





# Occupational Liver Disease

by: M.Saraei

# *Introduction*



- The liver is the target organ of many occupational and environmental chemicals and plays a central role in their detoxification and elimination
- Hepatic injury dose not differ clinically or morphologically from drug induced damage
- Occupational liver disease may be of secondary importance to damage
- Effects of multiple hepatotoxic exposure
- Lack of sufficiently sensitive and specific tests

- 
- 
- **Occupational history**
  - **result of personal or workroom air sampling**
  - **Remove the patient from exposure to suspected toxin**



# *Routes of Exposure*

- Inhalation
  - Ingestion
  - Percutaneous absorption
- 
- Inhalation the most important route particularly for volatile solvent
  - Percutaneous absorption is the most important for lipophilic agent

- 
- 
- Oral intake of hepatotoxic agents is usually of importance only in the rare case of accidental ingestion
  - Mouth breathing and gum and tobacco chewing can increase the amount of gaseous substances absorbed during the workday.



# ***Physical agent induced liver disorders***

- **Hyperthermia ( Heat Stroke)**
  - **Acute hepatic injury**
    - **Centrilobular necrosis**
    - **Cholestasis**
- **Ionizing radiation**
  - **A cumulative dose in excess of 3000 to 6000 Rad**
  - **Accidental intense exposure**
  - **Hepatitis 2-6 Week later**

# *Chemical agents associated with occupational liver disease*

<b>Arsenic</b>	<b>Cirrhosis, Hepatocellular carcinoma, Angiosarcoma</b>	<b>Pestiside</b>
<b>Beryllium</b>	<b>Granulomatosis disease</b>	<b>Ceramics workers</b>
<b>Ccl4</b>	<b>Acute hepatocellular injury, cirrhosis</b>	<b>Dry cleaning</b>
<b>Dimethylnitrosamine</b>	<b>Hepatocellular carcinoma</b>	<b>Rocket mfg</b>
<b>Dioxin</b>	<b>Porphyria cutanea tarda</b>	<b>Pestiside</b>
<b>Halothane</b>	<b>Acute hepatocellular injury</b>	<b>Anesthesiology</b>
<b>Hydrazine</b>	<b>Steatosis</b>	<b>Rocket mfg</b>
<b>Nitropropane</b>	<b>Acute hepatocellular injury</b>	<b>painter</b>
<b>PCB</b>	<b>Subacute injury</b>	<b>Electrical utility</b>
<b>TNT</b>	<b>Acute or Subacute hepatocellular injury</b>	<b>Munitions workers</b>
<b>Trichloroethylene</b>	<b>Acute hepatocellular injury</b>	<b>Cleaning solvent sniffing</b>
<b>Vinylchloride</b>	<b>Angiosarcoma</b>	<b>Rubber workers</b>

# *Acute hepatic injury*

- Cytotoxic injury or cholestatic injury
- Latent period 24 - 48 hours
- Clinical symptoms are often extra hepatic origin
- Anorexia, nausea, vomiting, jaundice, hepatomegaly
- In massive necrosis coffee-ground emesis, abdominal pain, reduction of liver size, ascites, edema, hemorrhagic diathesis
- During 24-48 hours somnolence and coma



# *Carbon Tetrachloride*

- Use as a liquid solvent, dry cleaning agent, fire extinguisher
- Dizziness, Headache, Visual disturbance, Confusion
- Hepatic disease occurs after 2-4 days
- Hepatomegaly, Splenomegaly, Jaundice
- Elevated serum transaminase, prolonged PT
- Hemorrhage, Hypoglycemia, Encephalopathy
- Renal failure may ensue a few days after the hepatic damage becomes manifest
- N- acetylcysteine



# Sub acute injury

- **Rare**
- **Most common: necrosis**
  - **TNT, Tetrachloroethane, PCB**
- **Symptoms:**
  - **Anorexia, Nausea, vomiting, Hepatomegaly & Jaundice**

# Chronic injuries

- **Asymptomatic until advanced stages**
- **Cirrhosis:**
  - Arsenicals, dimethylnitrosamine, CCL4, TNT, PCB
- **Steatosis:**
  - DMF, CCl4
- **Hepatoportal Sclerosis:**
  - VCM & Arsenic
- **Hepatic porphyria:**
  - Dioxin
- **Granuloma:**
  - Beryllium, Copper
- **Neoplastic changes:**
  - VCM, Arsenic

# Infectious Agents

HAV	Nursery & kindergarten staff Sewer workers
HBV & HCV	HCWs with blood and body fluid contact
Cytomegalovirus	Pediatric health care workers
Coxiella burnetti	Animal care workers, farm workers, slaughterhouse workers
Leptospira icterohaemorrhagiae	Sewer workers, farm workers



# Hepatitis A

١٣

- **High Risk Occupations:**
  - **HCWs**
    - Emergency rooms, surgery, laundry, children's psychiatry, dentists ,neonatal intensive care units
  - **Waste water treatment plant workers**
  - **Waste pickers,**
  - **home health workers**
  - **food handlers**
- **Incubation period:** 15-50 days (28-30 days)
- **Symptoms:** Abrupt onset, with fever, malaise, anorexia, nausea, abdominal discomfort, and jaundice
- **Transmission:** fecal-oral, Blood (rare)

# Hepatitis A

- **Highest concentration of virus excretion in Fecal:**
  - During the incubation period
  - Early in the prodromal phase
  - It diminishes rapidly once jaundice appears
- **Greatest infectivity:**
  - 2-week period immediately before the onset of jaundice or elevation of liver enzymes
- **Not chronic carrier**

- 
- 
- Fulminant hepatitis occurs rarely (<1% overall), but rates are higher with **increasing age** and in those with **underlying chronic liver disease**, including those with chronic hepatitis B or C infection.
  - Hepatitis A does not appear to be worse in HIV-infected patients when compared to HIV-negative persons

