

## **AWP 2-4 – Polymer Chemistry**

### **Polymers in Life Science – E) Regenerative medicine & implants**

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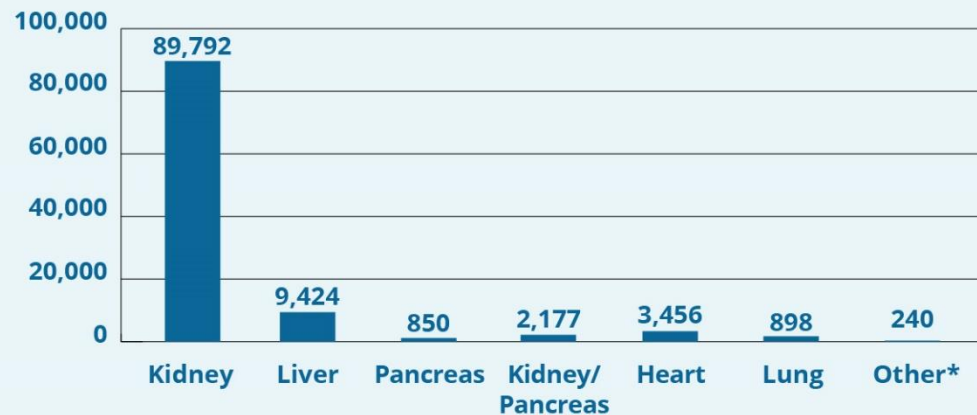
Summer term 2025

# Background

- Long waiting lists on transplants
- Also long waiting times and unclear time line!
- Below: some numbers from the US

**Patients on the Waiting List by Organ**

As of September 2024

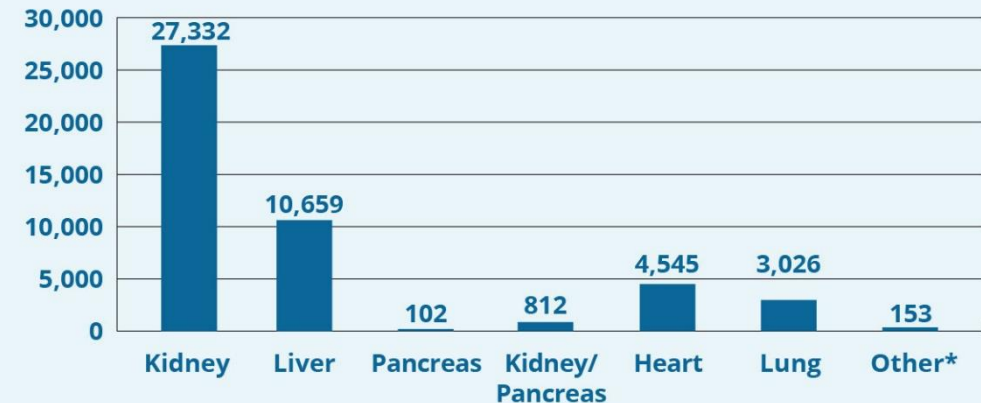


\*Other includes allograft transplants like face, hands, and abdominal wall.

Based on OPTN data as of September 15, 2024. Data subject to change based on future data submission or correction. Totals may be less than the sums due to patients included in multiple categories.

**Transplants Performed by Organ**

In 2023



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# Transplantation

- Successful transplantation of bone, soft tissue, and corneas in the 20th century
- 1954 first successful kidney transplant
- 1960s successful transplantation of pancreas/kidney, liver, isolated pancreas & heart
- Transplant surgery success continued into the 1980s with successful heart-lung, single lung, double lung, living-donor liver, and living-donor lung transplants
- Problems:
  - Eventual rejection of foreign organs by immune system
  - Immune suppressive medication necessary
  - Scarcity of available transplants

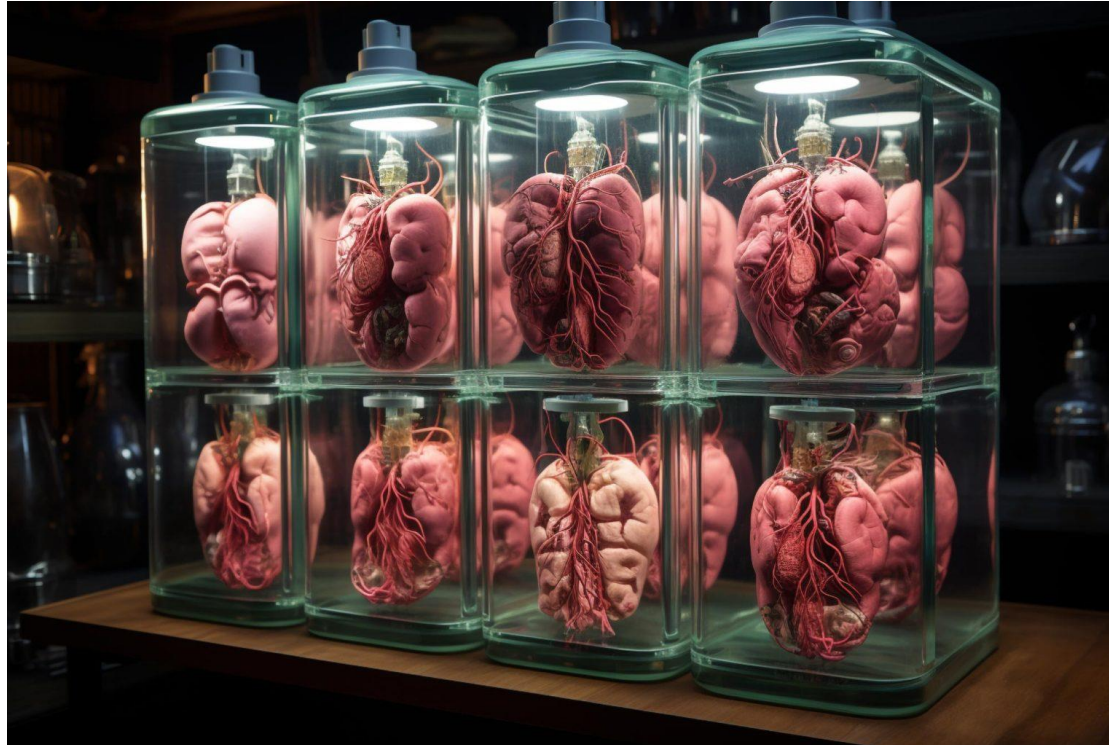
# Kidneys from genetically modified pigs

