the annual meeting of the American Academy of Orthopaedic Surgeons addressed this concern indirectly (258). There were 1,201 study participants without nonoccupational risk factors who had participated in procedures on patients with HIV infection or AIDS and had never used a “space suit” or other device to prevent inhalation of aerosols. Since power instruments are used frequently in orthopedic procedures, many of these participants may have been exposed to blood or tissue aerosols produced by these instruments; all were HIV seronegative (258).

The risk of HIV transmission after a percutaneous exposure appears to be influenced by several factors. To assess possible risk factors, the Centers for Disease Control and Prevention (CDC), in collaboration with international public health authorities, conducted a retrospective case-control study using data reported to national surveillance systems in the United States, France, Italy, and the United Kingdom. Based on logistic regression analysis, factors associated with HIV transmission after percutaneous exposure included a deep injury, a device visibly contaminated with the source patient’s blood, procedures involving a needle placed directly in the patient’s vein or artery, and a source patient who died from AIDS within 60 days of the exposure (39). The findings of the case-control study suggest that the risk for HIV infection likely exceeds 0.3% for percutaneous injuries involving a larger volume of blood and/or higher titer of HIV in the blood. Several laboratory studies support these findings. In vitro models have shown that increasing needle size and penetration depth are associated with increased blood transfer volume (182), that hollow-bore needles transfer greater volumes of blood than solid-suture needles, and that gloves reduce the amount of blood transferred (26). Studies also have shown that the level of infectious HIV present in the blood of most patients with symptomatic AIDS is significantly higher than the level present in patients with asymptomatic HIV infection (141). An additional finding of the case-control study was that postexposure use of zidovudine (ZDV) by HCWs was associated with a lower risk for HIV transmission (39). (This issue will be discussed in more detail in the section Postexposure Chemoprophylaxis for HIV [below]). It is also possible that host defense mechanisms influence the risk of HIV transmission. One study demonstrated an HIV-specific T-helper cellular immune response when peripheral blood mononuclear cells from a small number of HCWs exposed to HIV were stimulated in vitro by HIV. None of the HCWs seroconverted. One possible explanation for these observations is that host immune responses prevented establishment of HIV infection after exposure (75). Similar cytotoxic T-lymphocyte responses have been observed in other populations with repeated HIV exposure without resulting infection (70, 74, 160, 170, 225).

HIV Seroprevalence among Patients

In the United States, HIV seroprevalence rates vary widely by geographic area and patients’ demographic characteristics. The CDC’s Sentinel Hospital Surveillance System tested 195,829 anonymous patient blood samples at 20 hospitals in 15 cities between September 1989 and October 1991. The HIV seroprevalence at these institutions ranged from 0.2 to 14.2% and was highest among men aged 25 to 44 years and patients with infectious conditions (excluding symptomatic HIV infection) and drug-related conditions (153).

Similarly, seroprevalence data for unselected hospital admissions and for patients presenting to emergency departments, operating rooms, and obstetrical units have demonstrated considerable variation (Table 4). The lowest seroprevalence rates

---

Table 4. HIV seroprevalence in emergency, hospital, surgery, and obstetrics patients

<table>
<thead>
<tr>
<th>Authors (reference)</th>
<th>Yr</th>
<th>Setting</th>
<th>Location</th>
<th>No. of patients tested</th>
<th>No. of patients HIV positive (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Kelen et al. (158)</td>
<td>1987</td>
<td>Emergency department</td>
<td>Baltimore, Md.</td>
<td>2,302</td>
<td>119 (5.2)</td>
</tr>
<tr>
<td>Kelen et al. (157)</td>
<td>1988</td>
<td>Emergency department</td>
<td>Baltimore, Md.</td>
<td>254</td>
<td>152 (6.0)</td>
</tr>
<tr>
<td>Marcus et al. (178)</td>
<td>1989</td>
<td>Emergency department</td>
<td>Six high-AIDS areas</td>
<td>20,382</td>
<td></td>
</tr>
<tr>
<td>Nagachinta et al. (191)</td>
<td>1990</td>
<td>Emergency department</td>
<td>Los Angeles, Calif.</td>
<td>1,945</td>
<td>40 (2.1)</td>
</tr>
<tr>
<td>Mullins and Harrison (190)</td>
<td>1987–1991</td>
<td>Trauma center</td>
<td>Wichita, Kans.</td>
<td>2,004</td>
<td>3 (0.15)</td>
</tr>
<tr>
<td>Gordin et al. (119)</td>
<td>1987</td>
<td>Hospital</td>
<td>Washington, D.C.</td>
<td>616</td>
<td>23 (3.7)</td>
</tr>
<tr>
<td>Trepa et al. (251)</td>
<td>1993</td>
<td>Hospital</td>
<td>Denver, Colo.</td>
<td>2,825</td>
<td>155 (5.5)</td>
</tr>
<tr>
<td>Charache et al. (68)</td>
<td>1989</td>
<td>Elective surgery</td>
<td>Baltimore, Md.</td>
<td>4,087</td>
<td>18 (0.4)</td>
</tr>
<tr>
<td>Montecalvo et al. (187)</td>
<td>1992</td>
<td>Surgery-obstetrics</td>
<td>Valhalla, N.Y.</td>
<td>1,056</td>
<td>15 (1.4)</td>
</tr>
<tr>
<td>Kransinski et al.</td>
<td>1986–1987</td>
<td>Obstetrics</td>
<td>New York, N.Y.</td>
<td>1,192</td>
<td>28 (2.4)</td>
</tr>
<tr>
<td>Donegan et al. (94)</td>
<td>1987–1990</td>
<td>Obstetrics</td>
<td>Boston, Mass.</td>
<td>3,845</td>
<td>93 (2.4)</td>
</tr>
</tbody>
</table>

---

Footnotes:
1. 4.1 to 8.9 patients per 100 patient visits.
Incidence of Occupationally Acquired HIV Infection

As of 30 June 1999, a total of 191 U.S. workers had been reported to the CDC’s national surveillance system for occupationally acquired HIV infection (Table 5) (65). Fifty-five HCWs had known occupational HIV exposures, with a baseline negative HIV test and subsequent documented seroconversion. Fifty of these exposures were to HIV-infected blood, one was to visibly bloody fluid, one was to an unspecified fluid, and three were to concentrated virus in a laboratory. Of the 55 HCWs, 47 sustained percutaneous exposures, 5 had mucocutaneous exposures, 2 had both a percutaneous and a mucocutaneous exposure, and 1 had an unknown route of exposure. Twenty-five of these HCWs have developed AIDS.

Of the 191 U.S. workers reported to the CDC’s surveillance system, 136 have been reported as possible cases of occupationally acquired HIV infection. None of these HCWs reported behavioral or blood transfusion risk factors, and all reported occupational exposures to blood, body fluids, or laboratory specimens containing HIV. However, the time or source of infection was undocumented, usually because no baseline serum sample was available to establish seronegativity at the time of exposure.

The CDC’s surveillance system likely does not reflect the full extent of occupationally acquired HIV infection because of underreporting of known infections or underrecognition of HIV infection. Studies of HCWs in hospital settings suggest that many percutaneous injuries are not reported (129, 177).


