

Biology

/ Lecture 1 /

- Visualization technique
- Cell and tissue definition

Important notes:

Biology LECTURES – will be each week at 16:20. Only 28-9-2021 is state holiday (however the presentation „Membranes and organelles“ will be sent to your email or IS.MUNI profile)

Biology LAB EXERCISE – will be each second week, the protocols will be evaluated and you must have credits from each protocol. If somebody is ill in some week, the extra exercise will be prepared in December.

Today presentation (for day 21-9-2021) is focused to BASIC DEFINITION OF CELL BIOLOGY and BASIC OVERVIEW of Visualization technique in Medicine

What is visibility?

And what visibility is needed in medicine? :

Analogy:

If we fight against “**forest** disaster“, sometimes we need technique for macroscopic visibility, sometimes for detail (microscopic) visibility



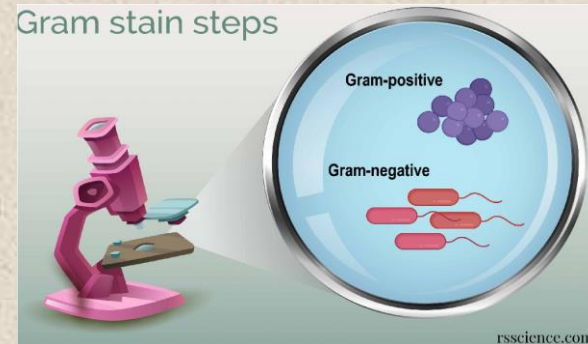
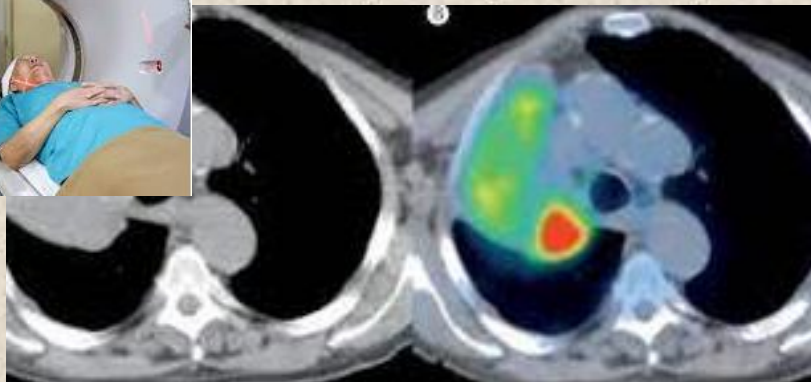
Good objective visualisation = key step for good fighting

What is visibility?

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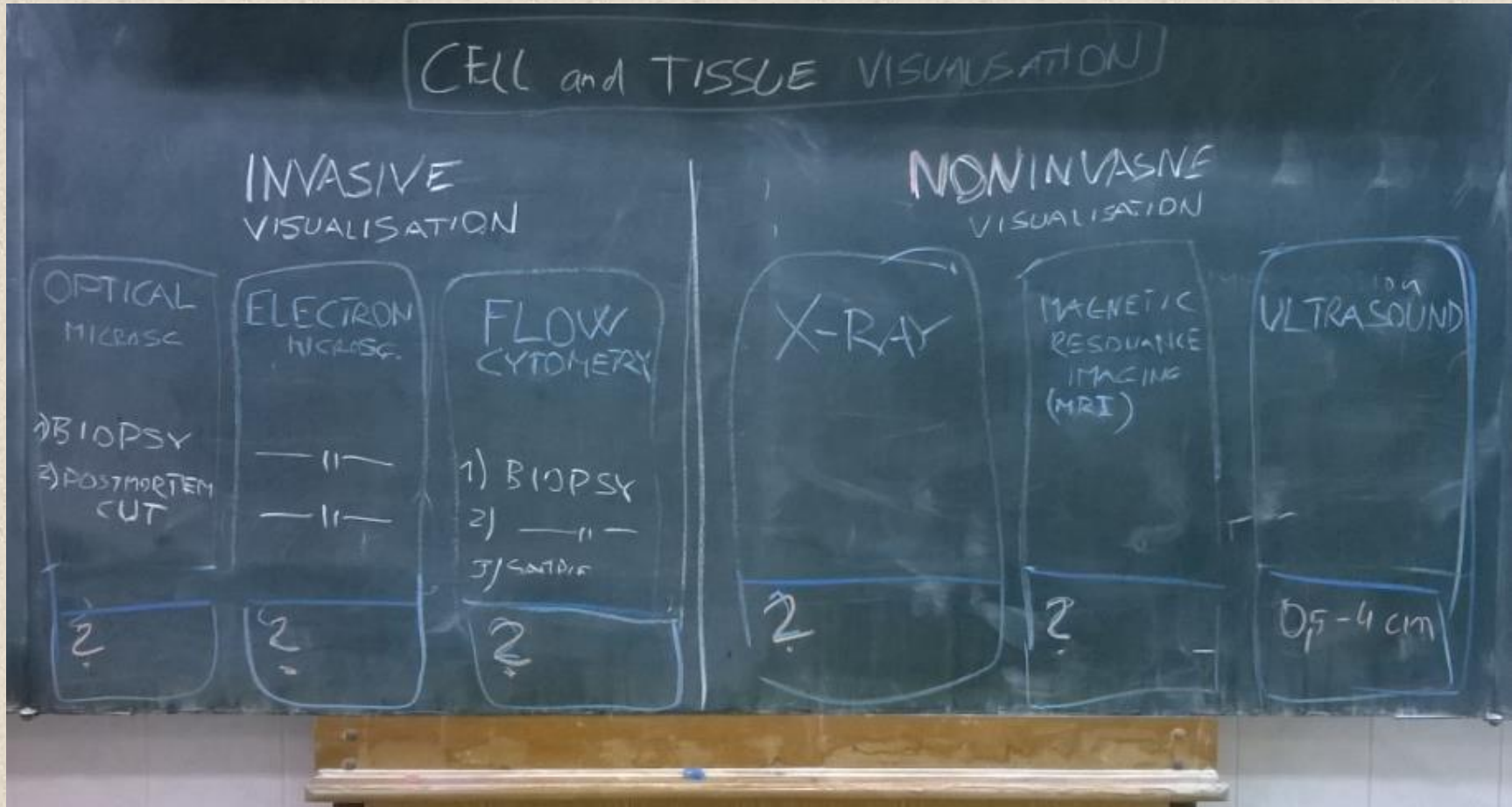
Good objective visualisation = key step for good fighting

The human and animal body is not o „bag of sugar water with smal soul inside“, however exact description of body and tissue structure had to wait to first „science-man“ **Aristotle** (384–322 BC). Before Aristotle, many Greek philosophers had speculated copartments of body and live organims but their theorizing was unsupported by empirical investigation.

TABLE 1–1 HISTORICAL LANDMARKS IN DETERMINING CELL STRUCTURE

1665	Hooke uses a primitive microscope to describe small chambers in sections of cork that he calls “cells.”
1674	Leeuwenhoek reports his discovery of protozoa. Nine years later, he sees bacteria for the first time.
1833	Brown publishes his microscopic observations of orchids, clearly describing the cell nucleus.
1839	Schleiden and Schwann propose the cell theory, stating that the nucleated cell is the universal building block of plant and animal tissues.
1857	Kölliker describes mitochondria in muscle cells.
1879	Flamming describes with great clarity chromosome behavior during mitosis in animal cells.
1881	Cajal and other histologists develop staining methods that reveal the structure of nerve cells and the organization of neural tissue.
1898	Golgi first sees and describes the Golgi apparatus by staining cells with silver nitrate.
1902	Boveri links chromosomes and heredity by observing chromosome behavior during sexual reproduction.
1952	Palade, Porter, and Sjöstrand develop methods of electron microscopy that enable many intracellular structures to be seen for the first time. In one of the first applications of these techniques, Huxley shows that muscle contains arrays of protein filaments—the first evidence of a cytoskeleton.
1957	Robertson describes the bilayer structure of the cell membrane, seen for the first time in the electron microscope.
1960	Kendrew describes the first detailed protein structure (sperm whale myoglobin) to a resolution of 0.2 nm using X-ray crystallography. Parutz proposes a lower-resolution structure for hemoglobin.
1965	Christian de Duve and his colleagues use a cell-fractionation technique to separate peroxisomes, mitochondria, and lysosomes from a preparation of rat liver.
1968	Petrán and collaborators make the first confocal microscope.
1970	Frye and Edidin use fluorescent antibodies to show that plasma membrane molecules can diffuse in the plane of the membrane, indicating that cell membranes are fluid.
1974	Lazarides and Weber use fluorescent antibodies to stain the cytoskeleton.
1994	Chalfie and collaborators introduce green fluorescent protein (GFP) as a marker to follow the behavior of proteins in living cells.