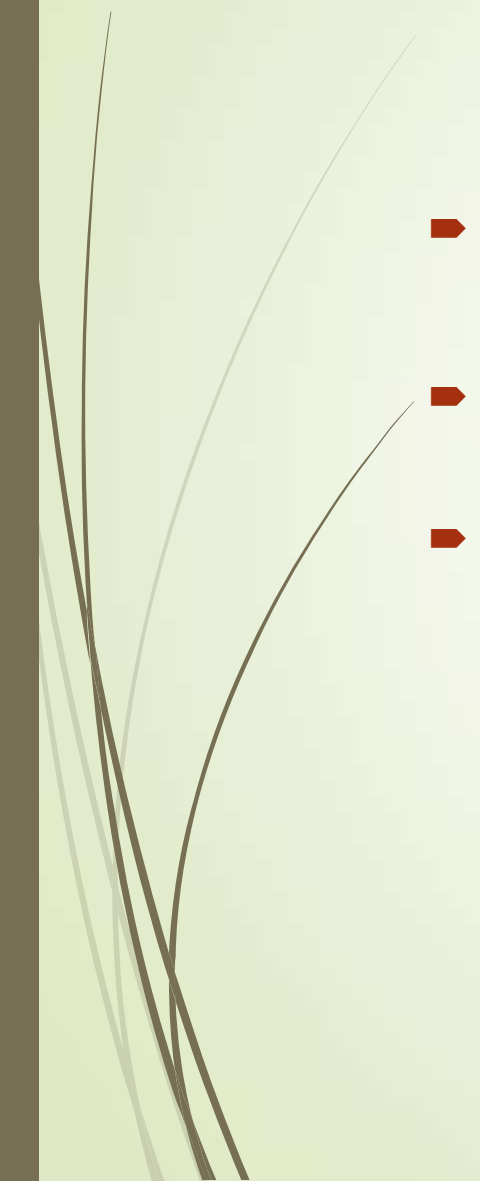
The diagnosis of acute hepatitis A is confirmed by:

- Presence of immunoglobulin IgM class anti-HAV in serum collected during the acute or early convalescent phase of the disease.
- IgG antibodies appear in the convalescent phase and remain positive for life
- The presence of IgG anti- hepatitis A antibody indicates either previous exposure or immunization





# Treatment

- symptomatic, with rest, analgesics, and fluid replacement
  - Fulminant hepatic failure occasionally follows acute HAV infection.
  - Orthotopic liver transplantation
- 





# Prevention

- Hand washing
- Avoiding tap water and raw foods in areas with poor sanitation.
- Heating foods appropriately  $>85^{\circ}\text{C}$  for 1 minute
- Chlorine, iodine, and disinfecting solutions (household bleach 1:100 dilution)



# Prevention

- A single intramuscular dose of 0.02 mL/kg of immune globulin (immune serum globulin, gamma globulin) given before exposure or during the incubation period

- 
- 
- **Pre exposure prophylaxis in persons who plan to travel in areas with high or intermediate hepatitis A endemicity depends on the duration of the travel:**
    - up to 1 month: 0.1 mL/kg**
    - up to 2 months: 0.2 mL/kg**
    - 2 months or longer: repeat dose of 0.2 mL/kg every 2 months**
  - **Post exposure prophylaxis, it is 0.1 mL/kg.**



# Prevention

- Once the diagnosis of acute infection is made, **close contacts** should be given HAV vaccine and/ or immune globulin promptly within **2 weeks** of exposure to prevent development of secondary cases.
- close contacts include:
  - staff of day-care facilities
  - food handlers (in establishments with a food handler diagnosed with hepatitis A)
  - institutions for custodial care
  - hospital staff if an unsuspected patient has been fecally incontinent

- Immune globulin can be used in cases where hepatitis A vaccination is contraindicated
- where travel is imminent
- It is less protective and only for short periods of time

## **Routine immune globulin administration is not recommended**

- **The usual office or factory conditions for persons exposed to a fellow worker with hepatitis A**
  - **Teachers with schoolroom contact**
- BUT**
- **Restaurant employees(food handlers) ,patrons**

- 
- 
- Food handlers should receive immune globulin when a common-source exposure is recognized
  - Restaurant patrons when the infected person is involved directly in handling uncooked foods without gloves.



This is especially the case when the patrons can be identified **within 2 weeks of exposure** and the food handler's hygienic practices are known to be deficient



- **Pregnancy or lactation is not a contraindication to immune globulin administration.**


# hepatitis A vaccine

- Persons traveling to or working in countries with intermediate or high HAV endemicity
- laboratory workers with exposure to live virus
- Animal handlers with exposure to HAV-infected primates
- Men who have sex with men (MSM),
- Illicit drug users (injections and non injection)
- Individuals with chronic liver disease,
- Individuals with clotting factor disorders
- Individuals with direct contact to others who have hepatitis A and homeless individuals.

- 
- 
- **Protective antibodies remain for as long as 4 years after two-dose vaccine series**
  - **There is no need for HAV booster vaccination after completion of the primary two-dose vaccination series**

## **Routine hepatitis A vaccination is not recommended for :**

- **Child-care workers**
- **Hospital workers**
- **Teachers**
- **Correctional workers**
- **Restaurant employees**
- **sewage treatment employees,**
- **Staff in institutions for the developmentally disabled.**




■ When outbreaks are recognized in these settings:


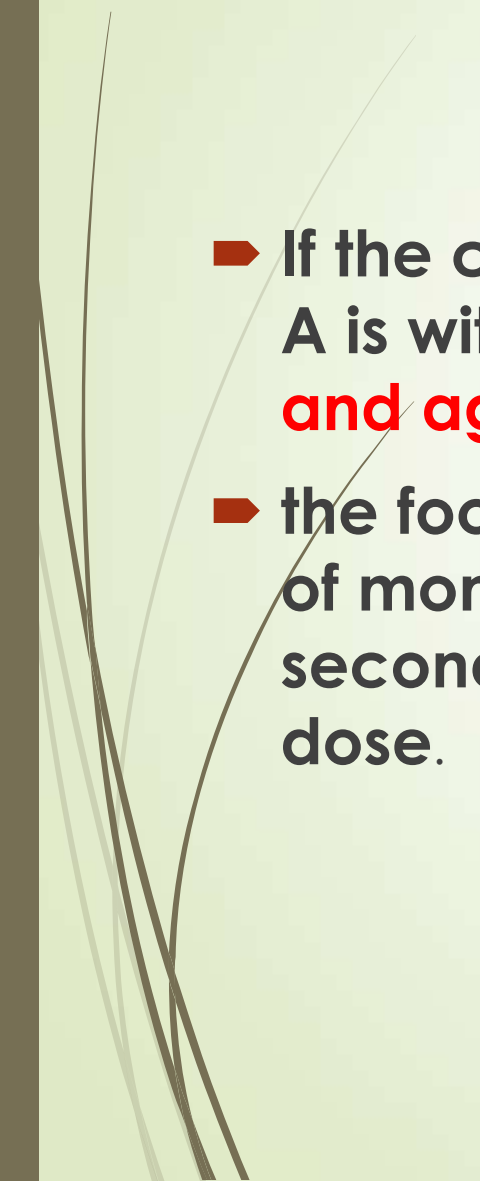
use of HAV vaccine and/ or immune globulin


