

## HEPATTS

(GIT block, Microbiology: 2018)

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#### **OBJECTIVES**;

viruses causing entericaly transmitted hepatitis

> HAV. HEV.

viruses that are causing
 hepatitis during their course of infection;
 e.g Cytomegalovirus (CMV)
 Epstein-Barr virus (EBV)
 Arbovirus (yellow fever virus )

- structure
- Epidemiology
- clinical presentations
- Lab diagnosis
- > Treatment
- prevention

## HEPATITIS

## Viral hepatitis

- As part of generalized infection (CMV, EBV, Yellow fever virus)
- Infect primarily the liver
  - Faecal-borne hepatitis (A & E)
  - Blood-borne hepatitis (B, C & D)

#### FECAL-BORNE HEPATITS

- **HAV**
- Picornaviridae

- **HEV**
- **4** Hepeviridae

- **4**Nonenveloped
- **4**Icosahedral
- $\pm ss$ , + sense RNA
- **4**One serotype

#### HEPATITIS A VIRUS

## Hepatitis A

Short incubation hepatitis
Infectious hepatitis
Epidemic hepatitis





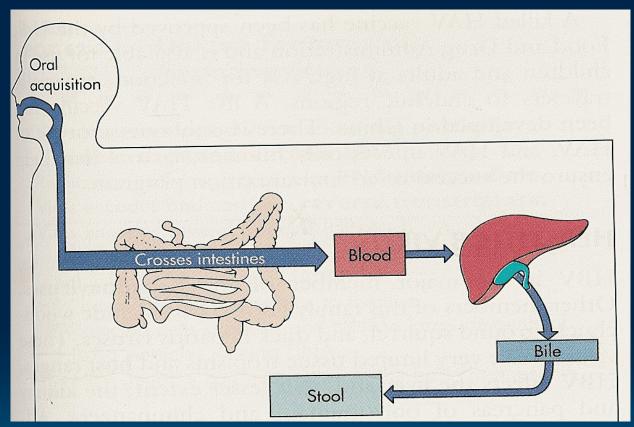
## **#**Epidemiology



- Distribution:
  - a worldwide, endemic in tropical countries
- Transmission:
  - Faecal-oral route [major route]
    Contaminated food &water
  - Sexual contact (homosexual men)
  - Blood transfusion (v.rarely)
- Age:
  - In developing countries; children\*
  - In developed countries; young adults

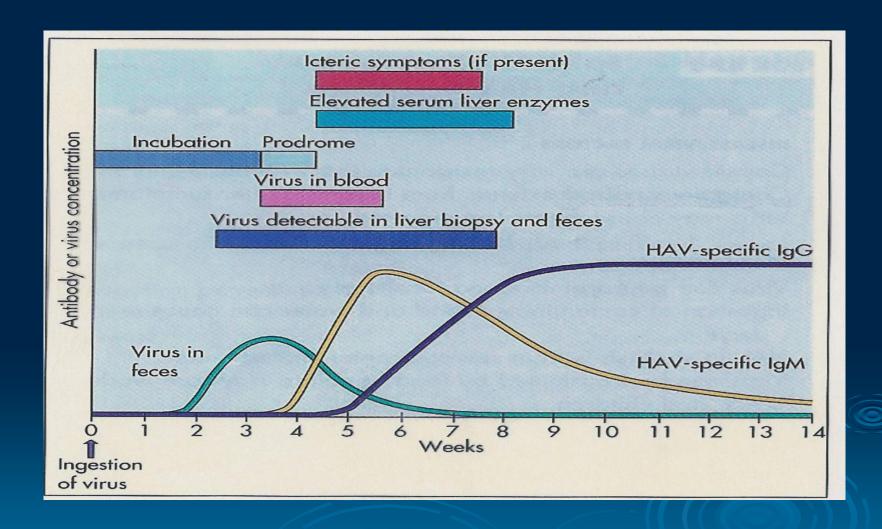
## #Pathogenesis





- CMI Damage of virus-infected hepatocyte
  - ALT ,AST & Bilirubin





#### **4** Manifestations



#### Hepatitis

- IP=2-6 Ws
- Pre-icteric phase: fever, fatique, N, V,& RUQP
- Icteric phase: dark urine, pale stool, jaundice





- Asymptomatic & anicteric inf —> common
- Symptomatic illness age



## #Prognosis

- Self-limited disease
- Fulminant hepatitis rare
- Mortality rate ~ 0.1 0.3%
- No chronicity or malignancy changes



## Lab. Diagnosis

#### Serology:

- Anti-HAV IgM —— Current inf
- Anti-HAV IgG —— previous inf
  - ---- immunity

## **4** Management



- Treatment:
  - Supportive therapy
- Prevention:
  - Sanitation & hygiene measures
  - HIg
  - Vaccine