We can divide the technique via the invasivity or the non-invasivity:



Homework: 1) add resolution to your exercisebook (information are on following pages)
2) describe basic components of optical microscopy.

Each technique have some advantages and disadvantages:

For example Hematoma of leg

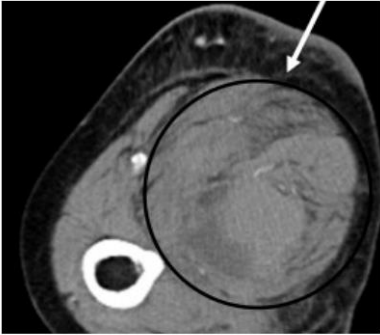
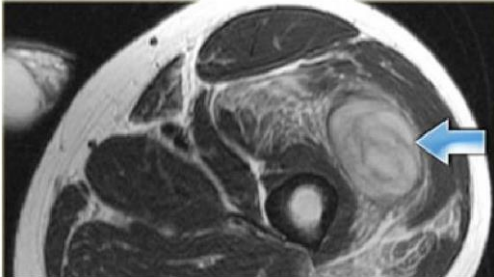
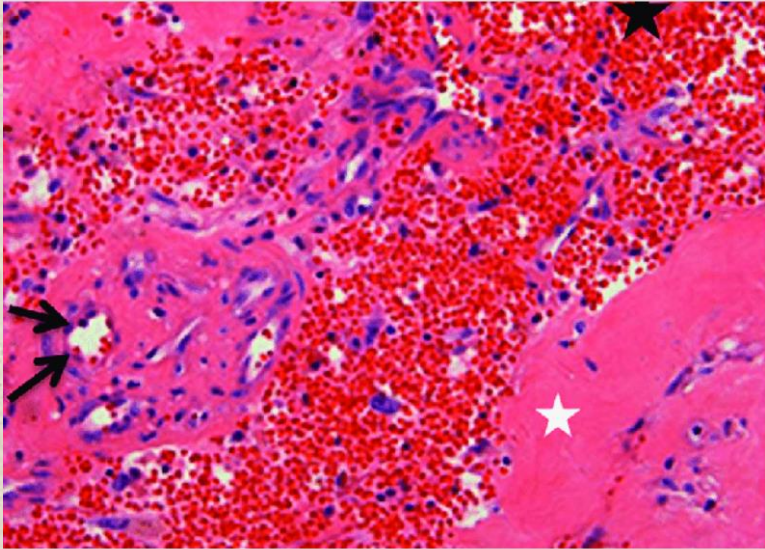


Hematoma by microscopy (focused inside the muscle)

Hematoma by X-RAY CT

Hematoma by NMRI

Hematoma by Ultra Sound

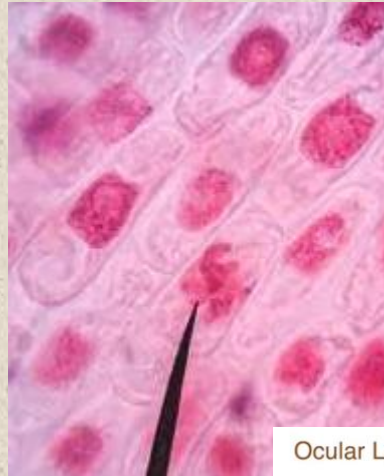


Microscopic view of organized hematoma showing angiogenesis (arrows), fibrosis (white asterisk) and extravasated red blood cells (black asterisk) (hematoxylin and eosin staining, original magnification, $\times 400$).

For medical and pharmacological curative strategy we need mostly combination of all these techniques.

Ad. Optical and Electron MICROSCOPY

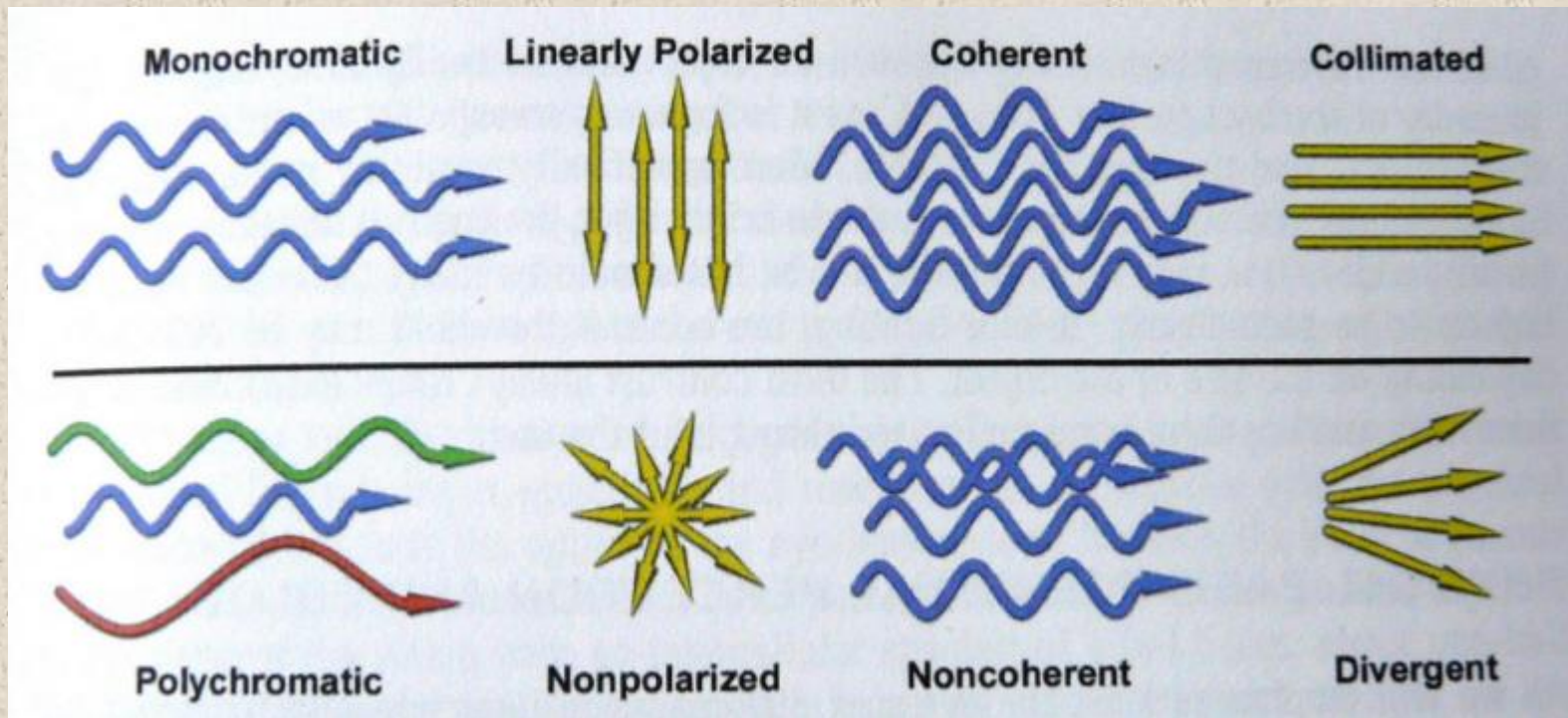
Looking to the tissue structure by microscopy is the similar problem like “visualisation“ of paving stone by blind man



We need identify where is the border of paving stone or cells (or another biological entities)

Basic tools for visualisation = fotons (light) and electron (particle)

Basic definition for diferent form of light, which are important for scinceman in microscopy:



Two important aspect of microscopy visualisation:

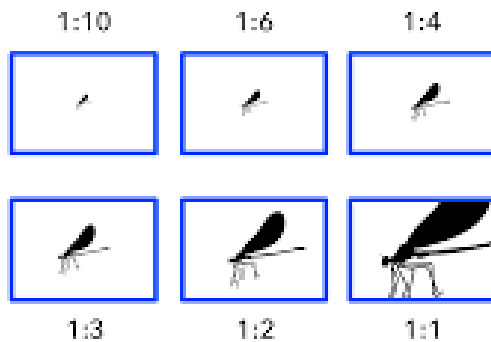
MAGNIFICATION

and

RESOLUTION

May be you know the theory from high school. Who not. There is very clear definiton: <https://www.youtube.com/watch?v=CVusz4wHaic>

