



Epidemiology of HBV Infection and Prevention Programs

Miriam J. Alter, Ph.D.

Division of Viral Hepatitis

Centers for Disease Control and Prevention



Outline

- **Clinical features**
- **Screening and diagnostic markers**
- **Epidemiology**
- **Progress in prevention**

Features of Hepatitis B Virus Infection

Incubation period	Average 8-12 weeks Range 6-26 weeks
Acute illness (jaundice)	10% <5 yrs old 30%-50% ≥5 yrs old
Case fatality rate	0.5%-1%
Chronic infection	30%-90% <5 yrs old 2%-6% ≥5 yrs old
Chronic hepatitis	66%
Premature mortality from CLD	15%-25%

Markers of HBV Infection

- **Serologic**
 - HBsAg and anti-HBs
 - Anti-HBc (HBcAg does not circulate in serum)
 - HBeAg and anti-HBe
- **Nucleic acid**
 - HBV DNA

Course of HBV Infection

HBsAg

- **Appears average of 6-8 weeks after exposure**
 - 1-3 weeks before ALT becomes abnormal
 - 3-5 weeks before onset of symptoms or jaundice
- **Reaches peak during acute stage of infection**
- **Declines to undetectable levels within 4-6 months indicating recovery**
- **Usually remains detectable in chronically infected persons**
- **Marker for transmission studies**

Course of HBV Infection

HBV DNA

- **Detected 2-5 weeks after infection and up to 40 days before HBsAg (mean of 6-15 days)**
- **Rises slowly at relatively low levels during seronegative period**
- **Also detected during chronic infection**

Course of HBV Infection

Anti-HBc

- **Appears after HBsAg, at onset of ALT abnormality**
 - Predominantly IgM class
- **Remains detectable lifelong**
 - Predominantly IgG after 6 months
- **Present in both resolved and chronic infections**
- **Isolated anti-HBc**
 - 2% of asymptomatic persons tested for HBV
 - Frequency directly related to frequency of infection
 - HBV DNA detected in <10%

Course of HBV Infection

Anti-HBs

- **Neutralizing antibody**
- **Develops during recovery**
 - Detectable along with anti-HBc
 - May become undetectable in up to 20% of patients after several years
- **Detected alone after immunization**
 - Becomes undetectable in 40% of persons by 5 years after vaccination, but protection continues

Interpretation of HBV Serologic Tests

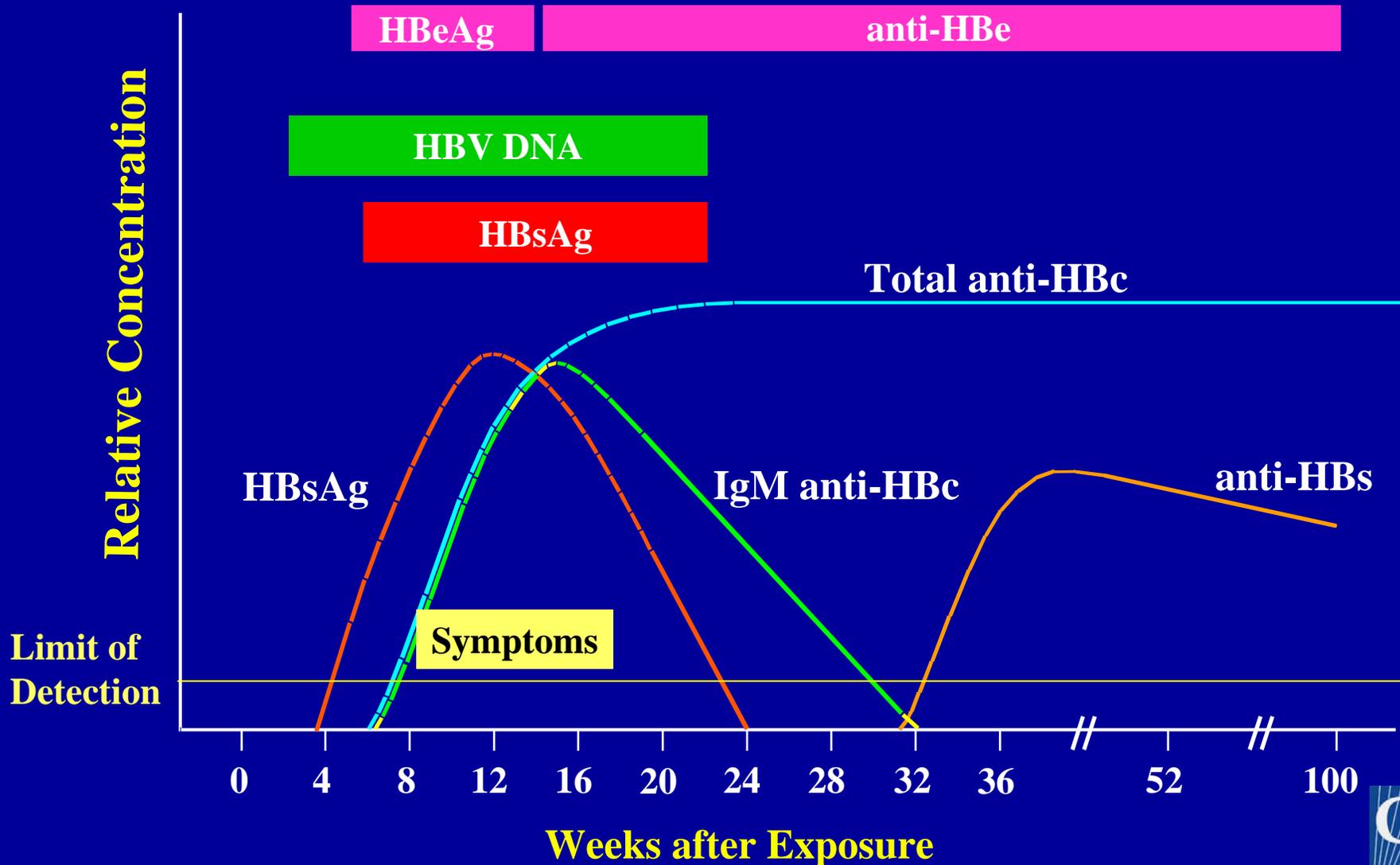
HBV					
DNA	HBsAg	Anti-HBc	Anti-HBs	Interpretation	
-	-	-	-	Susceptible	
+	-	-	-	Early acute, pre-seroconversion	
+	+	-	-	Early acute infection	
+	+	+	-	Acute or chronic infection	
-	-	+	+	Recovered/immune	
+	+	+	-	Chronic infection	

