

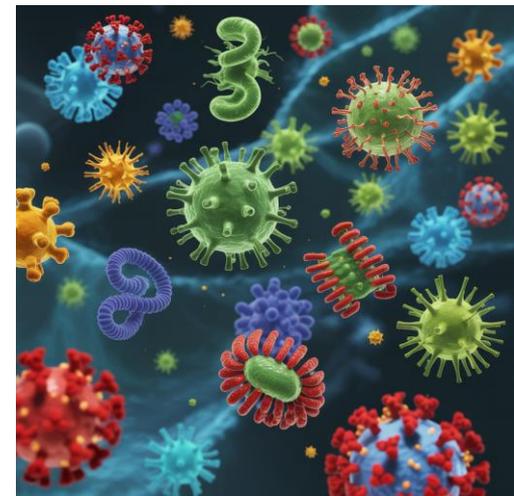
Virology I

Lecture 22

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Objectives

- What is a virus?
- Can we consider it to be alive or alive cell?
- What is the structure/function of the virus?
- How do viruses infect?
- What are prions?

History

- Louis Pasteur postulated that **rabies** was caused by a virus (1884).
- Ivanovsky showed a disease in **tobacco** was caused by a virus (1890s)
- 1950s virology was a multifaceted discipline
 - **Viruses**: noncellular particles with a definite size, shape, and chemical composition

The Position of Viruses in the Biological Spectrum

- Smallest infectious agents.
- There is no universal agreement on how and when viruses originated.
- Viruses are considered the most abundant microbes on earth.
- Viruses played a role in the evolution of Bacteria, Archaea, and Eukarya.
- Viruses are obligate intracellular parasites.

Virus properties

- Obligate intracellular **parasites** of bacteria, protozoa, fungi, algae, plants, and animals.
- Ultramicroscopic size ranging from 20 -750 nm(diameter) (**EM**); only **Pox viruses** by **LM**.
- Based on size viruses considered **Ultracentrifugation**.
- Not cellular in nature.
- Inactive macromolecules outside the host cell and active only inside host cells.
- Basic structure consists of protein shell (**capsid**) surrounding nucleic acid core.

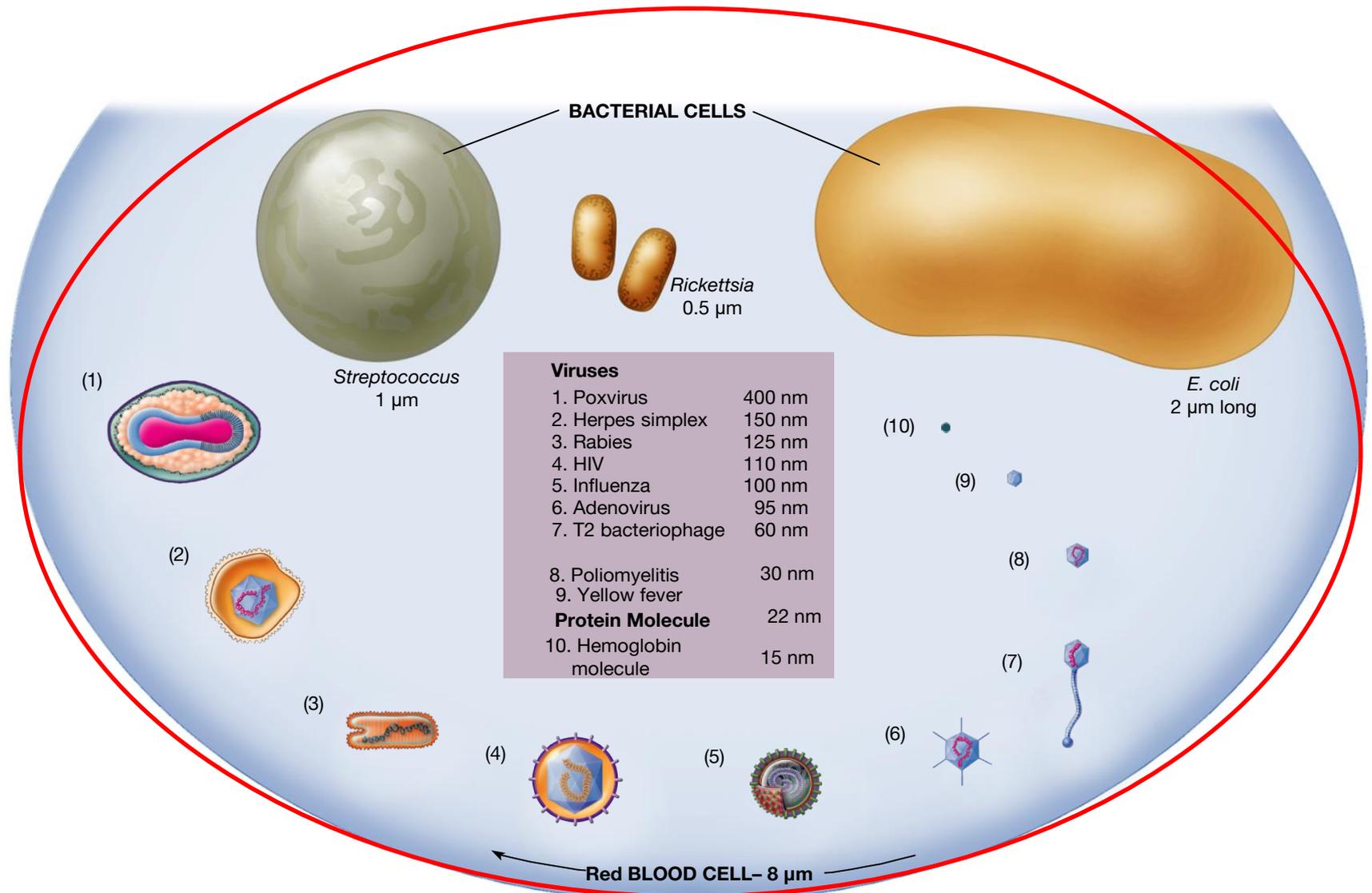
Con...