- Progress in xenotransplantation: pig organs are seen as a promising solution to the organ shortage.
- Genetic modification: Pigs are being genetically modified to minimise human rejection.
- Early success: Recently, pig hearts and kidneys have been successfully transplanted into humans.
- Challenges: Immune reactions, viral risks (e.g. porcine endogenous retroviruses) and ethical concerns remain.
- Future prospects: Research is focusing on CRISPR technology and immunosuppressants to improve long-term acceptance.



## 3D printed organs?

- Can we just grow new organs in a dish??



- Not quite, but research is ongoing

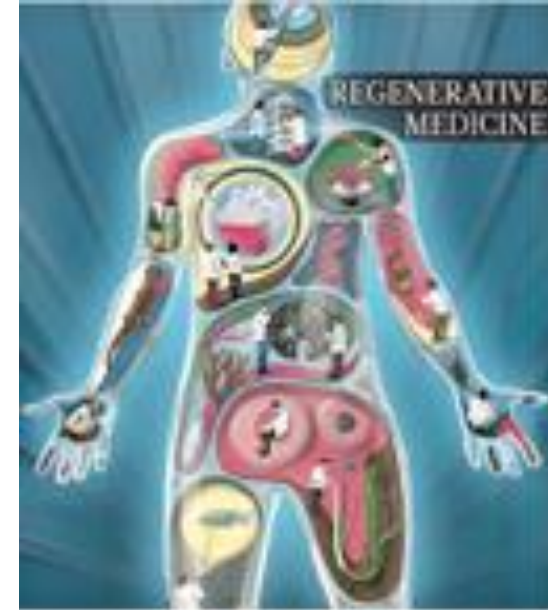


# Motivation

Regenerative medicine is concerned with the restoration, repair or replacement of damaged or diseased cells, tissues or organs.

## Key fields and technologies

- Stem cell therapy: Using embryonic or adult stem cells to regenerate tissue and organs.
- Tissue engineering: Mixing cells, biomaterials and bioactive molecules to create tissue and organs.
- 3D bioprinting: Printing living tissue or organ structures using bioinks from cells and biomaterials.
- Artificial organs: development of biohybrid or completely artificial organs
- Biological drugs and gene therapy: genetic modifications or growth factors
- Nanotechnology: nanomaterials for cell stimulation, drug release and tissue integration.
- Immunomodulation: influencing the immune system to promote healing



## More than science...



1997 Joseph Vacanti and his team includes an image of what appears to be a human ear growing out of a mouse's back. The team hopes that its research will lead to lab-grown ears for people whose ears are damaged. But the image sparks a backlash from animal rights activists and people who find the image shocking.

## Cells

- the smallest structural and functional unit of an organism, which is typically microscopic and consists of cytoplasm and a nucleus enclosed in a membrane

## Tissue

- distinct types of material of which animals or plants are made, consisting of specialized cells and their products

## Organ

- collection of tissues joined in a structural unit to serve a common function.

