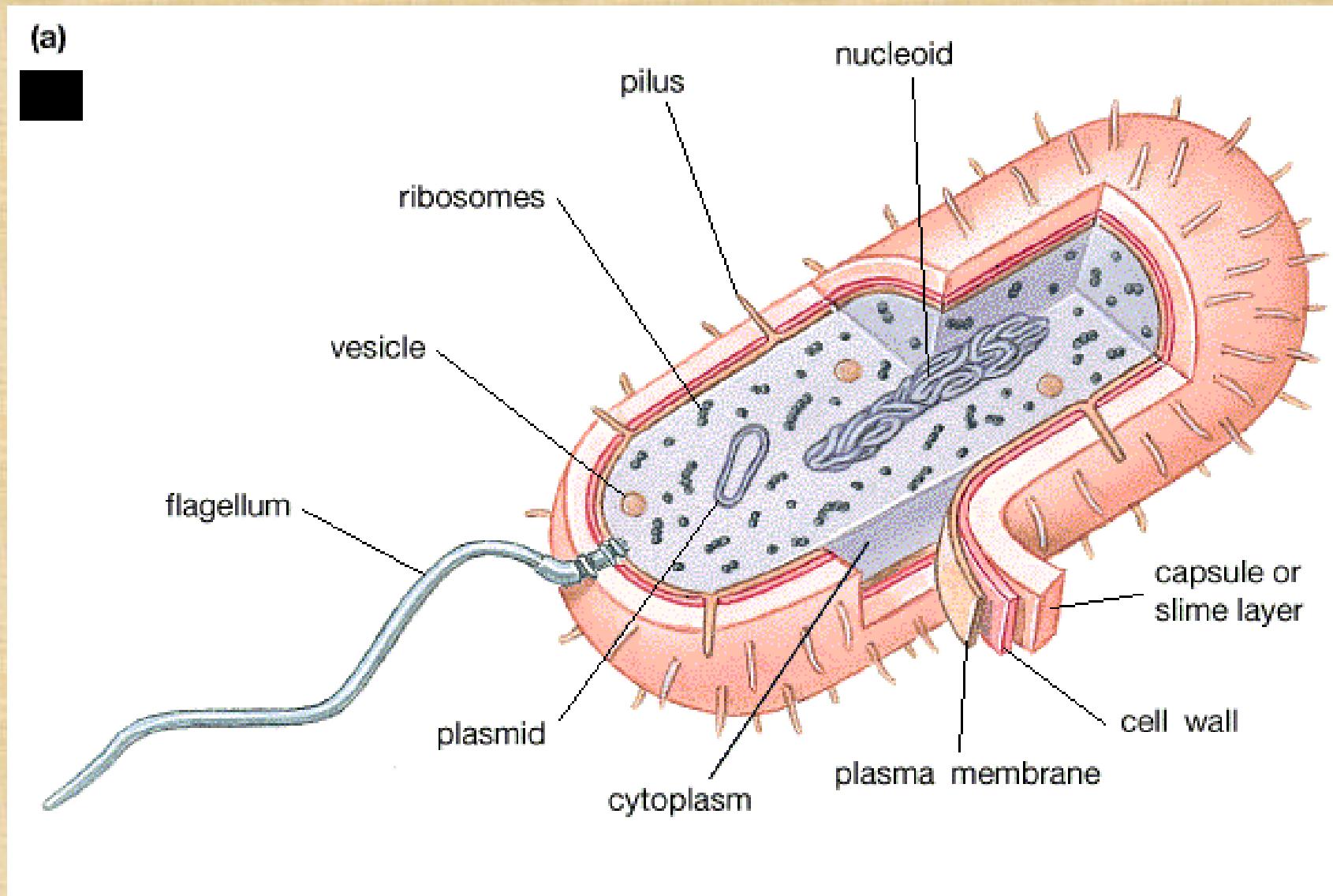


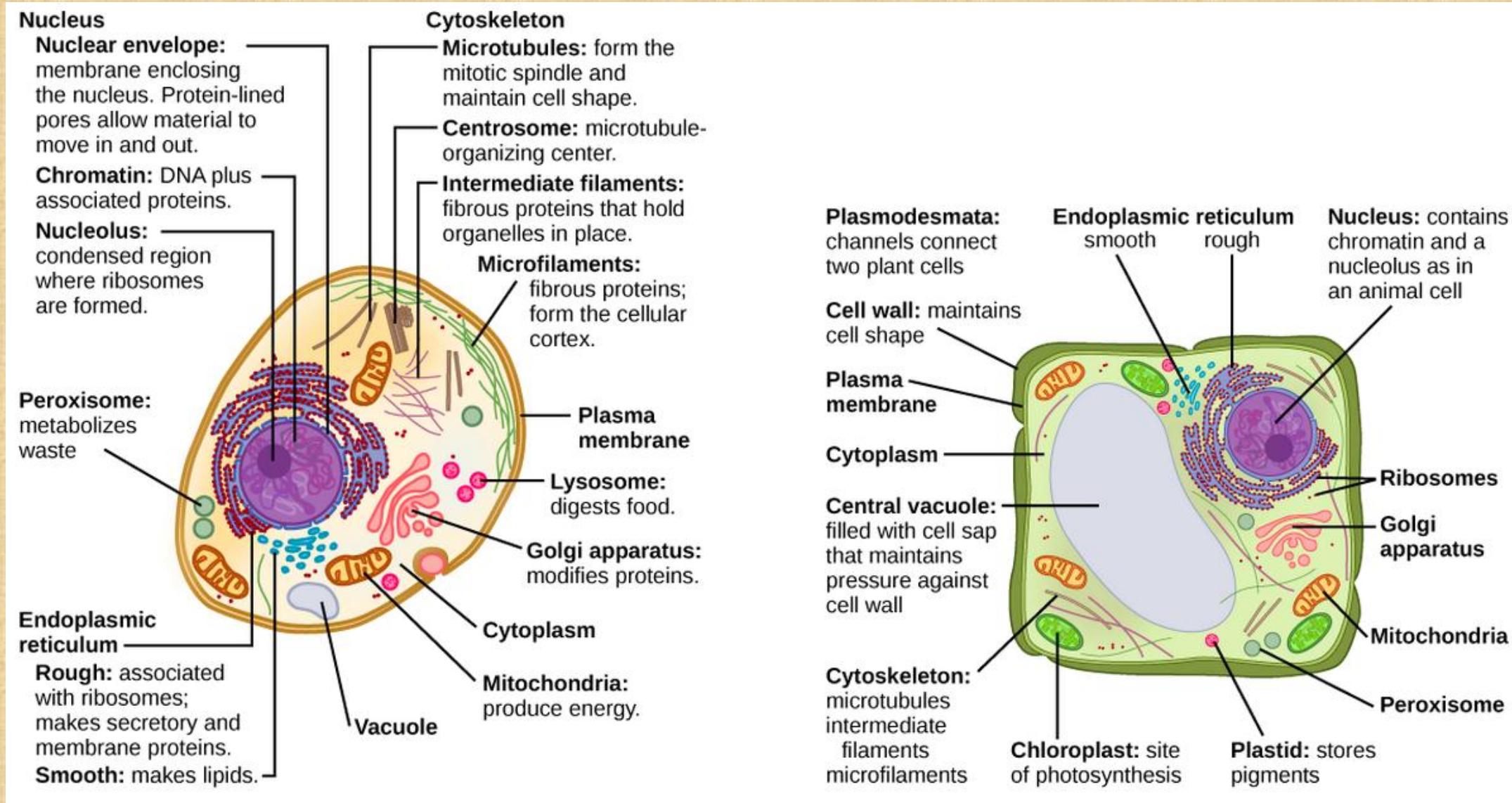
DNA: Genetic material

Dr. Ashok Kumar HG

Bacterium



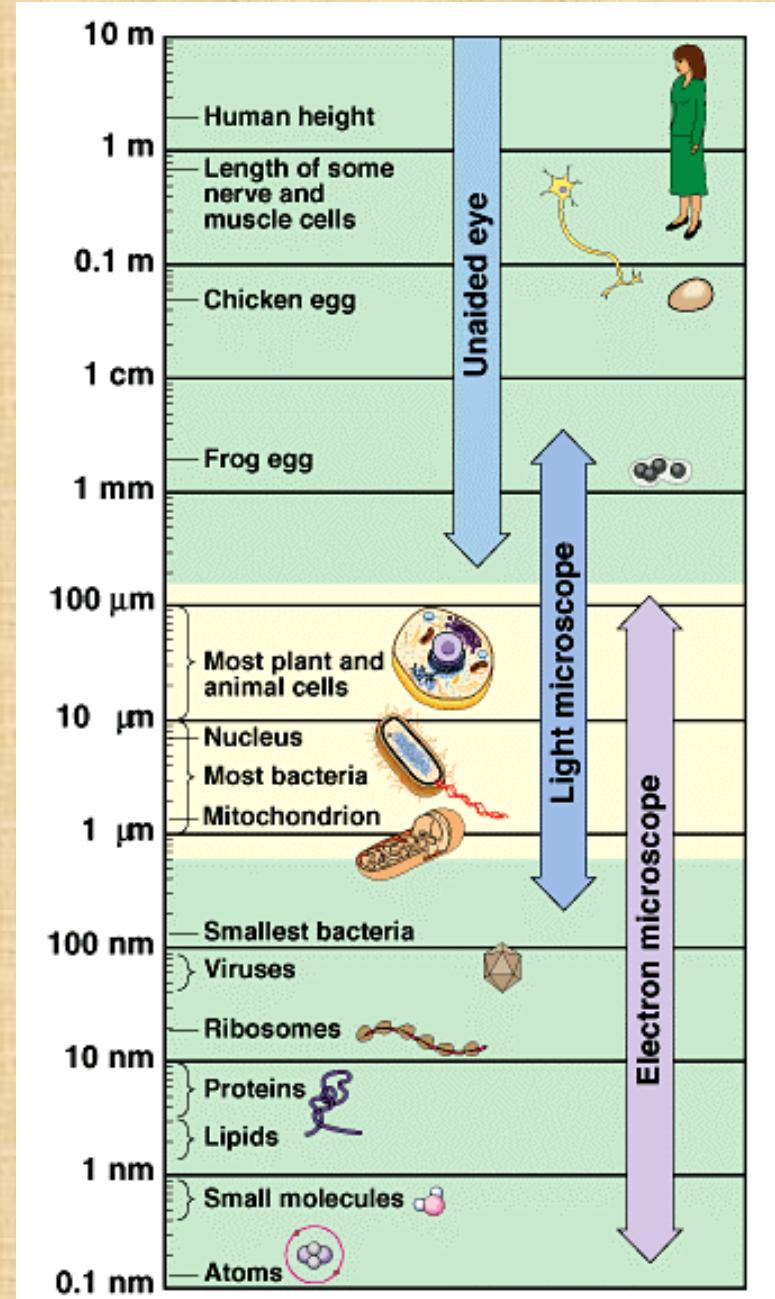
Animal and Plant Cells



Difference between Prokaryotic cell and Eukaryotic cell

	Prokaryotes	Eukaryotes
DNA	DNA is naked (no histones)	DNA associated with histones
	DNA is circular	DNA is linear
	Genes do not contain introns	Genes may contain introns
	DNA found in cytoplasm (nucleoid)	DNA found in nucleus
Internal Structures	No membrane-bound organelles	Have membrane-bound organelles
Ribosomes	Have 70S ribosomes	Have 80S ribosomes
Reproduction	Asexual (binary fission)	Asexual (mitosis) or sexual (meiosis)
	DNA is singular (haploid)	DNA is usually paired (diploid or more)
Average Size	Smaller ($\approx 1 - 5 \mu\text{m}$)	Larger ($\approx 10 - 100 \mu\text{m}$)

Cell size



(a)



Viruses

Proteins involved in DNA, RNA, protein synthesis
Gene regulation
Cancer and control of cell proliferation
Transport of proteins and organelles inside cells
Infection and immunity
Possible gene therapy approaches

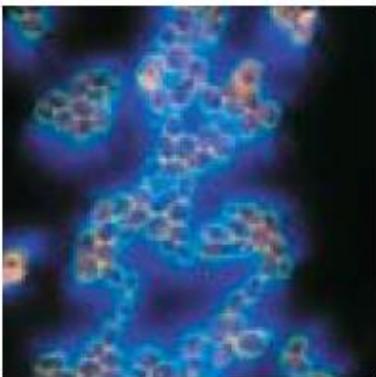
(b)



Bacteria

Proteins involved in DNA, RNA, protein synthesis, metabolism
Gene regulation
Targets for new antibiotics
Cell cycle
Signaling

(c)



Yeast (*Saccharomyces cerevisiae*)

Control of cell cycle and cell division
Protein secretion and membrane biogenesis
Function of the cytoskeleton
Cell differentiation
Aging
Gene regulation and chromosome structure

(d)



Roundworm (*Caenorhabditis elegans*)

Development of the body plan
Cell lineage