Virus properties

- Nucleic acid can be either DNA or RNA but not both.
- Nucleic acid can be double-stranded DNA, single-stranded DNA, single-stranded RNA, or double-stranded RNA.
- Molecules on virus surface impart high specificity for attachment to host cell. (Spike proteins)
- Multiply by taking control of host cell's genetic material and regulating the synthesis and assembly of new viruses.
- Lack enzymes for most metabolic processes. Only Ez for replication.
- Lack machinery for synthesizing proteins

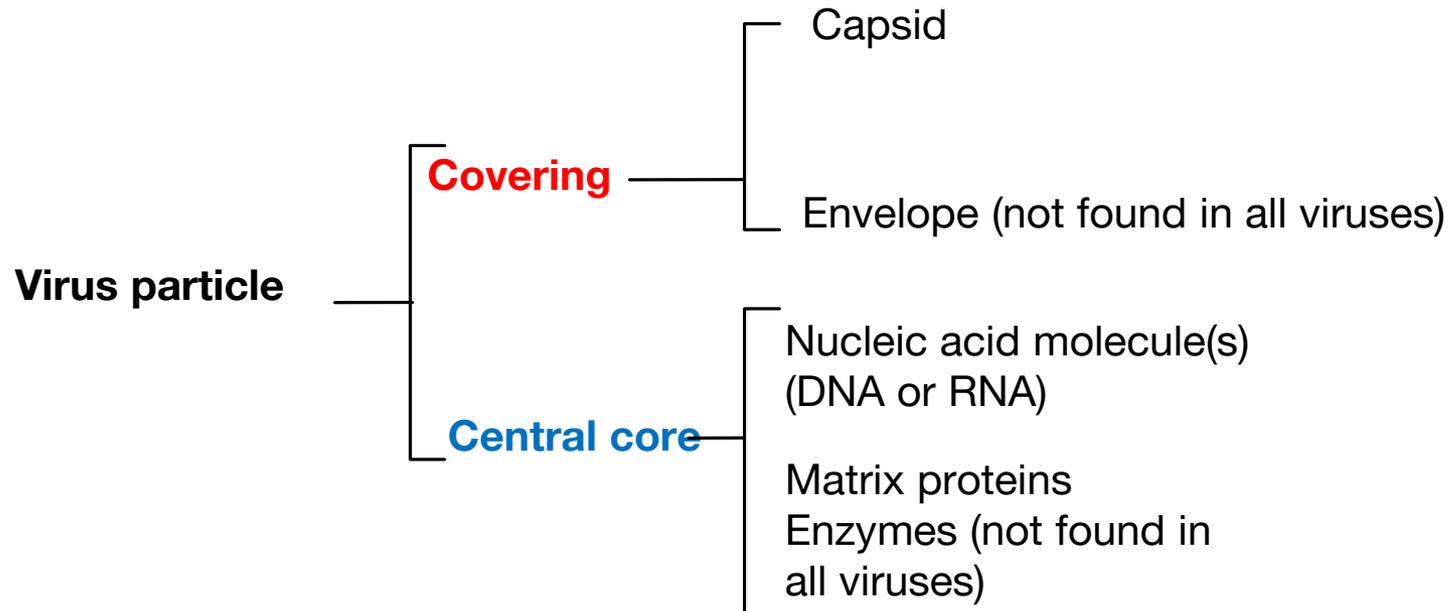
General Size of Viruses

- Size range – most $<0.2 \mu\text{m}$; requires **EM**



Viral Structure

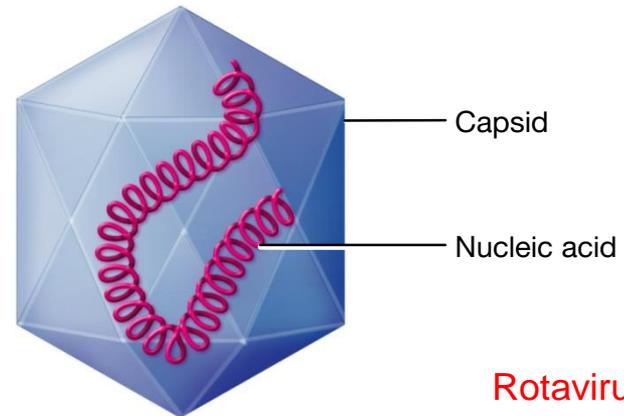
- Viruses bear no resemblance to cells
 - Lack protein-synthesizing machinery
- Viruses contain only the parts needed to invade and control a host cell



General Structure of Viruses

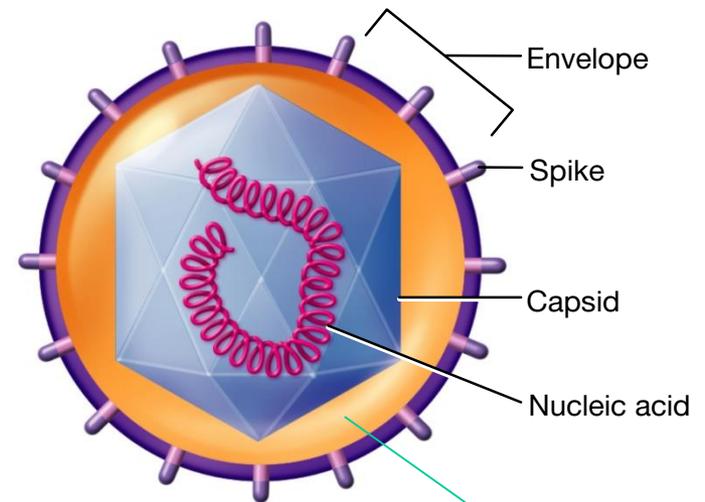
- **Capsids**

- All viruses have capsids (protein coats that enclose and protect their nucleic acid)
- The capsid together with the nucleic acid is the **nucleocapsid**
- **Some** viruses have an external covering called an **envelope**; those lacking an envelope are **naked**
- Each capsid is made of identical protein subunits called **capsomers**