Interpretation of HBV Serologic Tests

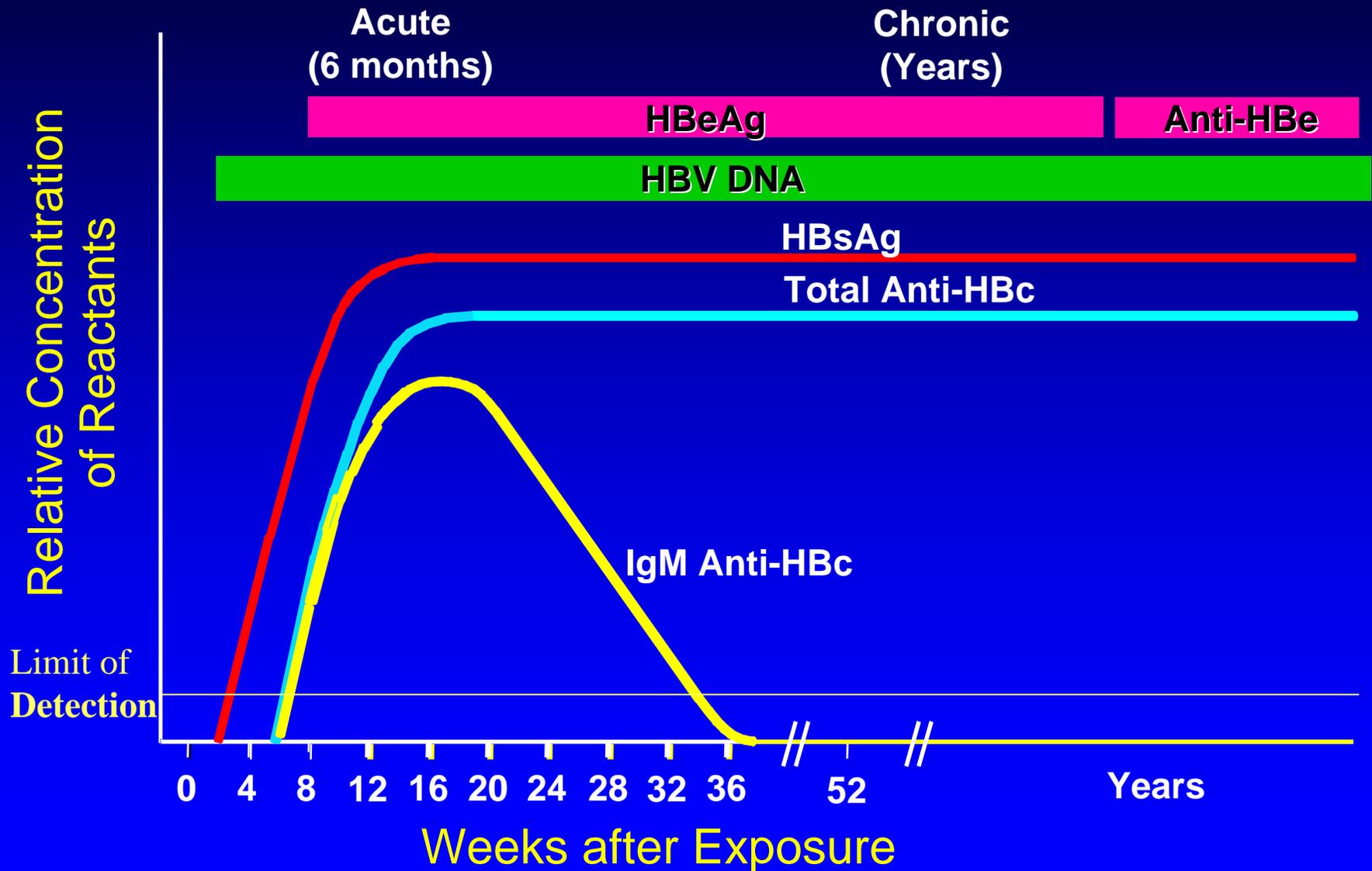
HBV

<u>DNA</u>	<u>HBsAg</u>	<u>Anti-HBc</u>	<u>Anti-HBs</u>	<u>Interpretation</u>
+/-	-	+	-	Resolving infection, past infection, "low-level" chronic infection, or false positive
-	-	-	+	Immune if >10 mIU/mL
-	+	-	-	Transient (<21 days) during vaccination