## System

- Functional groups of organs

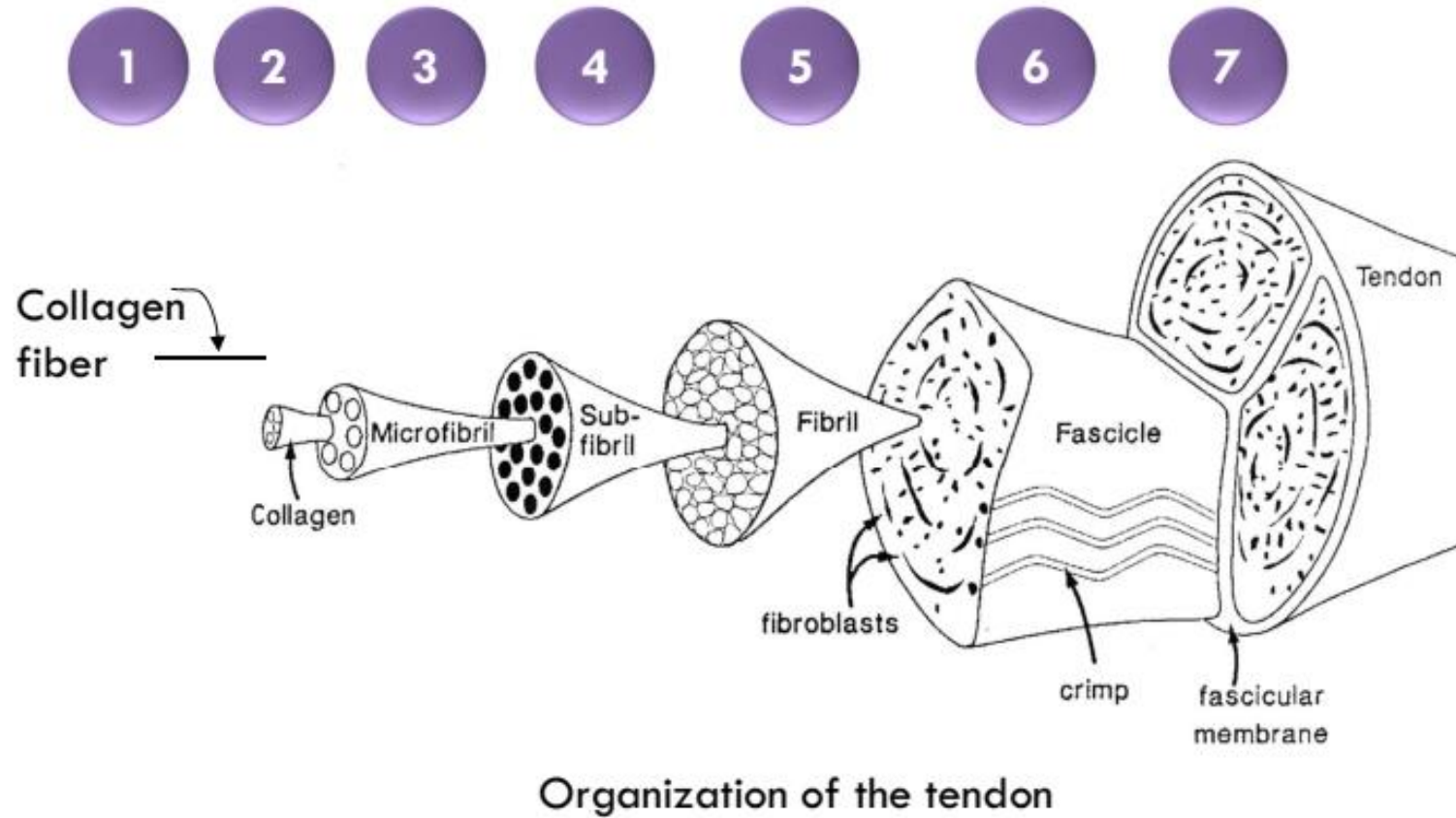
## Organism

- The basic living system, a functional grouping of the lower-level components, including at least one cell



# Complexity

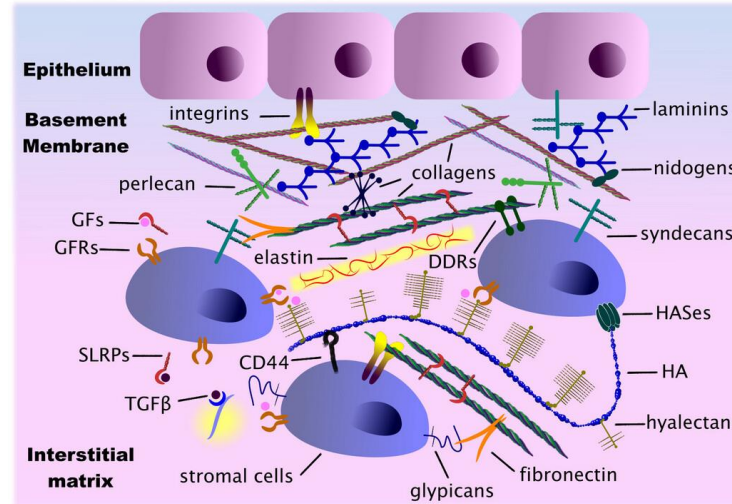
- 7-10 levels of structural organization can be defined in a tissue or organ



# Extracellular matrix

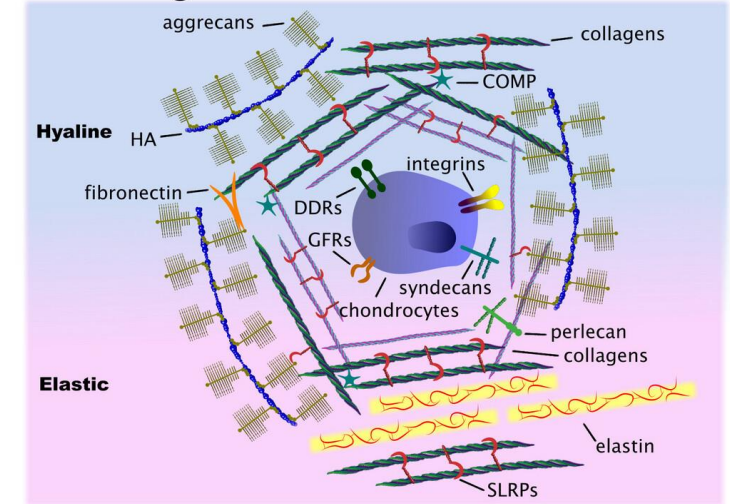
- Network of proteins and poly(saccharide)s
  - Collagen
  - Fibronectin
  - Glycoproteins
  - Proteoglycans
  - Glycoaminoglycans
- Key properties:
  - Mechanical modulus/viscosity
  - Growth factor concentration
  - Binds water & lubricates