Formation and function of the nervous system
Control of programmed cell death
Cell proliferation and cancer genes
Aging
Behavior
Gene regulation and chromosome structure

Each experimental organism used in cell biology has advantages for certain types of studies. Viruses and bacteria have small genomes amenable to genetic dissection. Many insights into gene control initially came from studies with these organisms. The yeast *Saccharomyces cerevisiae* has the cellular organization of a eukaryote but is a relatively simple single-celled organism that is easy to grow and to manipulate genetically. In the nematode worm *Caenorhabditis elegans*, which has a small number of cells arranged in a nearly identical way in every worm, the formation of each individual cell can be traced. [Lodish et al., 2008](#)

(e)



Fruit fly (*Drosophila melanogaster*)

Development of the body plan
Generation of differentiated cell lineages
Formation of the nervous system, heart, and musculature
Programmed cell death
Genetic control of behavior
Cancer genes and control of cell proliferation
Control of cell polarization
Effects of drugs, alcohol, pesticides

(f)



Zebrafish

Development of vertebrate body tissues
Formation and function of brain and nervous system
Birth defects
Cancer

(g)



Mice, including cultured cells

Development of body tissues
Function of mammalian immune system
Formation and function of brain and nervous system
Models of cancers and other human diseases
Gene regulation and inheritance
Infectious disease

(h)

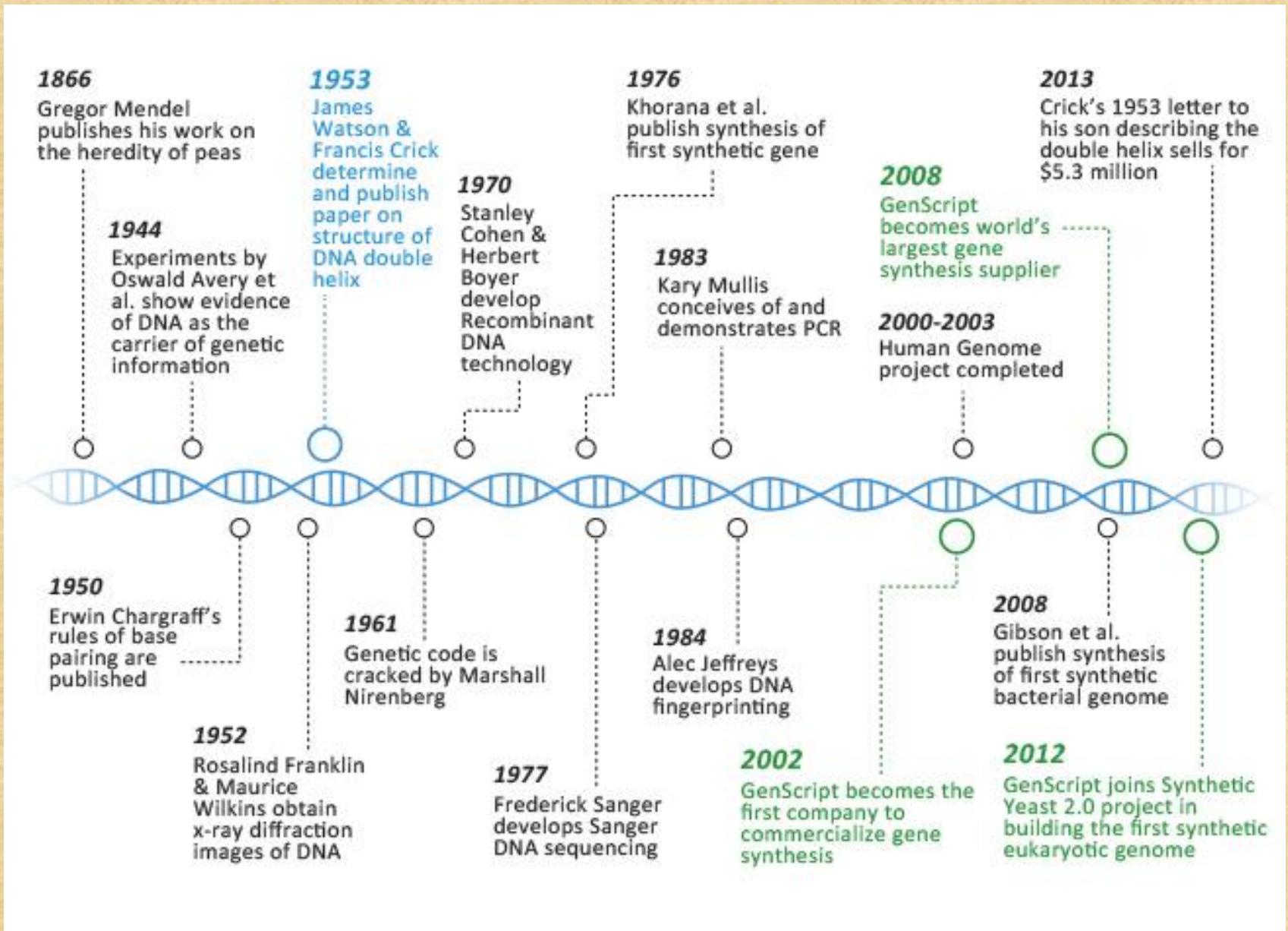


Plant (*Arabidopsis thaliana*)

