


Perinatal Hepatitis B



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1

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2

Hepatitis B Virus (Hep B or HBV)

- DNA hepadnavirus
- Antigenic components: HBsAg, HBcAg and HBeAg
- Remains infectious on environmental surfaces for at least 7 days at room temperature
- Transmittable in the absence of visible blood



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3

Epidemiology of Hepatitis B Infection

United States

- About 1,000 cases of perinatal HBV infection reported each year

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- Approximately 160 maternal-infant pairs are reported to public health each year
- 2015, 2 cases of perinatal Hepatitis B



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4

How is infection spread?

- Infected blood or body fluid
- Although HBsAg detected in breast milk, saliva, urine & tears, *ONLY* blood, serum, semen, vaginal fluids, CSF, synovial, pleural, pericardial, peritoneal and amniotic fluids are potentially infectious
- Primary reservoir of infection is the chronically infected
- Thus, transmission is most commonly through perinatal, household or sexual contact



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Risk factors

- Injection drug use
- Birth in an endemic country
- Birth to an infected mother
- Household contact of an acutely infected person or chronic carrier
- Multiple sexual partners or high risk sexual behavior
- Diabetic or on dialysis
- Occupational exposure to blood or body fluids

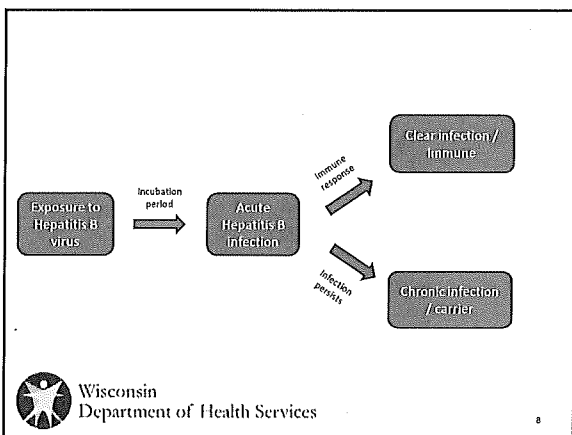
What about breastfeeding?

No increased risk to an infant who has received HBIG and Hep B vaccine per recommendations



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8

Acute Hepatitis B Infection

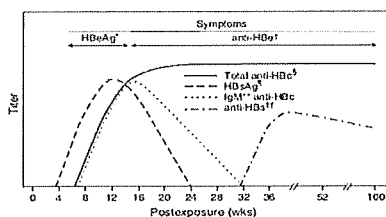
- Incubation period of 45-160 days
- May be asymptomatic – age dependent
 - decreased age, less likely to develop symptoms
 - 50% of adults and almost all children have no clinical symptoms
- Insidious onset of symptoms
- Clinical symptoms may include: anorexia, nausea, vomiting, lethargy, abdominal pain, joint pain or rash
- Jaundice and/or elevated LFTs (ALT > 100)



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9

Figure 1. Typical serologic course of acute Hepatitis B virus infection with recovery



* Hepatitis B e antigen.
 † Antibody to HBsAg.
 ‡ Antibody to hepatitis B core antigen.
 § Hepatitis B surface antigen.
 ¶ Immunoglobulin M.
 †† Antibody to HBsAg.

10

Chronic Hepatitis B Infection

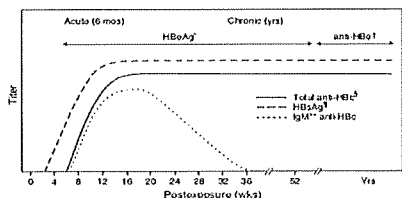
- Usually asymptomatic with no history of clinical hepatitis
- HBsAg, HBV DNA and/or HBeAg persist for at least 6 months
- May have elevated LFTs or evidence of liver damage on biopsy



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11

Figure 2. Typical serologic course of acute Hepatitis B virus (HBV) infection with progression to chronic infection

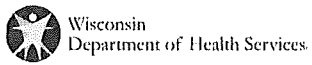


* Hepatitis B e antigen.
 † Antibody to HBsAg.
 ‡ Antibody to hepatitis B core antigen.
 § Hepatitis B surface antigen.
 ¶ Immunoglobulin M.