promptly within 2 weeks of exposure for persons

in close contact with infected patients or


students is recommended

- 
- If a food handler has been in contact with an individual who is acutely infected with hepatitis A. If he has been immunized against hepatitis A with documented evidence of a completed course of hepatitis A vaccine in the past 10 years, or one dose of monovalent vaccine within the past 12 months, they can be considered immune.
  - Those who have had laboratory- confirmed hepatitis A (previous anti- HAV IgG positive, or HAV RNA positive) can also be considered immune and then no further action is required

- 
- 
- If the contact with the source case of hepatitis A is within 14 days, provided they are **healthy and aged under 60**
  - the food handler should be given a first dose of monovalent hepatitis A vaccine and a second dose 6– 12 months after the initial dose.

- 
- Food handlers **aged 60 years and over** should be offered hepatitis A immunoglobulin in addition to monovalent hepatitis A vaccine.
  - A second dose of vaccine is recommended 6– 12 months after the first dose to ensure long- term protection.<sup>5</sup>



- 
- If the food handler has not been immunized within 14 days of exposure
  - they are at high risk of acquiring infection and should be **removed from activities** which involve preparing and handling ready- to- eat foods until 30 days post exposure.


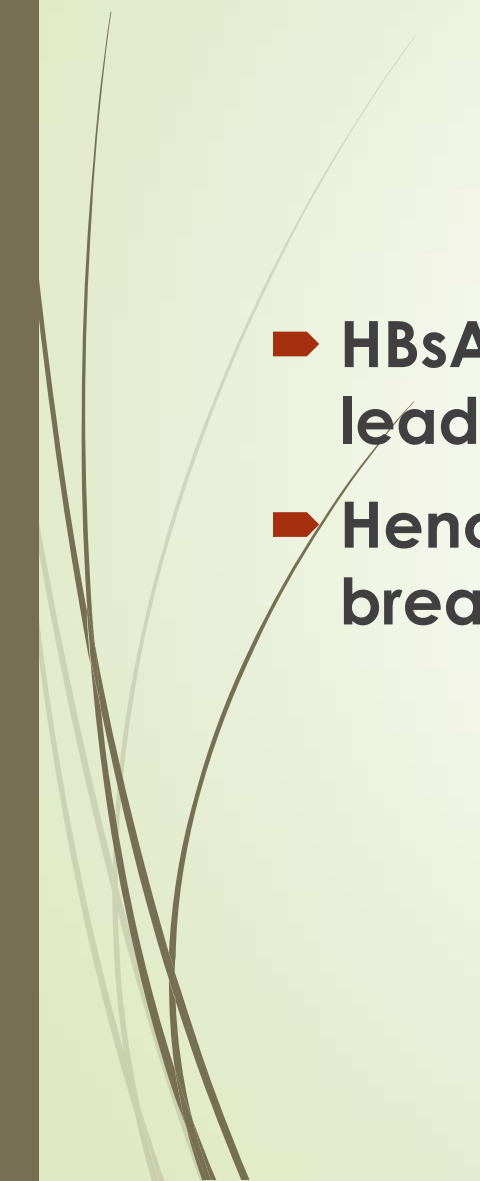
- 
- 
- If a worker is suspected of being infected with hepatitis A, should be excluded from work **until 7 days** after the onset of jaundice or
  - If there is no history of jaundice, 7 days after the onset of symptoms

# Hepatitis B

- Prevalence rate of HBS-Ag in hospital staff : **1-2%**
- Prevalence rate of Anti-HBS: **15-30%**
- Prior to hepatitis B vaccine:
  - Most frequent occupational infection among health care, laboratory, and public safety workers following human blood or body fluid exposure
  - 10 times higher than general population
- **By Standard Precautions and preexposure vaccination: sharp decline**

# Hepatitis B


- Blood contains the highest titers of virus in infected individual
- Low levels in other body fluids:
  - Cerebrospinal, Synovial, Pleural, Peritoneal, Pericardial, Amniotic
  - Semen and Vaginal secretions
- Viral titers in urine, feces, tears, and saliva are low enough not be routes of transmission
  - except in cases of human bites that usually involve some blood transmission

- 
- 
- HBsAg found in breast milk is also unlikely to lead to transmission,
  - Hence HBV infection is not a contraindication to breastfeeding.

- 
- The risk of infection with HBV depends on:  
the titer of virions in the infectious fluid

**And**

correlates with the presence or absence of  
hepatitis e antigen in the source patient.

- The risk of infection following percutaneous injury with both HBsAg- and hepatitis B e antigen (HBeAg)-positive blood is 22-31%;
  - The risk of developing serologic evidence of HBV infection is 37-62%.
- 

# Hepatitis B

- Risk for transmission of HBV after needlestick injuries: approximately **30%**
- HBV can remain viable for **at least 1 month** on dried surfaces at room temperature
  - Individuals with open cuts or abraded skin or mucous membranes contact contaminated surfaces
  - Most occupational infections have no clear percutaneous injury leading to HBV transmission
- Over **50%** of acute infections in adults are **asymptomatic**

# Hepatitis B

- Incubation period: 45 to 60 days after exposure
- HBS-Ag can be detected in serum 30-60 days after exposure
- Anti- HBS develops after resolve of infection