HIV Seroprevalence Surveys among HCWs

HIV seroprevalence surveys provide a way of indirectly assessing the risk of occupationally acquired HIV infection. The CDC has conducted two voluntary anonymous seroprevalence surveys of surgeons in different specialties. In 1992, a seroprevalence survey was done among general surgeons, obstetricians, gynecologists, and orthopedic surgeons practicing in moderate to high AIDS incidence areas. Of the 770 participating surgeons, one general surgeon, who reported nonoccupa-

### TABLE 5. HCWs with documented and possible occupationally acquired HIV infection reported through June 1999 in the United States

<table>
<thead>
<tr>
<th>Occupation</th>
<th>No. of documented cases of occupational transmission</th>
<th>No. of possible cases of occupational transmission</th>
</tr>
</thead>
<tbody>
<tr>
<td>Dental worker, including dentist</td>
<td>6</td>
<td></td>
</tr>
<tr>
<td>Embalmer or morgue technician</td>
<td>1</td>
<td>2</td>
</tr>
<tr>
<td>Emergency medical technician or paramedic</td>
<td>12</td>
<td></td>
</tr>
<tr>
<td>Health aide or attendant</td>
<td>1</td>
<td>15</td>
</tr>
<tr>
<td>Housekeeper or maintenance worker</td>
<td>1</td>
<td>12</td>
</tr>
<tr>
<td>Laboratory technician, clinical</td>
<td>16</td>
<td>16</td>
</tr>
<tr>
<td>Laboratory technician, nonclinical</td>
<td>3</td>
<td></td>
</tr>
<tr>
<td>Nurse</td>
<td>23</td>
<td>34</td>
</tr>
<tr>
<td>Physician, nonsurgical</td>
<td>6</td>
<td>12</td>
</tr>
<tr>
<td>Physician, surgical</td>
<td>6</td>
<td></td>
</tr>
<tr>
<td>Respiratory therapist</td>
<td>1</td>
<td>2</td>
</tr>
<tr>
<td>Technician, dialysis</td>
<td>1</td>
<td>3</td>
</tr>
<tr>
<td>Technician, surgical</td>
<td>2</td>
<td>2</td>
</tr>
<tr>
<td>Technician or therapist, other</td>
<td>10</td>
<td></td>
</tr>
<tr>
<td>Other health care occupations</td>
<td>4</td>
<td></td>
</tr>
<tr>
<td><strong>Total</strong></td>
<td><strong>55</strong></td>
<td><strong>136</strong></td>
</tr>
</tbody>
</table>

* HCWs are defined as those persons, including students and trainees, who have worked in a health care, clinical, or HIV laboratory setting at any time since 1978. Adapted from reference 65.

### TABLE 6. Published HIV seroprevalence in selected HCWs

<table>
<thead>
<tr>
<th>Occupation and authors (reference)</th>
<th>No. of HCWs tested</th>
<th>No. of HCWs HIV positive</th>
<th>% Prevalence</th>
</tr>
</thead>
<tbody>
<tr>
<td>Surgeon</td>
<td>770</td>
<td>1</td>
<td>0.13</td>
</tr>
<tr>
<td>Tokars et al. (258)</td>
<td>3,420</td>
<td>2</td>
<td>0</td>
</tr>
<tr>
<td>HCW blood donor</td>
<td>Chamberland et al. (66)</td>
<td>9,449</td>
<td>—</td>
</tr>
<tr>
<td>U.S. Army Reserve physician, dentist</td>
<td>3,347</td>
<td>3</td>
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<tr>
<td>Cowan et al. (82)</td>
<td></td>
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<tr>
<td>Dentist</td>
<td>Flynn et al. (107)</td>
<td>89</td>
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</tr>
<tr>
<td>Klein et al. (163)</td>
<td>1,132&lt;sup&gt;a&lt;/sup&gt;</td>
<td>1</td>
<td>0.09</td>
</tr>
<tr>
<td>Siew et al.&lt;sup&gt;c&lt;/sup&gt;</td>
<td>1,195</td>
<td>0</td>
<td>0</td>
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<tr>
<td>Gruninger et al. (123)</td>
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<td>0.09</td>
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<tr>
<td>Gruninger et al. (123)</td>
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<td>961</td>
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<td>40</td>
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<tr>
<td>Chirgwin et al. (71)</td>
<td>25</td>
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<td>Comodo et al. (80)</td>
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<tr>
<td>Goldman et al. (118)</td>
<td>49</td>
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<tr>
<td>Peterman et al. (215)</td>
<td>161</td>
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<tr>
<td>Mortician, embalmer</td>
<td>Gershon et al. (117)</td>
<td>130</td>
<td>1</td>
</tr>
<tr>
<td>Turner et al. (264)</td>
<td>129&lt;sup&gt;b&lt;/sup&gt;</td>
<td>0</td>
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</tr>
</tbody>
</table>

<sup>a</sup> One HCW lost to follow-up.

<sup>b</sup> Persons with nonoccupational risk excluded.


tional risk factors for HIV infection on an anonymous questionnaire, was HIV positive (209). In 1991, a seroprevalence survey was done among surgeons attending the annual meeting of the American Academy of Orthopaedic Surgeons. Of the 3,420 participants, two surgeons, both of whom reported nonoccupational risk factors, were HIV positive (258). Other seroprevalence studies similarly have shown low rates of HIV seropositivity among HCWs without nonoccupational risk factors for HIV infection (Table 6) (20, 66, 80, 107, 117, 118, 123, 163, 215, 264; P. Ebbensen, M. Melbye, F. Scheutz, A. J. Bodner, and R. J. Bigger, Letter, JAMA 256:2199, 1986; C. Siew, S. E. Gruninger, and S. A. Hoyat, Letter, N. Engl. J. Med. 318:1400–1401, 1988).

One limitation of seroprevalence studies is that the extent of occupational and nonoccupational exposure to HIV among tested workers is usually unknown. Also, the rates may be underestimates if individuals deferred testing because they knew they were or suspected they might be HIV positive. Nonetheless, these seroprevalence surveys indicate that there was not a high rate of undetected HIV infection among the HCWs studied, many of whom had substantial opportunity for occupational exposures.

**RISK OF OCCUPATIONAL TRANSMISSION OF HBV FROM PATIENTS TO WORKERS**

**Risk of HBV Infection Postexposure**

The probability of HBV transmission after an occupational exposure is dependent upon the concentration of infectious virions in the implicated body fluid, the volume of infective material transferred, and the route of inoculation (e.g., percutaneous or mucosal).

HBV is present in high titers in blood and serous fluids, ranging from a few virions to 10⁹ virions per ml (142). The virus is present in moderate titers in saliva, semen, and vaginal secretions (154). The titer in semen and saliva is generally 1,000 to 10,000 times lower than the corresponding titer in serum (44, 269). Other body fluids such as urine and feces contain very low levels of HBV unless contaminated with blood (91, 106, 149).

One of the most common modes of HBV transmission in the health care setting is an unintentional injury of an HCW from a needle contaminated with HBsAg-positive blood from an infected patient (5). The average volume of blood inoculated during a needlestick injury with a 22-gauge needle is approximately 1 μl (V. M. Napoli and J. E. McGowan, Letter, J. Infect. Dis. 155:828, 1987), a quantity sufficient to contain up to 100 infectious doses of HBV (243). The risk of transmission after a needlestick exposure to a nonimmune person is at least 30% if the source patient is HBeAg positive but is less than 6% if the patient is HBeAg negative (17, 120, 277). Blood from patients with HbsAg titers below the threshold of detection using routine serologic tests is rarely infectious (4). While overt percutaneous injuries are efficient modes of HBV transmission, other less-obvious exposures may also lead to occupation ally acquired HBV infection. In a case series of HBV-infected HCWs, fewer than 10% recalled a specific percutaneous injury, while 29 to 38% recalled caring for an HBsAg-positive patient within 6 months prior to their onset of illness (35; A. K. R. Chaudhuri and E. A. C. Follet, Letter, Br. Med. J. 284:1408, 1982).