12

Hepatitis B Surface Antigen (HBsAg)

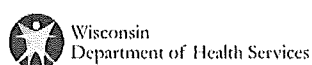
- Present in acute and chronic infection
- Anyone who is HBsAg-positive is potentially infectious
- Antigen used in the vaccine
 - may be detected up to one month following vaccination



13

Antibody to Hepatitis B Surface Antigen (anti-HBs or HBsAb)

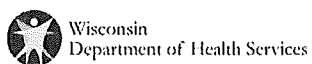
- Immune from resolved infection or vaccination
- Develops 1-2 months following vaccination or infection



14

Total Antibody to Hep B Core Antigen (total anti-HBc or total HBcAb)

- Present during acute, resolved or chronic infection
- NOT present following vaccination
- Passively transferred from a HBsAg-positive mother to her infant
 - may be detected up to 24 months following birth



15

IgM Antibody to Hepatitis B Core Antigen (IgM anti-HBc or IgM HBcAb)

- Acute or recent infection
- Specific for establishing a diagnosis of acute Hepatitis B
- Appears around time of illness onset
- Only marker positive during “window period” – time between disappearance of HBsAg and appearance of anti-HBs



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Hepatitis B e Antigen (HBeAg)

- Indicates infectiveness
- HBeAg positivity usually indicates a higher HBV viral load, and thus increased infectivity
- Eventually become undetectable in chronic infection



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17

Antibody to Hepatitis B e Antigen (anti-HBe or HBeAb)

- Presence of anti-HBe indicates lower risk of transmission / lower infectiveness




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Hepatitis B DNA or Viral Load (HBV DNA)


- Qualitative or quantitative
- Trends used to monitor effectiveness of treatment and/or eligibility for treatment



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19

Antigen	Significance	Corresponding antibody	Significance
HBsAg	Anyone with detectable antigen is infectious	anti-HBs	Indicates immunity
HBcAg	N/A – no lab test to detect	anti-HBc	Indicates acute, chronic or resolved infection
HBeAg	Viral replication is occurring	anti-HBe	Lower risk of transmission




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20

Comprehensive strategy to eliminate HBV transmission

1. universal immunization of infants at birth
2. prevent perinatal HBV infection by:
 - routine screening of all pregnant women
 - AND appropriate immunoprophylaxis to all infants born to HBsAg-positive and HBsAg-unknown mothers
3. routine immunization of children
4. immunization of all at risk adults



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21

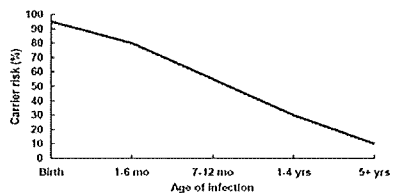
Why is perinatal Hepatitis B especially important?

- Perinatal transmission is highly efficient
- In the absence of appropriate prophylaxis, 70 - 90% infants born to HBsAg-positive mothers will develop infection
- In-utero accounts for < 2% all vertical transmission



22

Risk of Chronic HBV Carriage by Age of Infection



- 90% of infants infected at birth will develop chronic infection
- Conversely, 90% of people infected as adults will clear the infection

23

Children who acquire chronic infection as result of perinatal exposure...

- are usually asymptomatic with normal ALT
- have minimal to mild liver abnormalities
- are often HBeAg-positive with detectable viral loads (contagious)
- may have growth impairment
- 25% will die prematurely of cirrhosis or liver cancer



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Prophylaxis for Infant

HBIG (hepatitis B immune globulin)

- Passive, active immunoprophylaxis
- Short-term protection following exposure
- Prepared from serum with high anti-HBs titers
- Becomes less effective with increasing time since exposure
 - should be administered NO MORE than 7 days following exposure

Hepatitis B Vaccine

- Post-exposure
- Long-term protection

Administer at the same time, in separate sites, within 12 hours of birth

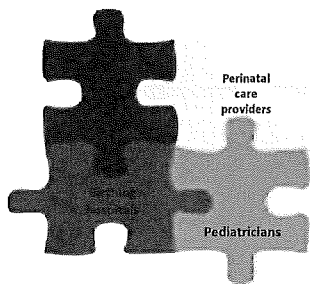
Together, HBIG and vaccine prevent 95% of all perinatal cases!