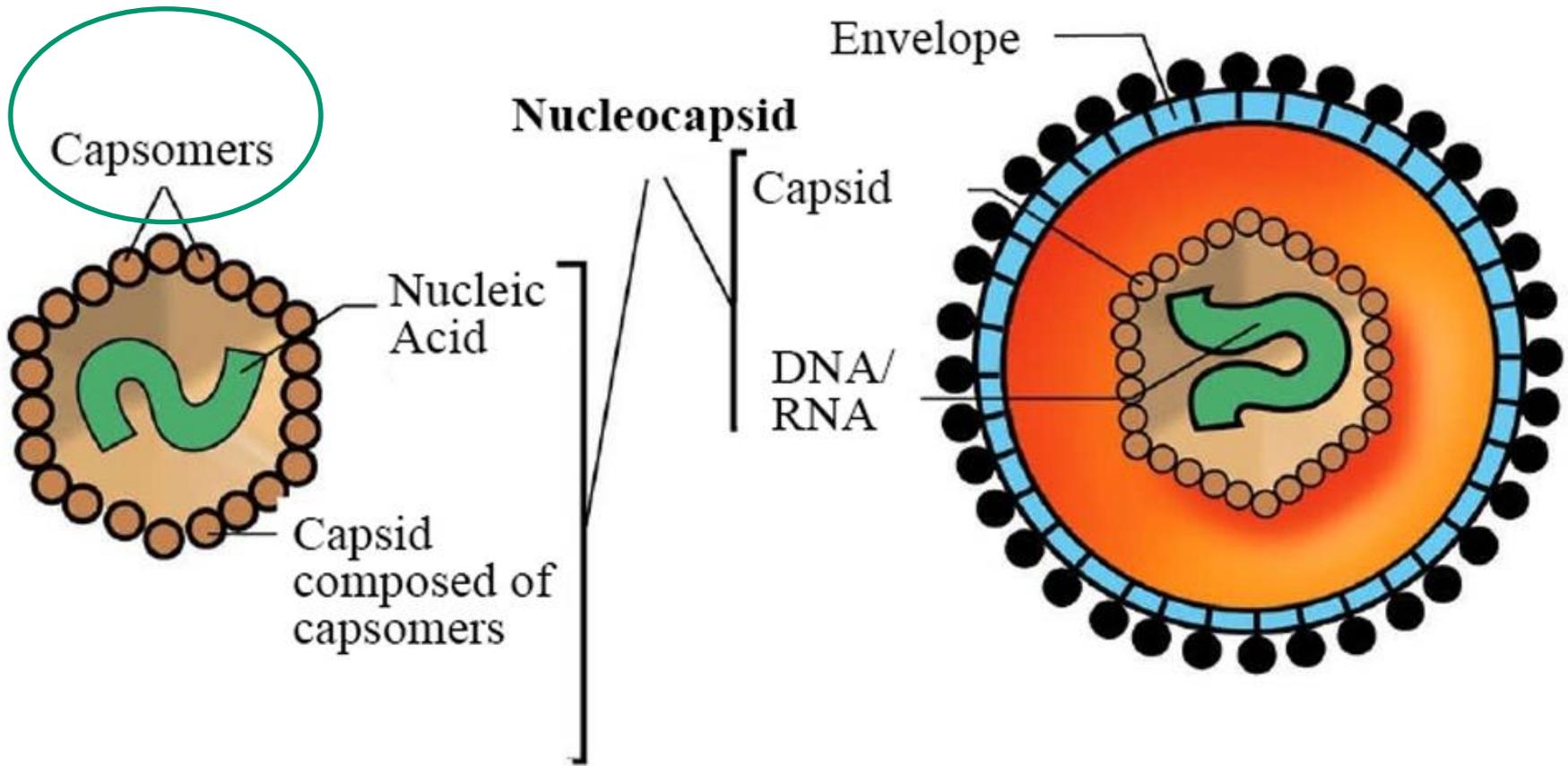


Rotavirus

(a) Naked Nucleocapsid Virus



(b) Enveloped Virus



NAKED VIRUSES

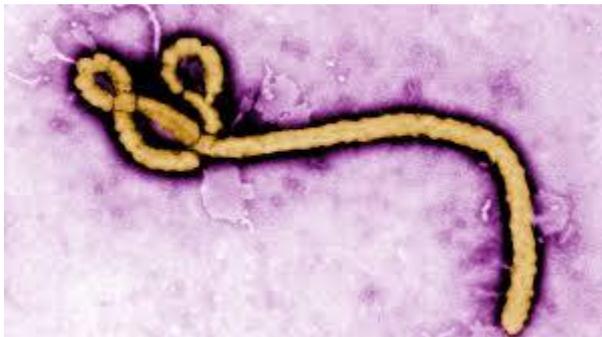
ENVELOPED VIRUSES

General Structure of Viruses

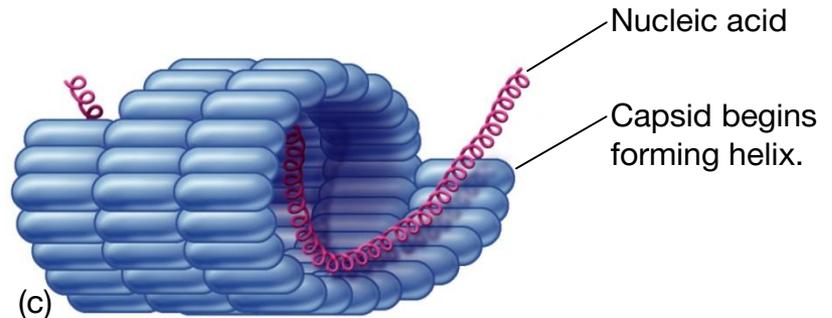
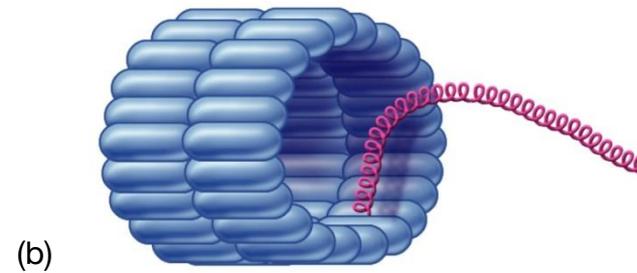
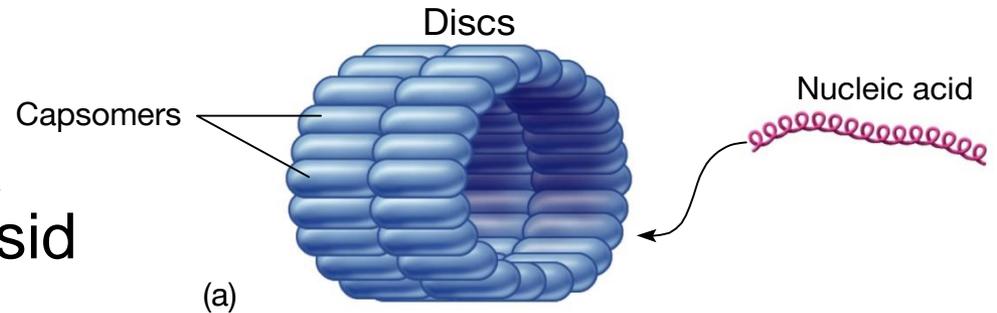
- Two structural capsid types:

A. Helical -

continuous helix of capsomers forming a cylindrical nucleocapsid



Ebolavirus



B. Icosahedral

General Structure of Viruses

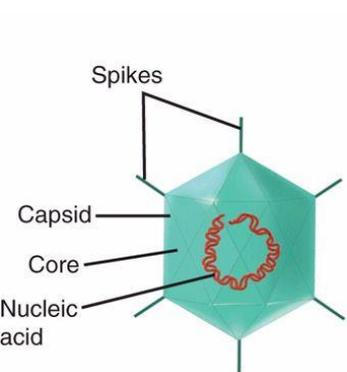