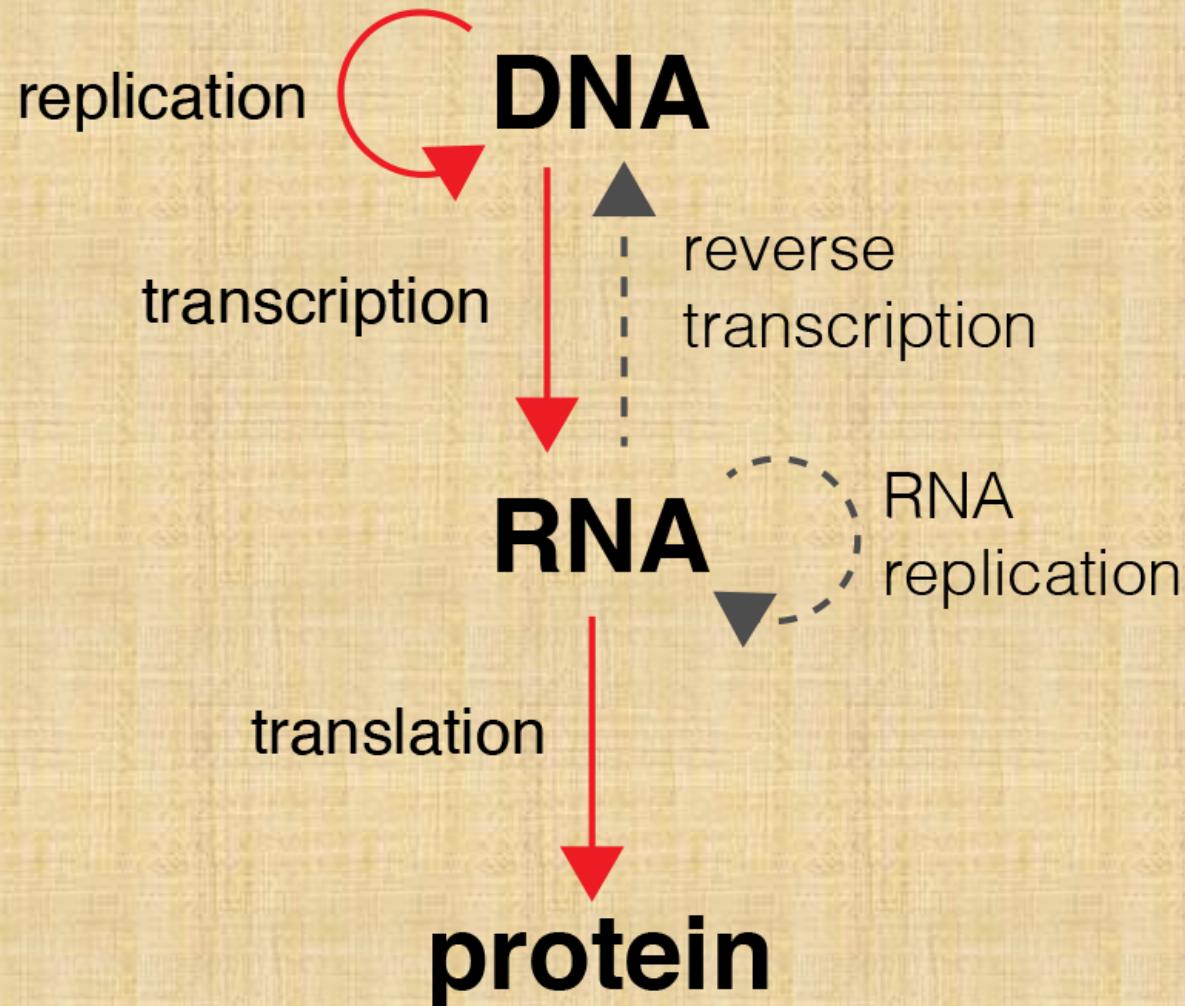
Development and patterning of tissues
Genetics of cell biology
Agricultural applications
Physiology
Gene regulation
Immunity
Infectious disease

The fruit fly *Drosophila melanogaster*, first used to discover the properties of chromosomes, has been especially valuable in identifying genes that control embryonic development. Many of these genes are evolutionarily conserved in humans. The zebrafish *Danio rerio* is used for rapid genetic screens to identify genes that control development and organogenesis. Of the experimental animal systems, mice (*Mus musculus*) are evolutionarily the closest to humans and have provided models for studying numerous human genetic and infectious diseases. The mustard-family weed *Arabidopsis thaliana*, sometimes described as the *Drosophila* of the plant kingdom, has been used for genetic screens to identify genes involved in nearly every aspect of plant life. Genome sequencing is completed for many viruses and bacterial species, the yeast *Saccharomyces cerevisiae*, the roundworm *C. elegans*, the fruit fly *D. melanogaster*, humans, and the plant *Arabidopsis thaliana*. It is mostly completed for mice and in progress for zebrafish. Other organisms, particularly frogs, sea urchins, chickens, and slime molds, continue to be immensely valuable for cell biology research. Increasingly, a wide variety of other species are used, especially for studies of evolution of cells and mechanisms.