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25

Hepatitis B Case Management



26

Local Public Health Departments (LHDs)

- Responsible for managing clients who reside in their jurisdiction
- Contact client
- Typically conduct a risk assessment
- Ensure household and sexual contacts are screened, vaccinated and referred as needed
- Provide education, resources and strategies to prevent transmission
- Remain in contact with mother throughout pregnancy
- Coordinate care with family, prenatal care provider, birthing hospital and/or pediatrician as needed
- Serve as a resource to medical providers
- Document case follow-up information in WEDSS

27

OB-GYN / perinatal care provider

- Routinely screen *all* pregnant women for HBsAg early in *each* pregnancy, regardless of vaccine history or risk factors
- Repeat screening at time of hospital admission if high risk or report new risk behaviors
- Report all positive labs per state statute (252.05)



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Birth Hospital

- Maintain stock of HBIG (or be able to obtain as needed)
- Develop policy to verify HBsAg status at admission
- Establish procedure to test women who present with unknown HBsAg status or new risk factors at time of delivery
- The administration of Hep B vaccine to ALL medically stable infants within 24 hours of birth is the standard of care
- Administer HBIG and Hep B vaccine within 12 hours of birth to all infants born to HBsAg-positive mothers
- Knowledge of and policy for special circumstances (i.e. infants weighing less than 2,000 grams)
- Implement a standard practice of administering vaccine and HBIG; standing orders are considered one of the 'best practices'
- Accurately document the mother's HBsAg status on the new born screening card



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29

Newborn Screening Cards

U000110
LABORATORY

For use only regarding please PRINT and COMPLETE the extra item

Baby's Name		SEX	Baby's Birthdate	
Baby's DOB (yy/mm/dd)		F	M	
Specimen Collection Site		Baby's Physician		
Mother's Name		Physician's Name		
Baby's Weight (grams)		Gender	Baby's Race	Physician's HPI
W		M		
Baby's NICU?		Percent of gestation?	Transvaginal?	Physician's HPI (signature)
N		N	Y	
Baby's Family		Newborn Hep B Surface Antigen		
		Neg		
Screening Screen Date		Right Ear	Left Ear	Right Hip
		<input type="checkbox"/> Pass <input type="checkbox"/> Fail	<input type="checkbox"/> Pass <input type="checkbox"/> Fail	<input type="checkbox"/> Pass <input type="checkbox"/> Fail
Pulse Ox Sat Sat Date		Free Oxygen?	Free	Fail
		<input type="checkbox"/> Pass <input type="checkbox"/> Fail	<input type="checkbox"/> Pass <input type="checkbox"/> Fail	<input type="checkbox"/> Pass <input type="checkbox"/> Fail
Not Screened (mark reason)		Refused	Obvious	Other
		<input type="checkbox"/> Refused <input type="checkbox"/> Obvious <input type="checkbox"/> Other	<input type="checkbox"/> Refused <input type="checkbox"/> Obvious <input type="checkbox"/> Other	<input type="checkbox"/> Refused <input type="checkbox"/> Obvious <input type="checkbox"/> Other
All Tests for Part of Screen		All Tests for Part of Screen		
Baby's Name		Baby's Name		

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30

Infants weighing less than 2,000 grams

- Have lower seroconversion rate following birth dose of vaccine
- However, by chronological age of 1 month, all medically stable infants are likely to respond regardless of birth weight or gestational age

Therefore, birth dose does not count as part of 3-dose series:

Infant will receive 3 additional doses of vaccine beginning at 1 month of age or hospital discharge, whichever is sooner



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31

Unknown HBsAg status at time of hospital admission

1. Test mother as soon as possible
2. Administer Hep B vaccine within 12 hours of birth*
3. Obtain lab results

HBsAg-positive or unavailable: administer HBIG within 7 days or at time of hospital discharge

Negative: complete Hep B vaccine series per ACIP recommendations

*If birth weight is <2000 grams, also administer HBIG within 12 hours



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32

Child's Pediatrician or Primary Care Provider

- Ensure timely completion of Hep B vaccine series
- Order PVST (HBsAg and anti-HBs)
- Follow-up and/or refer child as needed per lab results



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33

Hepatitis B vaccines

- Safe and highly effective
- Single antigen and part of combination vaccine (Pediatrix®)
 - only single antigen is used for birth to 6 weeks of age
- Brands are interchangeable
- If an infant receives Pediatrix®, they may receive 4 total doses of Hep B vaccine
- OK to administer concurrently with other vaccines
- None contain thimerosal
- No booster or routine post-vaccination testing recommendation - a complete 3-dose vaccine series confers long-term protection



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34

Post Vaccination Serologic Testing (PVST)



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35

PVST – who?