- Two structural capsid types:

A. Helical: All RNA viruses.

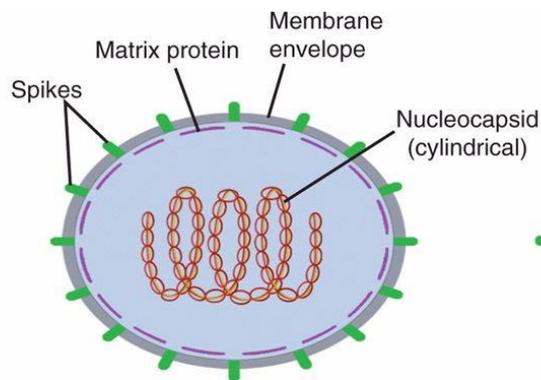
Except (*Picornaviridae*, *Flaviviridae*, *Togaviridae*, *Reoviridae*)

B. Icosahedral - 20-sided with 12 corners.

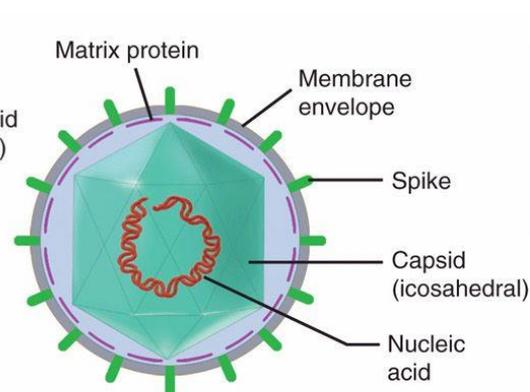
- During assembly of virus, the nucleic acid is packed into the center Icosahedral, forming **nucleocapsid**.
- All DNA viruses are **Icosahedral only** Poxviruses have complex structure.



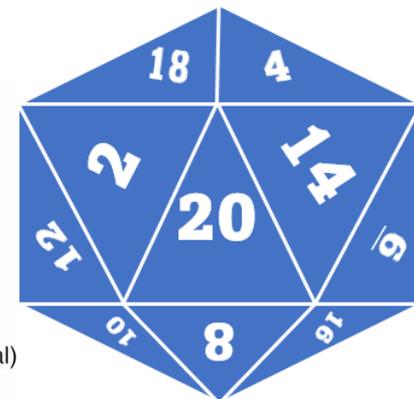
Naked capsid virus (icosahedral capsid)



Enveloped virus (helical nucleocapsid)