Acute HBV Infection with Recovery



Progression to Chronic Hepatitis B Virus Infection



Hepatitis B Virus Infection, United States

Newly acquired infections	73,000
Acute cases	21,000
Deaths from acute liver failure	290
Chronic infections	4,400
Persons ever infected (1990)	4.9%
Persons with chronic infection	1.25 million
HBV-related chronic liver disease	4% - 14%
Deaths from chronic disease/year	3,000-5000

Modes of HBV Transmission

Exposure to Blood or Body Fluids Containing Blood

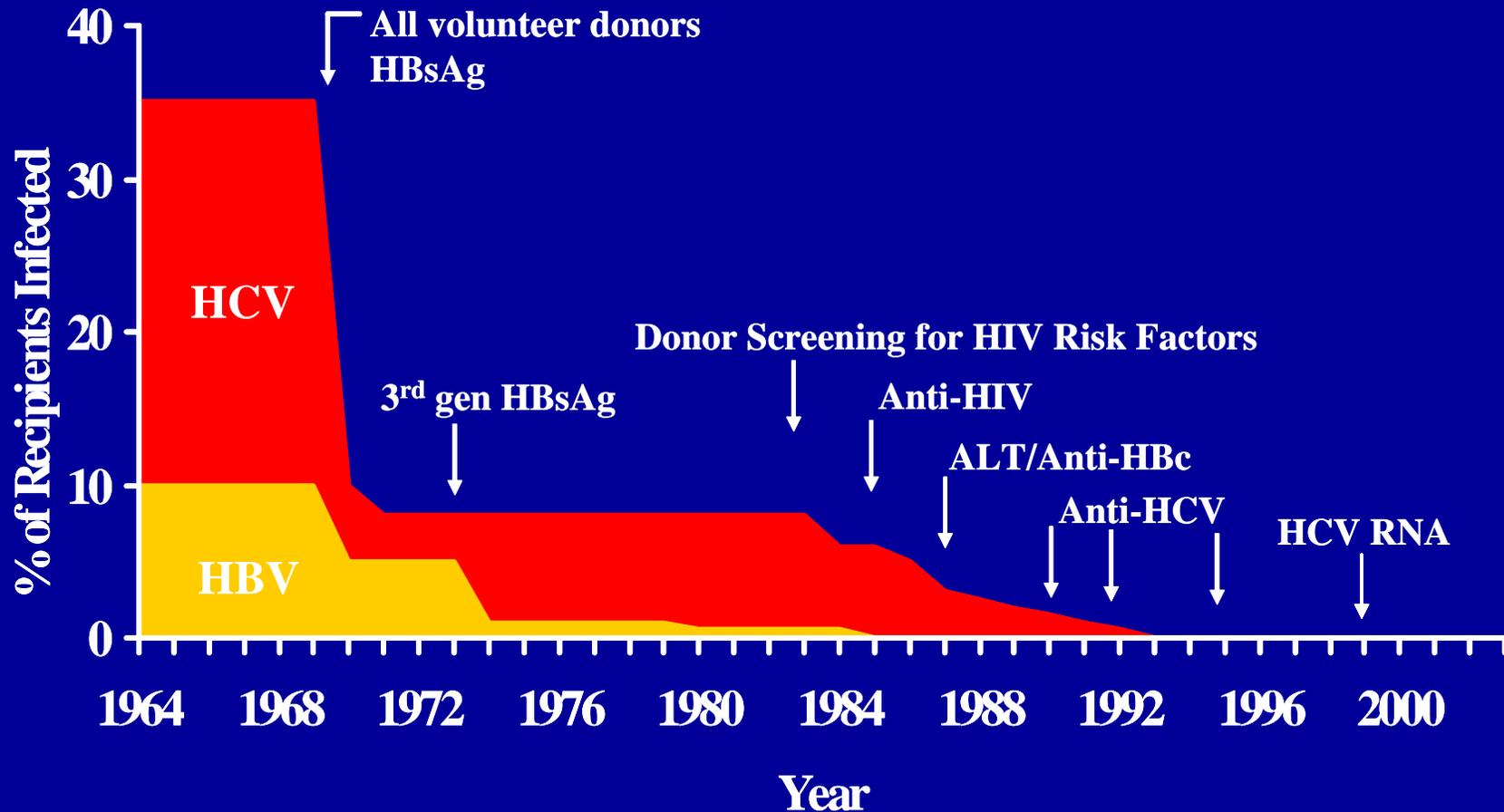
- **Percutaneous**

- Injecting drug use
- Occupational, household (needle stick, non-intact skin)
- Therapeutic (contaminated equip, unsafe injections)
- Transfusions and transplants from infectious donors

- **Permucosal**

- Sex with infected partner
- Birth to infected mother (perinatal)
- Household (exposure to infected contact)