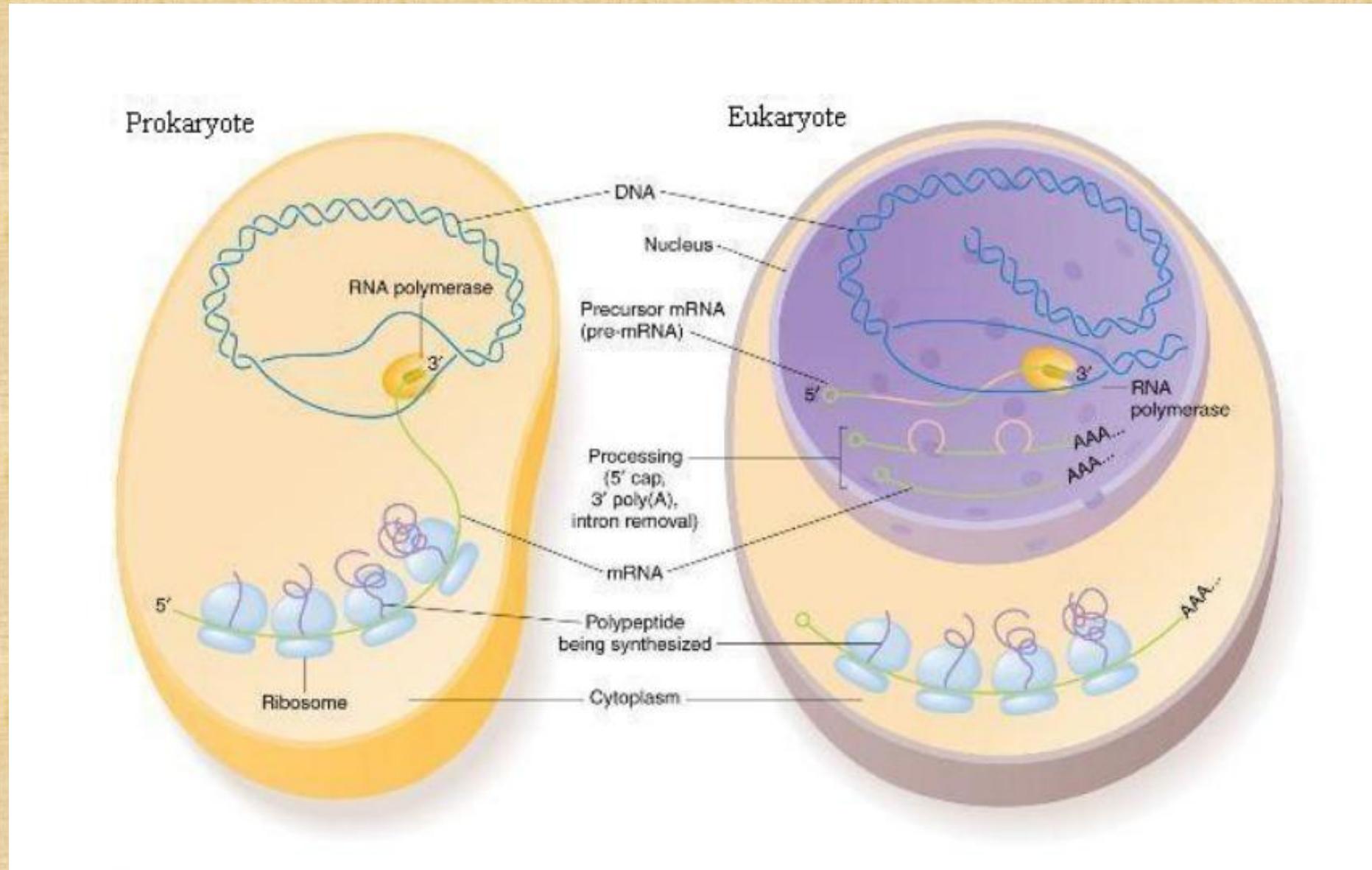
Lodish et al., 2008



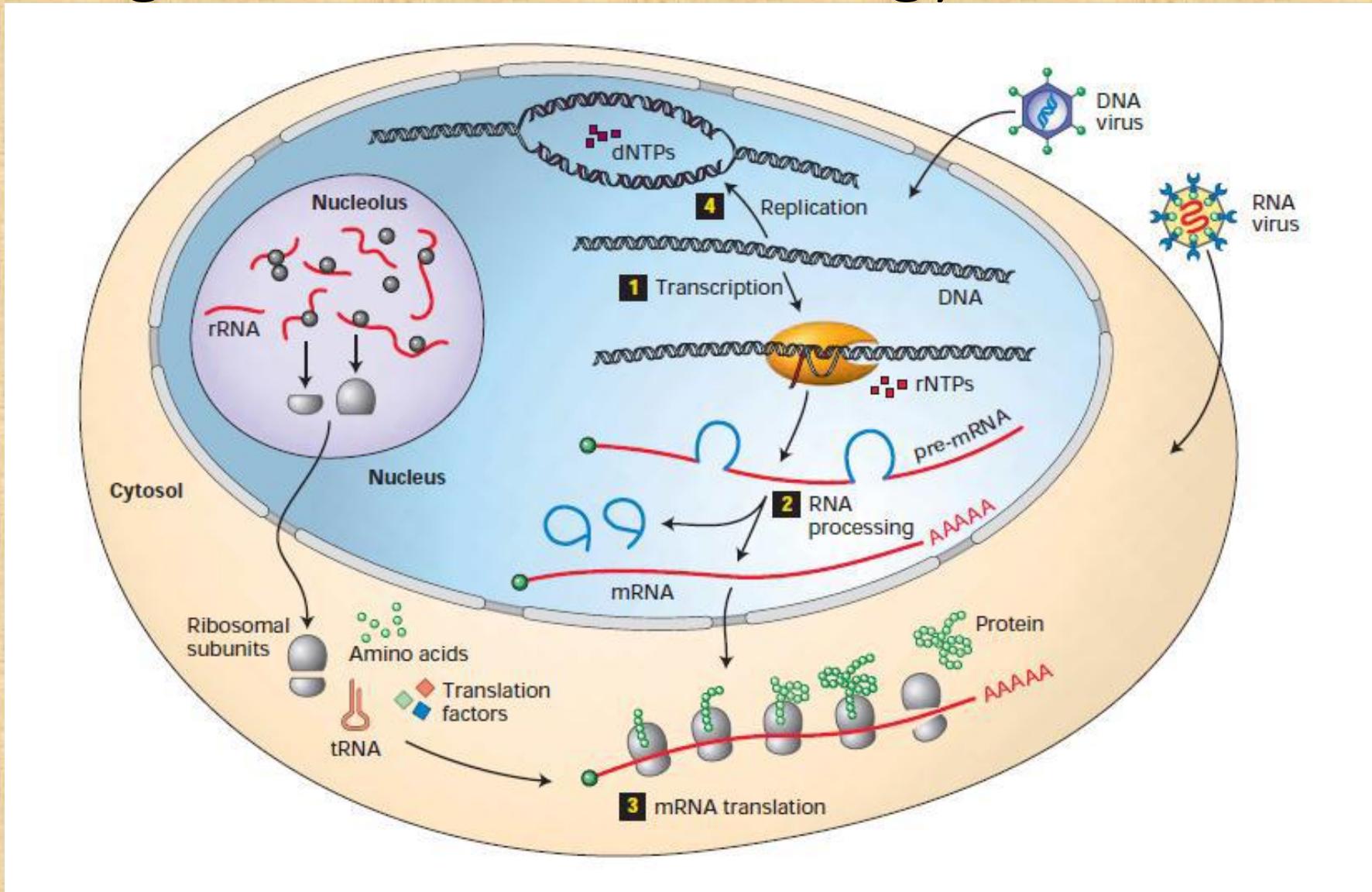
Central dogma of molecular biology



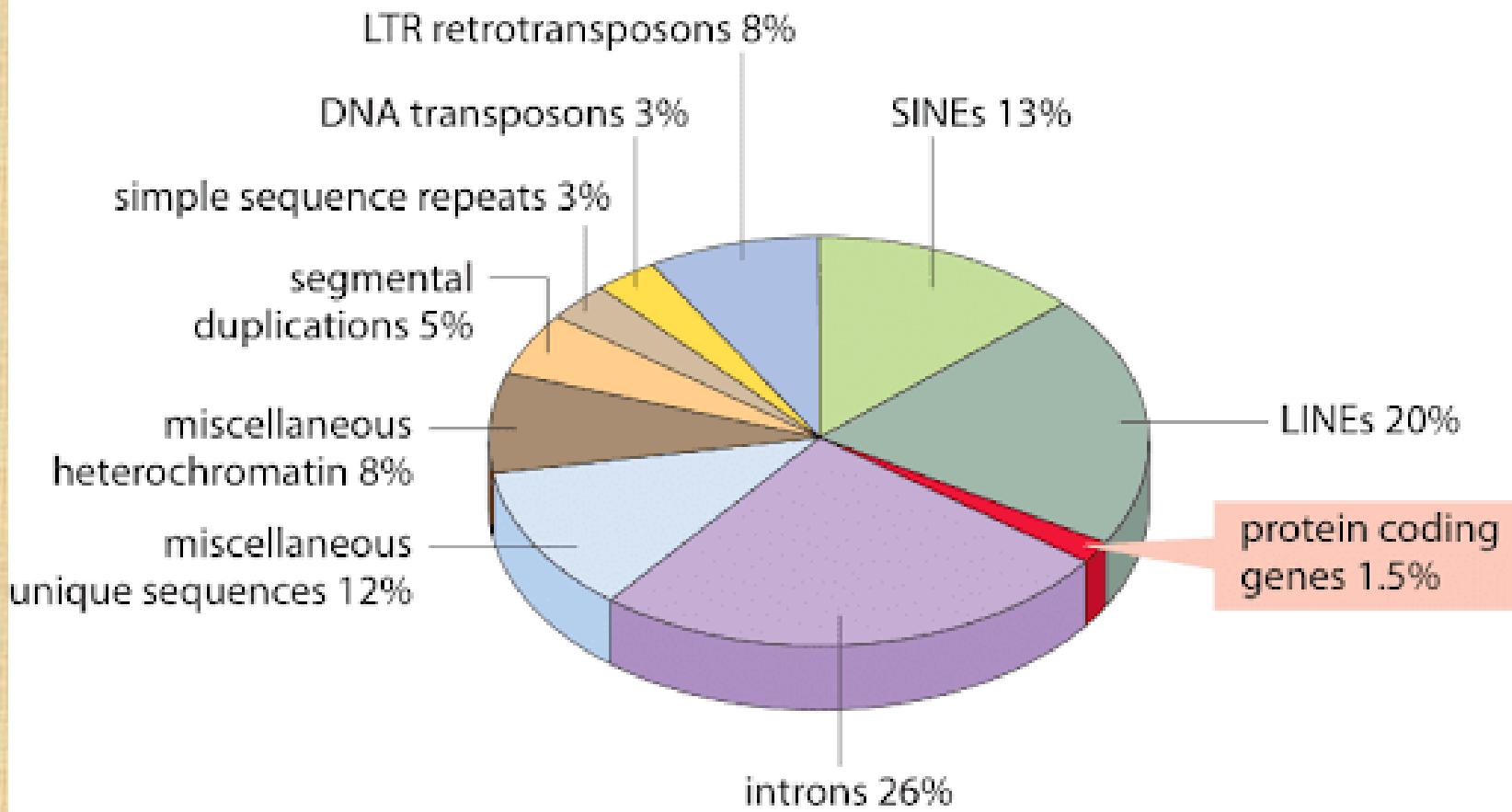
Prokaryotic and Eukaryotic Cells



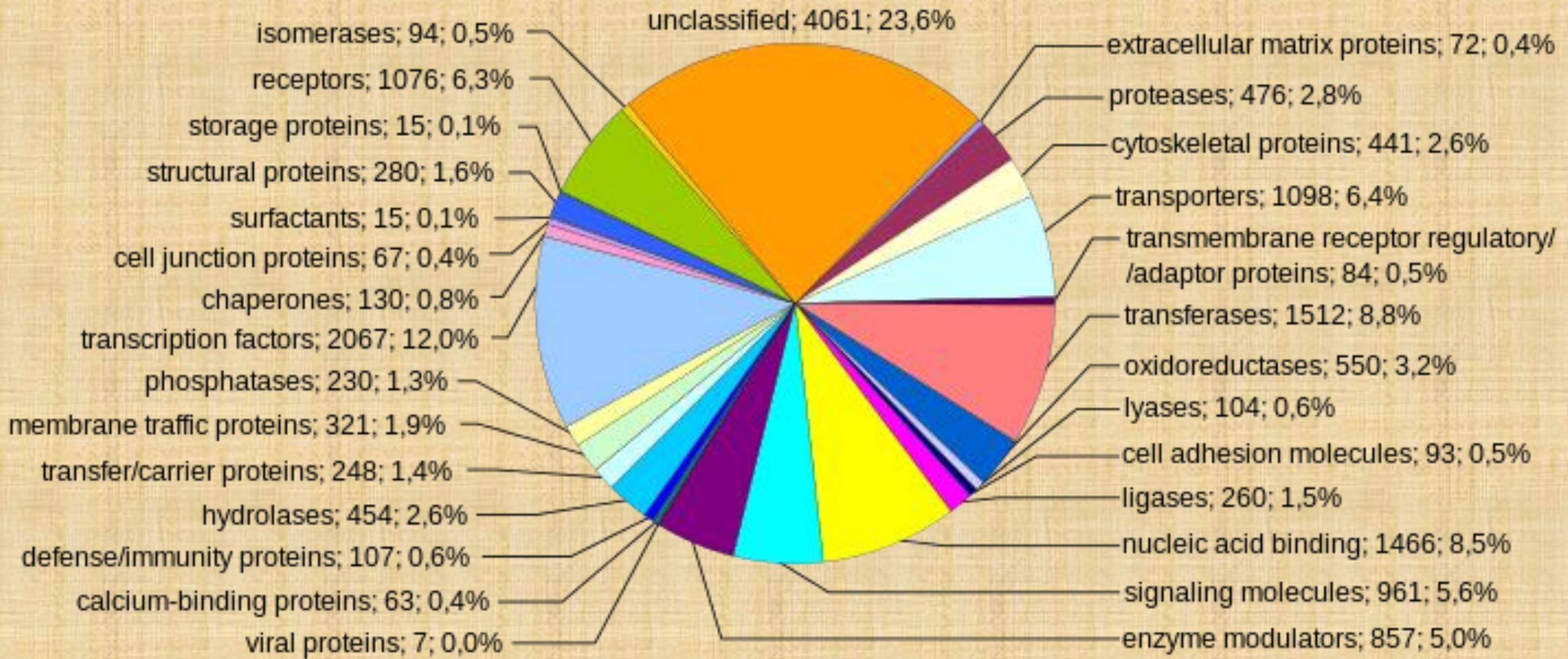
Central dogma of molecular biology



main components of the human genome



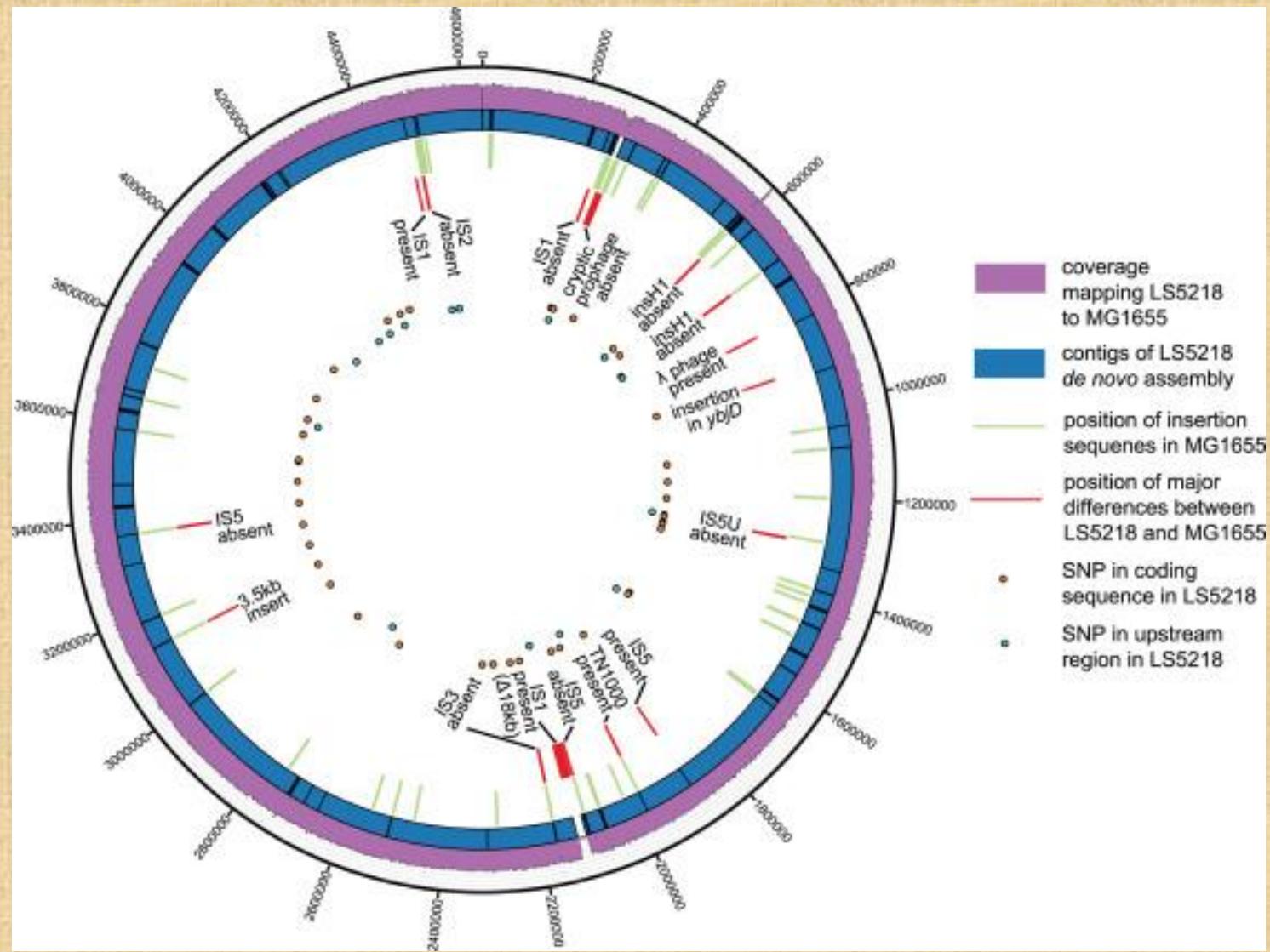
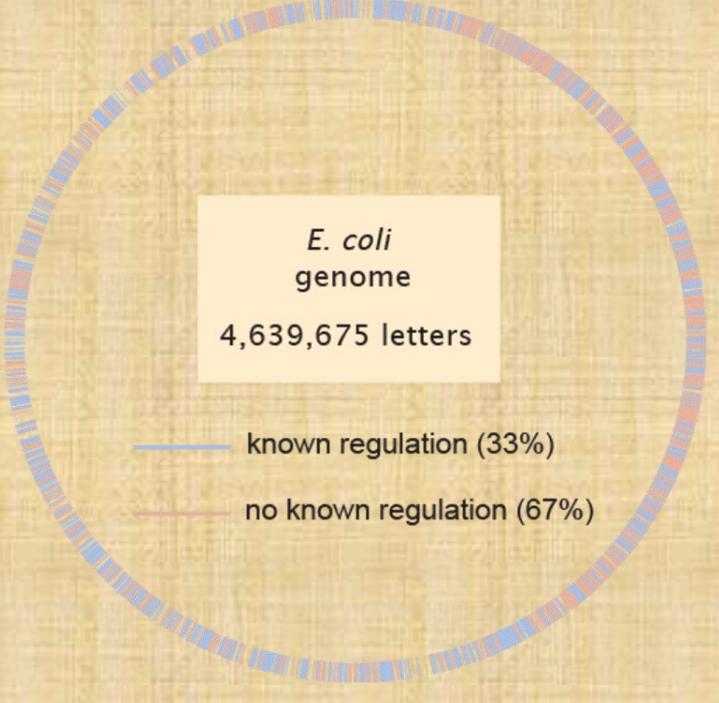
Short interspaced retrotransposable elements (SINEs), Long interspaced retrotransposable elements (LINEs)



Human genes categorized by function of the transcribed proteins, given both as number of encoding genes and percentage of all genes

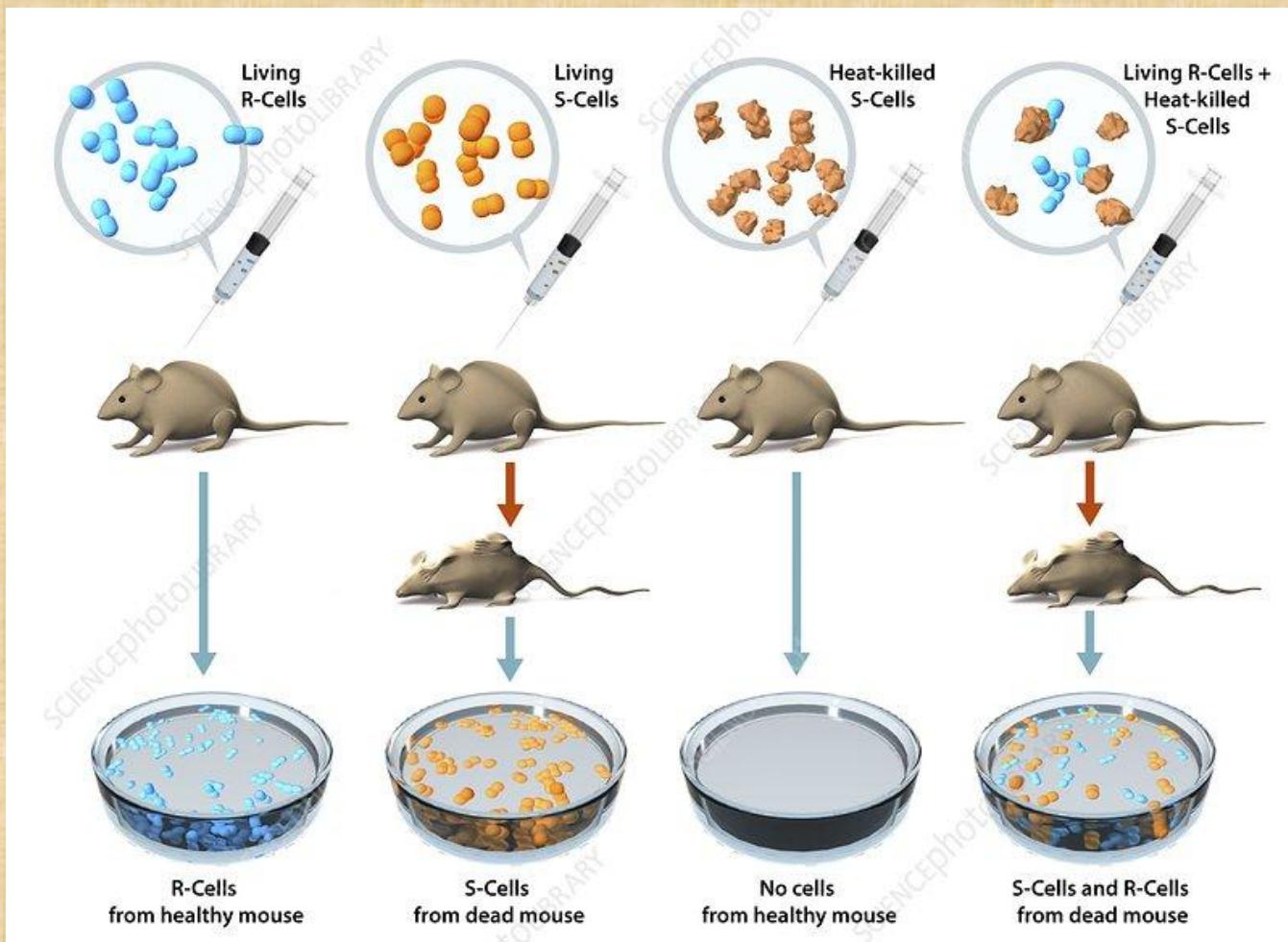
Examples of human protein-coding genes

Protein	Chrom	Gene	Length	Exons	Exon length	Intron length	Alt splicing
Breast cancer type 2 susceptibility protein	13	BRCA2	83,736	27	11,386	72,350	yes
Cystic fibrosis transmembrane conductance regulator	7	CFTR	202,881	27	4,440	198,441	yes
Cytochrome b	MT	MTCYB	1,140	1	1,140	0	no
Dystrophin	X	DMD	2,220,381	79	10,500	2,209,881	yes
Glyceraldehyde-3-phosphate dehydrogenase	12	GAPDH	4,444	9	1,425	3,019	yes
Hemoglobin beta subunit	11	HBB	1,605	3	626	979	no
Histone H1A	6	HIST1H1A	781	1	781	0	no
Titin	2	TTN	281,434	364	104,301	177,133	yes



Genome sequence and analysis of *Escherichia coli*

Griffith's experiment



Griffith's Experiment

In 1928, Frederick Griffith conducted one of the first experiments to show that cells possessed genetic material