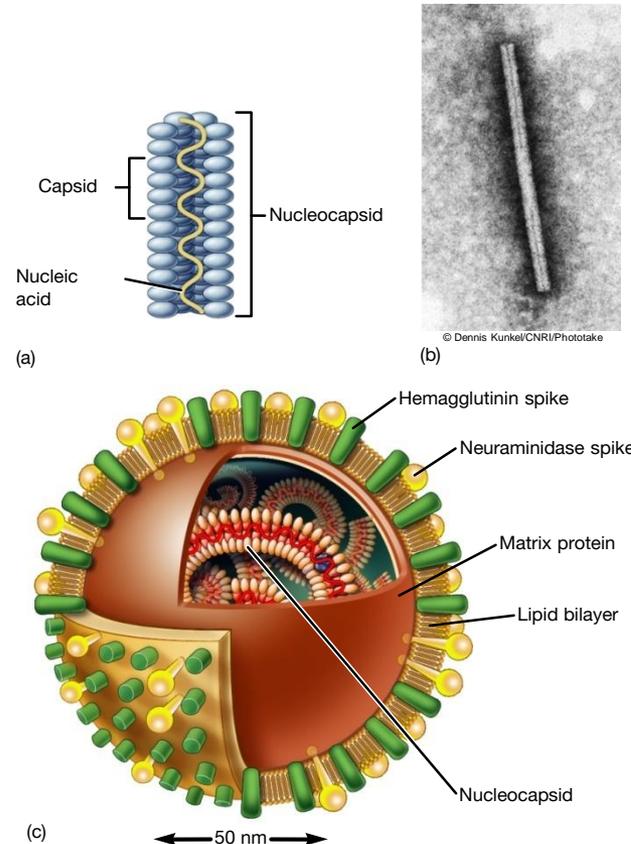
Enveloped virus (icosahedral capsid)



General Structure of Viruses

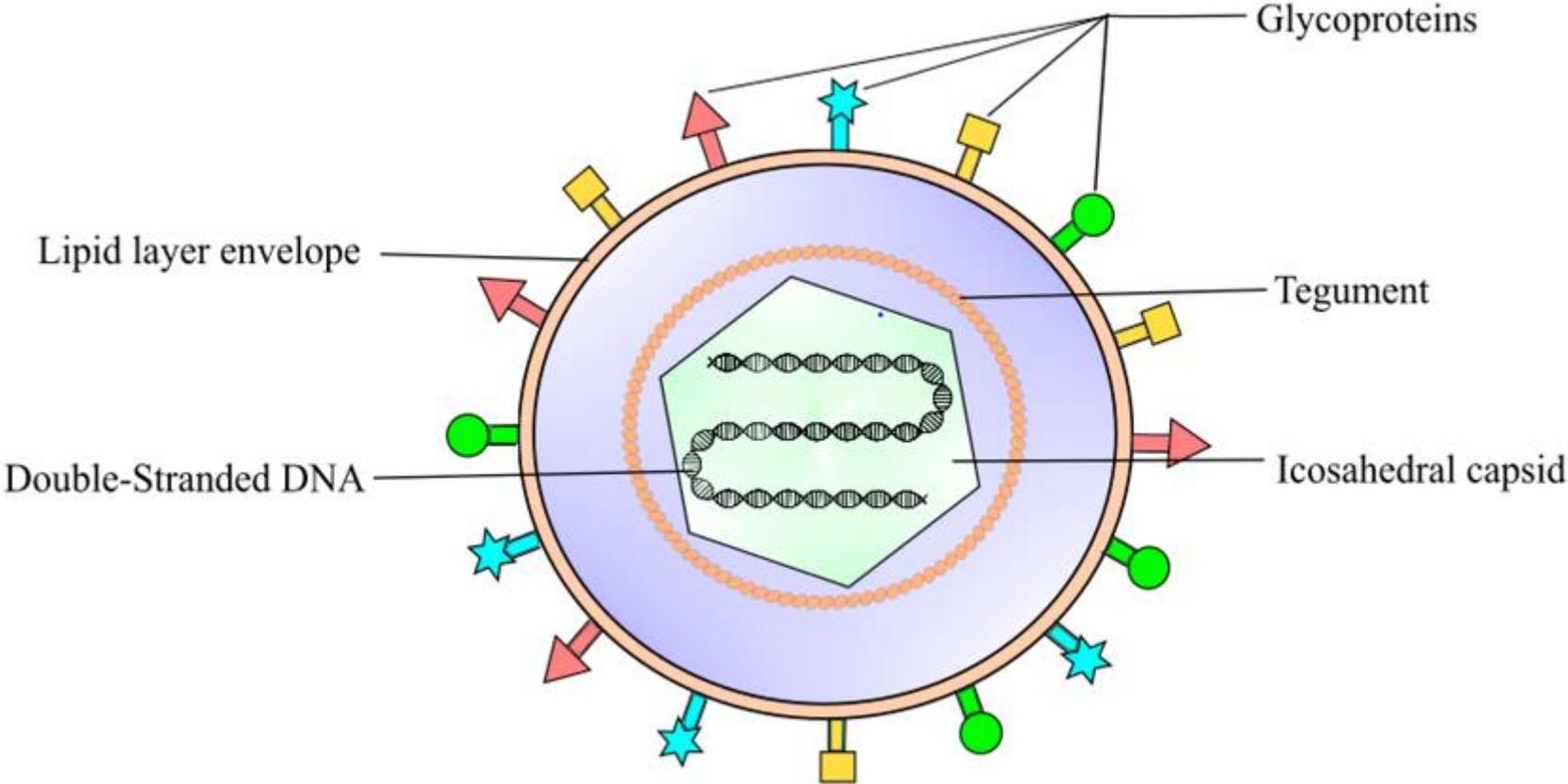
- **Viral envelope**

- Mostly animal viruses
- Acquired when the virus leaves the host cell
- Its lipid or lipoproteins.
- Exposed proteins on the outside of the envelope, called **spikes**, are essential for attachment of the virus to the host cell
- Ether sensitive
- E.g; **Herpes simplex**