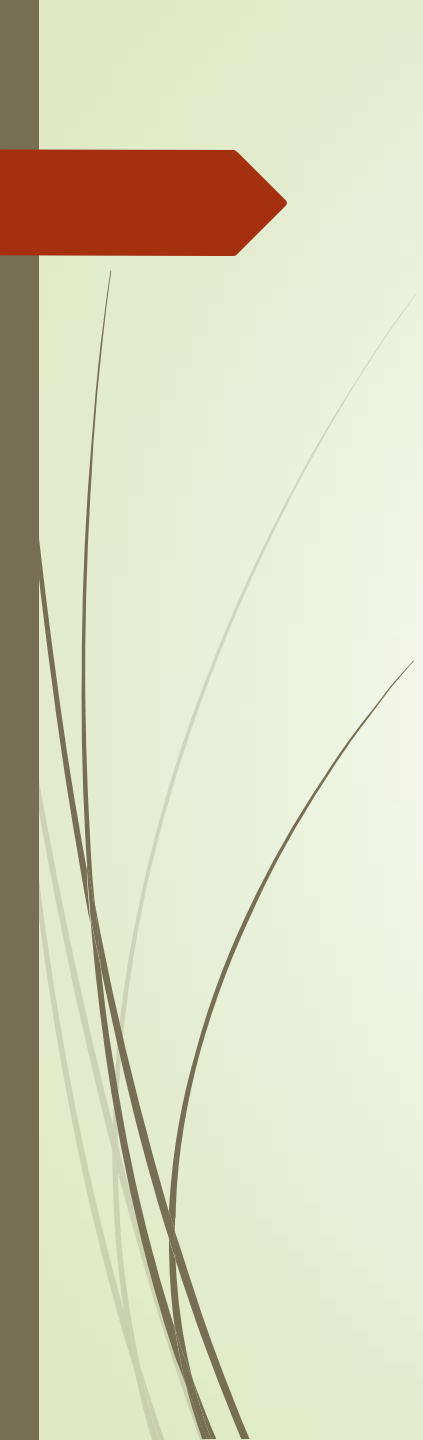
**Fulminant HBV infection is uncommon(<1%)**

- 10% of acute HBV infections lead to chronic infections
  - Significant number of those with occupational infections become chronic asymptomatic carriers



# Chronic Carrier State

- HBS Ag positive on at least two occasions **at least 6 months apart**
- High levels of HBS-Ag and anti-HBC
- Various levels of serum transaminases
- Progression to cirrhosis and hepatocellular carcinoma

- 
- Chronic infection develops more frequently in immunosuppressed persons :  
(eg, **hemodialysis** patients and persons with human immunodeficiency virus [**HIV**] infection) and persons with **diabetes**)

# Hepatitis B

Test	Result	Interpretation
HBS-Ag Anti-HBS Anti-HBC	Negative Negative Negative	Susceptible
HBS-Ag Anti-HBS Anti-HBC	Negative Positive Positive	Immune due to natural infection
HBS-Ag Anti-HBS Anti-HBC	Negative Positive Negative	Immune due to Hepatitis B vaccination

# Hepatitis B

Test	Result	Interpretation
HBS-Ag Anti-HBS Anti-HBC Anti-HBC (IgM)	Positive Negative Positive Positive	Acutely infected
HBS-Ag Anti-HBS Anti-HBC Anti-HBC (IgM)	Positive Negative Positive Negative	Chronically infected
HBS-Ag Anti-HBS Anti-HBC Anti-HBC (IgM)	Negative Negative Positive	Interpretation unclear: 1. Resolved infection 2. False positive Anti-HBC (Susceptible) 3. Low level chronic infection 4. Resolving acute infection

# Prevention

- Pre-exposure prophylaxis:
  - Vaccination in 3 doses: nearly 90% protection
- Who have received only one or two doses:
  - Not need to restart the series
  - Only need to complete the doses they did not receive
- Not immune after vaccination:
  - >45 years
  - obesity
  - smoking

# Prevention

- Check for surface antibodies:
  - 4 weeks to 6 months following the primary series
- Tests negative for antibodies:
  - one additional dose of vaccine will induce antibody protection in 15-25% of non responders
  - three additional doses (for a total of six doses) will induce antibodies in 30-50% of non-responders

# Prevention



- **Those who do not develop antibodies after six total doses should consider:**
  - Changing positions at work not involving blood or blood products
  - Three-dose series with 40 µg of antigen with either the Merck Recombivax HB formulated for hemodialysis patients
- **Fewer than 5% of persons receiving six doses fail to develop detectable anti-HBs antibody**
  - May have a low level of antibody that is not detected by routine serologic testing (“hyporesponder”)
  - Being chronically infected

# Prevention

- Loss of detectable anti-HB levels after immunization does not imply loss of protection
- **Positive antibody response should be verified**
- **Routine booster doses of hepatitis B vaccine are not recommended**





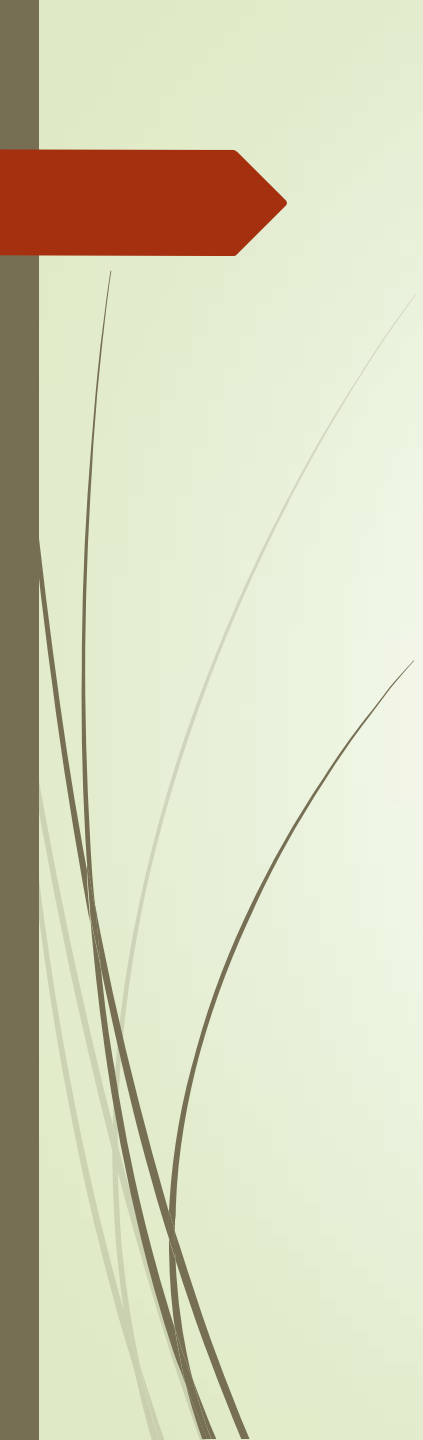
- 
- 
- **WHAT IS THE APPROPRIATE ADMINISTRATION **SITE FOR HEPATITIS B VACCINE** AND WHAT NEEDLE SIZE SHOULD BE USED?**



- 
- 
- ☐ A deep intramuscular (IM) injection into the deltoid muscle is recommended for adult hepatitis B vaccination.
  - ☐ A 22–25 gauge,” needle should be used, but a longer needle may be needed to reach deep into the muscle of obese persons.