**HBV Seroprevalence among Patients**

The risk of acquiring HBV is related to the prevalence of HBV infection in the patient population with which the HCW works. Patients who are HBsAg positive, either from acute or chronic infection, are potential sources of infection. Patients who are acutely infected may not be recognized since acute infection is symptomatic in only 10% of children and 30 to 50% of adults. Chronic HBV infection is often asymptomatic. HCWs who work in settings with patient populations with a relatively high prevalence of HBV infection, such as urban and tertiary-care hospitals (which more commonly serve groups at high risk for HBV infection, such as injecting drug users), have been shown to be at greater risk of occupational HBV infection than those who work in rural or community hospitals (133).

Prior to the implementation of guidelines for hepatitis B prevention, patients in hemodialysis centers had high rates of HBV infection, which posed an increased risk for workers in this setting (43, 189). However, between 1976 and 1993, the annual incidence of HBV infection decreased from 3.0 to 0.1% among hemodialysis patients and from 2.6 to 0.02% among staff members (254). Outbreaks of HBV infection in hemodialysis centers rarely occur today. When these outbreaks do occur, they are most often traced to failure to implement recommended infection control practices (11, 56, 198).

**Trends in the Incidence of Occupationally Acquired HBV Infection**

The number of HCWs infected annually with HBV in the United States is estimated from data reported to the CDC Viral Hepatitis Surveillance Program (VHSP). Annual estimates are derived by applying the proportion of people who acquired HBV occupationally in the health care setting in a given year as reported to the VHSP to the estimated number of HBV infections that occurred in that same year. For example, the CDC estimates that in 1985 about 12,000 HCWs became infected with HBV (48). This figure is derived from the proportion of people who acquired HBV occupationally in the health care setting in 1985 (6% of patients in the Viral Hepatitis Surveillance Program reported employment in a medical or dental field for 6 months prior to date onset of illness, and two-thirds of these patients were estimated to work in settings with potential exposure to blood or body fluids) and the estimated number of HBV infections that occurred in the United States in 1985 (300,000).

The incidence of HBV infection among HCWs has decreased substantially since the early 1980s (54). The estimated number of HBV infections among HCWs declined from 17,000 (386 per 100,000) in 1983 to 400 (9.1 per 100,000) in 1995 (176). The estimated incidence of HBV infections among HCWs in 1983 was about threefold higher than the incidence of HBV infections in the general U.S. population (122 per 100,000) and declined by 1995 to more than fivefold lower than the incidence in the general U.S. population (50 per 100,000).

The absolute decline in the number of HBV infections among HCWs is attributed to the implementation of standard precautions in health care settings, including the increasing use of barrier precautions and personal protective devices and increasing levels of hepatitis B vaccination coverage among HCWs (21, 126, 282) (see Hepatitis B Vaccination Coverage among HCWs [below]).

**HBV Prevalence among HCWs**

Prior to the availability of the hepatitis B vaccine, numerous cross-sectional surveys showed that HCWs had a three- to fivefold higher seroprevalence of HBV infection than the general U.S. population (48, 89, 93, 239, 241, 253). Prevalence rates of HBV infection of 13 to 18% have been demonstrated
among surgeons, and infection rates up to 27% have been demonstrated among dentists and oral surgeons (246, 278). By comparison, about 4% of first-time blood donors in the United States during the 1970s had serological markers of HBV infection (246).

Prevalence of previous infection with HBV has been found to increase with increasing age and to be directly related to the number of years employed as an HCW (78, 209, 241, 253). HCWs with frequent blood or needlestick exposures have a twofold higher prevalence of HBV infection than other HCWs (125). Physicians and dentists in specialties that involve frequent blood or needlestick exposure (e.g., obstetrician-gynecologists, pathologists, and oral surgeons) have a significantly elevated risk of HBV infection compared to specialists with less-frequent blood or needlestick exposure (e.g., pediatricians and psychiatrists) (278).

**RISK OF OCCUPATIONAL TRANSMISSION OF HCV FROM PATIENTS TO WORKERS**

**Risk of HCV Infection Postexposure**

HCV is transmitted efficiently by large exposures to blood such as through transfusion of blood or blood products from infectious donors. Overt percutaneous exposures to HCV (e.g., accidental needlestick injuries) also have been documented as means of HCV transmission.

The risk that an HCV-infected individual will transmit the virus may be related to the type and size of the inoculum and the route of transmission as well as the titer of virus, but data on the threshold concentration of virus needed to transmit infection are insufficient. Neither the presence of antibody nor the presence of HCV RNA is a direct measure of infectivity.


Follow-up studies of HCWs who sustained percutaneous exposures to blood from anti-HCV-positive patients have found variable rates of HCV transmission (30, 140, 161, 223, 247, 284). However, the average incidence of anti-HCV seroconversion after needlestick or sharps exposure from a known anti-HCV-positive source patient is 1.8% (range, 0 to 7%) (10, 64). In one study conducted in Japan, which included PCR testing for HCV RNA in source patients and HCWs, the risk of transmission after a needlestick exposure from a source patient with HCV RNA-positive blood was 10% (186). No infections have been associated with mucous membrane or nonintact skin exposures in prospective studies conducted to date; however, there have been two case reports of HCV transmission as a result of a blood splash to the conjunctiva (232; G. Ippolito, V. Puro, N. Petrosillo, G. De Carli, G. Micheloni, and E. Migliano, Letter, JAMA 280:28, 1998).

The importance of mucous membrane and inapparent par-

Several studies examining risk factors for HCV infection among HCWs have produced conflicting results. One study in New York found 2% of dentists and 9% of oral surgeons to be anti-HCV positive (162). In that study, the percentage of professional time spent practicing oral surgery was directly related to anti-HCV positivity. However, anti-HCV-positive dentists reported 50% fewer needlesticks during the previous 5 years than did anti-HCV-negative dentists. In contrast, anti-HCV positivity was associated with a reported history of frequent needlestick injuries in a survey of hospital-based HCWs in California (219). However, in studies among surgeons in several urban areas, no association was observed with recollection of skin, mucous membrane, or percutaneous exposure to blood during the last month or year (209; J. I. Tokars, M. Chamberland, C. Shapiro, C. Schable, A. Wright, D. Culver, M. Jones, P. McKibben, D. Bell, and the Serosurvey Study Committee, Proc. 2nd Annu. Meet. Soc. Hosp. Epidemiol. Am., p. 33, 1992).

PREVENTION OF OCCUPATIONAL EXPOSURES TO BLOOD

Standard Precautions

In 1987 the CDC developed universal precautions to help protect both HCWs and patients from infection with blood-borne pathogens in the health care setting (46). These recommendations stress that blood is the most important source of HIV, HBV, and other blood-borne pathogens and that infection control efforts should focus on the prevention of exposures to blood as well as the receipt of HBV immunizations. In 1995, the CDC’s Hospital Infection Control Practices Advisory Committee (HICPAC) introduced the concept of standard precautions, which synthesizes the major features of universal precautions and body substance isolation into a single set of precautions to be used for the care of all patients in hospitals regardless of their presumed infection status (111). Blood, certain other body fluids (e.g., semen, vaginal secretions, and amniotic, cerebrospinal, pericardial, peritoneal, and synovial fluids), and tissues of all patients should be considered potentially infectious (46, 47). Standard precautions apply to blood; all body fluids, secretions, and excretions (except sweat); non-intact skin; and mucous membranes (111). The core elements of standard precautions comprise (i) hand washing after patient contact, (ii) the use of barrier precautions (e.g., gloves, gowns, and facial protection) to prevent mucocutaneous contact, and (iii) minimal manual manipulation of sharp instruments and devices and disposal of these items in puncture-resistant containers (46, 47, 111).