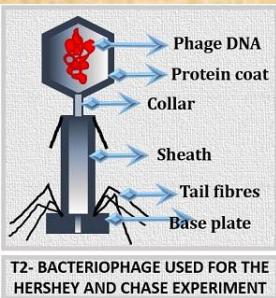
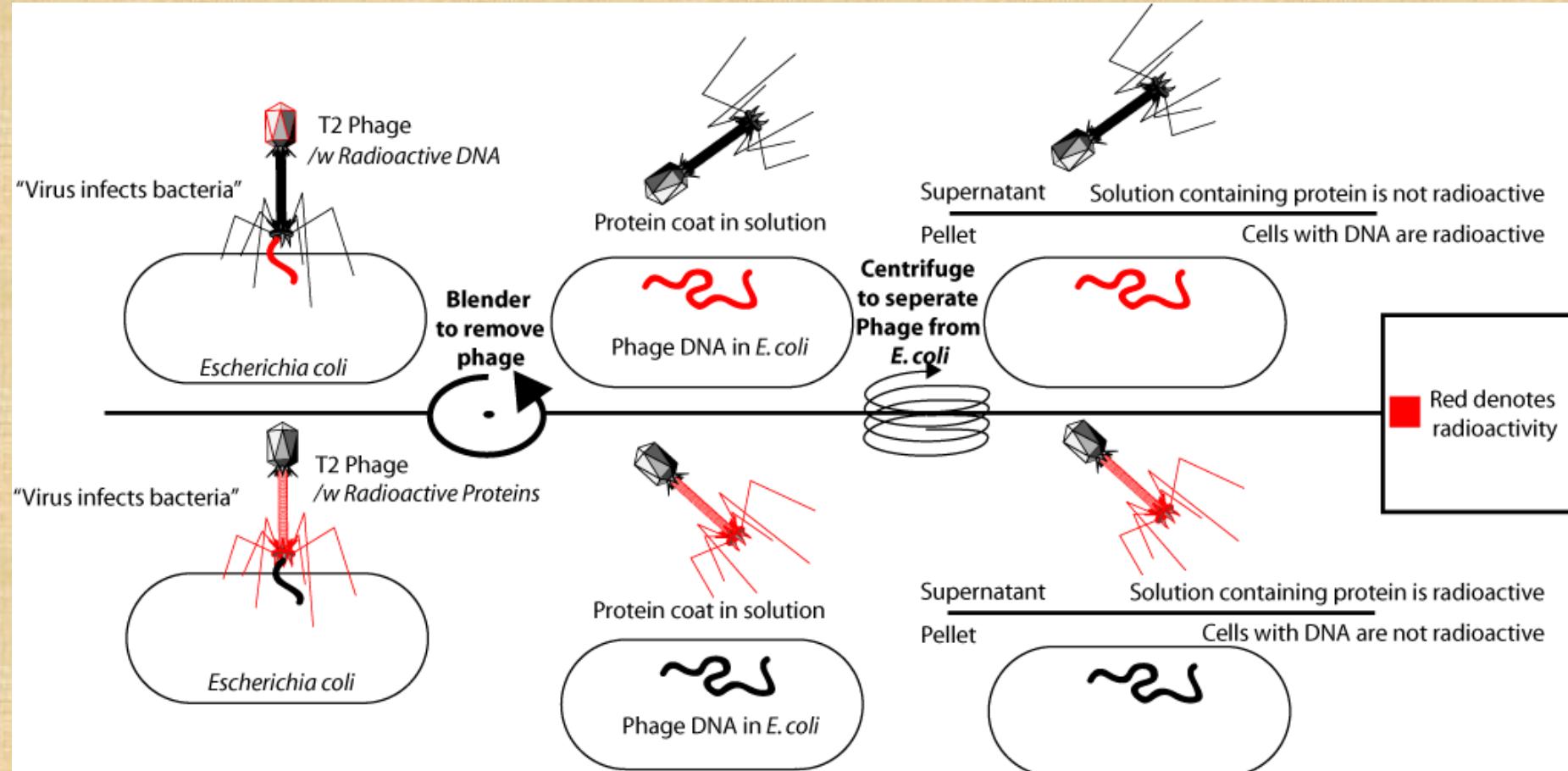
Griffith's experiment involved the use of two strains of pneumococcus – a deadly virulent strain (**S**) or a non-virulent strain (**R**)

- When Griffith infected mice with the non-virulent bacteria (strain **R**), the mice survived
- When Griffith infected mice with the virulent bacteria (strain **S**), the mice died
- When Griffith infected mice with heat-killed virulent bacteria (strain **R**), the mice survived as the bacteria had been killed
- When Griffith infected mice with a mix of heat-killed strain **S** and living strain **R**, the mice were found to have died

From this Griffith's concluded that the living **R** cells had somehow been transformed into virulent **S** cells

- This indicated that there was some form of transferrable genetic material present within the cells (i.e. DNA)

Hershey and Chase experiment

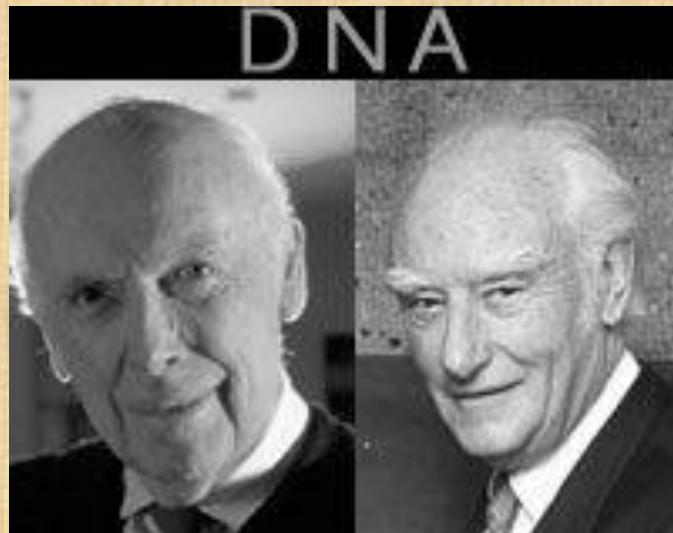


Hershey and Chase Experiment is a very popular experiment which provides evidence of **DNA** as “**Genetic material**”. It was introduced by the two scientists **A.D. Hershey** and **Martha Chase** in the year **1952**. After seven years of an experiment given by the Avery, Hershey and Chase gave the further proof of DNA as genetic material by the use of **radioactive bacteriophage**.

DNA: Franklin, Crick & Watson
1953



Structure of DNA

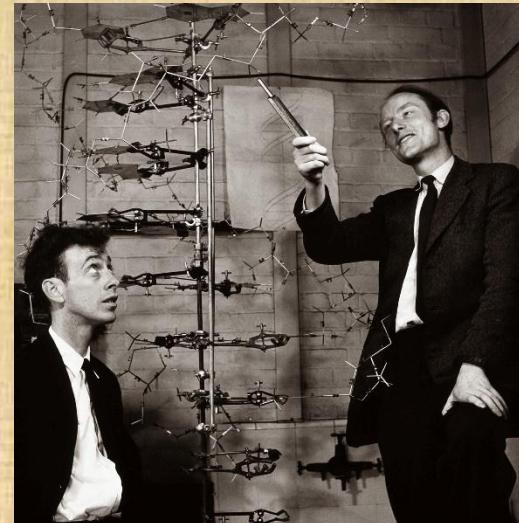


James Watson

Francis Crick



Maurice Wilkins Rosalind Franklin



Watson & Crick



Double helix structure
of DNA (1953)

X-ray diffraction pattern
from B form of DNA

Structure of DNA

No. 4286 April 25, 1953 NATURE 737

equipment, and to Dr. G. E. R. Deacon and the captain and officers of R.R.S. *Discovery II* for their part in making the observations.

¹ Young, B., Gerash, E., and Jevons, W., *Phil. Mag.*, **48**, 149 (1943).

² Luria, S. E., *Nature*, **153**, 534 (1944).

³ Cox, A. S., Woods Hole Papers in Phys. Curing, Metab., **13**, 10 (1940).

⁴ Wilkins, V. W., *Arch. Med. Austral. Pathol.* (Melbourne), **2** (1948).

MOLECULAR STRUCTURE OF NUCLEIC ACIDS

A Structure for Deoxyribose Nucleic Acid

We wish to suggest a structure for the salt of deoxyribonucleic acid (D.N.A.). This structure has novel features which are of considerable biological interest.

The novel feature of the structure is the manner in which the two chains are held together by the purine or pyrimidine bases. The planes of the bases are perpendicular to the fibre axis. They are joined together in pairs, a single base from one chain being hydrogen-bonded to a single base from the other chain, so that the two are side by side with identical purine rings facing each other. One base pair consists of a purine and the other a pyrimidine for bonding to a purine. The hydrogen bonds are made as follows: purine position 1 to pyrimidine position 1; purine position 6 to pyrimidine position 6.

If it is assumed that the bases can only occur in the structure in the planar purinoid tautomer form (that is, with the keto rather than the enol configuration) it is found that only specific pairs of bases can bond together. These pairs are: adenine (purine) with thymine (pyrimidine), and guanine (purine) with cytosine (pyrimidine).

In other words, if an adenine forms one member of a pair, on either chain, then all those assumptions about the other member must be thymine; similarly for guanine and cytosine. The sequence of bases on a single chain is therefore completely determined by the sequence of bases on the other chain. However, if only specific pairs of bases can be formed, it follows that if the sequence of bases on one chain is given, then the sequence on the other chain is also completely determined.

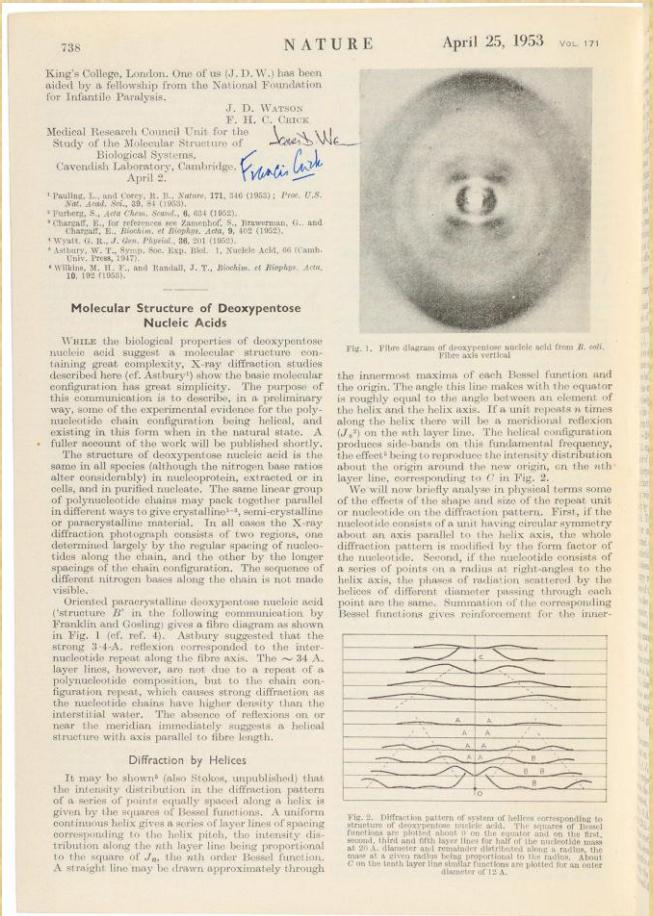
It has been found experimentally¹⁻⁴ that the ratio of the amounts of adenine to thymine, and the ratio of guanine to cytosine, are always very close to unity for deoxyribonucleic acid.