➤ **IS IT SAFE FOR HCWS TO BE VACCINATED  
DURING PREGNANCY?**

- 
- 
- ☐ **YES** . Pregnant women in occupations with a high risk of hepatitis B virus (HBV) infection (e.g., HCWs) should be vaccinated.
  - ☐ Hepatitis B vaccine contains no components that have been shown to pose a risk to the fetus at any time during gestation.

- 
- IF HCWS HAVE **NO DOCUMENTATION** SHOWING THEY RECEIVED HEPATITIS B VACCINE. HOWEVER, THEY ARE RELATIVELY SURE THEY RECEIVED THE DOSES MANY YEARS AGO. WHAT DO WE DO NOW?

- 
- 
- ☐ Unfortunately, inadequate documentation of vaccination is common.
  - ☐ Even if physicians think they may have been fully vaccinated, but it is not documented, the three-dose vaccination series should be administered.
  - ☐ There is no harm in receiving extra doses of vaccine.

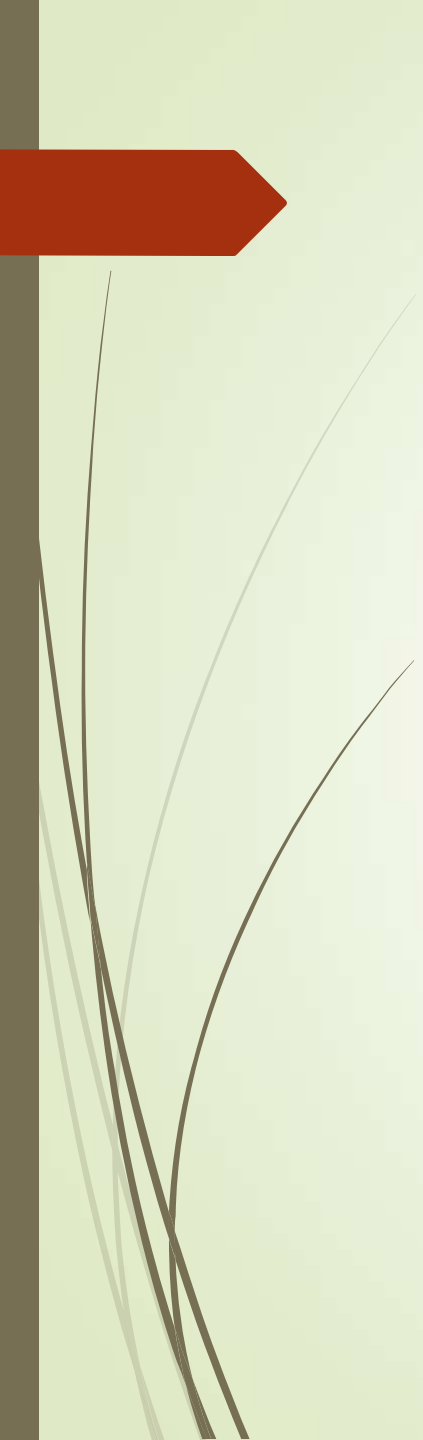
- 
- 
- A NURSE WHO RECEIVED THE HEPATITIS B VACCINE SERIES OVER **10 YEARS AGO** AND HAD A POSITIVE FOLLOW-UP TITER. AT PRESENT, THE TITER IS NEGATIVE. WHAT SHOULD SHE DO NOW?

- 
- 
- ☐ She doesn't need to do anything further.
  - ☐ For health care workers with normal immune status who have demonstrated an anti-HBs response following vaccination, booster doses of vaccine are not recommended nor is periodic anti-HBs testing.



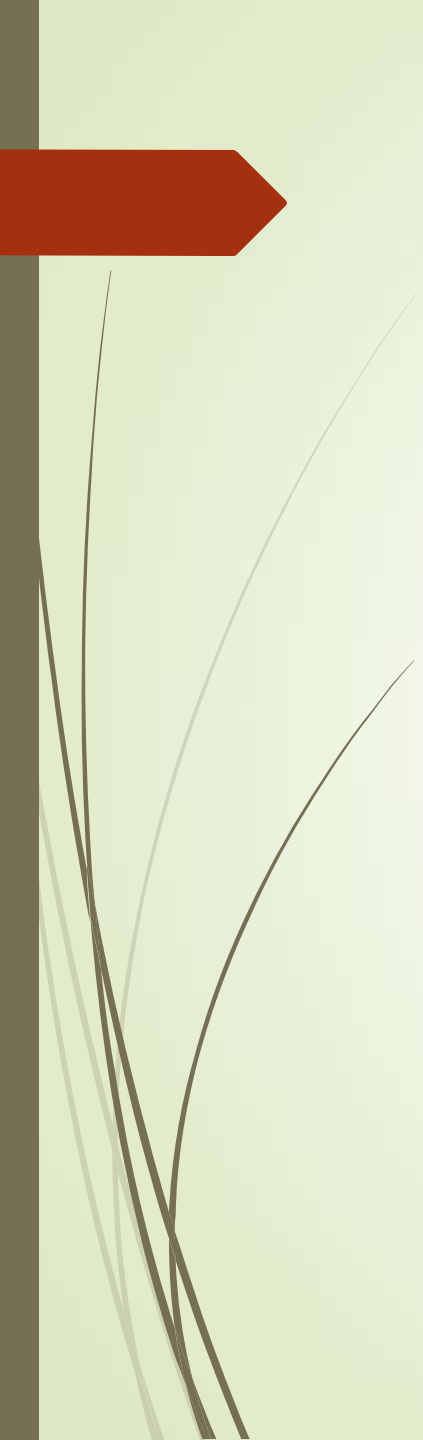
- 
- 
- A PERSON WHO IS A **KNOWN NON-RESPONDER** TO HEPATITIS B VACCINE HAS A PERCUTANEOUS EXPOSURE TO HBSAG-POSITIVE BLOOD. WHAT IS THE ACTION?