Example Magnifications

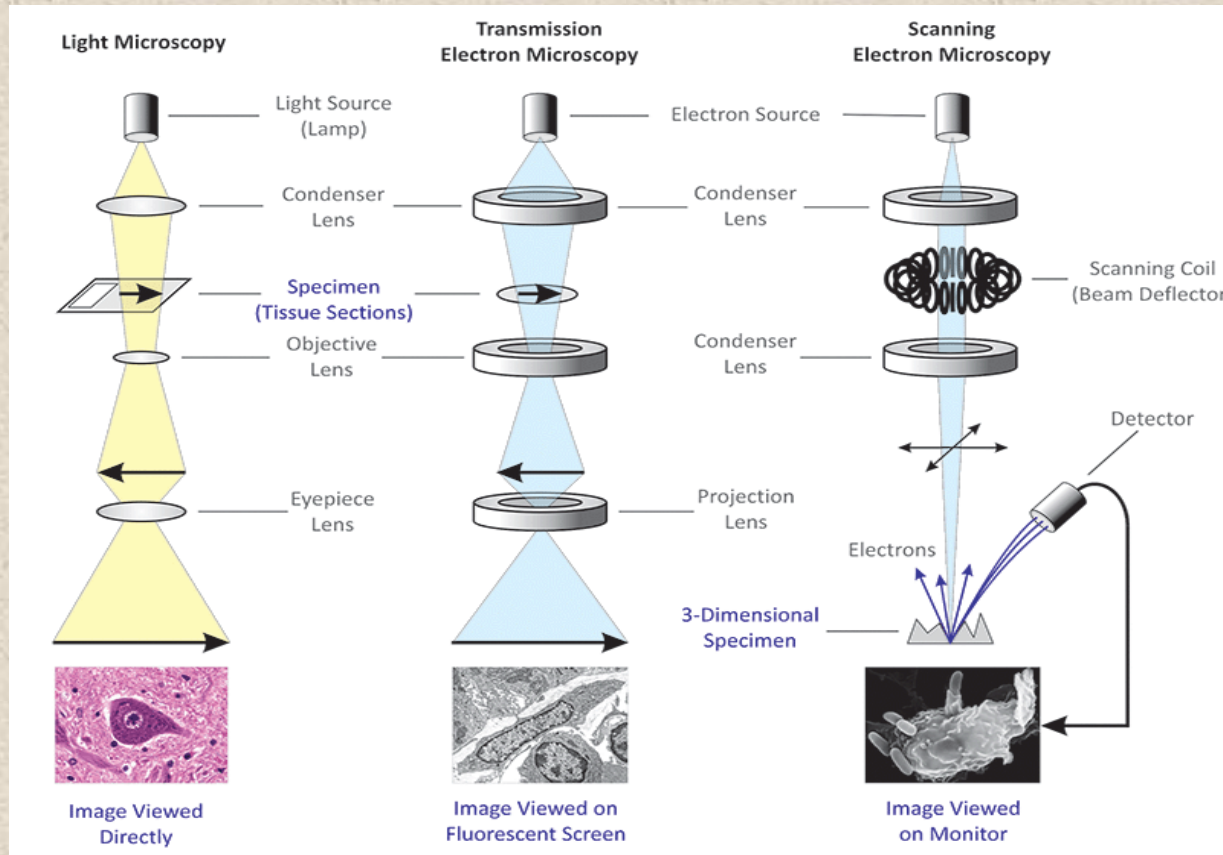


High resolution = ther is visibility of two leaflet

Low resolution = there is not, two leaflet seems like one violet flag

Basic construction of microscopies:

OPTICAL and ELECTRON



Resolution

200 nm

0.1 nm

0.5 nm

Magnific.

~ ×2000

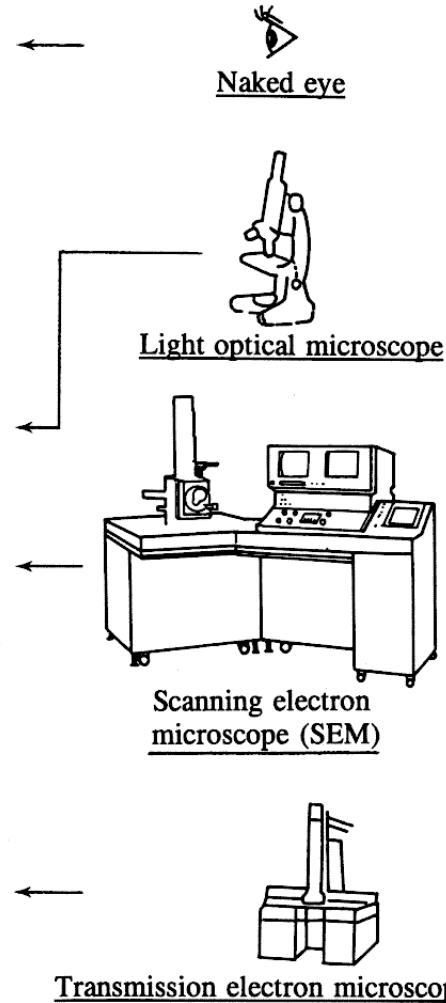
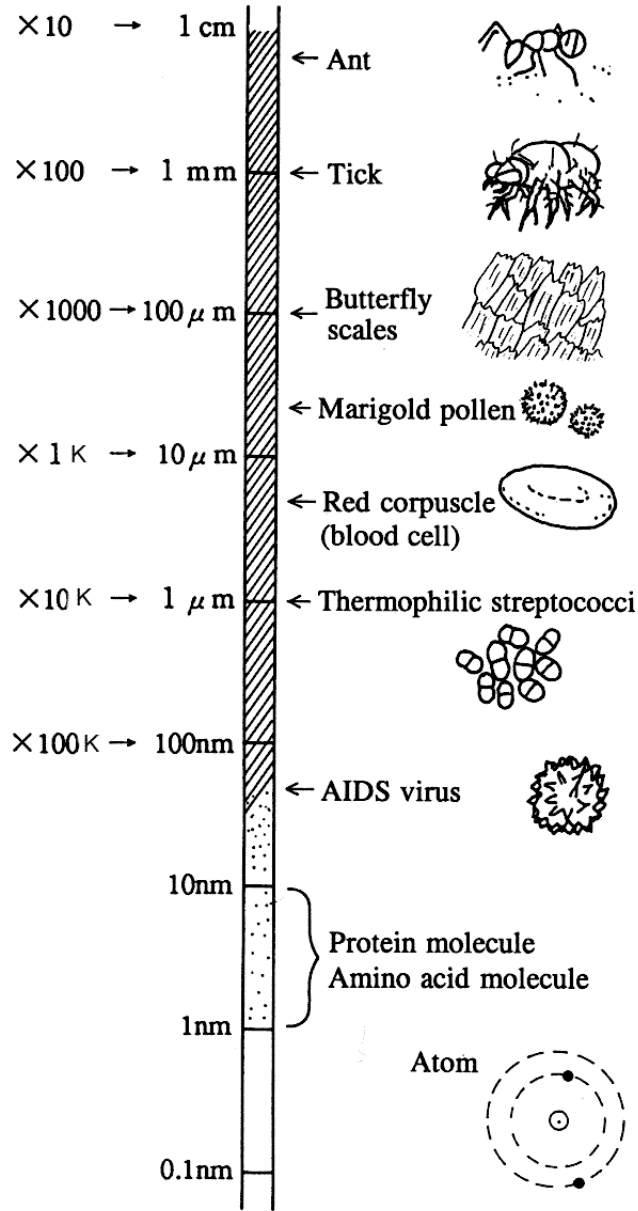
×50 ~ ×1,500,000