**A Connective Tissue**



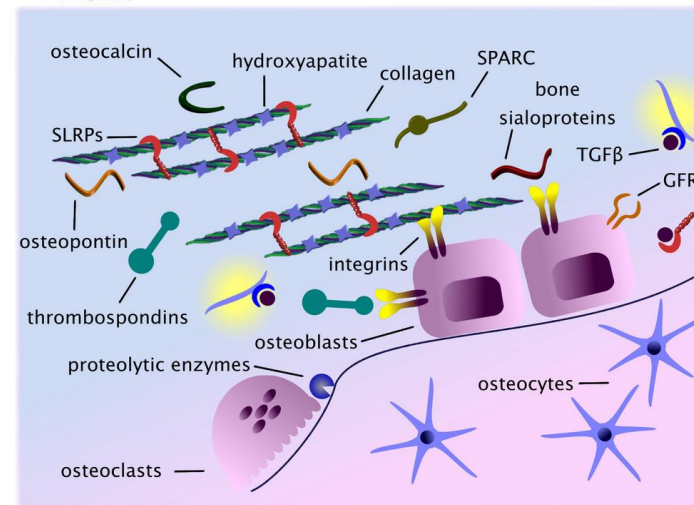
Connection, support, nourishment

**B Cartilage**



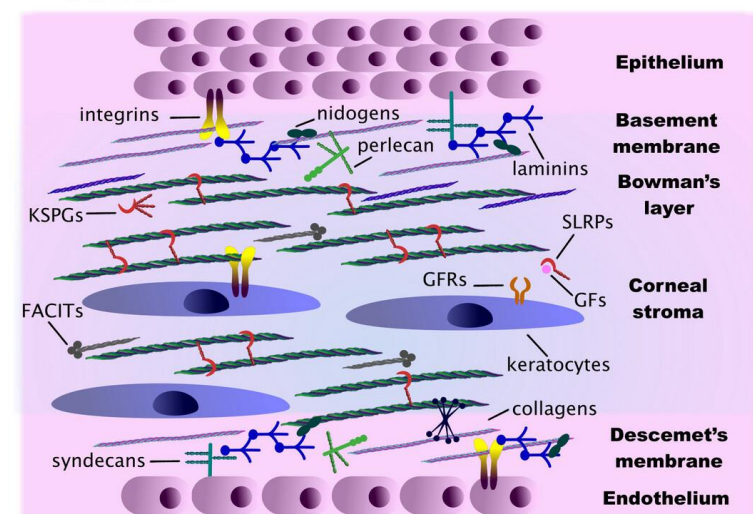
Mechanical support, viscoelasticity, lubrication

**C Bone**



Durable, strong, stiff structure with low elasticity for shock absorption

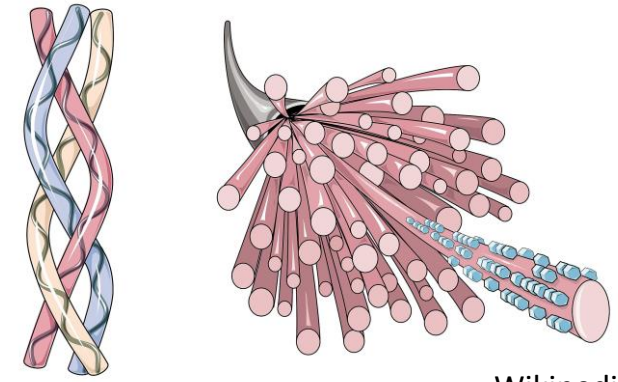
**D Cornea**



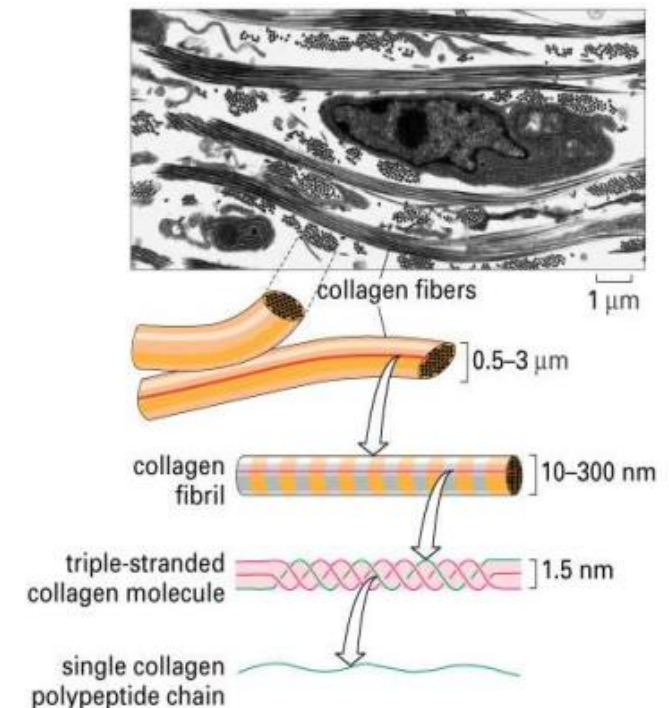
Transparency, refraction

## Collagen

- Protein (25% of Protein mass in human body)
- Very important in bones, tendons, ligaments and skin
- Degree of mineralization determines rigidity
- Procollagen is produced in the cell (e.g. fibroblasts) and shipped outside by exocytosis
- There it is cleaved, forming fibers which are chemically cross-linked