- 
- 
- ☐ The two-dose HBIG regimen would be the better choice.
  - ☐ The first dose of HBIG (0.06mL/kg) should be given as soon as possible after exposure and the second dose (same dosage) given one month later.

- 
- IF AN EMPLOYEE DOES NOT RESPOND TO HEPATITIS B VACCINATION, DOES HE NEED TO BE **REMOVED** FROM ACTIVITIES THAT EXPOSE HIM TO BLOOD-BORNE PATHOGENS? DOES THE **EMPLOYER** HAVE A RESPONSIBILITY IN THIS AREA BEYOND PROVIDING THE VACCINE?

- 
- 
- ☐ No regulations demand removal from the job situations described.
  - ☐ It is up to each organization to develop a policy concerning non-responders.

- 
- 
- DOES BEING CHRONICALLY INFECTED WITH HBV PRECLUDE ONE FROM **BECOMING A HEALTH PROFESSIONAL?**

- 
- ☐ **No.** All health professionals should practice standard precautions.
  - ☐ Those who are HBsAg-positive and HBeAg-positive should not perform exposure-prone invasive procedures (e.g., gynecologic, cardiothoracic surgery) unless they have been counseled by an expert review panel and been advised under what circumstances, if any, they may perform these procedures.
  - ☐ Such circumstances might include notifying prospective patients of the health professional's seropositivity before they undergo exposure-prone invasive procedures.

# Post exposure prophylaxis

## ➤ ☐ **Unvaccinated:**

- ☐ Source HBSAg+: HBIG & HBV vaccine series
- ☐ Source HBSAg-: initiate HBV vaccine series
- ☐ Source unknown: HBIG & HBV vaccine series

## ➤ ☐ **Previously vaccinated & Ab $\geq$ 10:**

- ☐ No treatment

## ➤ ☐ **Previously vaccinated & Ab less than 10:**

- ☐ Source HBSAg+:  
HBIG $\times$ 1 & revaccination or HBIG  $\times$ 2
- ☐ Source HBSAg-: No treatment
- ☐ Source unknown: if high risk source treat as HBSAg+

# Hepatitis C

## ➤ Transmission:

- Blood transfusion or intravenous drug abuse
- Sexual or mother to infant transmission is very low
- Risk of infection following occupational percutaneous exposure: 1.8% (0-7%)
- Mean incubation period is 6-8 weeks
- 80% cases being anicteric and asymptomatic
- Chronic hepatitis in 70% of infected person



# Hepatitis C

## ➤ Dx:

- Anti-HCV antibodies detectable 12 weeks-6 month following exposure
- HCV RNA by PCR (1-2 w after exposure)



# Prevention

- **Following percutaneous or mucosal occupational exposure:**
  - Baseline HCV Ab measurements
  - Follow-up HCV Ab measurements (6 w, 3 & 6 m)
- **Immune globulin or antiviral therapy for prophylaxis is not recommended**
- **No vaccine is currently available for HCV**



# Prevention

## ➤ **During this follow-up period:**

➤ Health care worker should refrain from:

➤ donating blood, plasma, organs, tissue, or semen

➤ **Health care worker should not need to:**

➤ modify sexual practices

➤ becoming pregnant

➤ breast-feeding



# Cytomegalovirus Infection

- Pediatric and immunosuppressed adult units
- kindergarten teachers
- Child-care workers
- Hepatitis
- Neonate with a congenital malformation



# *Coxiella burnetii*

- **Animal-care technicians**
- **Laboratory research personnel**
- **Abattoir workers**
- **Farmers**
- **Acute hepatitis occurs in up to 50% of cases and usually is self-limited.**


# *Malignant Liver Disease*

## ➤ **Hepatic Angiosarcoma**

- Vinyl chloride monomer
- Arsenic ( vineyard workers )
- Copper ( pesticide )
- Anabolic steroids
- Thorium dioxide( Thorotrast)



# Hepatocellular carcinoma

- ☐ vinyl chloride
  - ☐ Arsenic
  - ☐ Dimethylnitrosamine
- 





# Adjustments at work

- In patients with oesophageal varices, there is no restriction on occupation once the varices have been treated.
- Patients with ascites may experience difficulty with lifting, bending, or stooping.
- Patients with chronic or intermittent encephalopathy should not be employed in intellectually demanding work or jobs requiring a high degree of vigilance, including :

**Safety critical work or operating machinery**



- 
- 
- **Individuals suffering from hepatic cirrhosis with chronic encephalopathy or those who are cognitively impaired must not drive and must notify the Driver and Vehicle Licensing Agency (DVLA)**
  - **Group 1 and 2 licenses will be revoked or refused until recovery is satisfactory and other medical standards for fitness to drive (e.g. for psychiatric conditions) are satisfied.**



# Implications for employment

- **patients with chronic liver disease with ongoing inflammation or liver damage should not work with hepatotoxins.**
- **All patients working with hepatotoxins should avoid alcohol misuse and enzyme- inducing agents such as anticonvulsants, in particular phenobarbitate and phenytoin.**

# Medical Surveillance

- **Biochemical tests**
  - AST, ALT
  - ALP
  - LDH
  - Bilirubin
  - Urine bilirubin
- **Tests of synthetic liver function**
  - Alb
  - PT
  - Alpha fetoprotein
  - Ferritin
- **Clearance tests**
  - Sulfobromophthalein
  - Indocyanine green
  - Antipyrine test
  - Aminopyrine breath test
  - Serum bile acid
  - Urinary D-glucaric acid



# Transaminase

## ➤ **AST & ALT:**

- Most useful indicators of hepatocellular damage
- Sensitive not specific
- >8-10 folds: acute injury
- 2-3 folds chronic or mild acute
- In alcoholics ALT>300 is uncommon

➤ **Positive result:** Strongly suggest liver injury

➤ **Negative data:** Cannot rule out it

# Transaminase

## ➤ *High levels:*

- Viral
- alcoholic
- ischemic
- extrahepatic obstruction
- obese individuals

## ■ A serum AST:ALT ratio $> 1$ may suggest occupational liver disease

- Little Prognostic value
- Not correlate with extent of liver necrosis on biopsy



# Clinical Management of OCCUPATIONAL Liver Disease

- Occupational & medical Hx
  - Exposure to hepatotoxins
  - PMH of liver dis, medication
  - Review of symptoms(CNS Toxicity due to solvent exposure)
  - Travel to areas with endemic parasitic or viral disease
  - Steroid use, glue sniffing, recreational solvent use
  - Previous blood transfusion, tattoos, needle sticks, IV drug ...
  - Use of protective work practices
  - MSDS
  - Ask about other employees



