- o Fenestrated Have holes in endothelium but have continuous BM
  - Kidney and gut mucosa
- Sinusoids Incomplete BM and Large intracellular gaps
- Have a glycocalyx that is -vely charged has a barrier function
- **Endothelial BM** 
  - o Polarised cells with distinct expression of receptors on apical and luminal sides
  - o Endothelial luminal membrane resides on a BM and is associated with extracellular matrix collagen (4,3,1), fibronectin, laminin
- Endothelial intracellular junctions tighter on arterial side looser on post capillary venules
  - o 3 barriers to paracellular transport (cell to cell connection)
    - **Tight junctions** 
      - Claudins (Cldns) the more the tighter the junctions
      - Occludins (ocln)
      - Junctional adhesion molecules
    - Adherens junctions
    - Gap junctions connexions
  - Junctional proteins and integrins (cell to BM) are connected to the actin cytoskeleton of the EC allows some communication
- Continuous endothelium
  - Paracellular transport Water & small solutes (<3mm) pass between ECs</li>
  - Transcytosis Allows passage of larger solutes e.g. albumin (uses transendothelial channel)
  - Caveolae Smooth membrane invaginations and vesicles = highest density in capillary EC (vesicles move through the membrane and dump on the other side) .co.uk
  - Transendothelial channels
- Fenestrated continuous endothelium
  - o Glomerular endothelium, vessels within endocrine & exocrired
  - o Fenestrae/pores have a diaphragm may increate selectivity of the pore no albumin or peptide hormones (angiotensin)
  - Fenestrations permit greater it insoloothelial transport
- Sinusoidal endothelium
  - Liv in users and bone marrov
  - o Small ECs clear colloids and soluble waste macromolecules from the circulation
  - Large fenestrations
  - o BM has gaps
  - o High endocytic activity in clathrin-coated pits (receptor mediated and fluid phase endocytosis
  - For mass transport allow bulk flow
- Vesicular-Vacuolar organelles (VVO)
  - o A major route for transport of fluids and solutes across the endothelium particularly in inflammatory situations
  - From transcellular channels when they connect
  - o Particularly at post capillary venules
  - Vacuoles join together and allow large bulk transport (leukocytes, large molecules)

## **Endothelium and haemostasis**

- Provides a non-thrombogenic surface to maintain blood flow
  - Inhibits the activation of coagulation factors
  - o Breaks down clots that start to form
  - Inhibits platelet adhesion/activation
- Anti-coagulant properties of endothelium
  - o Tissue factor pathway inhibitor binds to FXa, FVIIIa, TFa to block activation of the extrinsic pathway
  - Anti-thrombin III localised with hepara sulphate proteoglycans (glycocalyx) binds and inactivates thrombin
  - Thrombomodulin converts thrombin into a Protein C activator