The CDC’s recommendations—along with the blood-borne pathogen standard issued by the Occupational Safety and Health Administration (OSHA), which requires that HBV vaccine be made available to HCWs with risk of occupational exposure, the development of written exposure control plans, the use of engineering and work practice controls to reduce exposures, and annual HCW training (266)—have caused widespread adoption of standard precautions in U.S. hospitals. Several investigators have attempted to assess the efficacy of standard precautions. For example, Beeckman et al. at the Clinical Center of the National Institutes of Health found a significant and sustained decrease in percutaneous injuries associated with the implementation of standard precautions (21). At the same institution, a comparison of the frequencies of HCWs’ blood exposures on self-reported questionnaires before and after standard precaution training found a decrease in the mean number of blood exposures per year among clinical HCWs, from 35.8 to 18.1 (102). Education of HCWs about needlestick prevention, along with effective communication and convenient placement of sharps containers, has been shown to decrease needlestick injuries by 60% among HCWs at a teaching hospital in California (126).

Personal Protective Barriers, Work Techniques, and Safety Devices

Skin and mucous membrane contacts frequently can be prevented with the use of barrier precautions, such as gloves, masks, gowns, and goggles, among HCWs in emergency room, operating room, and medical ward settings (102, 178, 259, 282). However, the greatest risk of blood-borne pathogen transmission comes from percutaneous injuries, which are not prevented by barriers but instead require changes in technique and/or use of safety devices. For instance, Tokars et al. noted that half of the percutaneous injuries during surgical procedures occurred when fingers, rather than instruments, were used during suturing, suggesting that the use of instruments or other changes in technique might reduce injuries (256). The use of blunt-tip suture needles during surgical procedures can significantly reduce suture-related percutaneous injuries. In a CDC study of blunt suture needle use during gynecological surgical procedures, researchers found no percutaneous injuries with blunt suture needles compared to 1.9 injuries per 1,000 conventional curved suture needles used and 14.2 injuries per 1,000 straight suture needles used (58).

Similarly, changes in the design of sharp instruments can prevent injuries in nonsurgical settings (151, 152). One study found that resheathable and bluntable needles reduced percutaneous injuries during phlebotomy by 23 to 76% (59). Many injuries in the health care setting are associated with intravenous (i.v.) tubing-needle assemblies. Studies have found that i.v.-related percutaneous injuries decreased approximately 72 to 100% following the introduction of needleless systems (112, 252; Skolnick et al., Letter, N. Engl. J. Med. 318:1400–1401); the greatest reductions were seen with those systems that did not permit needles to access i.v. lines. Although devices may be safer for HCWs, it is important that they be assessed for potential patient care complications. An outbreak of bloodstream infections associated with a needleless i.v. infusion system raised concerns regarding the potential for adverse patient outcomes associated with these devices (86). However, Adams et al. prospectively compared the incidences of various patient-related adverse outcomes for conventional and needleless i.v. access systems and found that the needleless system posed no greater risk of positive catheter tip or adapter fluid cultures, i.v. site complications, or nosocomial bacteremia (1).
Three intramuscular doses of hepatitis B vaccine induce a protective antibody response in 90% of healthy recipients. However, HBV has been shown to be inactivated by several intermediate-level disinfectants, including 0.1% glutaraldehyde and 500-ppm free chlorine from sodium hypochlorite (i.e., household bleach) (29, 105). Heating to 98°C for 2 min also inactivates HBV (165).

While specific animal infectivity studies have not been published, rapid degradation of HCV occurs when serum containing HCV is left at room temperature (83). Epidemiologic data also suggest that environmental contamination with HCV is not a significant route of transmission in the health care setting.

Standard sterilization and disinfection procedures recommended for patient care equipment are adequate to sterilize or disinfect items contaminated with blood or other body fluids from people infected with blood-borne pathogens, including HIV, HBV, and HCV. Because foreign material may interfere with the sterilization or disinfection procedure, devices must first be adequately cleaned. Cleaning before disinfection is particularly important for devices such as endoscopes that may become heavily soiled and cannot tolerate heat sterilization (180).

All spills of blood and blood-contaminated body fluids should be promptly cleaned by a person wearing gloves and using an Environmental Protection Agency-approved disinfectant or a 1:30 to 1:100 solution of household bleach. Visible material should first be removed with disposable towels or other means to prevent direct contact with blood. The area should then be decontaminated with an appropriate disinfectant (46).

**VACCINATION AGAINST HBV INFECTION**

**Prevention of HBV Infection Using Hepatitis B Vaccine**

Hepatitis B vaccine provides both preexposure and postexposure protection against HBV infection. Two types of hepatitis B vaccine, plasma-derived and recombinant, have been licensed in the United States, and both are very effective in preventing HBV infection. The plasma-derived vaccine is no longer available in the United States. The currently available vaccines are produced by recombinant DNA technology (51). Three intramuscular doses of hepatitis B vaccine induce a protective antibody response in >90% of healthy recipients. Adults who develop a protective antibody response are protected from clinically significant disease and chronic infection. Long-term studies of immunized adults and children indicate that immune memory remains intact for at least 12 years, even though anti-HBs levels may become low or undetectable (272, 279, 280). Routine booster doses of hepatitis B vaccine are not considered necessary (61).

Since it became available in 1981, hepatitis B vaccine has been recommended for HCWs who have anticipated exposure to blood or body fluids. It is preferable that HCWs be vaccinated during professional training or early in their careers, so that they are protected prior to being at risk of occupational HBV infection.

Persons at occupational risk of infection should be tested for anti-HBs after vaccination, since knowledge of a person’s HBV immune status allows for the most precise selection of a postexposure prophylaxis regimen, should an exposure occur. It is recommended that postvaccination testing be done for all HCWs who are at risk for having blood or blood-contaminated body fluid exposures (e.g., physicians, nurses, operating room technicians, dentists, dental hygienists, emergency medical technicians, phlebotomists, laboratory technologists and technicians, physician assistants, and nurse practitioners). Testing is not indicated for persons at low risk of mucosal or percutaneous exposure to blood or body fluids or HBV infection (i.e., public safety workers and HCWs without direct patient contact). When indicated, postvaccination testing should be done 1 to 2 months after completion of the three-dose series.

Persons who do not respond to the primary vaccine series should complete a second three-dose vaccine series or be evaluated to determine if they are HBsAg positive. Revaccinated persons should be retested at the completion of the second vaccine series. Nonresponders to vaccination who are HBsAg negative should be considered susceptible to HBV infection and should be counseled regarding precautions to prevent HBV infection and the need to obtain hepatitis B immunoglobulin (IG) prophylaxis for any known or probable parenteral exposure to HBsAg-positive blood.

**Hepatitis B Vaccination Coverage among HCWs**

Since 1982, hepatitis B vaccine has been recommended for HCWs with frequent blood or needle exposures (45). However, hepatitis B vaccine was not widely used among HCWs in the 1980s. In a survey of U.S. hospitals conducted during 1990 by OSHA, 91% of hospitals had hepatitis B vaccination programs for employees, and of these, 64% paid for the cost of vaccinating high-risk employees (i.e., those involved in direct patient care and laboratory work) (181). However, it was estimated that only 46% of high-risk employees had received the hepatitis B vaccine. Barriers to vaccine use among HCWs included the high cost of the vaccine, failure of employers to offer the vaccine at low or no cost, and a perception among some HCWs that they would not benefit from vaccination.

In 1991, OSHA issued a standard that required employers to offer hepatitis B vaccine at no cost to employees with reasonably anticipated contact with blood or other potentially infectious materials (266). This standard does not require the employer to conduct postvaccination testing or to provide booster doses of hepatitis B vaccine. Employees who administer first aid only as a collateral duty to their routine work assignment (e.g., teachers) do not need to be offered the hepatitis B vaccine until they give aid involving exposure to blood or other potentially infectious materials. If an exposure incident occurs, the employee should be evaluated for postexposure prophylaxis (PEP) in accordance with the recommendations of the Advisory Committee on Immunization Practices (ACIP) (60).