# Clinical Management of OCCUPATIONAL Liver Disease

## ➤ Physical Examination

### ■ *Acute liver disease*

RUQ tenderness

Hepatosplenomegaly

Jaundice

### ■ **Chronic liver disease**

Spider angioma

Palmar erythema

Testicular atrophy

Ascites

Gynecomastia



# Clinical Management of OCCUPATIONAL Liver Disease


- **Elevated serum transaminase level**
  - **R/O non-occupational causes**
  - **Remove for 3-4 weeks**
  - **Repeat**
  - **A persistently elevated serum transaminase concentration suggests a non-occupational cause of liver disease or, rarely, chronic occupational liver disease.**









# TUBERCULOSIS

- Transmitted by the airborne route and, depending on host factors, may lead to latent tuberculosis infection (LTBI) or tuberculosis disease (TB).
- 

- 
- 
- **Health care settings (especially hospitals, long term care facilities, and dialysis centers)**
  - **Refugee/immigration centers**
  - **Homeless shelters**
  - **Substance abuse treatment centers**
  - **Correctional institutions**

- 
- 
- After an incubation period of 4-12 weeks, infection usually remains subclinical and dormant without development of active disease,
  - But the Mantoux tuberculin skin test (TST) will become positive
  - Once infection occurs, the organism may disseminate from the lungs to other sites:  
GI, GU & bone.
  - The risk for reactivation is highest in the first year after exposure.





**The risk of development of clinical disease following infection is higher:**

- **In those younger than 5 years**
- **In states of undernutrition**
- **In certain immunopathologic states (eg, in persons with HIV)**
- **with certain genetic predisposition (persons with HLA-Bw15)**
- **In persons with some coexisting diseases (silicosis, end-stage renal disease, leukemia, lymphoma, upper gastrointestinal tract carcinoma, diabetes)**

# Tuberculin Skin Test

- **TST skin testing is an accepted method for screening high risk populations for TB infection**
- Prior exposure to TB
- Delayed hypersensitivity
- Neither 100 % sensitive nor specific
- 0.1 ml of 5 IU , intra dermally into the or volar surface of the forearm(48 -72 hr)
- Positive TST :  
exposed to TB in the past and is at risk for reactivation
- New TST (+):CXR, smear & culture sputum



## **5 mm of induration**

- **In close contacts of infectious patients**
- **Those with fibrotic changes on chest radiograph consistent with prior TB**
- **Organ transplant recipients**
- **Persons with (HIV) infection**
- **Persons with other types of significant immunosuppression**



# 10 mm or more

- Is considered positive in high-risk occupational groups (including employees in prisons or jails, health care facilities, or mycobacterial laboratories)
- Injection drug users
- Persons who have immigrated in the past 5 years from countries with high TB prevalence
- Children younger than 5 years
- Those with the coexisting diseases





# 15 mm or more

- In persons with none of the previously mentioned risk factors
- 

## کنترل اندیکاسیون انجام TST

TST فقط برای افرادی که واکنش شدید (مانند نکروز، تاول، زخم یا شوک آنافیلاکتیک نسبت به TST قبلی داشته اند) ممنوع است.

TST برای هیچ فرد دیگری از جمله نوزادان، کودکان، زنان باردار و شیرده یا افرادی که مبتلا به HIV یا دارای ضعف سیستم ایمنی هستند منع انجام ندارد.

تکرار TST در فواصل کوتاه هیچ خطری برای فرد ندارد مگر اینکه TST قبلی با واکنش شدید همراه بوده باشد.

# تفسیر کانورژن

- الف : افزایش مساوی یا بیش از ۱۰ میلی متر در یکدوره دو ساله برای افراد زیر ۲۵ سال
- ب : افزایش مساوی یا بیش از ۱۵ میلی متر در یکدوره ی دو ساله برای افراد بالای ۲۵ سال

# نتایج PPD در HCWs

■ الف : در HCWs هایی که خطر تماس با باسیل سل وجود ندارد و یا حداقل است و عوامل خطر دیگری ندارند اندوراسیون ۱۵ میلی متر و به بالا مثبت تلقی می شود .

■ ب : در HCWs هایی که خطر تماس با باسیل سل دارند و عوامل خطر دیگری وجود ندارد ، اندوراسیون ۱۰ میلی متر و به بالا مثبت تلقی می شود .

■ در HCWs ها معمولا افزایش ۱۰ میلی متری در قطر اندوراسیون ظرف مدت دوره ی ۲ ساله می تواند مثبت تلقی شود .

واکسیناسیون با ویروس های زنده از جمله سرخک ،اوریون،سرخجه ،فلج اطفال ،واریسلا و تب زرد ممکن است واکنش های TST را مختل کند . در این افراد اگر قرار است TST انجام شود به صورت زیر انجام می شود:

الف : یا در همان روز واکسیناسیون با واکسن ویروس زنده

ب : یا حداقل یک ماه پس از تزریق واکسن ویروس زنده



The TST test may be negative in the presence of :

- Overwhelming tuberculosis
- Measles
- Hodgkin disease
- Sarcoidosis
- Immunosuppressive states
- If the initial test is negative in individuals with suspected **reduced immune response** or in those who **will be screened annually** because of occupational or other risk, **it should be repeated**



# Two-step method

- If the first TST result in the two step baseline testing is positive, consider the person infected and evaluate and treat the person accordingly
- If the first test result is negative, the TST should be repeated in 1-3 weeks
- If the second test result is positive, consider the person infected and evaluate and treat the person accordingly
- If both steps are negative, consider the person uninfected and classify the TST as negative at baseline testing

استفاده از روش دو مرحله ای یا تک مرحله ای TST می تواند بر اساس متد زیر باشد:

۱- هیچ نتیجه ای از سابقه قبلی TST در دسترس نیست:

TST پایه ای دو مرحله ای انجام شود.

۲- سابقه TST منفی برای بیشتر از یکسال قبل از ورود به سیستم بهداشتی درمانی وجود دارد:

تست TST پایه ای دو مرحله ای انجام شود

۳- سابقه تست TST منفی در یکسال اخیر وجود دارد یا دو تست منفی TST در بیش از یکسال وجود دارد:

به عنوان تست پایه از TST یک مرحله ای استفاده می شود و این تست می تواند خودش به عنوان مرحله دوم باشد.



