Viral Hepatitis

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Background

• Viruses commonly affect the liver, resulting in a transient and harmless hepatitis.

• Viruses which target the liver are primarily described as hepatotropic viruses, and each of these can lead to clinically significant hepatitis and in some cases to the development of **chronic viral hepatitis** with **viral persistence**.

Background

- Five human viruses have been well described to date, including hepatitis
 - 1. A (HAV)
 - 2. B (HBV)
 - 3. C (HCV)
 - 4. D (HDV)
 - 5. E (HEV)

- Hepatitis A virus (HAV) is a major cause of acute hepatitis worldwide, accounting for up to 25% of clinical hepatitis in the developed world.
- HAV is an **enteric virus** and the **fecal oral route** is the main mechanism of transmission.
- The virus is particularly **contagious** and constitutes a public health problem throughout the world.

• The virus may go unnoticed by the patient particularly in children. However, HAV can cause acute liver failure in less than 1% of patients who are typically older adults.

• Types A is classically associated with an acute and sometimes severe hepatitis which is always **self-limited**, but **occasionally fatal**.

• The virus is particularly prevalent in areas of **poor sanitation**, and often associated with **water- and foodborne epidemics**.



Hepatitis A has a relatively short incubation period (2–7 weeks), during which time the virus replicates and abnormalities in liver function tests can be detected.



Is a viral infection of the liver spread when faecal matter enters the mouth



May last several weeks and can be debilitating but most people recover completely

Preventable with careful hand washing, keeping toilets and bathrooms clean, avoiding infected water sources

HEPATITIS A FACTS

SYMPTOMS INCLUDE



direct

contact



SPREAD BY





food & beverages cups & spoons

and any other objects handled by the infected person

• Up to **500 million** people worldwide are **chronically infected** with the hepatitis B virus (HBV).

• In endemic areas of Africa and the Far East, up to 15–20% of the population are chronic carriers of HBV.

• The single most important **risk factor for the development of chronic HBV** infection is **vertical transmission** (i.e., mother-to-child transmission) at birth.



 Hepatitis B causes acute hepatitis in adults and 5% of patients become chronic carriers, while
95% of patients infected in the neonatal period are chronically infected.



• Getting of HBV in adulthood is often via sexual transmission.

• HBV can also be transmitted **parenterally**, by the **transfusion of blood** or **blood products** and by **intravenous drug** use or **needle sharing**.



• Acute HBV infection has a long incubation period of 3–6 months, which is generally self-limiting and does not require antiviral therapy.

• Most patients recover within 1–2 months of the onset of jaundice.

- It is estimated that 15–40% of chronic HBV carriers will develop serious sequelae during their lifetime, namely
 - liver cirrhosis
 - hepatocellular carcinoma

- The primary goal in the management of chronic HBV infection is to prevent
 - cirrhosis
 - hepatic failure
 - hepatocellular carcinoma

• The eradication of HBV is near impossible because of the presence of extra-hepatic reservoirs of HBV and integration of HBV into the host genome.

• For this reason, the main goal of antiviral agents is continuous viral suppression.

• Potent oral antiviral agents **tenofovir** and **entecavir** as **first-line monotherapies**.

• While these two agents have emerged as the leading oral antivirals, the weaker and more outdated agents such as telbivudine, lamivudine and adefovir are still widely used.

• The use of lamivudine or adefovir as monotherapy is no longer recommended and should be avoided if at all possible, owing to the high rates of resistance reported with these drugs.

• While reported rates of **resistance** is **extremely low with entecavir**, and **none reported with tenofovir**.

• When start oral antiviral agents, the patient must be aware they are potentially goin on a lifelong course of treatment.



• **Pegylated interferon (peginterferon) alfa-2a** has re-emerged as a viable alternative to oral antiviral agents in the treatment of chronic HBV.

• Peginterferon alfa-2a dose is 180 mcg SC once weekly for 48 weeks.

- Interferon therapy advantages:
 - fixed treatment course
 - good rates of surface antigen loss in selected patients

• The earlier stated advantages must be weighed against the disadvantages associated with an injection-based therapy (48 weeks) and the interferon side effect.



- Peginterferon alfa-2a side effects include
 - influenza-like symptoms
 - decrease in hematological parameters (haemoglobin, neutrophils, white blood cell count and platelets)
 - gastrointestinal complaints
 - psychiatric disturbances (anxiety and depression)
 - hypo- or hyperthyroidism

 Over 170 million people worldwide are chronically infected with hepatitis C virus (HCV).

• Hepatitis C rarely causes an acute hepatitis but up to 85% of patients become chronic carriers.

• It is estimated that more than 2.7 million people in the USA are chronically infected with HCV, where it is the leading cause of death from liver disease.

• HCV is a predominantly **blood borne virus** with **greater infectivity** than the **human immunodeficiency virus** (HIV).





• HCV is transmitted **parenterally**, most commonly through **intravenous contaminated blood** and **blood products**, and **the sharing of contaminated needles**.

• The vertical transmission rate from HCV infected mother to child is **less** than 3%.

• There remains a small risk of HCV infection associated with tattooing, electrolysis hair removal, ear piercing, acupuncture and sexual contact.



• The **primary aim** of treating patients with chronic HCV is **viral clearance** that defined as the absence of viraemia 6 months after antiviral therapy has been discontinued.

- Viral clearance
 - improves the patient's quality of life
 - reduces the risk of cirrhosis and hepatocellular carcinoma

• **Pegylated interferon** and **ribavirin** combination therapy are now the standard care of chronic HCV.

- The treatment duration, however, can be individualized based on the
 - baseline viral load
 - virological response during treatment

- Patients failing to achieve a significant reduction in viral load after 12 weeks will normally have therapy discontinued.
- The current standard of care combination therapy is limited by
 - the side effect profile (already stated in HBV)
 - complications of therapy
 - poor patient tolerability

- It is accepted that these side effects are a major obstacle preventing completion of therapy by
 - hindering compliance
 - enforcing significant dose reductions

• While erythropoietin and antidepressants may alleviate some side effects.

• Hepatitis D virus (HDV) is an incomplete virus that can establish infection only in patients simultaneously infected by HBV.

• It is estimated that 5% of HBV carriers worldwide are infected with HDV.

• It is endemic in the Mediterranean area and is transmitted sexually.

 In other geographical areas, it is confined to blood transfusions and intravenous drug users.
HEPATITIS

- Hepatitis E virus (HEV) is endemic in
 - India
 - Asia
 - Middle East
 - parts of Latin America

- It is transmitted **enterically** (i.e., via contaminated water and food) and leads to **acute hepatitis**.
- The symptoms of HEV are no different from other causes of viral hepatitis, with an average incubation **period of 42 days**.
- Hepatitis E infection during pregnancy, especially in the third trimester, is characterized by a more severe infection that sometimes results in increasing maternal and fetal mortality and morbidity.

• A vaccine to prevent hepatitis E virus infection has been developed and is licensed in China, but is not yet available elsewhere.

• As an acute disease is usually self-limiting.

• Immunosuppressed people with chronic hepatitis E benefit from specific treatment using ribavirin, an antiviral drug. In some specific situations, interferon has also been used successfully.