Subsequent to the issuance of the OSHA guidelines, hepatitis B vaccination coverage substantially increased among HCWs, especially among younger HCWs. In 1991 and 1992, surveys indicated that approximately 90% of orthopedic and hospital-based surgeons aged 20 to 29 years from urban areas had received the hepatitis B vaccine. However, among surgeons who had practiced more than 10 years, 25% had not received the hepatitis B vaccine and were susceptible to HBV infection (209; Tokars et al., Proc. 2nd Annu. Meet. Soc. Hosp. Epidemiol. Am., p. 33, 1992). A survey conducted among 150 hospitals in 1992 found that 51% of the employees who were eligible to receive hepatitis B vaccine had completed the vaccination series (2). By 1994, a telephone survey of 113 hospitals found that 67% of eligible employees had completed the hepatitis B vaccination series (176). Vaccination coverage was highest among personnel with frequent exposure to infectious body fluids and lowest for employees at low risk for exposure. Coverage levels among eligible employee groups surveyed in 1994 were 81% among phlebotomists, 72% among nurses, 71% among physicians and residents, 63% among nurse aides, 59%
among custodial and security personnel, 44% among clerical administrative staff, and 44% among food service workers.

MANAGEMENT OF OCCUPATIONAL EXPOSURES

Although exposure prevention remains the best strategy for protecting HCWs from occupationally acquired infection, exposures are nevertheless likely to occur. Employers should have in place a system that includes written protocols for prompt reporting, evaluation, counseling, treatment, and follow-up of occupational exposures that may place a worker at risk of blood-borne pathogen infection. Employers also must establish exposure control plans and comply with incident reporting requirements mandated by OSHA (50, 266). Access to clinicians who can provide postexposure care should be available during all work hours, including nights and weekends. Persons responsible for providing postexposure counseling should be familiar with evaluation and treatment protocols and the facility's procedures for obtaining drugs for PEP (63).

Exposure Reporting

The prompt reporting of exposures is important, not only for management of the exposure but also for identification of workplace hazards and evaluation of preventive measures. Reporting systems should include ready access to expert consultants as well as safeguards to protect the confidentiality of the exposed worker. Unfortunately, a significant proportion of percutaneous injuries are not reported to hospital surveillance systems (range, 5 to 60%) (59, 129, 177, 183, 204). Timely and complete reporting of exposures can be facilitated by education of HCWs and a supportive, nonpunitive response by employers. HCW education, including orientation and in-service programs, should familiarize HCWs with their personal risk of occupational blood-borne pathogen exposure, measures to prevent such exposures, and the principles of postexposure management. HCWs must understand the importance of reporting exposures immediately after they occur, since certain indicated interventions (e.g., PEP for HIV and HBV) must be initiated promptly to be effective (38, 50, 114).

Exposure Assessment and Emergency Management

Upon reporting an exposure, the HCW should be evaluated and counseled regarding the risk of blood-borne pathogen infection, the potential usefulness of PEP for HIV and/or HBV, the need for follow-up evaluation, and precautions to prevent possible HIV transmission to others during the follow-up period (50). Risk evaluation should include an assessment of factors that may increase or decrease the probability of infection transmission.

First aid, if necessary, should be administered as quickly as possible. Puncture wounds and other cutaneous injury sites should be washed with soap and water, and exposed oral and nasal mucous membranes should be vigorously flushed with water. Eyes should be irrigated with clean water, saline, or sterile irrigants (50, 115). Although there is no evidence that antiseptics for wound care reduce the risk of blood-borne pathogen transmission, their use is not contraindicated. The use of bleach or other caustic agents that cause local tissue trauma is not recommended (38).

After any exposure, efforts should be made to identify and evaluate clinically and epidemiologically the source patient for evidence of HIV, HBV, and/or HCV infection. The source patient should be informed of the incident and consent should be obtained for HIV, HBV, and HCV testing.

The circumstances of the exposure should be recorded in a confidential medical record. Data collection should include demographic information about the exposed worker, details about the exposure itself (including date, time, job duty being performed, type of exposure, amount and type of fluid or material involved, type of device used, and severity of exposure), a description of infection control precautions used, information about the source patient, and details about postexposure management, counseling, and follow-up (50, 114).

Postexposure Chemoprophylaxis for HIV

Background. Information from a retrospective case-control study of HCWs from France, the United Kingdom, and the United States suggesting that ZDV PEP may reduce the risk for HIV transmission after occupational exposure to HIV-infected blood (55), along with data on ZDV efficacy in preventing perinatal transmission (81) and evidence that PEP prevented or ameliorated retroviral infection in some studies in animals (27), prompted the Public Health Service (PHS) to publish a statement on management of occupational exposures to HIV in 1996 (57). The PHS subsequently published expanded and updated recommendations for occupational HIV exposure management for HCWs in May 1998 (63). These guidelines have been supported by groups such as the International AIDS Society—USA (42).

ZDV and other reverse transcriptase inhibitors may be important for PEP by preventing early viral dissemination. Studies of HIV-infected patients have shown that other antiretroviral agents, such as the reverse transcriptase inhibitor lamivudine, and the class of protease inhibitors that includes saquinavir and indinavir (IDV) significantly decrease plasma HIV levels, especially when used in combination with ZDV (100). Protease inhibitors may be useful for prophylaxis based on the site of activity in the replication cycle (i.e., after viral integration has occurred) in addition to demonstrated effectiveness in reducing viral load.

Animal studies. PEP has prevented or ameliorated retroviral infection in some studies with animals, particularly when it was administered soon after exposure (199, 230, 242, 251, 262). However, the application of animal studies, especially those using nonhuman retroviruses, is uncertain. In addition to the use of nonhuman retroviruses, many variables, such as viral inoculum size, antiretroviral dose, administration route, time to onset of treatment, and dose interval, may influence the apparent effectiveness of the treatment under study (27).

Human studies. There are few data with which to assess the efficacy of PEP in humans. The optimal study design for determining the efficacy of ZDV for PEP would be a prospective, placebo-controlled trial. However, this has not been possible because of the requirement for a large number of HCWs and the relatively low rate of HIV seroconversion following occupational exposure (S. W. LaFon, B. D. Moorey, J. P. McScullen, K. H. Pattishall, M. L. Smiley, M. D. Rodgers, and S. N. Lehrman, Program Abstr. 30th Intersci. Conf. Antimicrob. Agents Chemother., abstr. 489, 1990). In the absence of such a trial, other sources of data have been used to assess the use of ZDV for PEP.

In a multicenter, double-blind, placebo-controlled clinical trial of ZDV to prevent perinatal HIV transmission, ZDV therapy was associated with a 67.5% reduction in the risk of mother-to-infant HIV transmission (53). The protective effect of ZDV was only partly explained by reduction of the HIV titer in maternal blood, suggesting a possible direct prophylactic effect of ZDV (248). Additionally, a recent placebo-controlled study in Thailand showed that a short-term antenatal regimen of ZDV reduced the risk for perinatal HIV transmission by...
51% (62). Also, the CDC retrospective case-control study found that PEP with ZDV was associated with a decrease of approximately 81% in the risk for HIV seroconversion among HCWs who had a percutaneous exposure to HIV-infected blood (39).