×10 ~ ×1,000,000

MAGNIFICATION

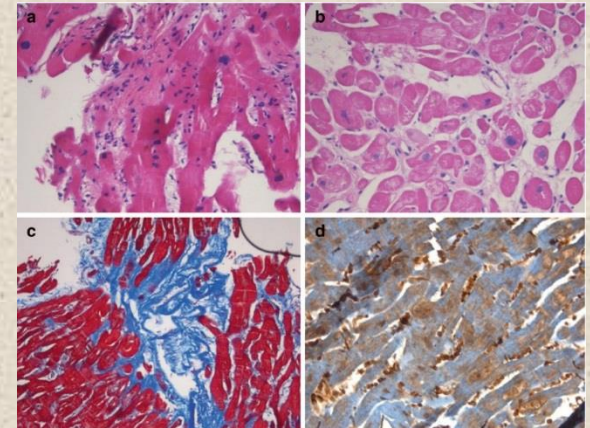
Object

RESOLUTION

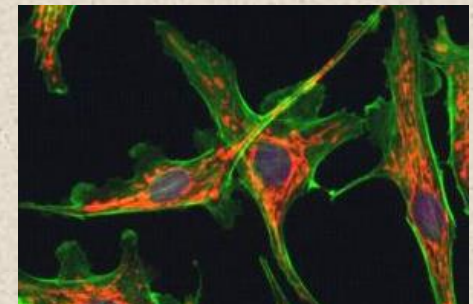
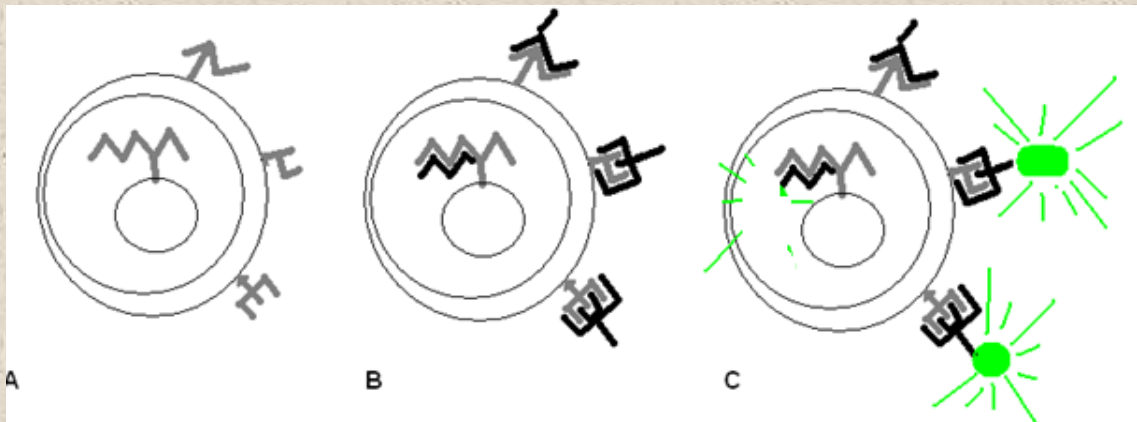


How we can upgrade some structure of cells for better contrast ?
Use staining.

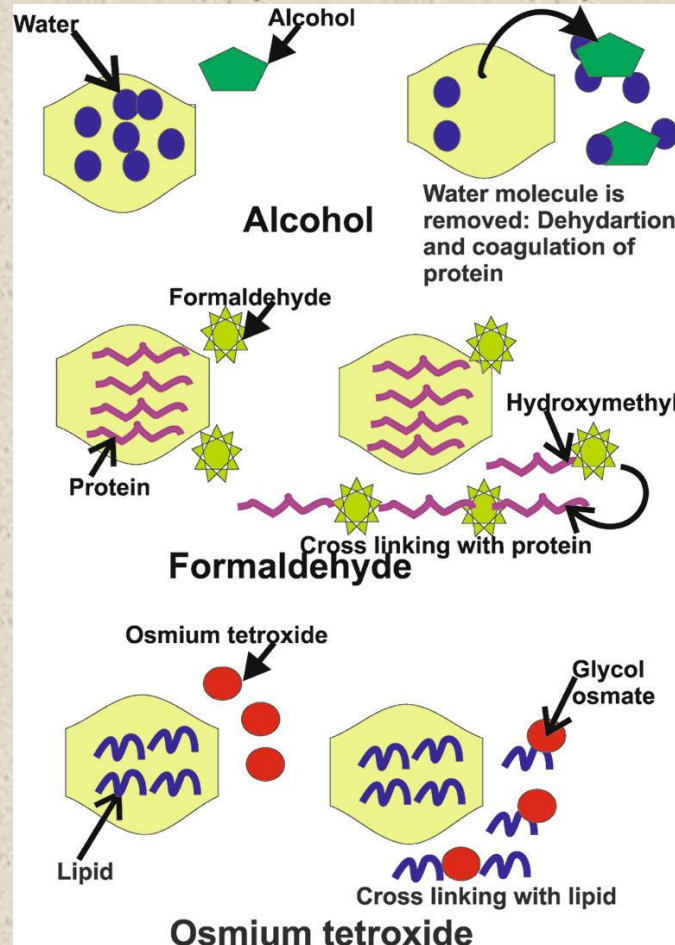
1) Traditional Histochemistry Staining (used chemicals which have specific affinity to some part of cell or tissue, for example DNA, collagen etc.



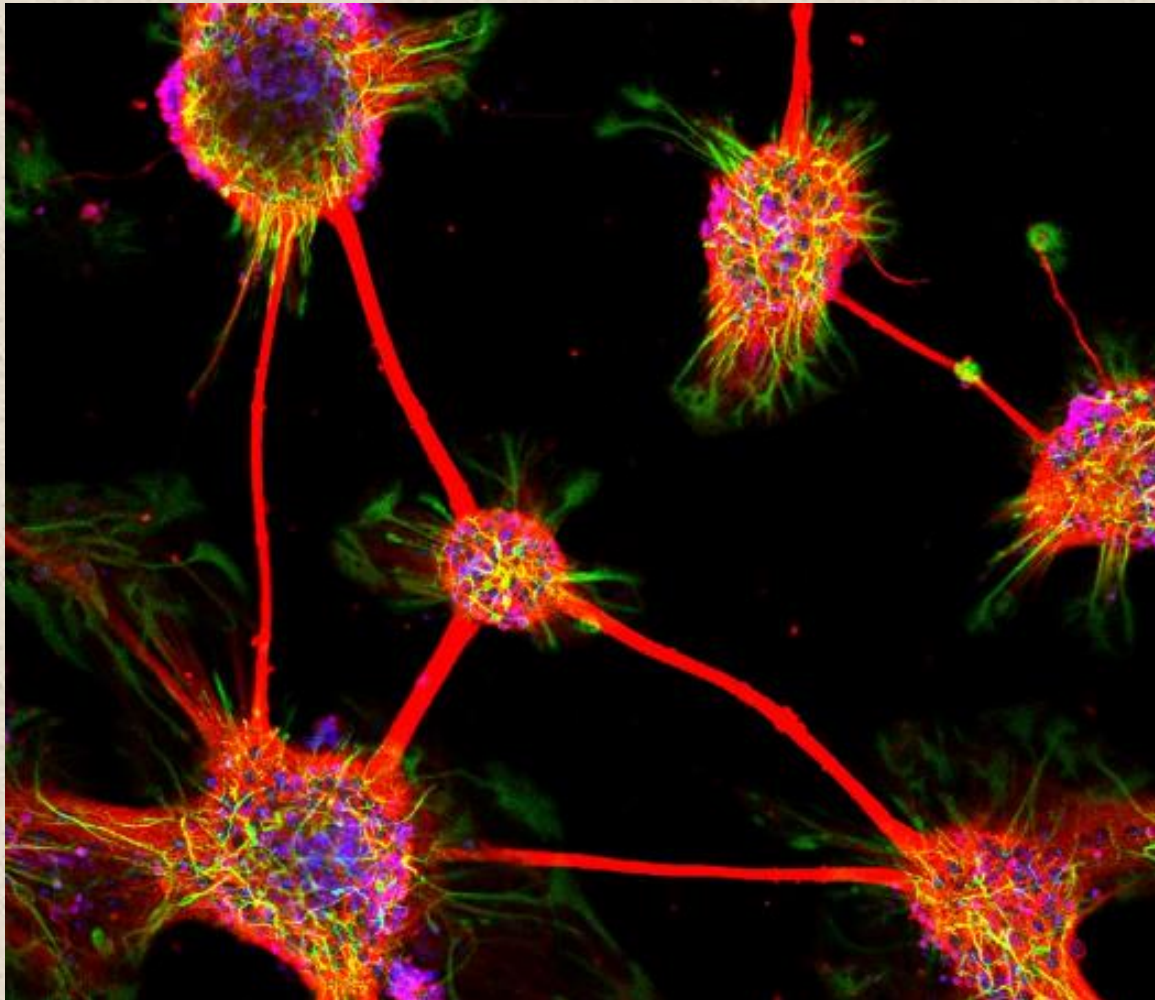
2) Antibody staining (best way: primary and secondary antibody which makes some structure fluorescent)



3) Variety of staining for Electron microscopy (mostly heavy metal like Osmium)