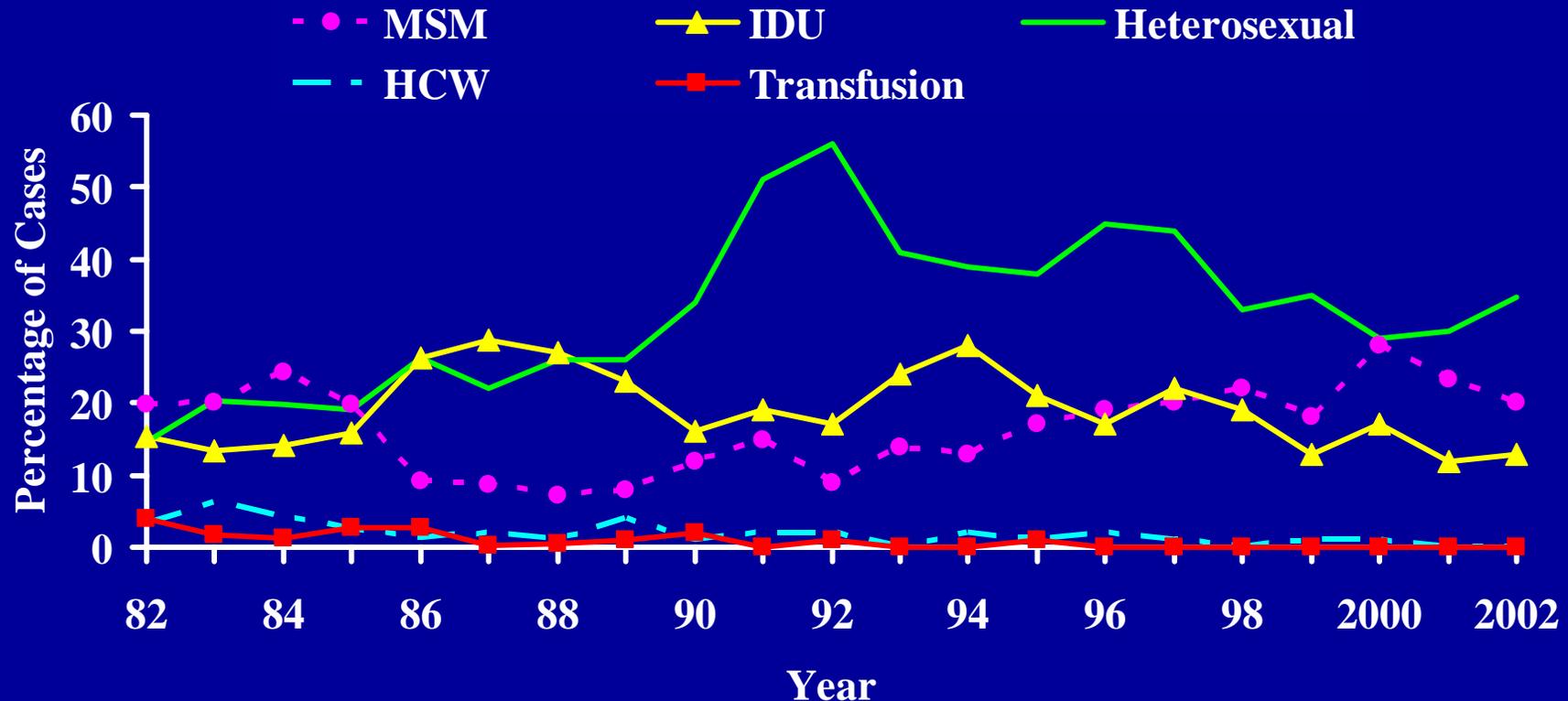
Posttransfusion Hepatitis



Adapted from HJ Alter



Reported Cases of Acute Hepatitis B by Selected Risk Factors, Sentinel Counties, United States, 1982-2002



Source: Goldstein S et al. JID 2001; 185:713-719 and CDC, unpublished data



Patients with Acute Hepatitis B Transfused During Exposure Period, 1982-2003, Sentinel Counties

- **51 reported - 8 with other risk factors**
 - 2 IDUs, 3 infected sex partners, 3 multiple partners
- **43 reported no other risk factors**
 - 34 (79%) 1982-1988
 - 6 (14%) 1989-1993
 - 3 (7%) 1994-1998
 - Follow-up testing of donors – HBV seronegative
 - All 3 cases hospitalized during incubation period
 - 1 for entire 6 months prior to onset of illness

Source: Goldstein S et al. JID 2001; 185:713-719 and CDC, unpublished data.