However, any protection afforded is not absolute. Failure of ZDV PEP to prevent HIV infection in HCWs has been reported (156, 174; G. Weisburd, J. Biglione, M. M. Arbulu, J. C. Terrazzino, and A. Pesiri, Program Abstr. XI Int. Conf. AIDS, abstr. pub. C. 1141, 1996). Additional failures of ZDV PEP have been described among individuals exposed to an inoculum of HIV-infected blood larger than what would be expected from a needlestick. These non-HCW cases included one blood transfusion, one suicidal self-inoculation, one assault with a needle-syringe, and two instances of accidental intravenous infusion of HIV-infected blood components during nuclear medicine procedures (156). Possible factors that may have contributed to the apparent failures in these instances include exposure to a ZDV-resistant strain of HIV, a high-titer and/or large-inoculum exposure, delayed initiation and/or short duration of PEP, and possible factors related to the host (e.g., cellular immune system responsiveness) and/or to the source patient’s virus (e.g., presence of syncytium-forming strains) (63).
PHS recommendations for chemoprophylaxis. Chemoprophylaxis should be recommended to exposed workers after occupational exposures associated with a known risk for HIV transmission, should be considered for exposures with a negligible risk, and may not be warranted for exposures that do not pose a known risk for HIV transmission (Fig. 1). For exposures for which PEP is considered appropriate, exposed workers should be informed that (i) knowledge about the efficacy and toxicity of drugs used for PEP is limited; (ii) only ZDV has been shown to prevent HIV transmission in humans; (iii) there are no data to address whether adding other antiretroviral drugs provides any additional benefit for PEP, but some ex-
peptoids recommend combination drug regimens because of increased potency and concerns about drug-resistant virus; (iv) data regarding toxicity of antiretroviral drugs in persons without HIV infection or who are pregnant are limited; and (v) any or all drugs for PEP may be declined by the exposed worker. HCWs who have HIV occupational exposures for which PEP is not recommended should be informed that the potential side effects and toxicity of taking PEP outweigh the negligible risk of transmission posed by the type of exposure.

Most HIV exposures will warrant only a two-drug regimen, using two nucleoside analogue reverse transcriptase inhibitors, usually ZDV and lamivudine. The addition of a third drug, usually a protease inhibitor (i.e., IDV or nelfinavir), should be considered for exposures that pose an increased risk for transmission or when resistance to the other drugs used for PEP is known or suspected. ZDV-resistant strains of HIV can be transmitted and have been reported to cause primary infections (99; G. Ippolito, P. Del Poggio, C. Arici, G. P. Gregis, G. Antonelli, and E. Riva, Letter, JAMA 272:433–434, 1994). If the exposure source is unknown or the HIV status of the source patient cannot be tested, information about the circumstances of the exposure should be assessed to determine the risk for transmission of HIV. Certain situations, as well as the type of exposure, may suggest an increased or decreased risk; an important consideration is the prevalence of HIV in the population group (i.e., institution or community) from which the contaminated source material was derived. Decisions regarding appropriate management should be individualized based on the risk assessment (63).

PEP should be initiated as soon as possible (i.e., within hours of the exposure). The interval within which PEP should be started for optimal efficacy is not known. The optimal duration of PEP also is unknown. Because 4 weeks of ZDV appears sufficient to be protective in HCWs (39), PEP probably should be administered for 4 weeks, if tolerated. When PEP is used, drug toxicity monitoring should include a complete blood count and renal and hepatic chemical function tests at baseline and 2 weeks after starting PEP.

Counseling and follow-up. All HCWs with occupational exposure to HIV should receive follow-up counseling, postexposure testing (Table 7), and medical evaluation, regardless of whether they receive PEP. HIV antibody testing should be performed for at least 6 months postexposure (e.g., at 6 weeks, 12 weeks, and 6 months). HIV testing using EIA should be performed on any HCW who has an illness that is compatible with an acute retroviral syndrome. HIV antibody testing using EIA should be used to monitor for seroconversion. The routine use of direct virus assays (e.g., PCR for HIV RNA) to detect infection in exposed HCWs generally is not recommended (34, 113, 114).

The psychological impact of an occupational HIV exposure may be considerable and should be addressed during counseling and follow-up (236). Experts have found that supportive counseling is an important part of management (98, 114, 135, 136). To prevent the possibility of further transmission to others, the HCW should be advised to refrain from donating blood, semen, or organs during the follow-up period and to refrain from breast-feeding when safe and effective alternatives are available. To prevent HIV transmission to sexual contacts, all exposed HCWs should abstain from, or use latex condoms during, sexual intercourse throughout the follow-up period, especially during the first 6 to 12 weeks after the exposure, when most HIV-infected persons are expected to seroconvert (63).

Toxicity. An important goal of PEP is to encourage and facilitate compliance with the prescribed regimen. Therefore, the toxicity profile of antiretroviral agents is a relevant consideration. All of the antiretroviral agents have been associated with side effects (63). Side effects associated with many of the nucleoside analogue reverse transcriptase inhibitors are chiefly gastrointestinal (e.g., nausea or diarrhea). Rare but serious side effects, such as seizures, have been reported with ZDV PEP (M. D’Silva, D. Leibowitz, and J. P. Flaherty, Letter, Lancet 346:452, 1995). The use of protease inhibitors has been associated with new onset of diabetes mellitus, hyperglycemia, diabetic ketoacidosis, and exacerbation of preexisting diabetes mellitus (266a; M. D. Dubé, D. L. Johnson, J. S. Cumer, and J. M. Leedon, Letter, Lancet 350:713–714, 1997). Nephrolithiasis has been associated with IDV use (including in HCWs using the drug for PEP) (S. A. Wang and the HIV PEP Registry Group, Program Abstr. Infect. Dis. Soc. Am. 35th Annu. Meet., abstr. 482, 1997); however, the incidence of this potential complication may be limited by drinking at least 48 oz (1.5 liters) of fluid per 24-h period (19). Rare cases of hemolytic anemia also have been associated with the use of IDV. Nelfinavir, saquinavir, and ritonavir have all been associated with the development of diarrhea; however, this side effect usually responds to treatment with antimotility agents that can be

### TABLE 7. Recommended serologic testing for HCWs following occupational exposures to HIV, HBV, and HCV

<table>
<thead>
<tr>
<th>Infection status of source patient</th>
<th>Recommended serologic tests at:</th>
</tr>
</thead>
<tbody>
<tr>
<td>HIV-positive</td>
<td>Baseline</td>
</tr>
<tr>
<td>Anti-HBs-positive</td>
<td>HIV antibody testing using EIA</td>
</tr>
<tr>
<td>Anti-HCV-positive</td>
<td>HCV antibody testing using EIA; ALT measurement</td>
</tr>
<tr>
<td>Unknown</td>
<td>HIV antibody testing using EIA; anti-HBs if previously vaccinated to HBV and response to vaccination unknown; HCV antibody testing using EIA; ALT measurement</td>
</tr>
</tbody>
</table>

* Confirmation by Western blot testing of all anti-HIV results reported as reactive by EIA.

* Confirmation by supplemental anti-HCV (i.e., recombinant immunoblot assay [RIBA]) testing of all anti-HCV results reported as repeatedly reactive by EIA.

* If earlier diagnosis of HCV infection is desired, testing for HCV RNA may be performed at 4 to 6 weeks.
prescribed for use, if necessary, at the time any one of these drugs is prescribed for PEP. The manufacturer’s package insert should always be consulted for questions about potential drug interactions.

**Postexposure Prophylaxis for HBV**

PEP with hepatitis B vaccine and hepatitis B IG among persons susceptible to HBV is highly effective in preventing infection after an exposure. Management of HCWs after percutaneous (e.g., needlestick, laceration, or bite) or mucosal (e.g., mucous membrane or ocular) exposure to potentially infectious body fluids must include consideration of (i) the HBsAg status of the source of exposure and (ii) the hepatitis B vaccination and vaccine response status of the exposed HCW. The ACIP of the PHS and HICPAC have provided detailed advice on postexposure immunoprophylaxis (60). Table 8 provides a guide to recommended management for various HBV exposures. Ideally, immunoprophylaxis should be initiated as soon as possible after a percutaneous or percutaneous exposure; its value beyond 7 days after exposure is unclear.