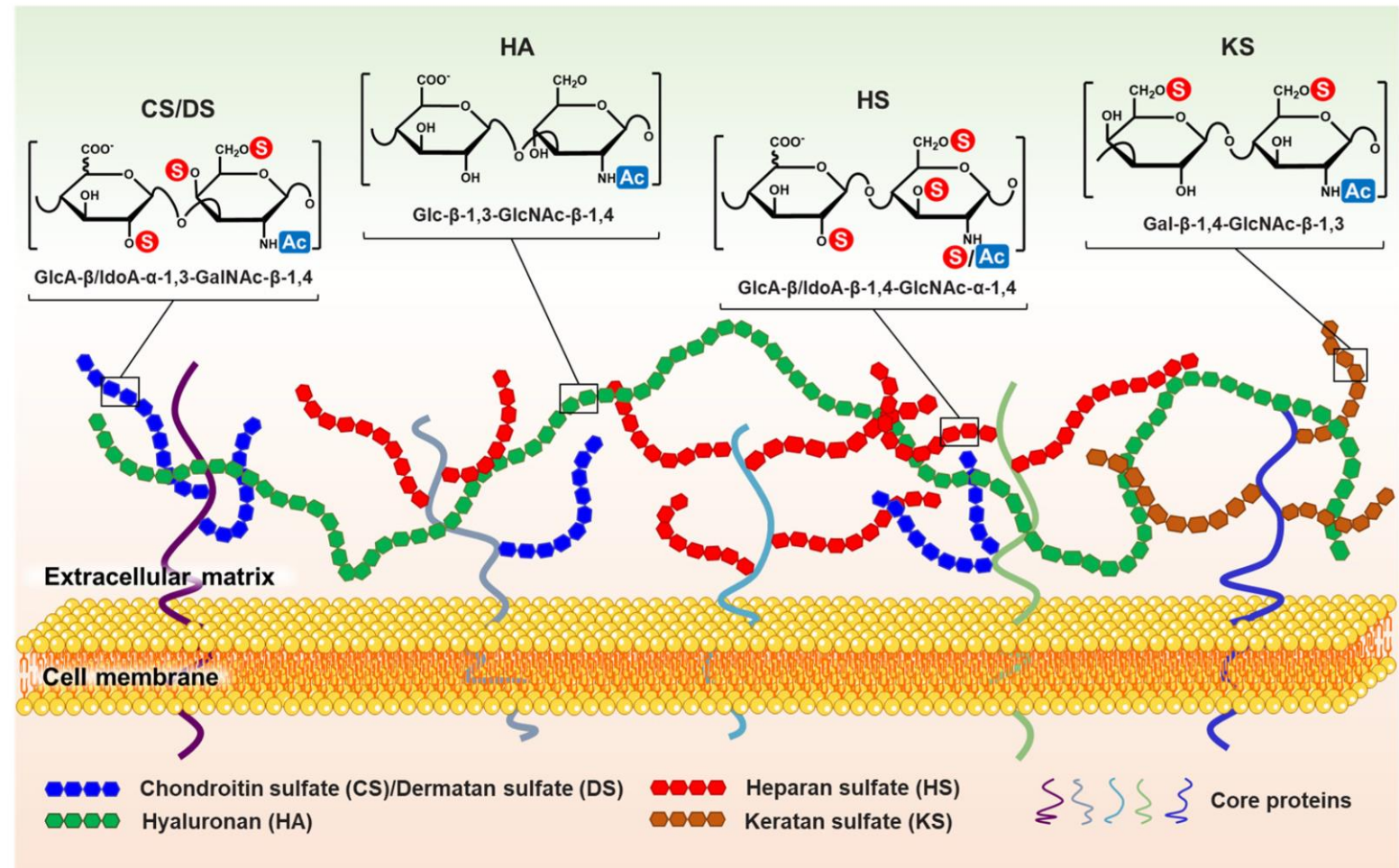
Wikipedia.de





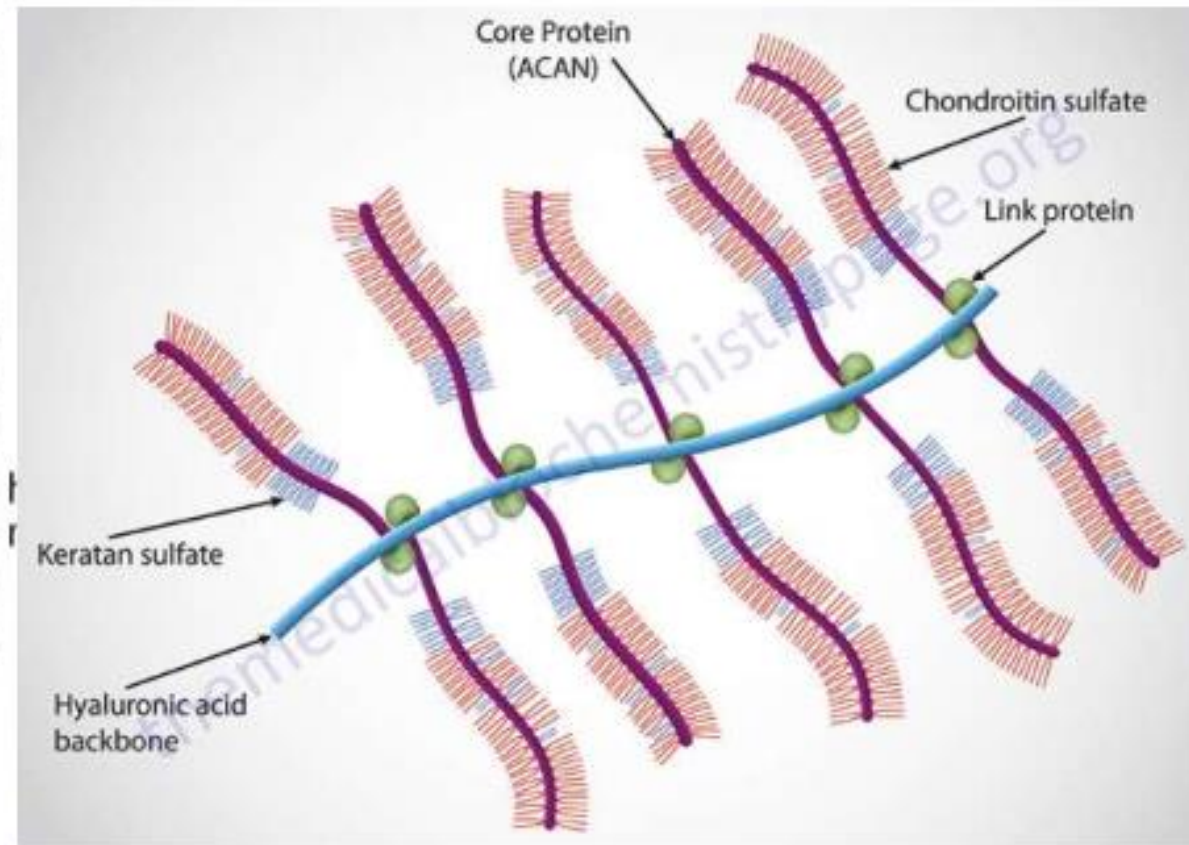
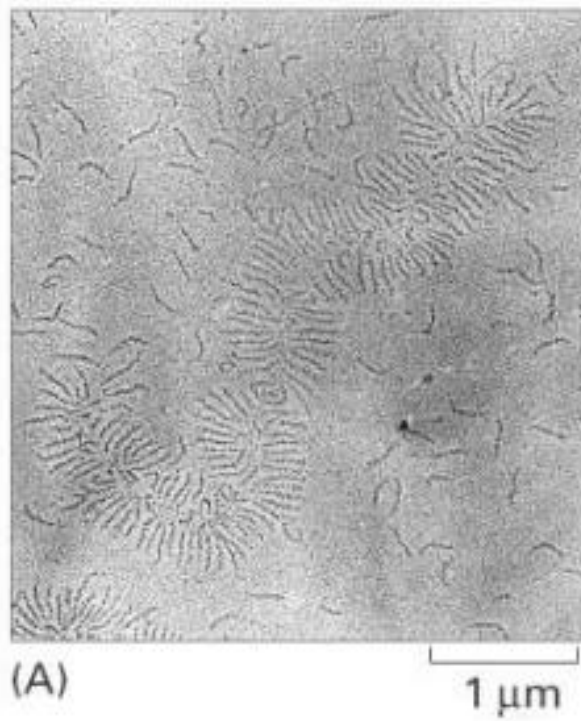
## Glycoaminoglycans (GAG)