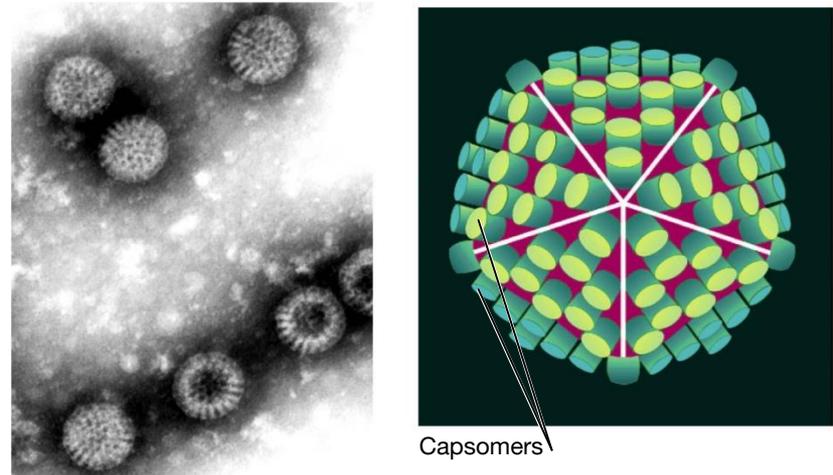
Flu virus

Herpes simplex virus

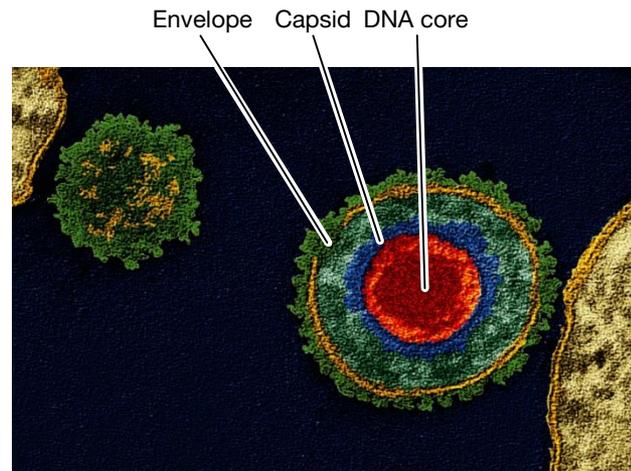


Functions of Capsid/Envelope

- Protects the nucleic acid when the virus is outside of the host cell
- Helps the virus bind to a cell surface and assists the penetration of the viral DNA or RNA into a suitable host cell.
- Immunogenic.



(a) Fred P. Williams, Jr./EPA

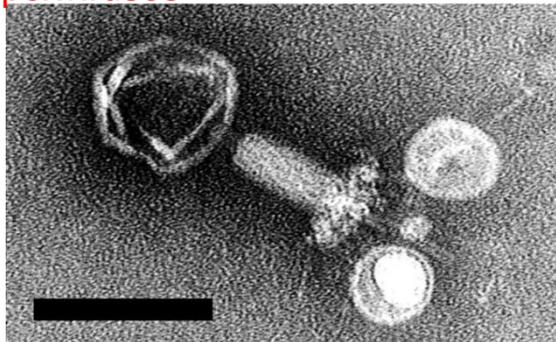
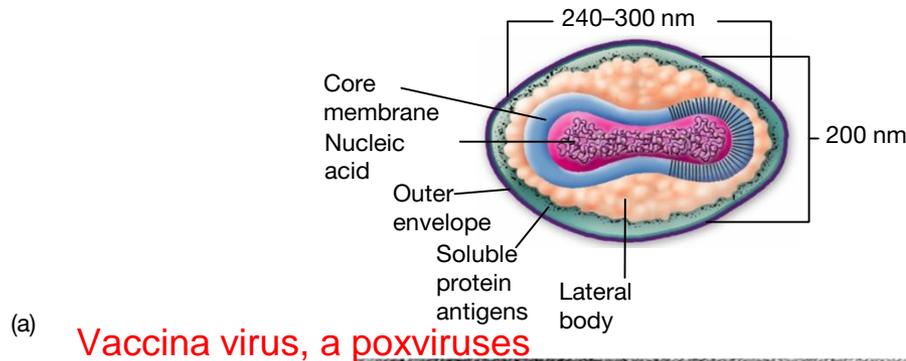


(b)

General Structure of Viruses

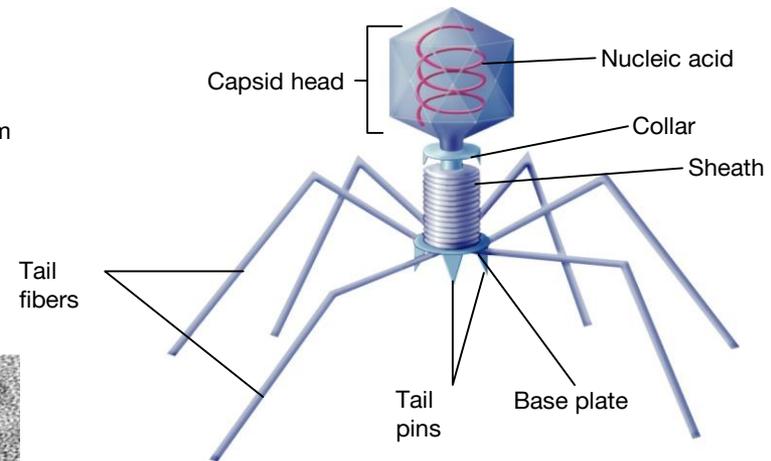
C. Complex viruses: atypical viruses

- Poxviruses lack a typical capsid and are covered by a dense layer of lipoproteins
- Some **bacteriophages** have a polyhedral nucleocapsid along with a helical tail and attachment fibers