Nosocomial Transmission of HBV

- **Rare relative to other sources for infection**
 - Recognized primarily in context of outbreaks
- **Unsafe injection practices**
 - Spring loaded finger stick devices
 - Multiple dose medication vials
 - Therapeutic injections
 - Contaminated jet injector
 - Re-use of needles and syringes
 - Contaminated medication preparation area

Cases of Acute Hepatitis B Reporting Transfusion Enhanced Surveillance, NNDSS, 2003

Total cases reported	7,381
Reported transfusion	49
Not acute hepatitis	(15)
Never transfused	(13)
Transfused >6 months ago	(11)
Acute cases transfused	10
Donor infected	1*

***Single donor pre-seroconversion**

Source: National Notifiable Diseases Surveillance System, CDC, unpublished

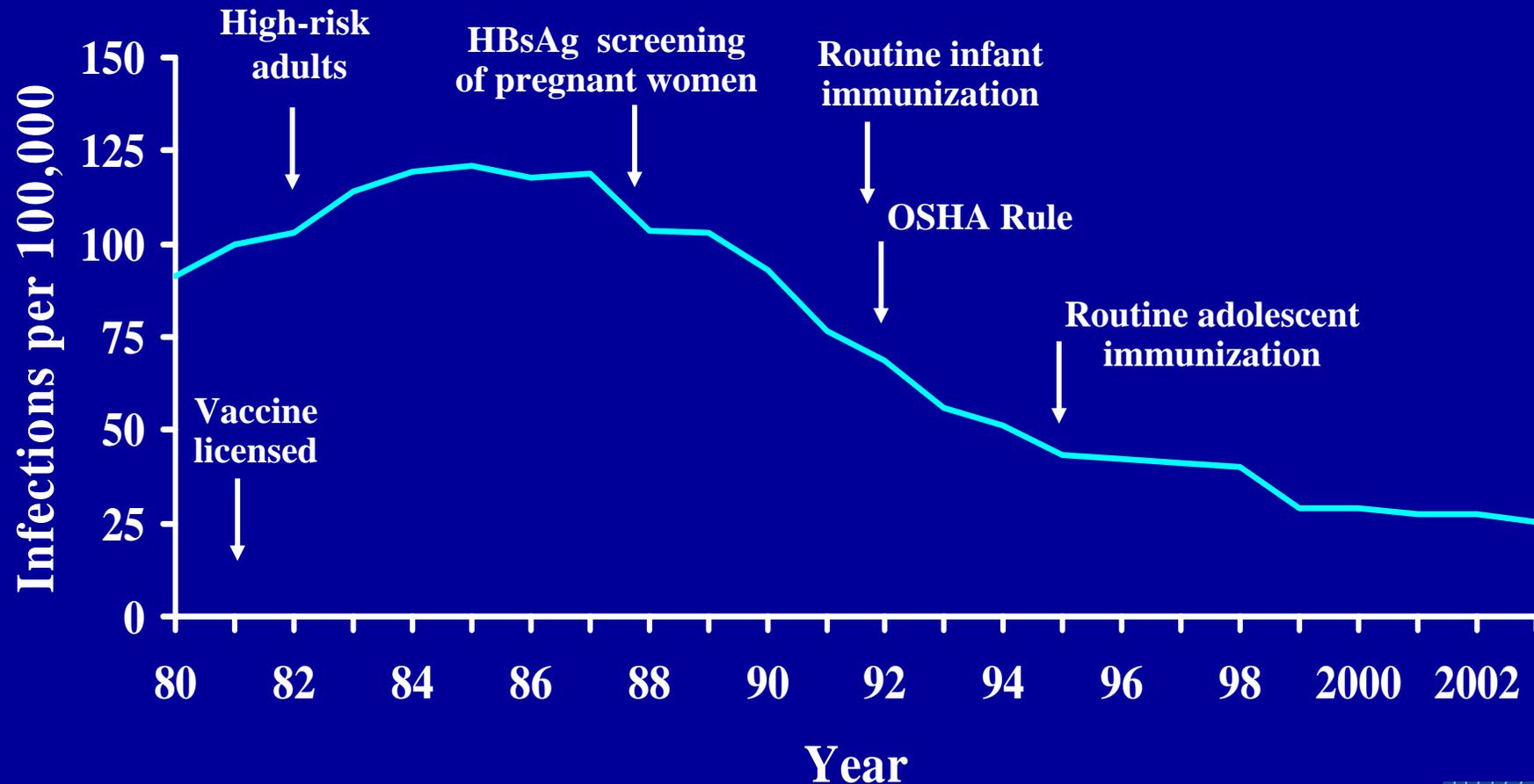


HBV Prevention and Control

- **Donor screening and testing**
 - At least 50% of acute HBV infections have histories deferred by screening
- **Infection control**
- **Preexposure vaccination**
 - Infants, children, adolescents
 - Adults at high-risk
- **Postexposure prophylaxis (vaccine, HBIG)**
 - Infants born to infected mothers
 - Screen all pregnant women for HBV
 - Ensure all newborns receive first vaccine dose
 - Occupational exposures
 - Sex and household contacts of HBsAg-positives
- **Harm reduction counseling and services**