## #Prevention

## HIg:

- Given before or within 2 Ws of exposure
- Indication: travelers
  unvaccinated, exposed p

## #Prevention



#### Vaccine:

- inactivated
- **4**Given IM at [0,6-12 M]
- ♣ >1 Y of age
- ♣S/E : mild local reaction





P at high risk of severe dis

**4**A combination vaccine (HAV &HBV)



## HEPATITIS E VIRUS

- Hepeviridae
- **#** Epidemiology:
- outbreak of waterborne & sporadic cases of VH
- Age; young adults
- **4** 4 routes of transmission;
  - Waterborne\*
  - Zoonotic foodborne
  - Bloodborne
  - Perinatal

#### **HEPATITIS E VIRUS**

#### **4** Clinical features:

- ~ HAV infection & exceptions:
  - Longer IP =4-8 Ws
  - Chronic hepatitis and cirrhosis (not HCC)
  - Fulminant disease
  - Mortality rate ~10 times > HAV
    - $\sim (1-3\%)$
    - ~ 20% in pregnancy

#### **HEPATITIS E VIRUS**

- Lab diagnosis:
  - ELISA Anti-HE IgM
- Treatment:
  - Not specific
- **#** Prevention:
  - Sanitation & hygiene measures
  - No Ig
  - No vaccine

## Herpesviridae

1-Herpes simplex virus type -1

2-Herpes simplex virus type -2

3- Varicella –Zoster virus

4-Epstein-Barr virus

5-Cytomegalovirus

6-Human herpes virus type-6

7-Human herpes virus type-7

8-Human herpes virus type-8

HSV-1

HSV-2

**VZV** 

EBV

**CMV** 

HHV-6

HHV-7

**HHV-8** 



dsDNA, Icosahedral & Enveloped Virus



#### <u> Epstein – Barr Virus EBV</u>

- > It is lymphotropic.
- > It has oncogenic properties;

Burkitt's lymphoma Nasopharyngeal carcinoma

#### **Epidemiology**

- Distribution : worldwide
- Transmission:
  - Saliva [kissing disease]
- Age:

Socio-economic status: SE

- Low SE class early childhood
- High SE class adolescence

#### **Clinical Features:**



#### <u> 1-Immunocompetent host</u>

- Asymptomatic
- Infectious mononucleosis [glandular fever]
  - Mainly in teenagers & young adults
  - $\triangleright$  IP = 4-7 weeks
  - Fever, pharyngitis, malaise, LAP, hepatosplenomegaly & abnormal LFT  $\pm$  hepatitis.
  - Complications( acute air way obstruction, splenic rupture, CNS inf)
- Chronic EBV infection

#### 2- Immunocompromised host

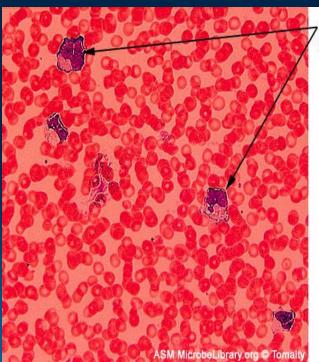
Lymphoproliferative disease (LD)

#### Dx:

#### Hematology:

■ Î WBC

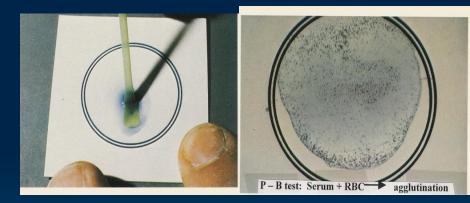
lymphocytosis
( Atypical lymphocytes )



Atypical lymphocyte with deformed nucleus and dark rimmed cytoplasm

#### Serology:

- Non-specific AB test;
  - Heterophile Abs +ve
  - Paul-Bunnell or mono-spot test



EB\

EBV-specific AB test:IgM Abs to EBV capsid antigen



#### **Management:**

- > Treatment:
  - Antiviral drug is not effective in IMN
- > Prevention:
  - No vaccine

## Cytomegalovirus CMV

- Special features ;
- Infected cell enlarged with multinucleated .
  - [cyto=cell, megalo=big]
- Resistant to acyclovir.
- Latent in monocyte ,lymphocyte & other .

- Distribution: worldwide .
- Transmission;
  - Early in life:
    - Transplacental
    - Birth canal
    - Breast milk
  - Young children: saliva
  - Later in life: sexual contact
  - Blood transfusion & organ transplant .