**Postexposure Management of HCV**

**HCV prophylaxis.** Several studies have attempted to assess the effectiveness of prophylaxis with IG for the prevention of posttransfusion NANB hepatitis. However, the results from these studies are difficult to compare and interpret because of lack of uniformity in diagnostic criteria, varied study designs (including some lacking blinding and control groups), and administration of the first dose of IG prior to rather than after exposure in all but one study (164, 231, 238). Data from these studies have not been reanalyzed since anti-HCV testing became available. Beginning in 1992, IG has been manufactured from plasma that has been screened for anti-HCV. Therefore, if protective antibody does exist, screening and removal of anti-HCV-positive units may reduce the effectiveness of IG as PEP for HCV. An experimental study conducted with chimpanzees showed IG’s lack of efficacy in preventing infection after exposure (168).

Substantial challenges exist in the development of an effective vaccine against HCV, including the significant heterogeneity of the HCV genome and the lack of protective immunity elicited by HCV in the host (222). The development of an effective vaccine against HCV awaits a better understanding of the molecular biology of and immune response to this virus.

**Follow-up of HCWs after an occupational exposure to HCV.** There is currently no effective PEP for HCV infection. In the absence of effective prophylaxis, persons who have been exposed to HCV may benefit from knowing their infection status so they can seek evaluation for chronic liver disease and treatment. The CDC recently issued recommendations that individual institutions implement policies and procedures for follow-up after percutaneous or percutaneous exposure to anti-HCV-positive blood (64a). The purpose of follow-up testing is to address individual workers’ concerns about their risk and outcome and to identify persons who might benefit from antiviral therapy. At a minimum, such policies should include (i) baseline testing of the source for anti-HCV, (ii) baseline and follow-up (e.g., at 4 to 6 months) testing of the exposed person for anti-HCV and ALT activity (Table 6) (if earlier diagnosis of HCV infection is desired, testing for HCV RNA may be performed at 4 to 6 weeks), (iii) confirmation by supplemental anti-HCV testing of all anti-HCV results reported as positive by EIA, and (iv) education of HCWs about the risk for and prevention of transmission of all blood-borne pathogens, including HCV, in occupational settings, with the information routinely updated to ensure accuracy.

While interferon has a proven efficacy in treating chronic hepatitis C, there is no specific therapy of proven benefit for acute hepatitis C (109). Several studies have suggested that

### Table 8. Recommended PEP for percutaneous or percutaneous exposure to HBV in the United States

<table>
<thead>
<tr>
<th>Vaccination and antibody response status of exposed worker</th>
<th>Treatment when source is found to be:</th>
</tr>
</thead>
<tbody>
<tr>
<td>HBSAg positive</td>
<td>HBSAg negative</td>
</tr>
<tr>
<td>Unvaccinated</td>
<td>Initiate HB vaccine series</td>
</tr>
<tr>
<td>Previously vaccinated</td>
<td>Initiate HB vaccine series</td>
</tr>
<tr>
<td>Known responder</td>
<td>No treatment</td>
</tr>
<tr>
<td>Known nonresponder</td>
<td>Treat with one dose of HBIG&lt;sup&gt;b&lt;/sup&gt; and initiate HB vaccine&lt;sup&gt;c&lt;/sup&gt;</td>
</tr>
<tr>
<td>Antibody response unknown</td>
<td>No treatment</td>
</tr>
<tr>
<td>Test exposed person for anti-HBs (if adequate, no treatment; if inadequate, treat with one dose of HBIG and vaccine booster)</td>
<td>No treatment</td>
</tr>
<tr>
<td>If known high risk source, treat as if source were HBSAg positive</td>
<td>Test exposed for anti-HBs (if adequate, no treatment; if inadequate, treat with vaccine booster and recheck titer in 1 to 2 mo)</td>
</tr>
</tbody>
</table>

<sup>a</sup> Adapted from reference 60.  
<sup>b</sup> HBIG, hepatitis B IG (dose, 0.06 ml/kg [intramuscularly]).  
<sup>c</sup> Hepatitis B vaccine.  
<sup>d</sup> A responder is defined as a person with adequate levels of anti-HBs in serum (i.e., anti-HBs is ≥10 mIU/ml); inadequate response to vaccination defined as level of anti-HBs in serum of <10 mIU/ml.
early therapy with alpha interferon may be effective in preventing progression from acute to chronic disease (36, 200, 270). However, there are no data indicating that treatment begun early in the course of chronic infection is less effective than treatment begun in the acute phase of infection.

**MANAGEMENT OF INFECTED HCWS**

Transmission of HIV from Infected HCWs to Patients

There have been two reported instances of HIV transmission from HCWs to patients. In July 1990, the CDC reported a case of transmission of HIV from a Florida dentist to a patient during an invasive dental procedure (49). Subsequent epidemiologic investigation and molecular genetic sequencing identified five additional patients who were infected while receiving care from the dentist. Each of the six patients had no other identified risk factors for acquiring HIV, and each was infected by a strain of HIV that closely matched that of the dentist by genetic sequencing analysis. Although the specific incidents that resulted in HIV transmission to these patients are uncertain, evidence strongly supports dentist-to-patient rather than patient-to-patient transmission (72, 206). A second case, reported in 1997, involved an orthopedic surgeon in France, who probably became infected with HIV in 1983 and had performed surgical procedures on 3,004 persons since that time. An epidemiologic investigation found one person among these patients who was HIV seronegative before a prolonged invasive procedure performed by the surgeon in 1992 and who subsequently was HIV seropositive. No other risk factors were documented for the patient, and nosocomial transmission of HIV from the surgeon to the patient was confirmed by an evaluation of viral sequences from both persons (28).

Retrospective investigation data. Even before reports of the Florida dentist case were published, many health departments, hospitals, and other agencies were conducting investigations of HIV-infected HCWs and notifying patients who had received care from these providers. Retrospective studies of a number of HIV-infected dentists, surgeons, and physicians revealed no evidence of HCW-to-patient HIV transmission during patient care (Table 9). A summary of all published and unpublished investigations of which the CDC was aware through January 1995 showed no documented cases of HIV transmission among 22,171 patients treated by HIV-infected HCWs, including a breast surgeon, a general surgeon, two obstetrics and gynecology residents, two orthopedic surgeons, and several dentists (227). An epidemiologic and laboratory follow-up of 110 of the 113 identified seropositive patients showed that the majority (90 of 110 [82%]) were either documented as having been infected before receiving care from the infected HCW or had established risk factors for acquiring HIV; 15 did not have clearly established risks but had had opportunities for potential exposure to HIV; and five had no identified risk (227). Genetic sequence analysis was done on HIV strains from three HCWs and 30 seropositive patients, including three of the five patients who had no identified risk and 13 of the 15 patients with potential but undocumented risks for exposure. In no instances were the viral strains of patients and HCWs found to be related (227). Although limited by the lack of complete availability of HIV test results, procedure records, and information about the stage of the HCW’s HIV infection during the time the worker did procedures, these results are consistent with conclusions by the CDC and others that the risk of HIV transmission from HCWs to patients is very low (25, 52).