- Negatively charged polysaccharides
- Example hyaluronic acid



## Proteoglycans

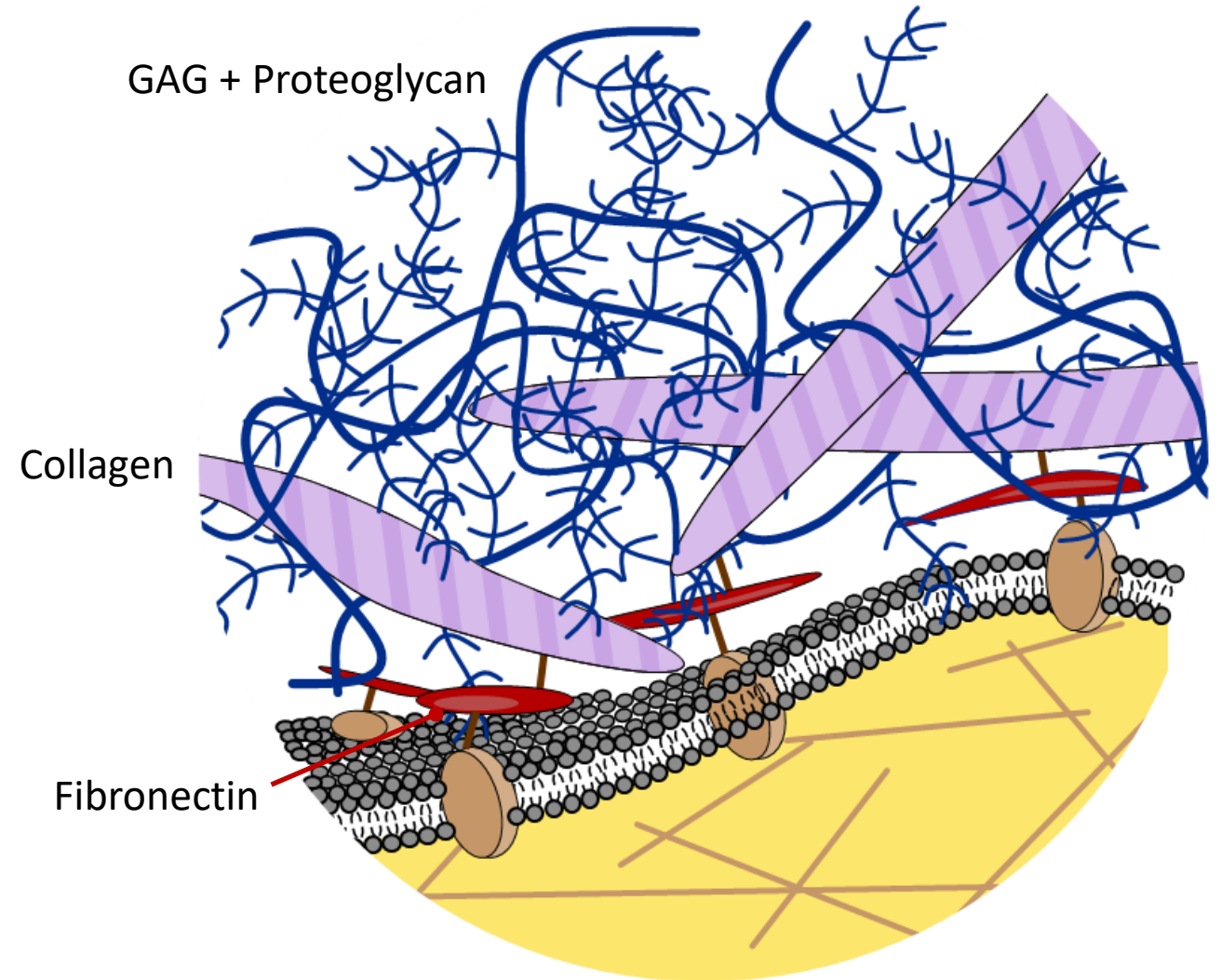
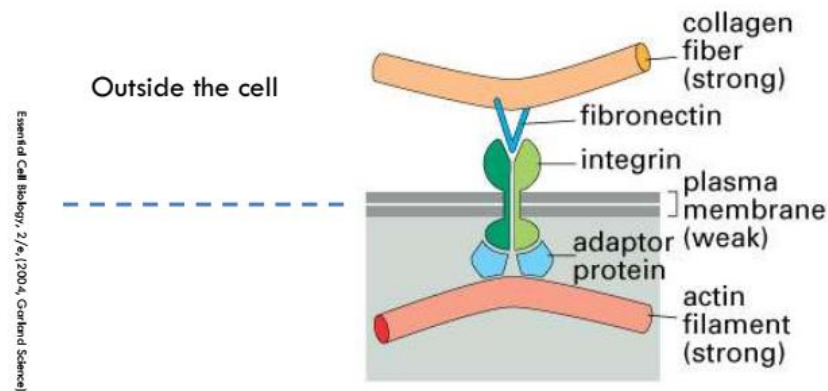
- Proteins linked to the side chain of GAG