It is probably impossible to build this structure with the bases in place in the deoxyribose, as the extra oxygen atom would make too close a van der Waals contact.

The previously published X-ray data⁵ on deoxyribonucleic acid are insufficient for a rigorous test of our structure, but we note that each chain consists of phosphate diester groups joining β -D-deoxyribofuranose residues with $3\text{-}5'$ linkages. The two chains (but not the bases) are linked by hydrogen bonds perpendicular to the fibre axis. Both chains follow right-handed helices, but owing to the dyad the sequences of the atoms in the two chains run in opposite directions. Each chain loosely resembles Furberg's model No. 1; that is, the bases are on the inside of the helix, and the sugar and phosphate groups on the outside. The configuration of the sugar and the atoms near it is close to Furberg's "standard configuration", the sugar being roughly perpendicular to the attached base. There

This figure is purely diagrammatic. The two chains are represented by two phosphate-ribose chains, each consisting of two phosphate groups and two ribose groups. The vertical line marks the fibre axis.

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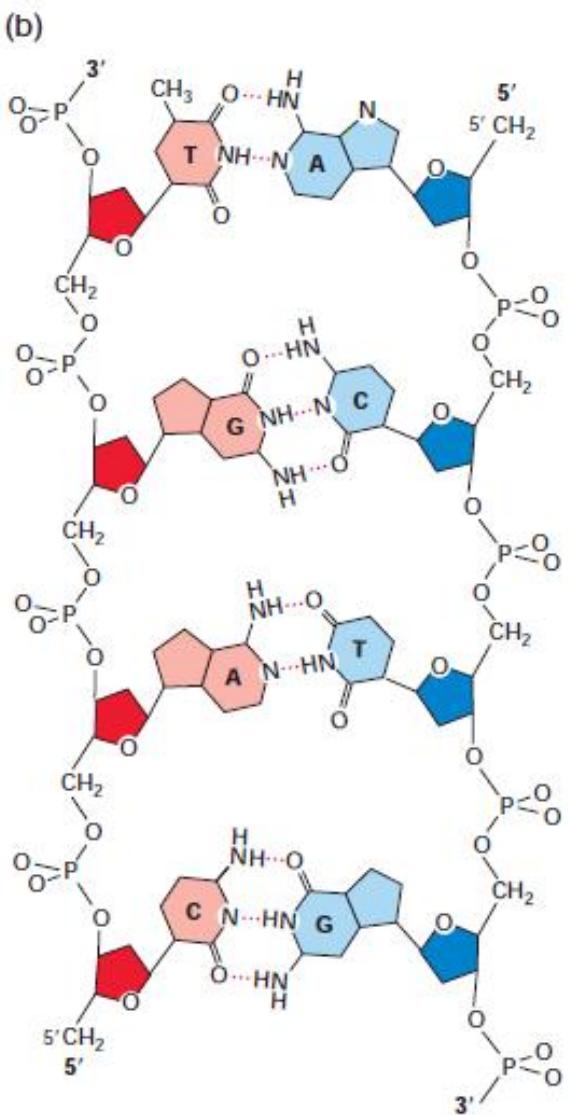
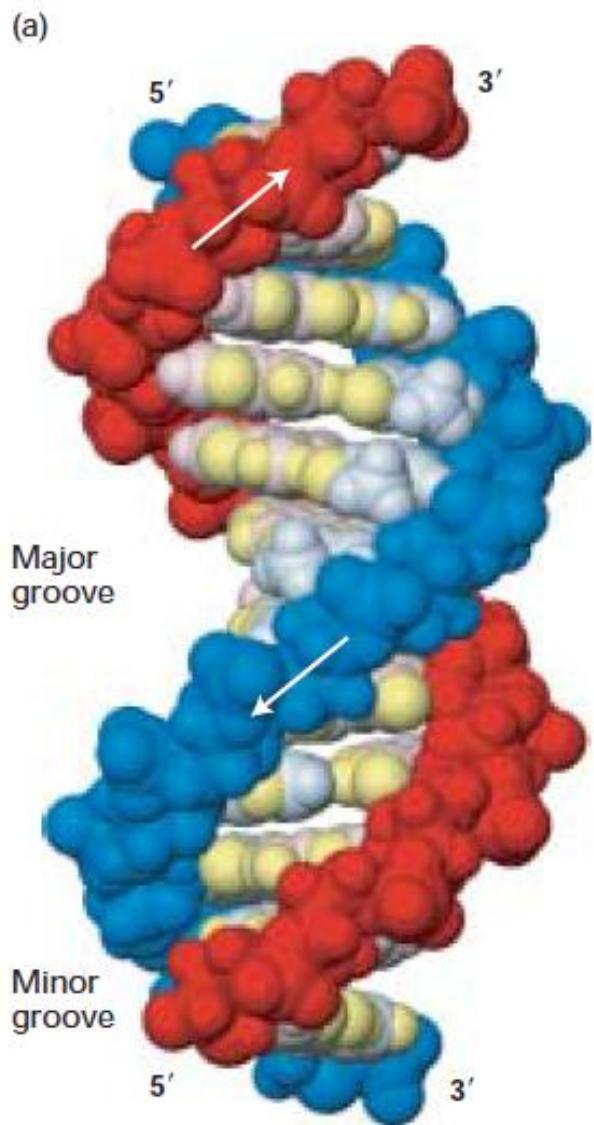


Watson and Crick (1953) developed a DNA double helix model based on Franklin's X-ray crystallography analysis and Erwin Chargaff's experiment (ratio of A and t are 1:1, and G and C are 1:1).

Watson and Crick model of DNA

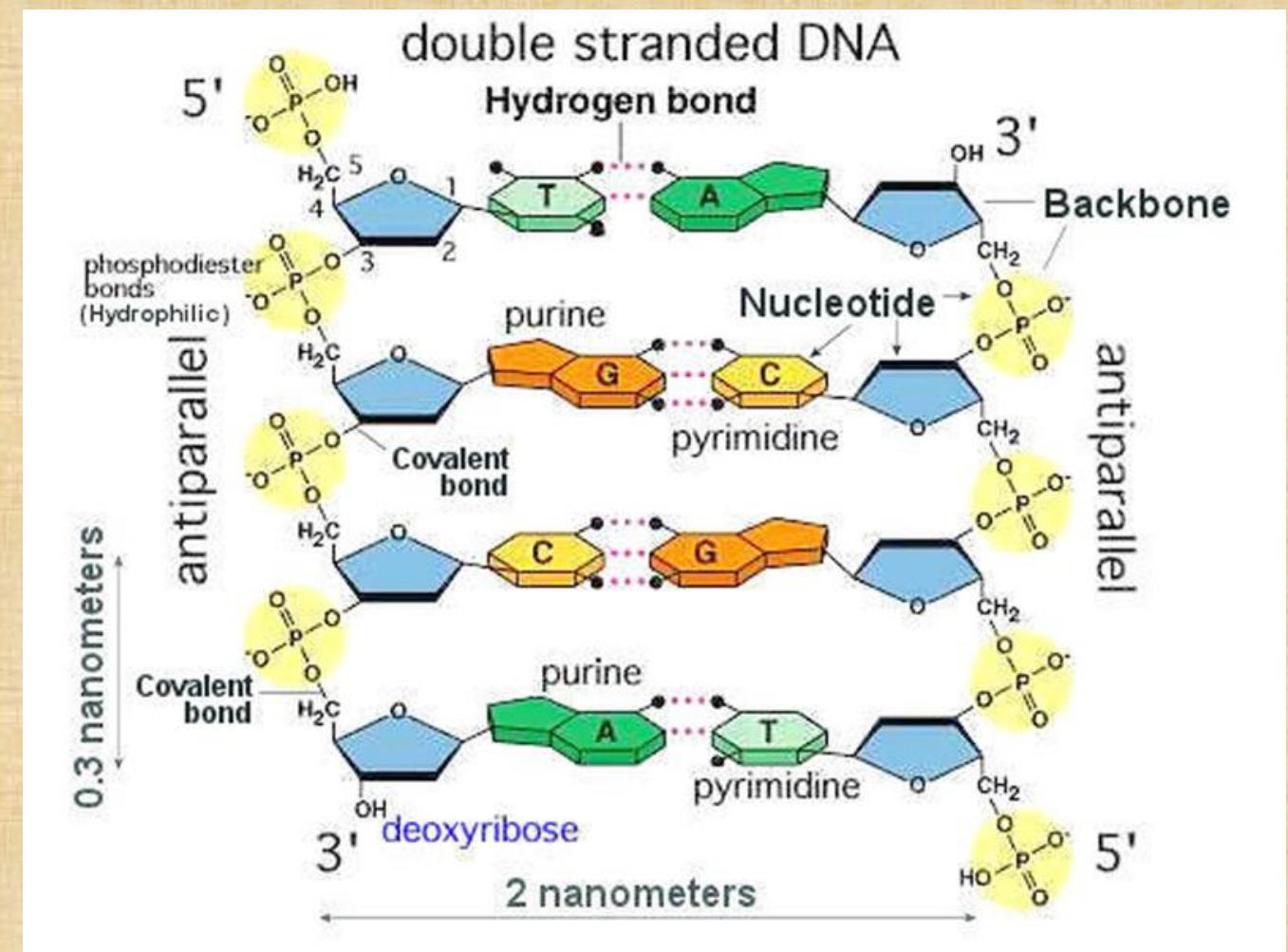
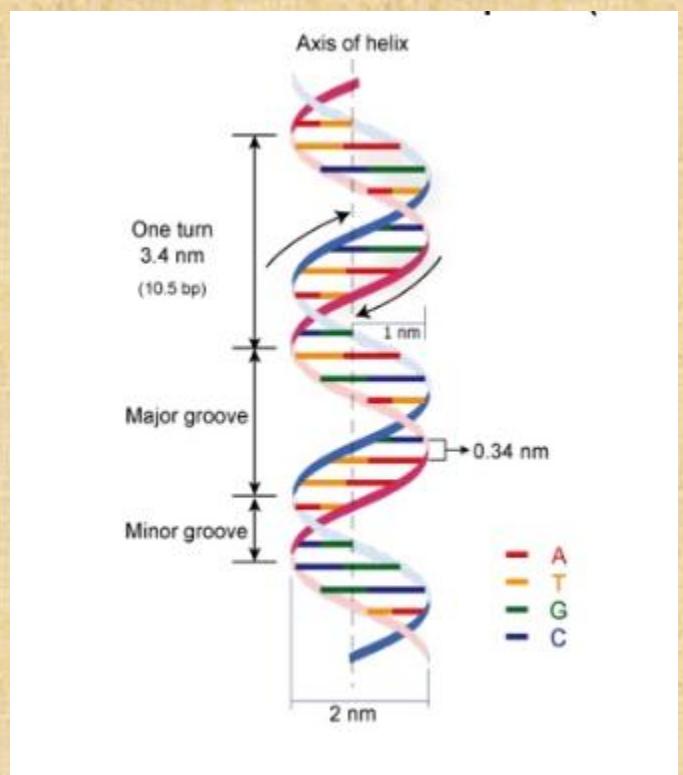
Some Basic

