### EPSTEIN BARR EBV VIRUS & PARVOVIRUS





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**HLS Module** 

# EPSTEIN BARR VIRUS EBV







Infectious mononucleosis.
Burkitt's lymphoma.
B-cell lymphomas.
Nasopharyngeal carcinoma.
Hairy leukoplakia.

#### **Important Properties:**

- EBV is also called human herpesvirus 4 (HHV-4).
- Infects mainly lymphoid cells, primarily B lymphocytes.
- Infects the epithelial cells of the pharynx.
- In latently infected cells, EBV DNA is in the nucleus,

not integrated. Some genes are transcribed.

## Structure and genome:

- Ds linear DNA enveloped virus.
- Approximately **122 180 nm** in diameter.
- The nucleocapsid is surrounded by an envelop made of protein.



### **EBV** antigens and proteins:

### A. Latent proteins:

- 1. EB viral nuclear antigen complex (EBNA)
- 2. Latent membrane protein (LMP)
- 3. Terminal protein
- 4. Lymphocyte-detected membrane antigen (LYDMA)

### <u>B. Lytic cycle proteins:</u>

- 1. Membrane antigen (MA)
- 2. Early antigen complex (EA)
- 3. Viral capsid antigen complex (VCA)

### **Transmission & Epidemiology**

- Exchange of **saliva** (e.g., during kissing).
- Blood transmission  $\rightarrow$  very rare.
- Objects such as drinking glass,....
- EBV infection is worldwide.
- In the first few years of life  $\rightarrow$  asymptomatic.
- Clinically apparent infectious mononucleosis is highest in those who are exposed to the virus later in life (e.g., college students).

## **Pathogenesis & Immunity**

- Oropharynx →blood→ B lymphocytes (Latent). Cytotoxic T lymphocytes "atypical lymphs" react against the infected B cells.
- (Atypical lymphocytes are lymphocytes that have been activated to respond to a viral infection (occasionally a bacterial or parasitic infection).
- The immune response : IgM abs. to the VCA, IgG abs. to the VCA follows and persists for life.
- Lifetime immunity is based on antibody to the viral membrane antigen.
- Nonspecific heterophil antibodies → disappear within 6 months after recovery.

#### **Pathogenesis & Immunity**



## **Clinical Findings:**

#### Infectious mononucleosis :

- High risk group: 17-25 ys
- Acute, self-limiting
- Fever, sore throat, lymphadenopathy, anorexia, lethargy and splenomegaly. Hepatitis is frequent; encephalitis in some patients.
- Spontaneous recovery in 2 to 3 weeks.

Severe pharyngitis with diffuse pharyngeal inflammation and tonsillar swelling.

why is it called infectious mononucleosis? an acute infectious disease accompanied by atypical large peripheral blood lymphocytes.





## Hairy leukoplakia:

A <u>whitish</u>, <u>non-malignant</u> lesion with an irregular "hairy" surface on the lateral side of the tongue, in immuno-compromised individuals, especially AIDS patients.

- Burkitt's lymphoma, Hodgkin's lymphoma, nasopharyngeal carcinoma.
- post-transplant lymphoproliferative disorder (B-cell lymphoma).





#### Diagnosis

#### EBV is suspected when patients having

- fever
- pharyngitis
- lymphadenopathy

#### <u>Lab diagnosis</u>



- Full blood count
- Lymphocytosis
- - Atypical lymphocytes.
- Heterophile Abs
- (monospot method)
- Agglutination



#### EBV special Abs

- VCA- IgM
- VCA- IgG
- EBNA-IgG



RT-PCR for viral DNA



## **Laboratory Diagnosis:**

#### 1. Hematologic :

- Absolute lymphocytosis with >10% abnormal lymphocytes.
- Atypical lymph: are enlarged cells, expanded nucleus, and an abundant vacuolated cytoplasm.



### **Heterophile Antibodies**

- Are Abs produced by human immune cells in response to EBV infection.
- Negative in the incubation period and after the active infection has subsided.
- Heterophile means: they react with antigens other than the antigens that stimulated it.



### 2. immunologic:

- The heterophil antibody: early diagnosis of IM usually positive by week 2 of illness.
- The EBV-specific antibody tests: The IgM VCA for early illness; IgG VCA for prior infection, antibodies to EA and EBNA.

#### Molecular assays:

 Nucleic acid hybridization is the most sensitive means of detecting EBV in patient materials.

#### **Treatment:**

- None! Self-limiting
- Treat symptoms
- Misdiagnosis with streptococcal infection
- Acyclovir in high doses may be useful in life threatening EBV infections.





## PARVOVIRUS

## Parvovirus B19



#### Diseases:

- Erythema infectiosum (slapped cheek syndrome, fifth disease)
- > Aplastic anemia.
- > Hydrops fetalis.

#### Important properties:

A very small (22 nm), non-enveloped, icosahedral, with a single-stranded DNA genome, negative-sense, no virion polymerase. There is one serotype.





## **Transmission:**

- Respiratory route.
- Trans-placental.
- Blood donation.

## Pathogenesis & Immunity:

●B19 virus infects two types of cells: Erythroblasts in the BM → aplastic anemia. Endothelial cells →rash.

Immune complexes of virus and IgM or IgG  $\rightarrow$  rash, arthritis.

Infection provides lifelong immunity against reinfection.

#### **Clinical Findings:**

#### (1): Erythema Infectiosum (Slapped Cheek Syndrome, Fifth Disease):

 Mild disease, childhood, lowgrade fever, coryza, and sore throat, rash





 Children with chronic anemia → severe aplastic anemia (aplastic crisis)

#### (3): Fetal Infections:

- First trimester → fetal death.
- Second trimester →hydrops fetalis.
  (4): Arthritis:
  - Adults, women, small joints of the hands and feet bilaterally.

#### (5): Chronic B19 Infection:

 In immunodeficiencies: chronic anemia, leukopenia, or thrombocytopenia.



## Laboratory Diagnosis:

- IgM antibodies.
- In immuno-compromised patients, viral DNA by PCR.
- Fetal infection : PCR of amniotic fluid.



## **Treatment & Prevention:**

- No specific treatment.
- Immune globulins in immunodeficiencies.
- No vaccine.