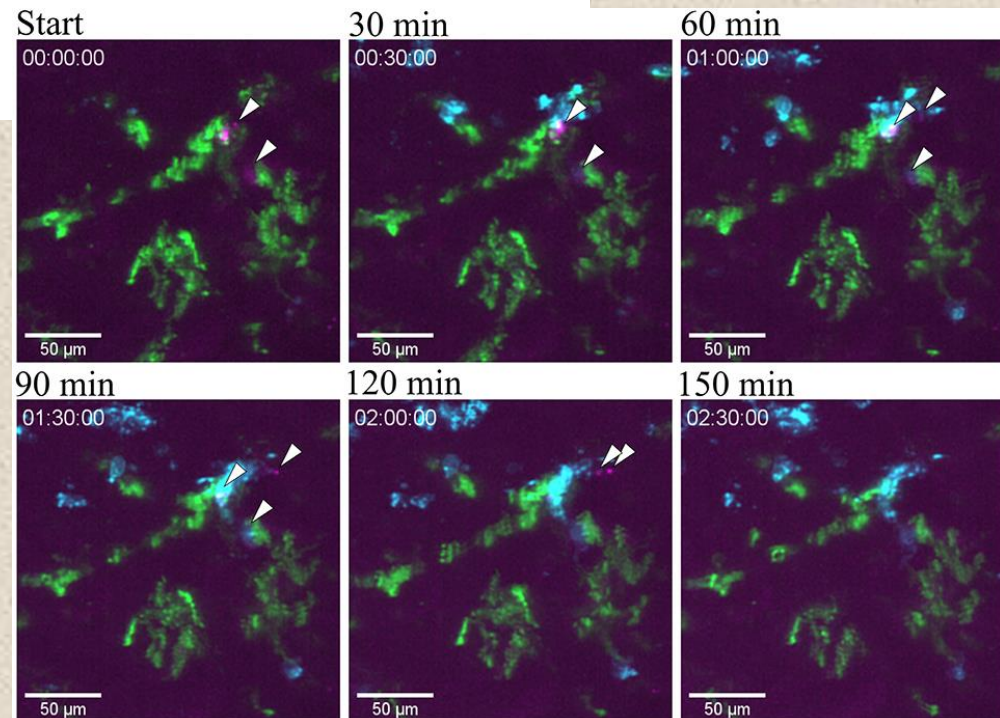
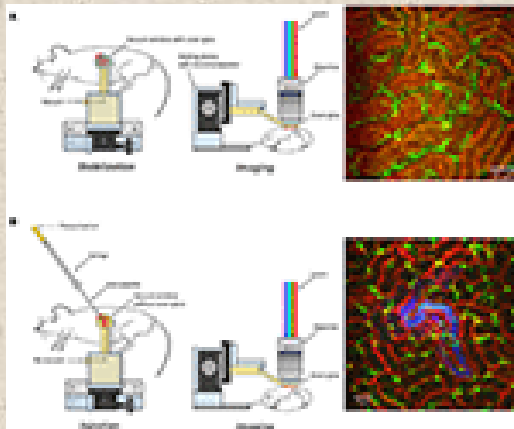
Example of final photo by optical microscope. Multicolor image (fluorescence microscopy of fluorescent probe adhered to specific part of neuron cells)



Most of the microscopy scan needs separation of sample from live body. However there are also futuristic and very modern machines, for example INTRAVITAL MICROSCOPY:

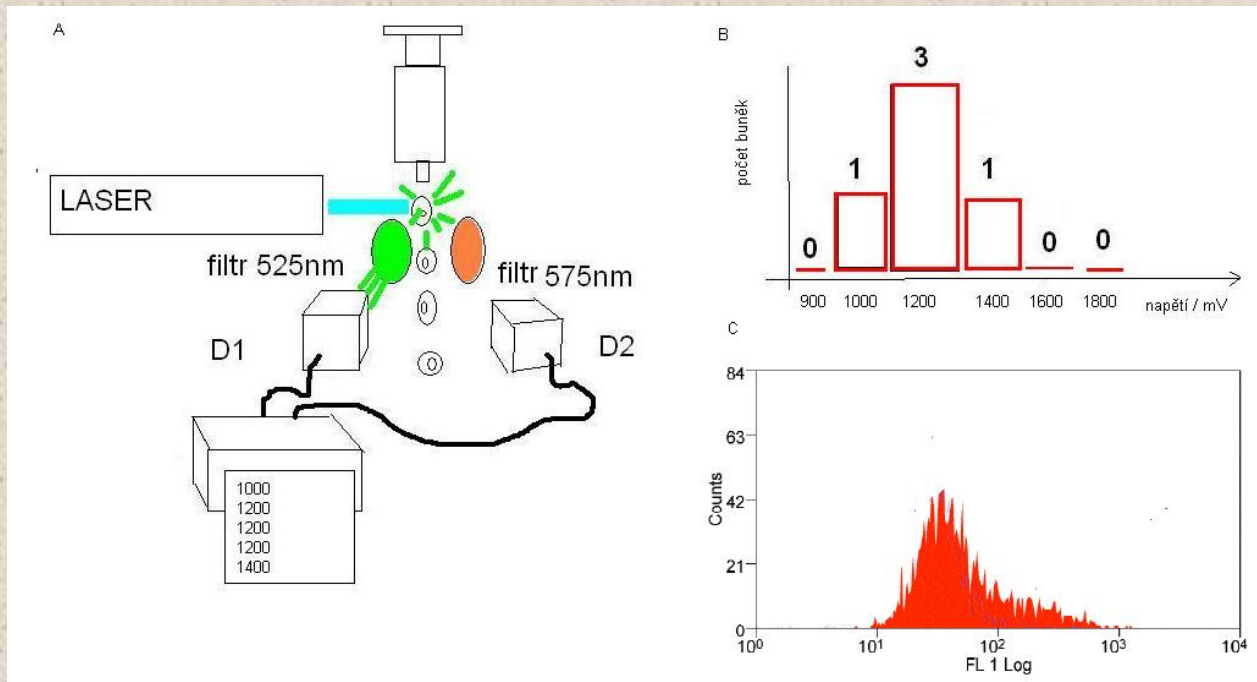
High Resolution Intravital Imaging of the Renal Immune Response to Injury and Infection in Mice

[John Sedin](#)



Ad. FLOW-CYTOMETRY

Flow cytometry is a technology that **provides rapid multi-parametric analysis of single cells in solution**. Flow cytometers have started be used in hematology to speed analysis of blood cells and analysis of surface protein on these cells

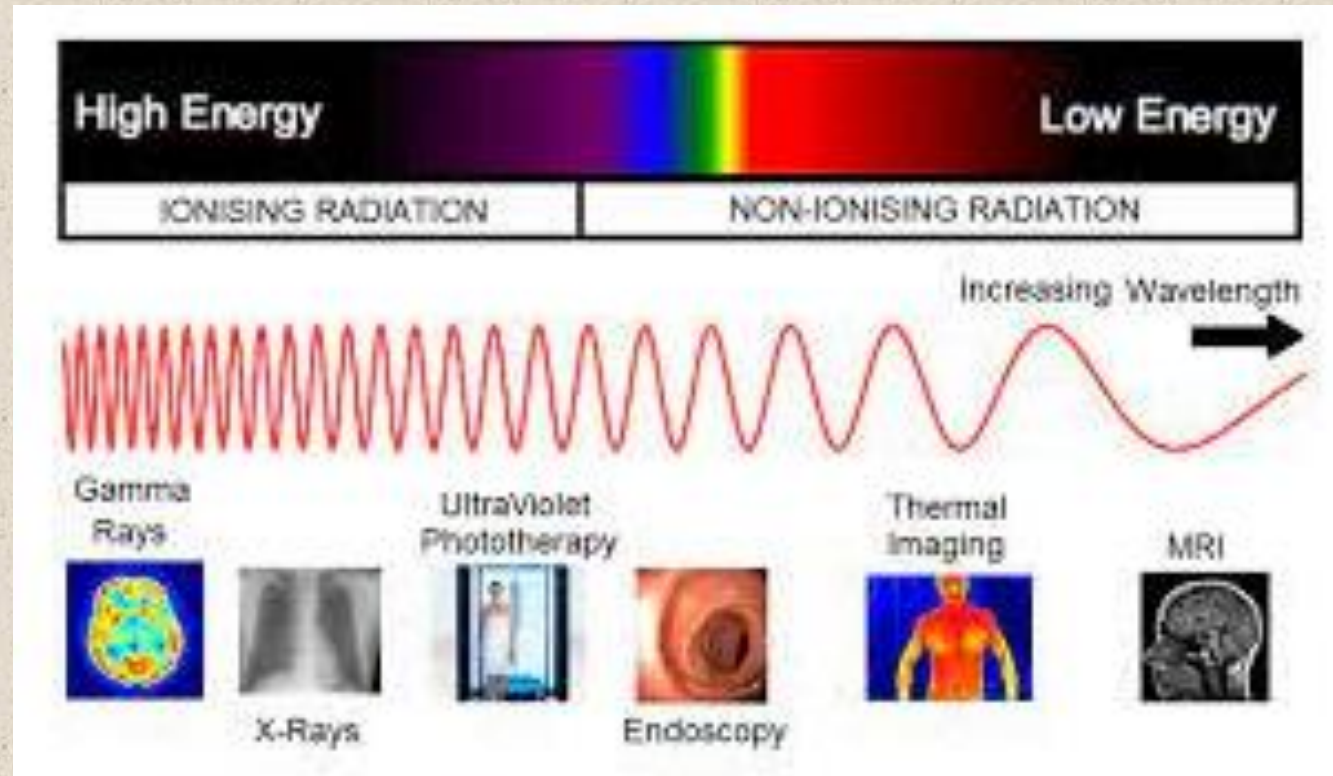
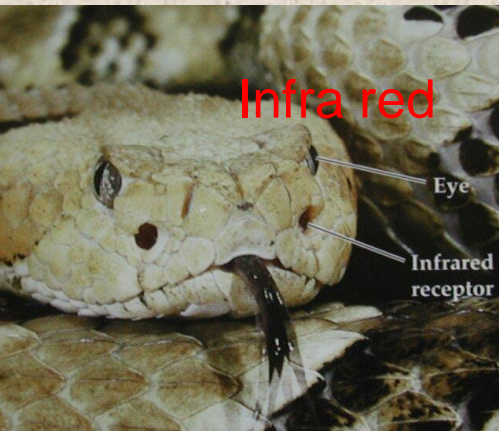