<table>
<thead>
<tr>
<th>Occupation</th>
<th>Authors and reference</th>
<th>No. of patients tested</th>
<th>No. of patients HIV positive</th>
<th>No. of HIV-positive patients linked to HCWs</th>
</tr>
</thead>
<tbody>
<tr>
<td>Family physician</td>
<td>Danila et al. (85)</td>
<td>325</td>
<td>0</td>
<td>0</td>
</tr>
<tr>
<td>Dentist</td>
<td>Dickinson et al. (92)</td>
<td>900</td>
<td>5</td>
<td>0</td>
</tr>
<tr>
<td>Dentist</td>
<td>Jaffe et al. (150)</td>
<td>665</td>
<td>28</td>
<td>0</td>
</tr>
<tr>
<td>Surgeon</td>
<td>Mishu et al. (185)</td>
<td>1,279</td>
<td>1</td>
<td>0</td>
</tr>
<tr>
<td>Breast surgeon</td>
<td>Rogers et al. (228)</td>
<td>468</td>
<td>2</td>
<td>0</td>
</tr>
<tr>
<td>Orthopedic surgeon</td>
<td>von Reyn et al. (271)</td>
<td>1,174</td>
<td>2</td>
<td>0</td>
</tr>
</tbody>
</table>

Transmission of HBV from Infected HCWs to Patients

Since the introduction of serologic testing in the 1970s, there have been at least 46 reports worldwide of HBV-infected HCWs transmitting HBV to patients during invasive procedures (24). The number of patients infected by a single HCW ranged from 1 to more than 50. Most reports of HCWs transmitting HBV to patients occurred prior to the widespread use of barrier precautions in some HCW groups (e.g., gloves by dentists), and many have involved readily apparent deficiencies in infection control practices. Although clusters in which there were no apparent deficiencies have occasionally occurred, investigations indicate that when HCWs adhere to recommended infection control procedures the risk of HBV transmission from HCW to patient is low.

A combination of factors is believed to be responsible for HBV transmission from HCWs to patients (52). One factor associated with increased risk of transmission is the HCW being HBeAg positive, indicating a higher level of infectivity (137, 138, 217, 218, 221, 275). In the United Kingdom, several episodes of HBeAg-negative surgeons transmitting HBV have been reported (127, 128, 145, 250). These surgeons were found to be carriers of a precore mutant strain of HBV that prevents expression of HBeAg but allows the expression of infectious virus. No such transmission has been reported from other parts of Europe or Japan, where the frequency of this strain appears more common (31, 201), or from the United States, where the frequency of this strain is unknown. Other factors believed to be responsible for HBV transmission from infected HCWs to patients include contamination of surgical wounds or traumatized tissue either from unintentional injury to the HCW during invasive procedures and/or a major break in standard infection control practices (e.g., not wearing gloves during an invasive procedure).

Clusters of HBV transmission from HCW to patient have been reported even when deficiencies of surgical technique or infection control practice could not be identified. In 1991, an HBeAg-positive cardiothoracic surgeon was determined to have transmitted HBV to 19 (13%) of 142 patients (132). In a simulation in which the surgeon tied surgical knots continuously for 1 h, visible skin separations were observed on his index fingers and HBsAg was detected in the saline used to
rinse out his gloves. Whether these phenomena contributed to transmission is not clear.

Transmission of HCV from Infected HCWs to Patients

There are no reported cases of HCV transmission from infected surgeons or dentists to patients in the United States. Worldwide, there are three reports of HCV transmission from infected health care providers (95, 101; P. Brown, News, Br. Med. J. 319:1219, 1999). Between 1988 and 1993 in Spain, five patients developed acute HCV infection after undergoing heart valve replacement. On investigation, the cardiovascular surgeon was determined to have chronic hepatitis C and was implicated in the transmission of HCV to these patients (101). However, the factors responsible for transmission could not be identified.

In the United Kingdom, an HCW was found to be the probable source of infection during the investigation of a patient who developed acute HCV after cardiothoracic surgery (95). A retrospective investigation of 277 (91%) of the 304 other patients who had undergone invasive procedures performed by this surgeon found no additional cases of transmission. A third case, also in the United Kingdom, involving HCV transmission to one patient from a gynecologist, is currently under investigation (P. Brown, News, Br. Med. J. 319:1219, 1999).

The CDC also is aware of a retrospective investigation of an HCV-infected plastic surgeon whose infection was diagnosed during a routine physical examination. HCV testing of 85% of the surgeon’s patients was performed more than 6 months after their surgeries. Although three patients had evidence of HCV infection as indicated by the presence of anti-HCV antibodies, no provider-to-patient transmission was detected. One of these patients was known to have anti-HCV antibody before surgery, and the other two (one of whom had a history of injection drug use) were infected with strains that had a viral genotype and/or serotype different from that of the surgeon’s strain (CDC, unpublished data).

In summary, specific factors related to an increased likelihood of transmission from HCV-infected HCWs to patients have yet to be identified. Available data indicate that the risk of HCV transmission from an infected HCW to a patient is extremely low.

Prevention of Infection Transmission from Infected HCWs to Patients

The CDC has looked for episodes of blood-borne virus transmission to patients in health care settings, and the accumulated data have shown that the overall risk of blood-borne virus transmission from infected health care providers to patients is very low and that those for HIV and HCV specifically are extremely low.

To minimize the risk of blood-borne pathogen transmission from HCWs to patients, all HCWs should adhere to standard precautions, including the appropriate use of hand washing, protective barriers, and care in the use and disposal of needles and other sharp instruments (52). Technique changes and safer needle devices that potentially reduce percutaneous injury and recontact rates during surgery may also help reduce risks (256). Currently available data provide no basis for recommendations to restrict the practice of HCWs infected with HIV, HBV, or HCV who perform duties or procedures not identified as exposure-prone, provided the infected HCWs practice recommended surgical or dental technique and comply with standard precautions and current recommendations for sterilization and disinfection (52).

Prevention of HIV and HBV transmission during invasive procedures. The CDC has characterized exposure-prone procedures as those that include digital palpation of a needle tip in a body cavity or the simultaneous presence of the HCW’s fingers and a needle or other sharp instrument or object in a poorly visualized or highly confined anatomic site. During these procedures the routine use of gloves may not prevent injuries caused by sharp instruments and does not eliminate the potential for exposure of a patient to the HCW’s blood.

To minimize the risk of HIV and HBV transmission from infected HCWs to patients during invasive procedures, the CDC issued recommendations in 1991 (52). HCWs who perform exposure-prone invasive procedures and who do not have serologic evidence of immunity to HBV from vaccination or previous infection should know their HBsAg status (and, if positive, their HBeAg status). HCWs who are infected with HBV and are HBeAg positive and HCWs who are infected with HIV should not perform exposure-prone procedures unless they have sought counsel from an expert review panel and been advised under what circumstances, if any, they may continue to perform these procedures. Such circumstances would include notifying prospective patients of the HCW’s seropositivity before they undergo exposure-prone invasive procedures.

Since these recommendations have been issued, there has been no report of HIV transmission and only one report of HBV transmission from an infected HCW to patients in the United States (132).

HCWs with blood-borne viruses. The Society for Healthcare Epidemiologists of America has recently issued guidelines that include recommendations for management of HCWs infected with HIV, HBV, and HCV (3). Specifically, the society recommended that all HCWs use double gloving for procedures and that infected providers not be excluded from any aspect of patient care unless epidemiologically incriminated in the transmission of infections despite adequate precautions. The American College of Surgeons has stated that surgeons infected with HBV and HCV have no reason to alter their practice but should seek expert advice and appropriate treatment to prevent chronic liver disease (16).

No episode of HCV transmission from an infected HCW to a patient during surgical or dental care procedures has been observed in the United States. The CDC does not recommend restriction of HCWs with hepatitis C from performing invasive procedures.

CONCLUSION

Future directions in the area of management of blood-borne pathogen infections in HCWs include more systematic surveillance of occupationally acquired HIV, HBV, and HCV infection; better definition of the epidemiology of blood contact and the efficacy of preventive measures; development and evaluation of new safety devices and protective barriers; evaluation of PEP; and development and evaluation of vaccines for HIV and HCV.

A sustained commitment to the occupational health of HCWs will ensure maximum protection for HCWs and patients and the availability of optimal medical care for all who need it.

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