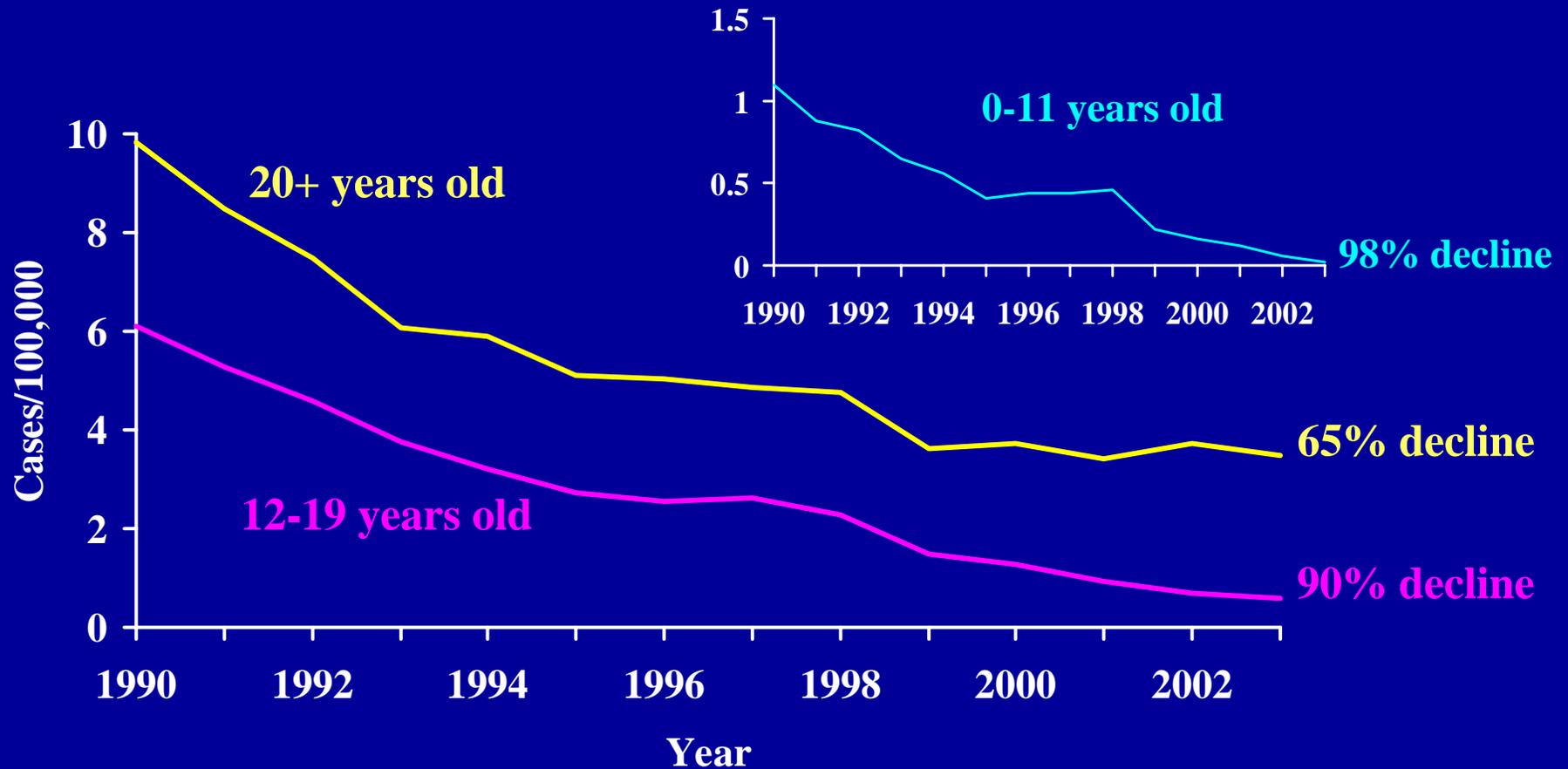
Estimated Incidence of Acute HBV Infection United States, 1980-2003



Updated 8/04



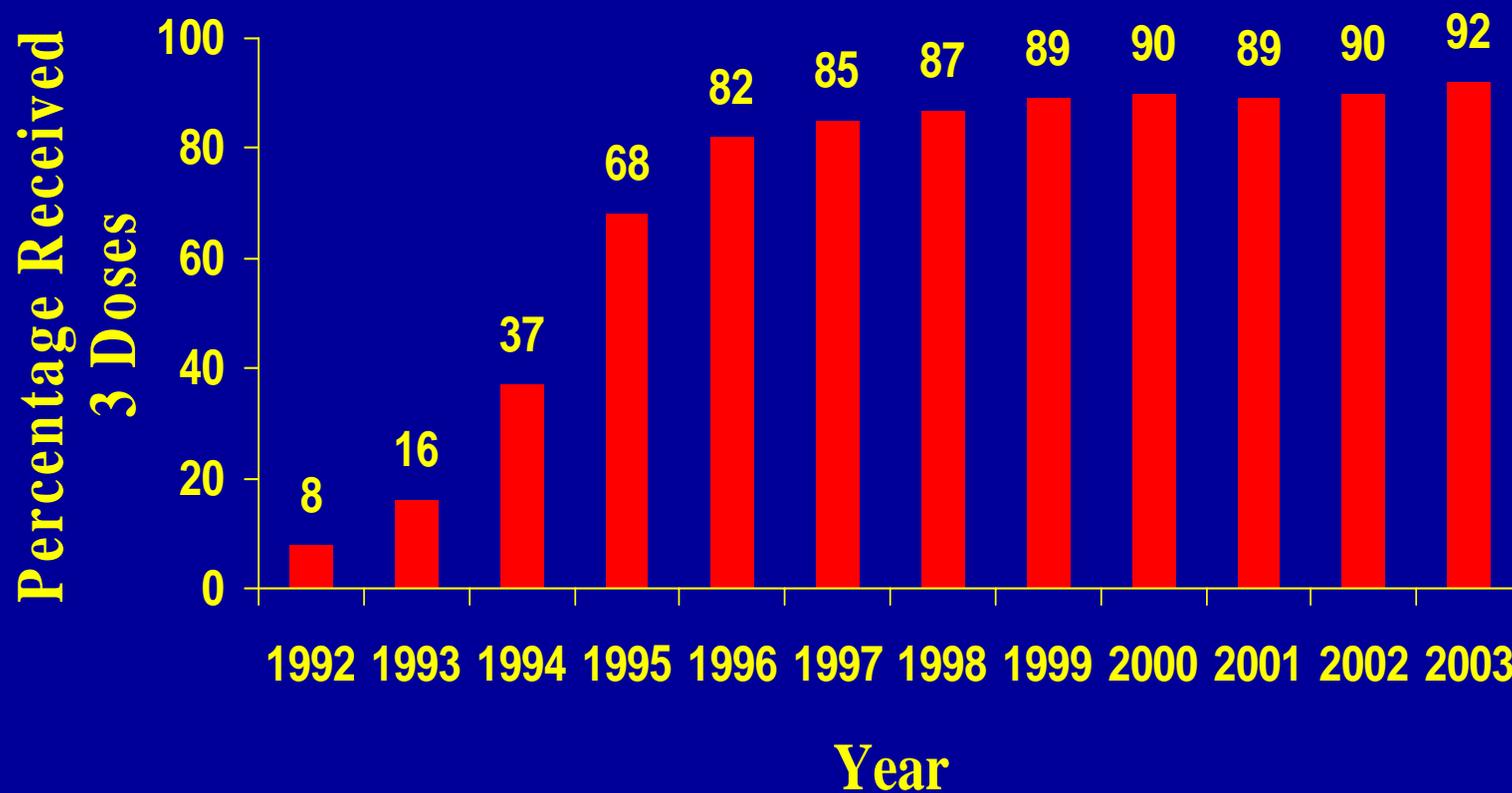
Incidence of Acute Hepatitis B by Age, United States, 1990-2003