Good illustrative video:

<https://www.youtube.com/watch?v=B2zreF2dnWk>

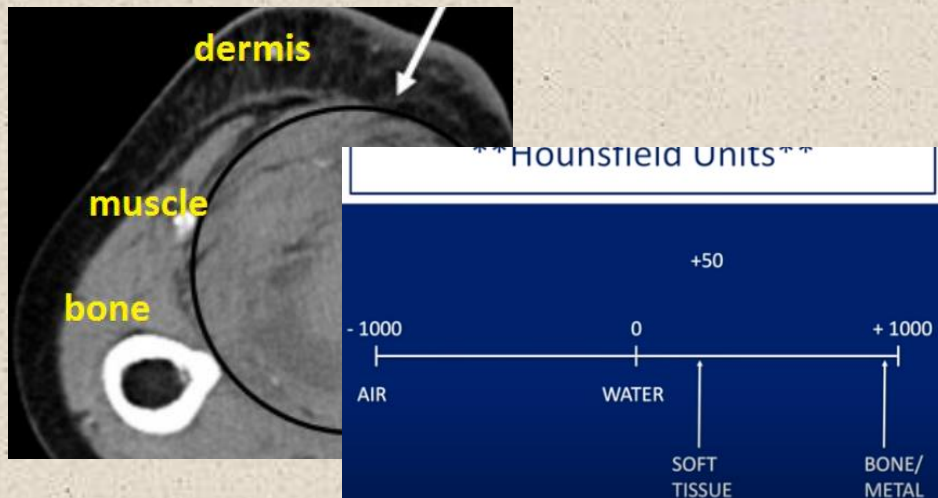
Ad. X-RAY and MRI TOMOGRAPHY

People have only one visibility detector – eye. Half a century ago, the light was only detecting tool for all doctors and biology specialist. However visualisation of some animal and some machine is based on another electromagnetic :

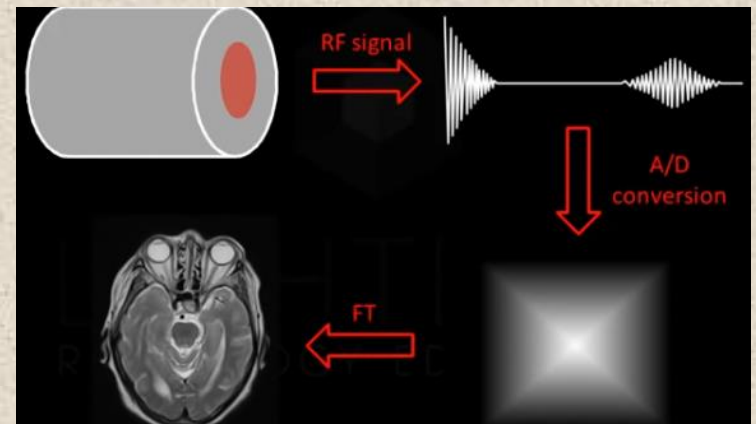


X-RAY CT and also MRI have only grey scale output. We can say that each millimeter of tissue is precomputed to one pixel (whit, grey or black). So microscopy has variety of color and resolution of cells ant biological entities could be higher thanks to specific staining, on the oposite the CT has contrast which correlates to „DENSITY OF MATERIAL“ (bones – the highest density, air – the lowest), MRI has contrast which correlates to H2O concentration in the tissue (bone – low concentration, soft tissue – higher concentration). For aditive contrast we use for example stining of intestine or blood by metal nanoparticles.

X-RAY CT



MRI



CT and MRI are very sophisticated technical apparatus, where great physical theoretic background is needed from quantum physics and nuclear physics theory. Very illustrative videos for medical and biological workers is here:

CT image quality and

<https://www.youtube.com/watch?v=qsHTrQ0lb2s>

MRI basic principles and resolution

<https://www.youtube.com/watch?v=Ok9ILlYzmaY>

<https://www.youtube.com/watch?v=aQZ8tTZnQ8A>

<https://www.youtube.com/watch?v=VnpqylFYtql>

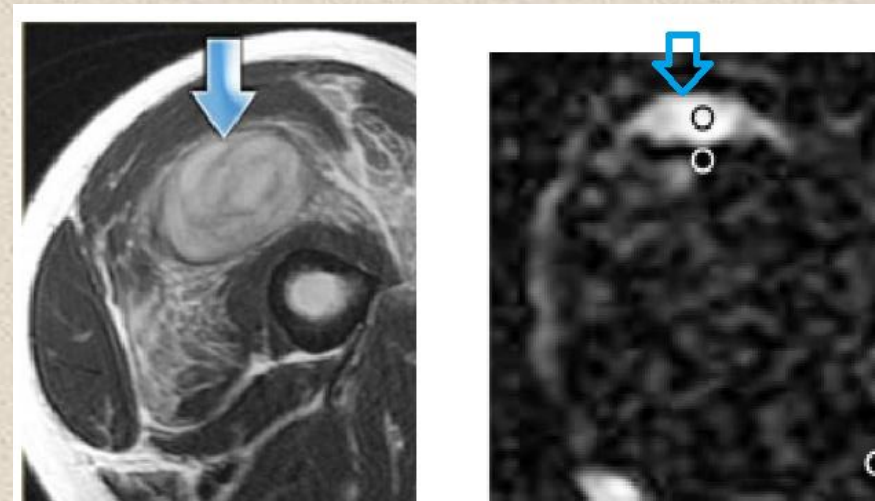
X-RAY CT



The spatial resolution of CT is excellent and the primary strength of the modality. Current CT scanners have a spatial resolution of **0.5–0.625 mm in the z-axis**, and approximately 0.5 mm in the x- to y-axes