(b)

© Bin Ni, Chisholm Lab, MIT



(c) **T4 bacteriophage**

Types of Viruses

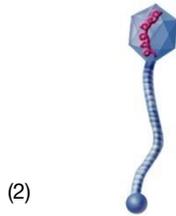
A. Complex Viruses

A. Complex viruses:

- (1) poxvirus, a large DNA virus
- (2) flexible-tailed bacteriophage



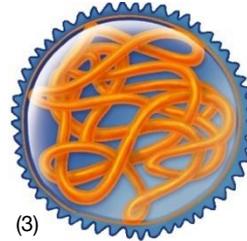
(1)



(2)

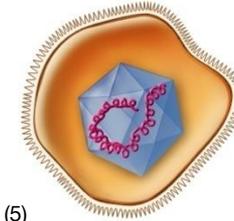
B. Enveloped Viruses

Helical



(3)

Icosahedral



(5)



(4)



(6)

C. Nonenveloped Naked Viruses

Helical



(7)

Icosahedral



(8)



(9)

B. Enveloped viruses:

With a helical nucleocapsid:

- (3) mumps virus
- (4) rhabdovirus (Rabies)

With an icosahedral nucleocapsid:

- (5) herpesvirus
- (6) HIV (AIDS)

C. Naked viruses:

Helical capsid:

- (7) plum poxvirus

Icosahedral capsid:

- (8) poliovirus
- (9) papillomavirus

Nucleic Acids

- **Viral genome** – either **DNA** or **RNA** but never both.
- Carries genes necessary to invade host cell and redirect cell's activity to make new viruses.
- Number of genes varies for each type of virus – few to hundreds.

Nucleic Acids

- DNA viruses

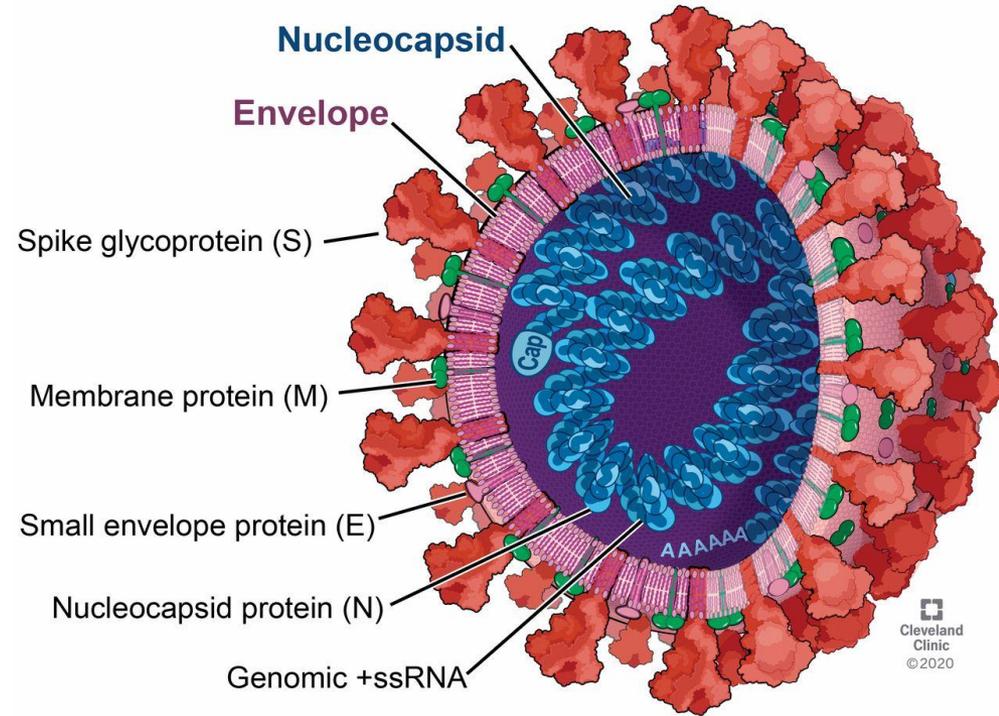
- Usually double stranded (**ds**) but may be single stranded (**ss**)
- Circular or linear

- RNA viruses

- Usually (**ss**), may be (**ds**), may be **segmented** into separate RNA pieces.
- ssRNA genomes ready for immediate translation are **positive-sense RNA**
- ssRNA genomes that must be converted into proper form (positive-sense RNA) are **negative-sense RNA**; that required a viral RNA-dependent RNA polymerase. e.g: Influenza virus, Ebola virus, Rabies virus, Measles virus.

CORONAVIRUSES

- Containing a single strand of positive-sense RNA.
- **Helical** nucleocapsid structure.



General Structure

- Pre-formed enzymes may be present
 - **Polymerases** – DNA or RNA
 - **Replicases** – copy RNA
 - **Reverse transcriptase** – synthesis of DNA from RNA (AIDS virus)

How Viruses Are Classified

- Main criteria presently used are structure, chemical composition, and genetic makeup
- Currently recognized: 59 orders, 189 families, and 2224 genera of viruses
 - Family name ends in -**viridae**, i.e. *Herpesviridae*
 - Genus name ends in -**virus**, *Simplexvirus*
 - Species Herpes simplex virus I (**HSV-1**)

Classification & Taxonomy of viruses

1. **According to nucleic acid type**
2. **According to host**
3. **According to symmetry**
4. **According to presence of envelope**
5. **According to clinical or affected organ systems** (e.g., respiratory, enteric, hepatic, or arboviruses).
6. **Baltimore Classification** (based on how viruses produce mRNA).