Source: CDC, National Notifiable Diseases Surveillance System (NNDSS)



Hepatitis B Vaccine 3 Dose Coverage Among 19-35 Month Old Children, by Year of Survey, 1992-2003



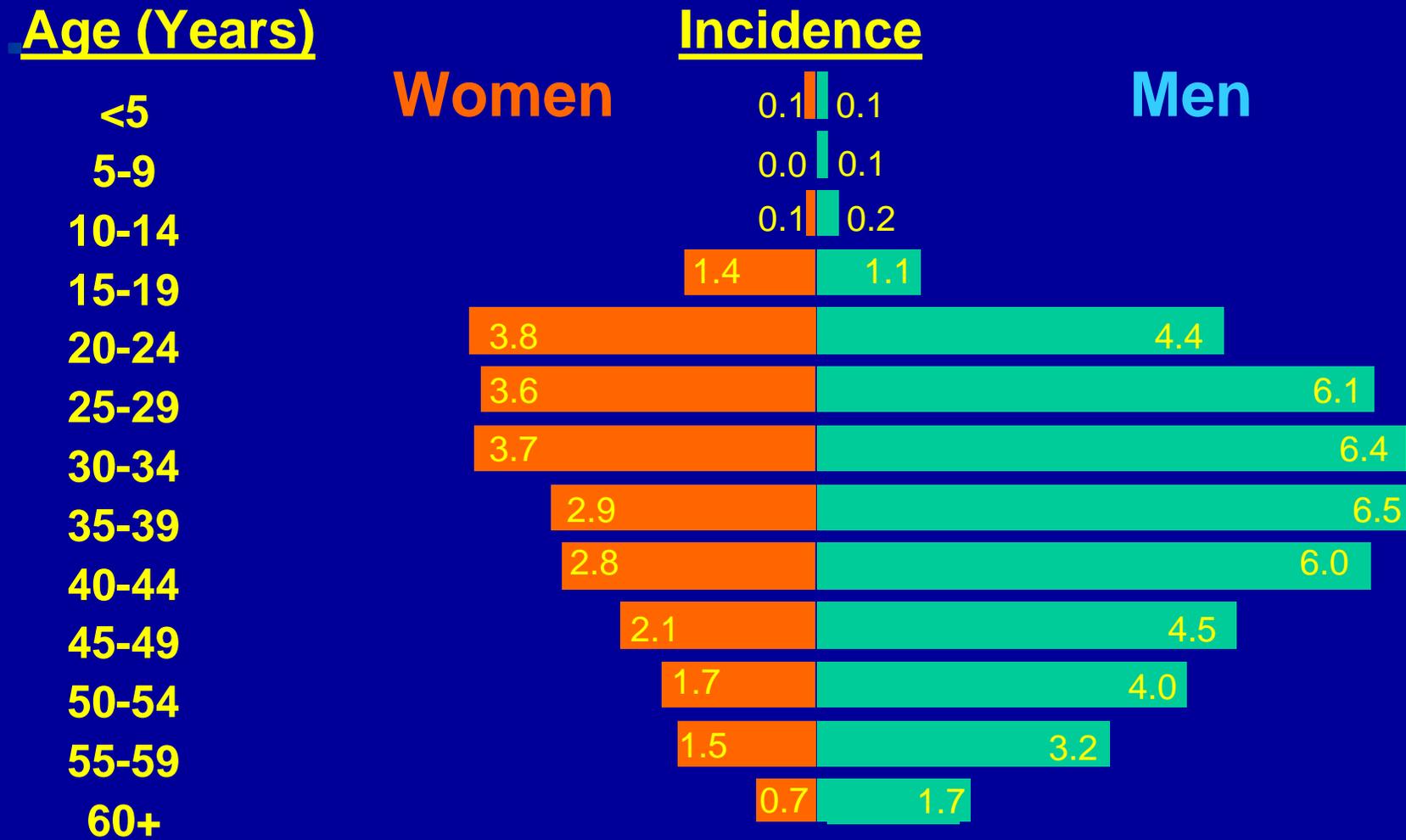
*Source: National Immunization Survey, CDC