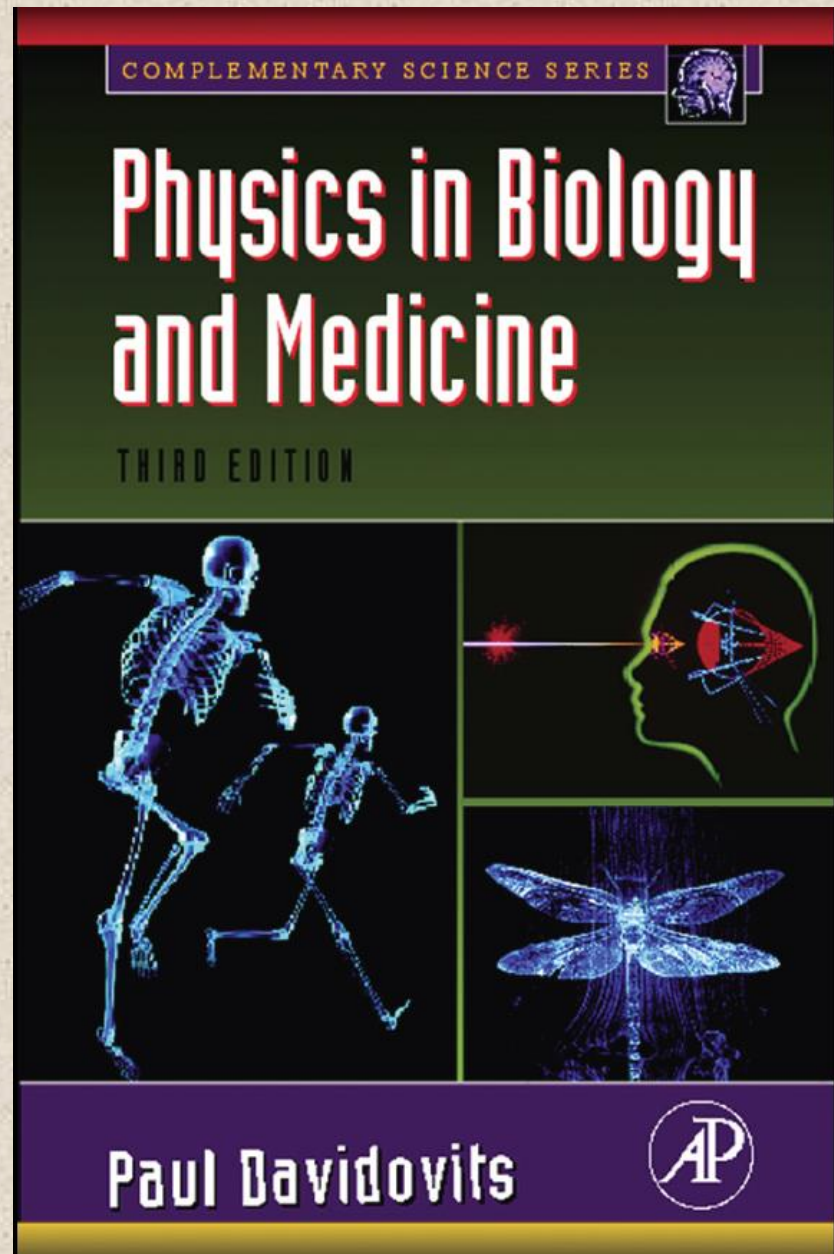
MRI

High resolution vs. Bad resolution



Nowadays, most MRI scanners used for medical purposes have B_0 values of 1.5 or 3 T and can reach typical resolutions of around **$1.5 \times 1.5 \times 4$ mm**. In parallel, ultra-high magnetic field MRI scanners with $B_0 = 11.7$ T are developed for research purpose and resolutions of $80 \times 80 \times 200$ μm

Detail
literature:



CELLS and CELL BIOLOGY

Cells are the fundamental units of life. Thus it is to *cell biology*—the study of cells and their structure, function, and behavior—that we must look for an answer to the question of what life is and how it Works.

Medicinal experts should have good overview not only about human cells, but also about another historical cells and viruses, because their interaction with human body is critical development of many pathologies (flu, diabetic wound, pathology of intestine micro-organism, ...**aerobic or anaerobic environment could induce different bacterial activity etc**)



Bacteria

Archaea

Eukarya



Prokaryotes



Eukaryotes

The Prokaryotic Cell

Of all the types of cells revealed by the microscope, *bacteria* have the simplest structure and come closest to showing us life stripped down to its essentials. Indeed, a bacterium contains essentially no organelles—not even a nucleus to hold its DNA. This property—the presence or absence of a nucleus—is used as the basis for a simple but fundamental classification of all living things. Organisms whose cells have a nucleus are called eukaryotes (from the Greek words *eu*, meaning “well” or “truly,” and *karyon*, a “kernel” or “nucleus”). Organisms whose cells do not have a nucleus are called prokaryotes (from *pro*, meaning “before”).

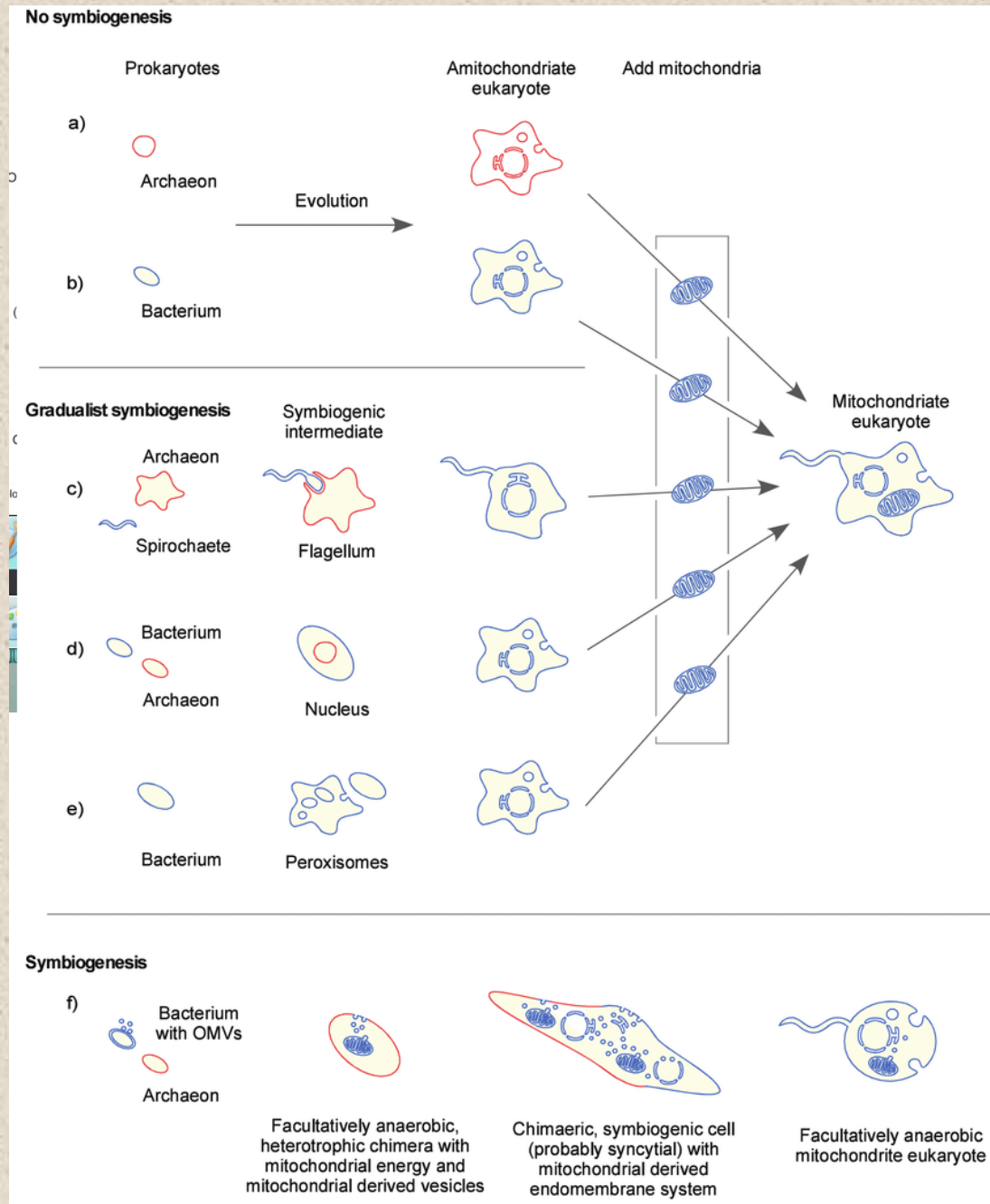
The Eukaryotic Cell

Eukaryotic cells, in general, are bigger and more elaborate than bacteria and archaea. Some live independent lives as single-celled organisms, such as amoebae and yeasts ([Figure 1–13](#)); others live in multicellular assemblies. All of the more complex multicellular organisms—including plants, animals, and fungi—are formed from eukaryotic cells. By definition, all eukaryotic cells have a nucleus. But possession of a nucleus goes hand-in-hand with possession of a variety of other organelles,

Notes to relationship of Prokaryotic and Eukaryotic

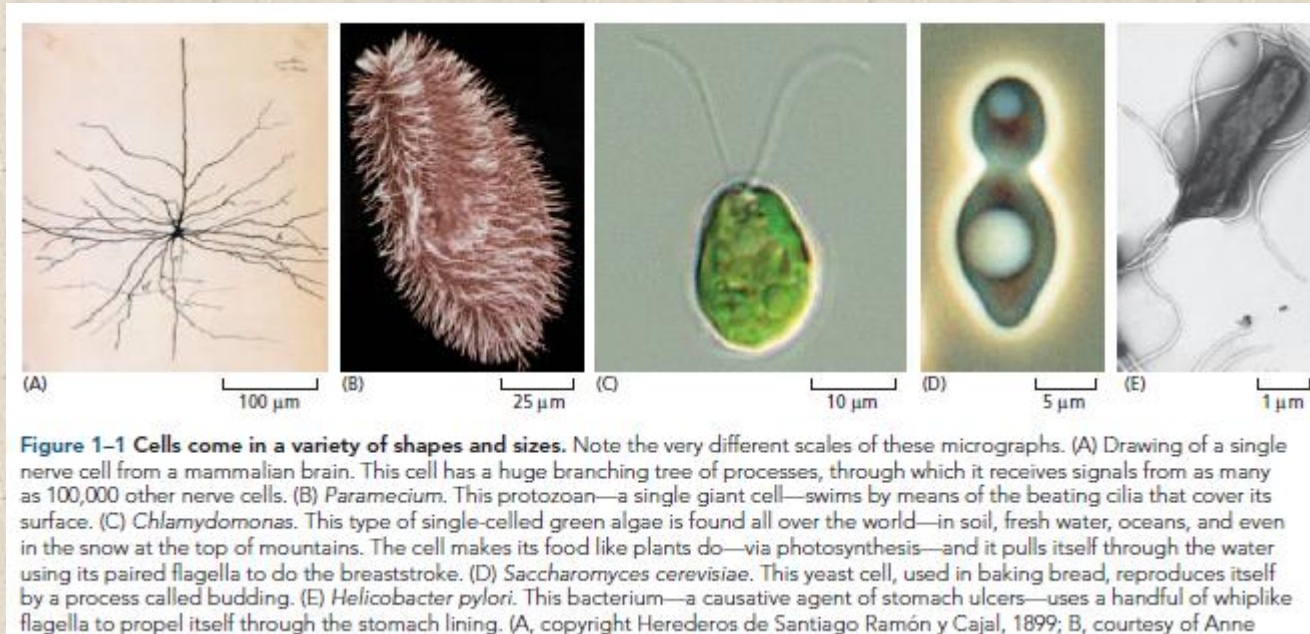
....Schematic comparison of selected theories for eukaryote origin.