Human Viruses & Viral Diseases

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TABLE 6.2 Important Human Virus Families, Genera, Common Names, and Types of Diseases

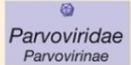
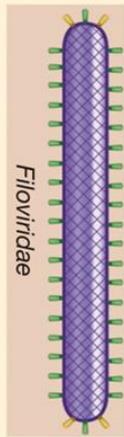
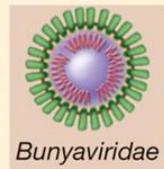
Nucleic Acid Type	Family	Genus of Virus	Common Name of Genus Members	Name of Disease
DNA Viruses				
 Poxviridae Chordopoxvirinae	Poxviridae	<i>Orthopoxvirus</i>	Variola and vaccinia	Smallpox, cowpox
	 Herpesviridae	Herpesviridae	<i>Simplexvirus</i>	Herpes simplex 1 virus (HSV-1)
<i>Varicellovirus</i>			Herpes simplex 2 virus (HSV-2)	Genital herpes
<i>Cytomegalovirus</i>			Varicella zoster virus (VZV)	Chicken pox, shingles
<i>Mastadenovirus</i>			Human cytomegalovirus (CMV)	CMV infections
 Adenoviridae	Adenoviridae	<i>Mastadenovirus</i>	Human adenoviruses	Adenovirus infection
	Papillomaviridae	<i>Papillomavirus</i>	Human papillomavirus (HPV)	Several types of warts
 Papillomaviridae	Polyomaviridae	<i>Polyomavirus</i>	JC virus (JCV)	Progressive multifocal leukoencephalopathy
	 Hepadnaviridae	Hepadnaviridae	<i>Hepadnavirus</i>	Hepatitis B virus (HBV or Dane particle)
 Polyomaviridae		Parvoviridae	<i>Erythrovirus</i>	Parvovirus B19
	 Parvoviridae Parvovirinae			
RNA Viruses				
 Picornaviridae	Picornaviridae	<i>Enterovirus</i>	Poliovirus	Poliomyelitis
			Coxsackievirus	Hand-foot-mouth disease
 Caliciviridae	Caliciviridae	<i>Rhinovirus</i>	Hepatitis A virus (HAV)	Short-term hepatitis
			<i>Calicivirus</i>	Human rhinovirus
 Togaviridae	Togaviridae	<i>Alphavirus</i>	Norwalk virus	Viral diarrhea, Norwalk virus syndrome
			Eastern equine encephalitis virus	Eastern equine encephalitis (EEE)
 Flaviviridae	Flaviviridae		Western equine encephalitis virus	Western equine encephalitis (WEE)
			Yellow fever virus	Yellow fever
			St. Louis encephalitis virus	St. Louis encephalitis

TABLE 6.2 (Continued)



100 nm

Flaviviridae	<i>Rubivirus</i> <i>Flavivirus</i>	Rubella virus Dengue fever virus West Nile fever virus	Rubella (German measles) Dengue fever West Nile fever
Bunyaviridae	<i>Bunyavirus</i> <i>Hantavirus</i> <i>Phlebovirus</i> <i>Nairovirus</i>	Bunyamwera viruses Sin Nombre virus Rift Valley fever virus Crimean–Congo hemorrhagic fever (CCHF) virus	California encephalitis Respiratory syndrome Rift Valley fever Crimean–Congo hemorrhagic fever
Filoviridae	<i>Filovirus</i>	Ebola, Marburg virus	Ebola fever
Reoviridae	<i>Coltivirus</i> <i>Rotavirus</i>	Colorado tick fever virus Human rotavirus	Colorado tick fever Rotavirus gastroenteritis
Orthomyxoviridae	<i>Influenza virus</i>	Influenza virus, type A (Asian, Hong Kong, and swine influenza viruses)	Influenza or “flu”
Paramyxoviridae	<i>Paramyxovirus</i> <i>Morbillivirus</i> <i>Pneumovirus</i>	Parainfluenza virus Mumps virus Measles virus Respiratory syncytial virus (RSV)	Parainfluenza Mumps Measles (red) Common cold syndrome
Rhabdoviridae	<i>Lyssavirus</i>	Rabies virus	Rabies (hydrophobia)
Retroviridae	<i>Oncornavirus</i> <i>Lentivirus</i>	Human T cell leukemia virus (HTLV) HIV (human immunodeficiency viruses 1 and 2)	T-cell leukemia Acquired immunodeficiency syndrome (AIDS)
Arenaviridae	<i>Arenavirus</i>	Lassa virus	Lassa fever
Coronaviridae	<i>Coronavirus</i>	Infectious bronchitis virus (IBV) Enteric corona virus SARS virus	Bronchitis Coronavirus enteritis Severe acute respiratory syndrome

Effect physical and chemical agents on viruses

1. Cold:

- Storage at -40 to -70c, or by lyophilization at 4c, via freezing then vacuum to remove water and O₂ to be powder.
- Stable at cold temp.

2. **Heat** at 50-60c/ 30min will be destroyed. Only **HBV**.

3. **Fluctuating temp**; will destroy viruses.

4. **Radiation**

5. **pH** 5-9 stable.

- *Enteroviruses* are resistant to acidic.: **Polioviruses**

6. **Salts** (Mgcl₂); stabilization used in Polio vaccine

Effect physical and chemical agents on viruses

7. **Phenols** compounds: Viruses resist

8. **Glycerol 50%** (destroys bacteria) ; and virus resist, so it used as preservative for viruses.

9. **Antibiotics**: Resistant

10. **Ether** and detergents : destroy viruses envelopes.

11. **Oxidizing agents** (H₂O₂, Chlorine, Iodine): destroyed virus envelope.

12. **Formaldehyde**: Destroy Nucleic acid without affecting viral antigenicity (Capsid). Used in vaccination