Immunity to HBV Infection Among Adolescents and High-Risk Adults

- **Adolescents 13-15 years old - 60% vaccinated**
- **General population adults (30-60 years old)**
 - **Natural immunity**
 - Whites 4%
 - Blacks 15%
 - Asians 60-80%
- **High-risk adults vaccinated**
 - HCWs and PSWs - 70%-80%
 - IDUs, MSMs, STD clients, inmates - <10%

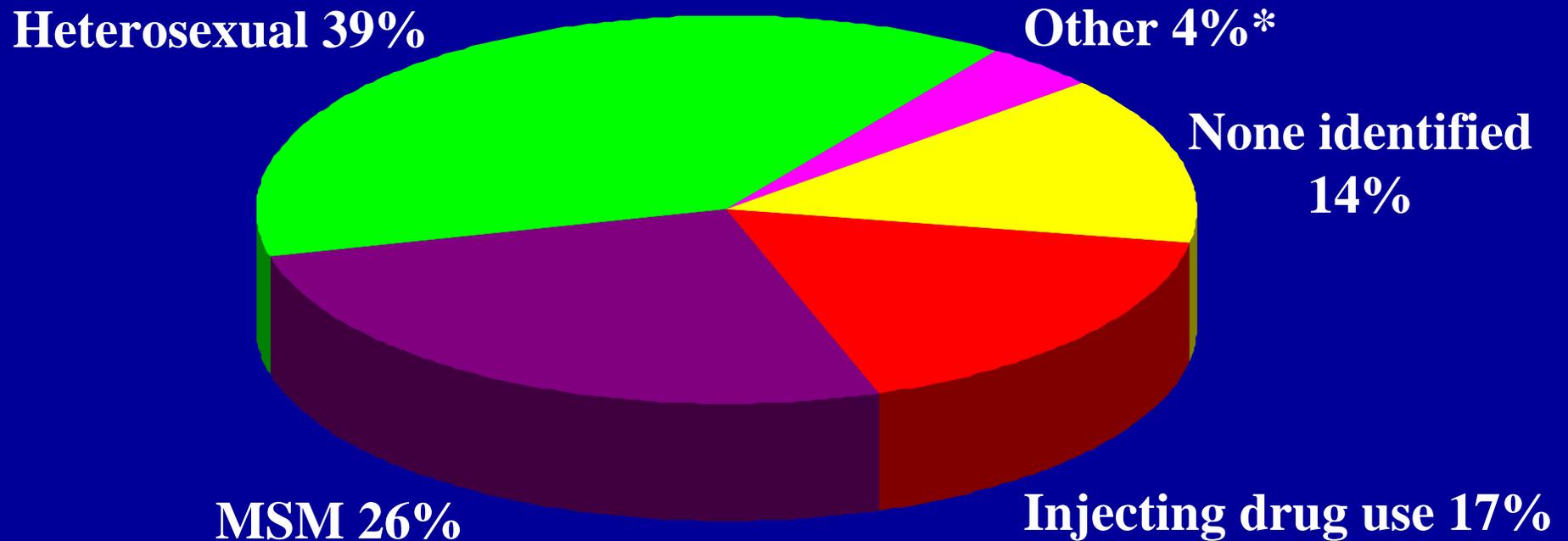
Incidence of Hepatitis B, by Age and Sex, United States



Source: National Notifiable Diseases Surveillance System (NNDSS)



Reported Risk Factors for Acute Hepatitis B United States 2000-2003



* Other - Household contact, occupational exposure, hemodialysis, institutionalization, transfusion

Source: Adapted from Sentinel Counties and NNDSS, CDC



Acute Hepatitis B Cases Which Could Have Been Prevented if Vaccination Offered as Recommended

Prior Opportunity

Percent of Cases

Known infected contact	9%
History STD rx or incarceration	55%
Total	64%

Source: Goldstein ST et.al., JID 2002;185:713-9



Elements of a Successful Vaccination Program

<u>Elements Required for Success</u>	Present in	
	<u>Childhood Program</u>	<u>Adult Program</u>
Evidence-based recommendations	X	X
Implementation strategy/partners	X	X
Provider/patient education	X	X
Vaccine purchase	X	
Infrastructure for vaccine delivery	X	



A National High-Risk Adult Hepatitis B Immunization Program

- **Integrate vaccination into existing public health and correctional health programs**
- **Strengthen infrastructure to deliver vaccine**
- **Purchase vaccine**
- **Further determine methods to improve vaccination coverage**