ALL infants born to HBsAg-positive
mothers



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36

PVST – what?

HBsAg
and
anti-HBs



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37

PVST – when?

9-12 months of age
(or 1-2 months following final dose of vaccine, if the series is delayed)

No sooner than 9 months of age to avoid detection of passive anti-HBs from HBIG and to detect late-onset infection

No maximum age
(if child is over 2 years of age, anti-HBc should also be tested to rule out resolved infection)

Update: Shortened Interval for Postvaccination Serologic Testing of Infants Born to Hepatitis B-Infected Mothers (PVST, 2015)
<http://www.cdc.gov/mmwr/preview/mmwrhtml/mm6433a5.htm>



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38

PVST – where?

- Child's primary care provider
- Lab
- Some local health departments



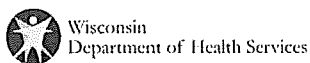
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39

PVST – why?

HBsAg – ensure child is not infected as a result of birth or continuous household exposure

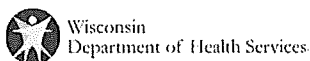
anti-HBs (HBsAb) – ensure that child is immune as a result of vaccination



40

Why both tests?

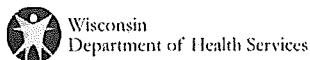
- Ensure child is not infected
- Ensure child will receive care in a timely manner if they are infected
- Ensure child is immune following vaccination
- Ensure child receives timely re-vaccination if not immune
- Minimize the amount of time a child remains vulnerable to infection



41

PVST Results

HBsAg	anti-HBsAg	Interpretation	Action
+	-	Infected	Monitor and/or refer
-	+	Protected, not infected	None
-	-	Unprotected, susceptible	Re-immunize, repeat PVST
+	+	Unlikely	Repeat tests



42

Why is public health calling about these test results?

Ideally, PVST results will be:

- HBsAg – negative
- anti-HBs – positive

Neither are reportable results (labs will not submit results electronically)

Perinatal Case managers must collaborate with medical provider to obtain results



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43

What happens if the child is infected?

- Child then becomes a case of perinatal Hepatitis B
- Monitor or referral to specialist if appropriate



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44

What happens if not immune?

- Administer a second 3-dose vaccine series as soon as possible
 - no minimum interval between first and second series
- Repeat PVST 1-2 months following second vaccine series
- If still do not sero-convert, considered a non-responder
 - counsel on risk of transmission/exposure
 - receive HBIG if have a known exposure

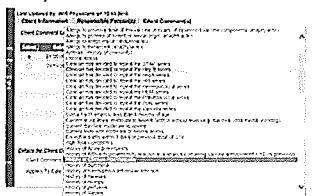


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45

Repeat Series in WIR

- The WIR functionality allows the client comment (on the Personal Information Window) "Repeat a Series"
- When the comment is entered with a date, the forecast will be repeated



Questions?

Resources

- WI Local Health Departments
<https://www.dhs.wisconsin.gov/lh-depts/counties.htm>
- DPH Perinatal Hepatitis B Manual
<https://www.dhs.wisconsin.gov/publications/p4/p44502.pdf>
- CDC resources
<http://www.cdc.gov/hepatitis/hbv/perinatalxmtn.htm>

- <http://www.cdc.gov/hepatitis/resources/professionals/training/serology/training.htm>
- Give Birth to the End of Hep B
<http://www.immunize.org/protect-newborns/>
- Pink Book
<https://www.cdc.gov/vaccines/pubs/pinkbook/downloads/hepb.pdf>
- American Academy of Pediatrics [Hepatitis B.] In: Kimberlin DW, Brady MT, Jackson MA, Long SS, eds. *Red Book: 2015 Report of Committee on Infectious Diseases*. 30th ed. Elk Grove Village, IL; American Academy of Pediatrics; 2015 [400-423]
- Heymann, David L. (2015) *Control of Communicable Disease Manual*. Washington, DC.: APHA



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49