Details in
Symbiogenesis, gradualism, and mitochondrial energy in eukaryote origin
September 2017
[Periodicum Biologorum](#) 119(3)



Date (million years ago)	Organisms	Events	Atmospheric oxygen (~%)
3800	Prokaryote chemoautotrophs	Origin of life	0
3500–3000	Prokaryote heterotrophs; precursors of cyanobacteria. Stromatolites. Sulfur bacteria	Beginning of <u>photosynthesis</u>	Traces
2100	Filamentous spirally curled organisms, (Grypania)	Major land masses; shallow seas, Iron deposits, BIFs	0.1%
2000	Cyanobacteria tolerant to O ₂	Sterols in bitumen (fossil organisms)	0.2%
1700	Spheromorph Acritarchs, primitive unicellular eukaryotes	Atmosphere oxidising Endosymbiosis. <u>Aerobic respiration</u>	0.3%
1200	Red algae and metaphytes	Large cells. Endosymbiosis. Aerobic respiration. Meiosis. Genetic recombination	0.5%
1000–550	Various primitive multicellular eukaryotes in precambrian fossils, some mineralized. Green algae dominant. Early land plants	Fossils and tracks. Oxygen and ozone accumulating	1–4%
450–present	Full flourishing multicellular <u>eukaryotes</u> ; land living organisms	<u>Ozone layer completed.</u> Crust movements more pronounced. Super continents formed. Ocean basins altered	10–21%

Cells vary enormously in appearance and Function

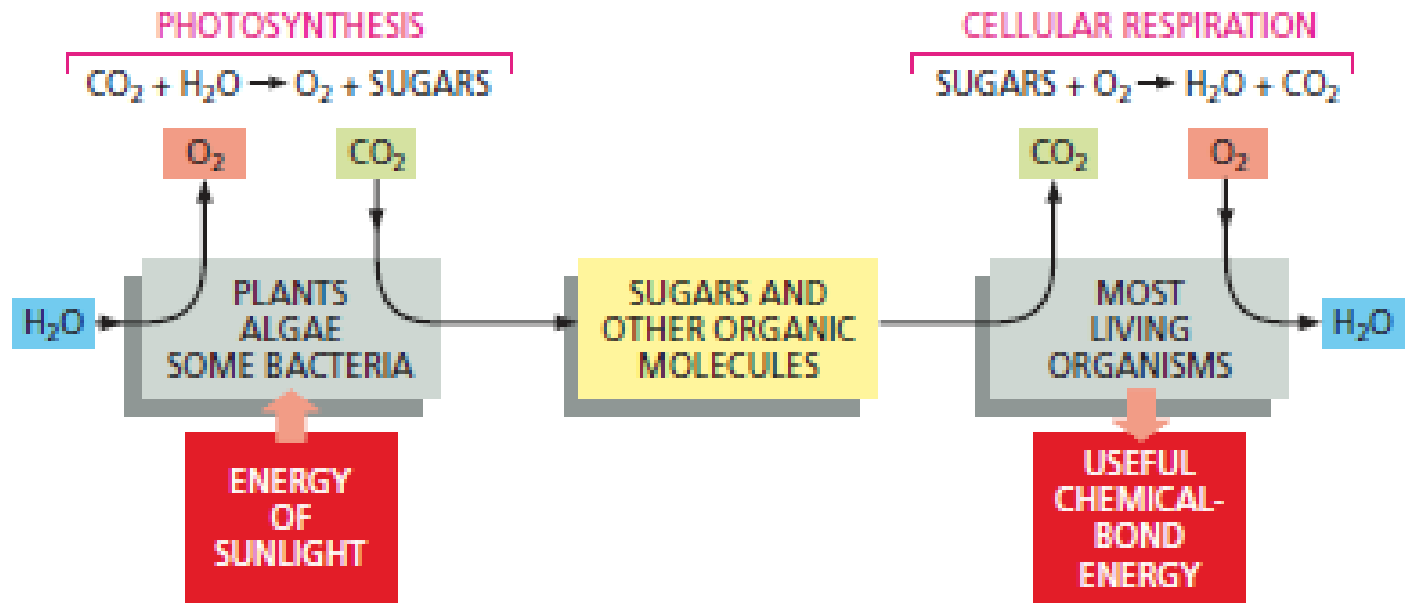


However biologist during centuries of modern science made basic identification of basal cell principles:

Fundamental principles of all known cells are:

(A) CHEMICAL MACROMOLECULES and STRUCTURE

(B) BIOENERGETIC



Ha ha ha
our cells are different



Essential Concepts

- Cells are the fundamental units of life. All present-day cells are believed to have evolved from an ancestral cell that existed more than 3 billion years ago.
- All cells are enclosed by a plasma membrane, which separates the inside of the cell from its environment.
- All cells contain DNA as a store of genetic information and use it to guide the synthesis of RNA molecules and proteins.
- Cells in a multicellular organism, though they all contain the same DNA, can be very different. They turn on different sets of genes according to their developmental history and to signals they receive from their environment.
- Animal and plant cells are typically 5–20 μm in diameter and can be seen with a light microscope, which also reveals some of their internal components, including the larger organelles.c

The electron microscope reveals even the smallest organelles, but specimens require elaborate preparation and cannot be viewed while alive.

- Specific large molecules can be located in fixed or living cells with a fluorescence microscope.
- The simplest of present-day living cells are prokaryotes: although they contain DNA, they lack a nucleus and other organelles and probably resemble most closely the ancestral cell.
- Different species of prokaryotes are diverse in their chemical capabilities and inhabit an amazingly wide range of habitats. Two fundamental evolutionary subdivisions are recognized: bacteria and archaea.
- Eukaryotic cells possess a nucleus and other organelles not found in prokaryotes. They probably evolved in a series of stages, including the acquisition of mitochondria by engulfment of aerobic bacteria and (for plant cells) the acquisition of chloroplasts by engulfment of photosynthetic bacteria.
- The nucleus contains the genetic information of the eukaryotic organism, stored in DNA molecules.

- The cytoplasm includes all of the cell's contents outside the nucleus and contains a variety of membrane-enclosed organelles with specialized functions: mitochondria carry out the final oxidation of food molecules; in plant cells, chloroplasts perform photosynthesis; the endoplasmic reticulum and the Golgi apparatus synthesize complex molecules for export from the cell and for insertion in cell membranes; lysosomes digest large molecules.
- Outside the membrane-enclosed organelles in the cytoplasm is the cytosol, a very concentrated mixture of large and small molecules that carry out many essential biochemical processes.
- The cytoskeleton is composed of protein filaments that extend throughout the cytoplasm and are responsible for cell shape and movement and for the transport of organelles and other large molecular complexes from one location to another.
- Free-living, single-celled eukaryotic microorganisms are complex cells that can swim, mate, hunt, and devour other microorganisms.
- Animals, plants, and some fungi consist of diverse eukaryotic cell types, all derived from a single fertilized egg cell; the number of such cells cooperating to form a large multicellular organism such as a human runs into thousands of billions.
- Biologists have chosen a small number of model

Conclusion:

Cell definition

The cell is the structural and functional elementary unit of all living organisms, conserving the features of the organism, having the ability of self-control, self-regulation, and self-reproduction, being the result of a long time of evolution

Key terms, which everybody had to understand from actual lection of Biology

KEY TERMS

archaeon
bacterium
cell
chloroplast
chromosome
cytoplasm
cytoskeleton
cytosol
DNA
electron microscope

eukaryote
evolution
fluorescence microscope
genome
homologous
micrometer
microscope
mitochondrion
model organism

nucleus
organelle
photosynthesis
plasma membrane
prokaryote
protein
protozoan
ribosome
RNA