#### Acquired Infections;

- Immunocompetent host
  - Asymptomatic
  - Self-limited illness
    - Hepatitis
    - Infectious mononucleosis like syndrome [Heterophile AB is –ve]
- Immunocompromised host
  - Encephalitis , Retinitis , Pneumonia ,
  - Hepatitis\*, Esophagitis, Colitis



## Lab. Dx



#### \* Histology:

Intranuclear inclusion bodies

[Owl's -eye]

#### \* Culture:

In human fibroblast

1-4 wks → CPE

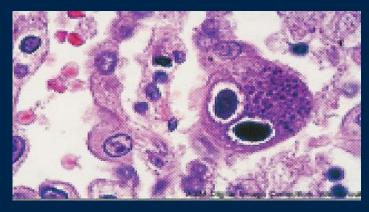
► Shell Vial Assay → 1-3 days

#### \* Serology:

► AB → IgM : current inf

→ IgG: previous exposure

Ag CMV pp65 Ag by IFA



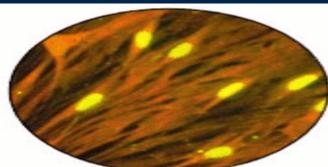


Fig. 2, CMV centrifugation culture fixed and stained 16 hrs after inoculation showing viral proteins in nuclei of infected human fibroblast cells



#### Rx.



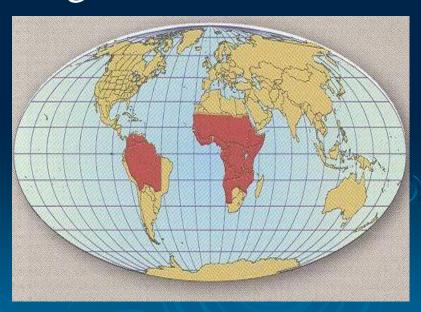
- Ganciclovir
  - is effective in the Rx of severe CMV inf.
- Foscarnet: the 2nd drug of choice.

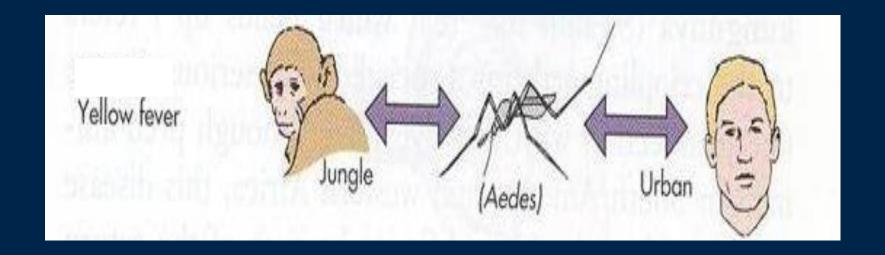
#### Prevention:

- Screening;
  - Organ donors
  - Organ recipients
  - Blood donors
- Leukocyte-depleted blood.
- Prophylaxis: Ganciclovir, CMVIG.
- No vaccine.

## <u>Arthropod – borne Viruses</u> (Arboviruses) **Yellow Fever virus**

- > Flaviviridae
- Asymptomatic to Fever ± Jaundice ±
   hemorrhage ± renal failure
- EpidemiologyTropical Africa& South America
  - 1. Jungle Yellow Fever
  - 2. Urban Yellow Fever





#### Jungle Yellow Fever:

- Vector: mosquito
- Reservoir: Monkey
- Accidental host: human
- It is a disease of Monkeys

#### **Urban Yellow Fever:**

- Vector: mosquito
- Reservoir: human
- It is a disease of humans

#### Dx.

- Lab. Methods:
  - A- Isolation
  - B IgM -AB\* ELISA, IF: (most used)
  - C YFV- RNA by RT-PCR

#### **Prevention**

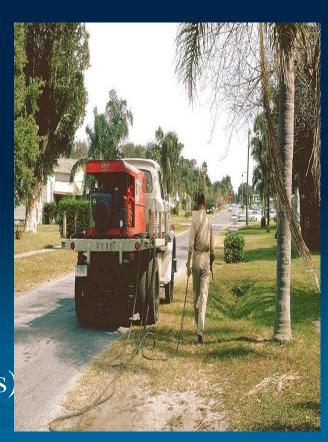
#### 1-Vector Control:

- Elimination of vector breading sites
- Using insecticides
- Avoidance contact with vectors

(repellants, net)

#### 2-Vaccine:

Yellow Fever vaccine (LAV, one dose /10 yrs)





## Reference books

## Review of Medical Microbiology and Immunology.

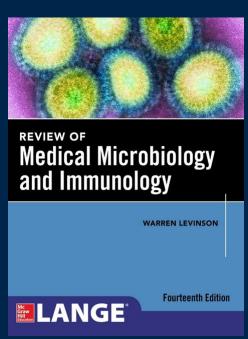
By: Warren Levinson.

14<sup>th</sup> Edition, 2014.

#### <u>Medical Microbiology.</u>

By: David Greenwood, Richard Slack,
John Peutherer and Mike Barer.

17<sup>th</sup> Edition, 2007.





# Thank gou