- 
- 
- Occupational candidates who should be given high priority for LTBI treatment include :

Employees of high-risk congregate setting including :

- Correctional facilities
- Nursing homes
- Homeless shelters
- Hospitals, other health care facilities
- Mycobacteriology laboratory personnel

## پیشگیری

- داروی انتخابی جهت درمان عفونت نهفته سل **ایزونیازید** است و برای یک **دوره ۹ ماهه** تجویز می شود، البته گاهی در یک دوره ۶ ماهه تجویز می شود که در افراد زیر دوره ۶ ماهه **توصیه نمی شود** :

- HIV مثبت

- وجود ضایعات فیبروتیک در قفسه سینه

- کودکان

- در صورت عدم تحمل ایزونیازید، ریفامپین تجویز می شود که طول درمان با ریفامپین ۴ ماه است.

- در مبتلایان به **HIV** جهت درمان عفونت نهفته سلی از ایزونیازید بمدت ۹ ماه یا ریفامپین بمدت ۴ ماه استفاده می شود.

- در افراد HIV مثبت که تحت درمان دارویی با داروهای آنتی رتروویرال هستند از **ریفابوتین** بجای ریفامپین

استفاده می شود.

## پیشگیری

- داروی انتخابی جهت درمان عفونت نهفته سل **ایزونیازید** است و برای یک **دوره ۹ ماهه** تجویز می شود، البته گاهی در یک دوره ۶ ماهه تجویز می شود که در افراد زیر دوره ۶ ماهه **توصیه نمی شود** :

- HIV مثبت



- وجود ضایعات فیبروتیک در قفسه سینه



- کودکان

- در صورت عدم تحمل ایزونیازید، ریفامپین تجویز می شود که طول درمان با ریفامپین ۴ ماه است.

- در مبتلایان به **HIV** جهت درمان عفونت نهفته سلی از ایزونیازید بمدت ۹ ماه یا ریفامپین بمدت ۴ ماه استفاده می شود.

- در افراد HIV مثبت که تحت درمان دارویی با داروهای آنتی رتروویرال هستند از **ریفابوتین** بجای ریفامپین استفاده می شود.

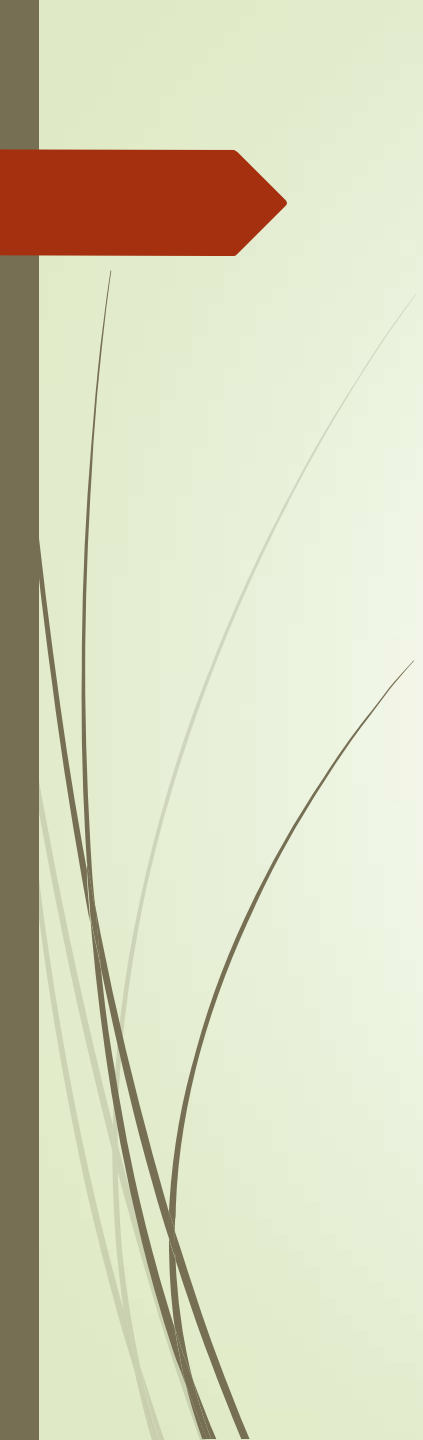
- 
- 
- Isoniazid (INH) plus rifapentine dosed weekly for 12 doses, rifampin dosed daily for 4 months, or INH dosed daily for 6 or 9 months.

- 
- 
- **Persons not known to have a previously positive skin test who have substantial contact with a person with infectious TB should be skin tested immediately**
  - **Then retested 8-12 weeks after the infectious contact**
  - **If conversion occurs, physical examination and chest radiography should occur to rule out acute clinical infection before beginning therapy for LTBI**



# **HIV/AIDS in the Workplace**

- **How is HIV transmitted:**  
from an infected person by body fluids  
such as blood or other blood  
containing secretions
- **Preventive measures:**  
wearing protective clothing, gowns,  
gloves, masks and goggles

- 
- **US: 5 % of individuals with AIDS have been employed in healthcare setting.**
  - **< 0.3 % of HCW were documented to have become HIV positive following occupational exposure most are nurses and laboratory workers.**
  - **Increase risk**
    - **Deep injury**
    - **Visible contamination of the device with blood**
    - **Needle placement directly into an artery or vein**
    - **Exposure to an individual with elevated viral titers**



# Post Exposure Management

**If exposure occurs :**

➤ **Skin**

Wash with soapy water

Do not use caustic agent or bleach

➤ **Eye, nose, mouth**

Rinse with water for 10 minutes

➤ **Needle stick or cut**

Wash with soapy water

Allow to bleed freely

Apply first aid





# Post Exposure Management

- Test healthcare worker for HIV after exposure at baseline.
- Treatment, if started, should be initiated immediately after exposure, within 1- 2 hours.
- Continue treatment for 4 weeks.



# **Follow up Testing and Appointments**

- **Follow up testing**

**HIV testing at baseline , 6 wk. 12 wk, and 6 months post exposure**

- **4 th generation combination p 24 antigen antibody HIV test :**

**HIV testing :at baseline , 6 wk. , and at 4 months post exposure.**

**CBC, Renal and Hepatic Function Tests(At baseline and 2 weeks post exposure)**

**Thank you for your attention**  
**Any Questions?**