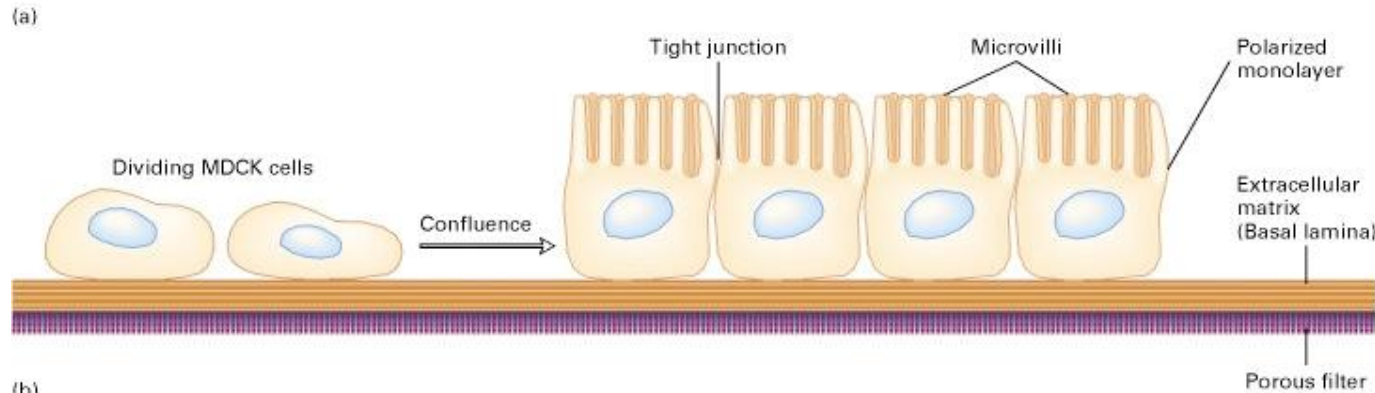


## Fibronectin

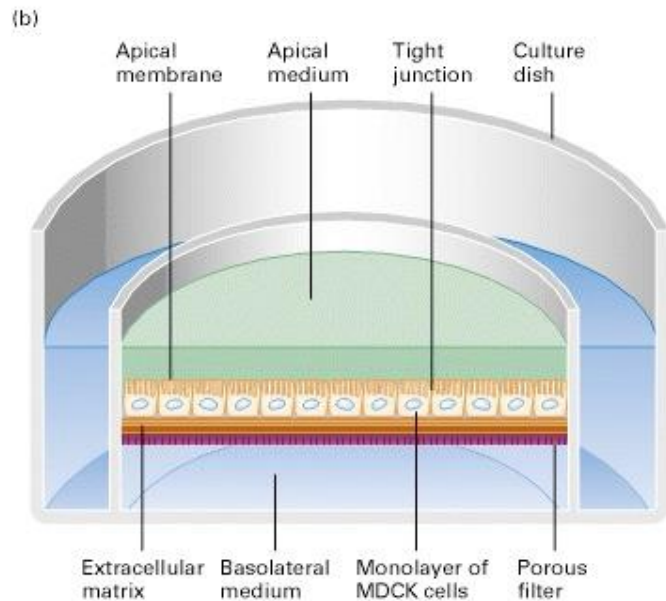
- High molar mass glycoprotein
- Binds to cellular receptors (integrins) and connects them to the ECM
- Key role in cell migration, growth, differentiation and wound healing



# Cells need a „home“



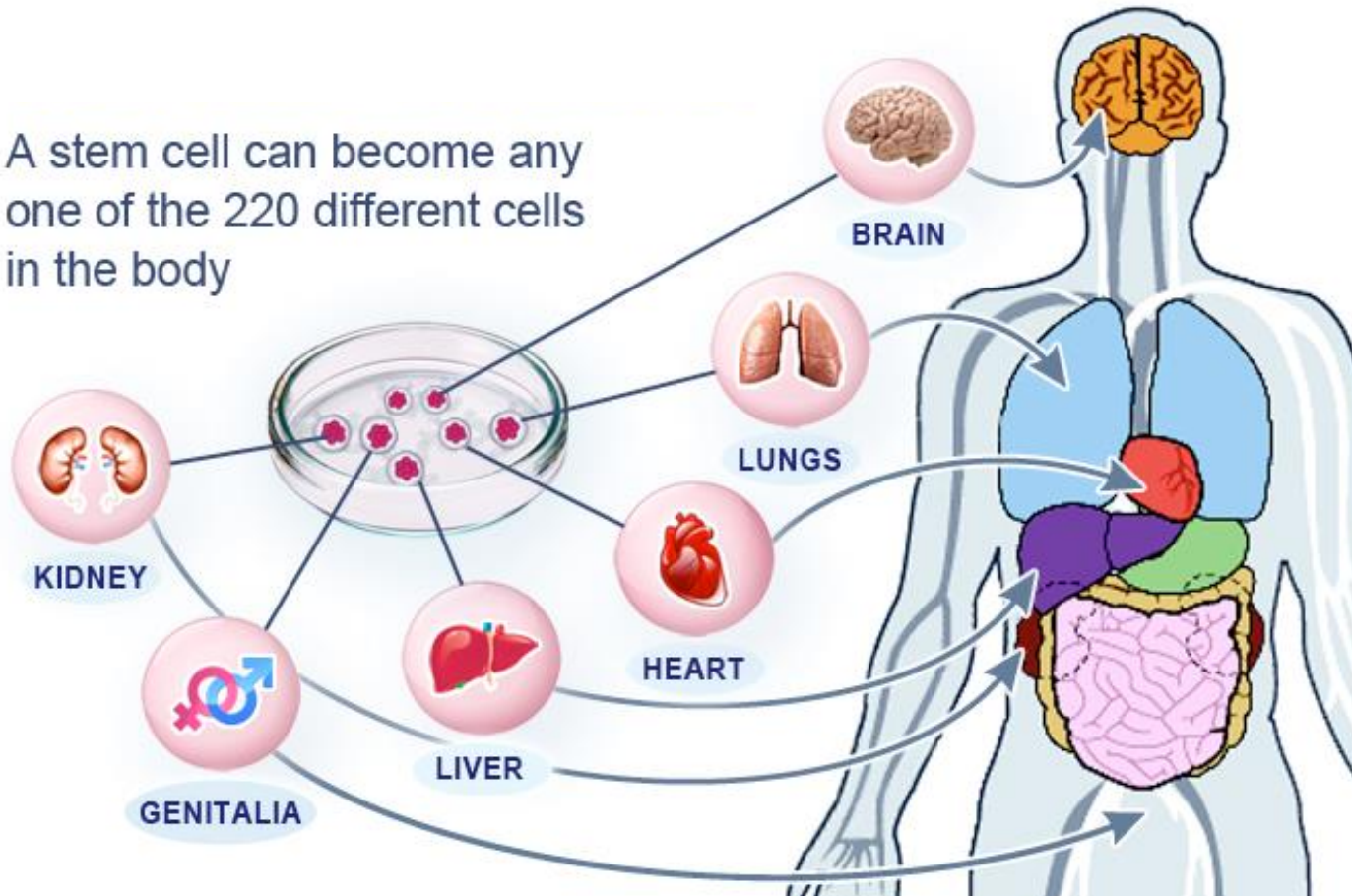
ECM has a key role in the cellular environment!



The tendency of animal cells in vivo to interact with one another and with the surrounding extracellular matrix is mimicked in their growth in culture. Unlike bacterial and yeast cells, which can be grown in suspension, most cultured animal cells require a surface to grow on.

# Stem cells

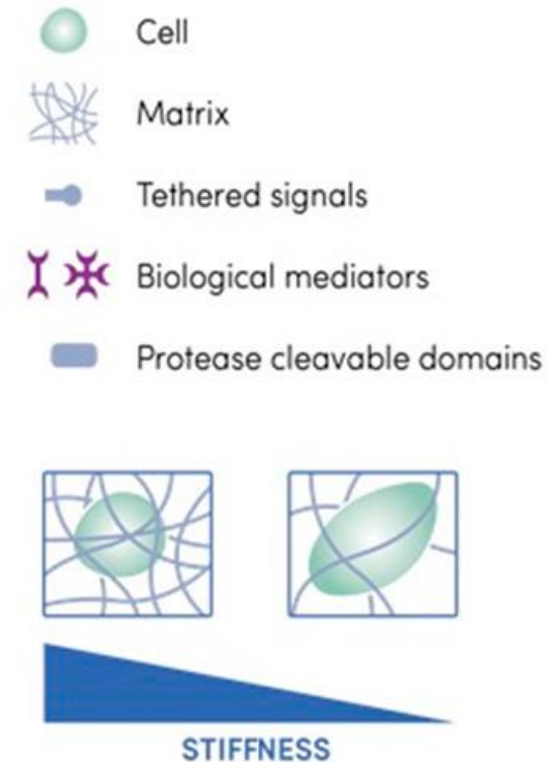
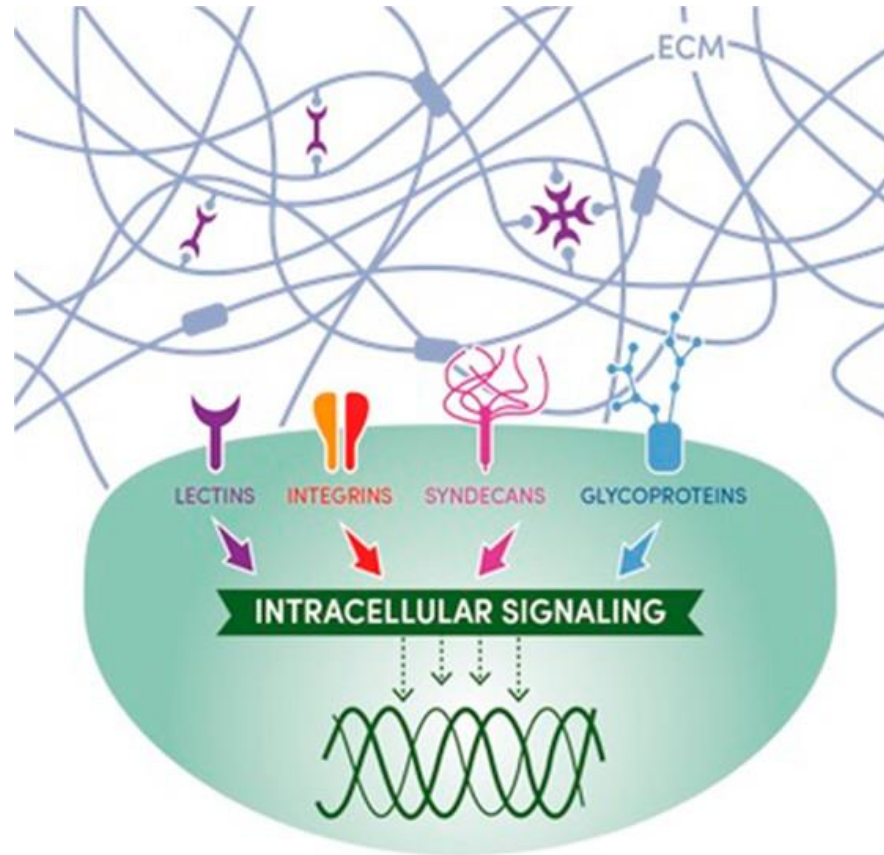
A stem cell can become any one of the 220 different cells in the body



- Stem cells are undifferentiated biological cells that can differentiate into specialized cells.
- Induced pluripotent stem cells (iPS cells) are a type of pluripotent stem cell that can be generated directly from adult cells.
- 4 transcriptional factors (Myc, Oct3/4, Sox2 and Klf4) were sufficient to convert mouse embryonic or adult fibroblasts to pluripotent stem cells

# Cell Differentiation in the ECM

- ECM guides differentiation of stem cells based on
- mechanical properties
- Mechanical stress
- Shape of niche
- Signaling ! (growth factors)





# Natural or synthetic ECM?