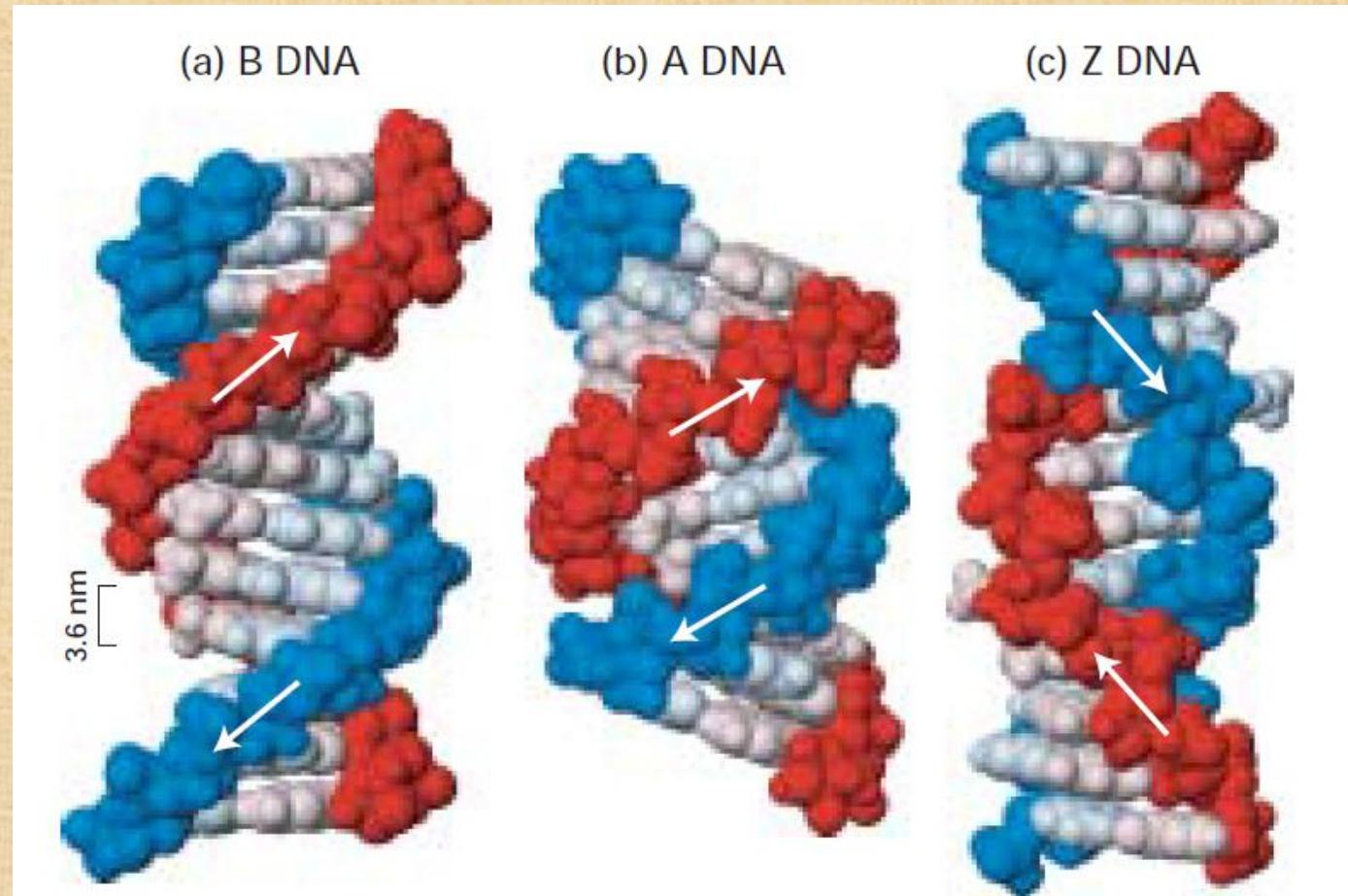
- Nucleoside- is a compound formed by the combination of a pentose sugar and nitrogen base.
- Nucleotide-is a compound formed by the combination of nucleoside and phosphate group.
- Nucleotides building blocks of nucleic acids.
- Nucleotide have three characteristic components.
 1. **A nitrogenous base**
 2. **A pentose sugar**
 3. **A phosphate group**



The DNA double helix. (a) Space-filling model of B DNA, the most common form of DNA in cells. The bases (light shades) project inward from the sugar-phosphate backbones (dark red and blue) of each strand, but their edges are accessible through major and minor grooves. Arrows indicate the 5'n3'direction of each strand. Hydrogen bonds between the bases are in the center of the structure. The major and minor grooves are lined by potential hydrogen bond donors and acceptors (highlighted in yellow). (b) Chemical structure of DNA double helix. This extended schematic shows the two sugar-phosphate backbones and hydrogen bonding between the Watson-Crick base pairs, AT and GC. [Part (a) from R. Wing et al., 1980, *Nature* **287**:755; part (b) from R. E. Dickerson, 1983, *Sci. Am.* **249**:94.]

Structure of DNA

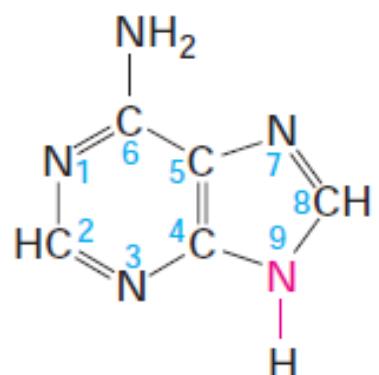




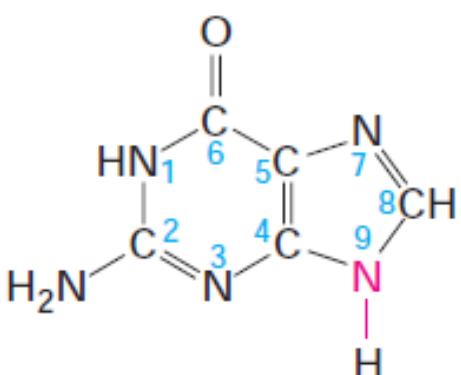
Models of various known DNA structures. The sugar-phosphate backbones of the two strands, which are on the outside in all structures, are shown in red and blue; the bases (lighter shades) are oriented inward. (a) The B form of DNA has \approx 10.5 base pairs per helical turn. Adjacent stacked base pairs are 0.36 nm apart. (b) The more compact A form of DNA has 11 base pairs per turn and exhibits a large tilt of the base pairs with respect to the helix axis. (c) Z DNA is a left-handed double helix.

<u>Property</u>	<u>B-DNA</u>	<u>A-DNA</u>	<u>Z-DNA</u>
Strand	Antiparallel	Antiparallel	Antiparallel
Type of Helix	Right-handed	Right-handed	Left-handed
Overall shape	Long and narrow	Short and wide	Elongated and narrow
Base pair per turn	10	11	12
Distance between adjacent bases	0.34 nm	0.23 nm	0.38 nm
Pitch/turn of helix	3.40 nm	2.82 nm	4.56 nm
Helical Diameter	2.0 nm	2.3 nm	1.8 nm
Tilt/inclination of bp to axis	10°	20°	90°

PURINES

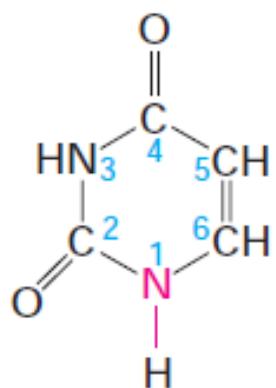


Adenine (A)

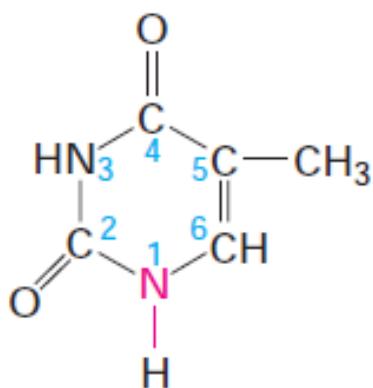


Guanine (G)

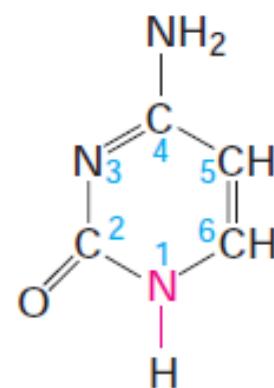
PYRIMIDINES



Uracil (U)

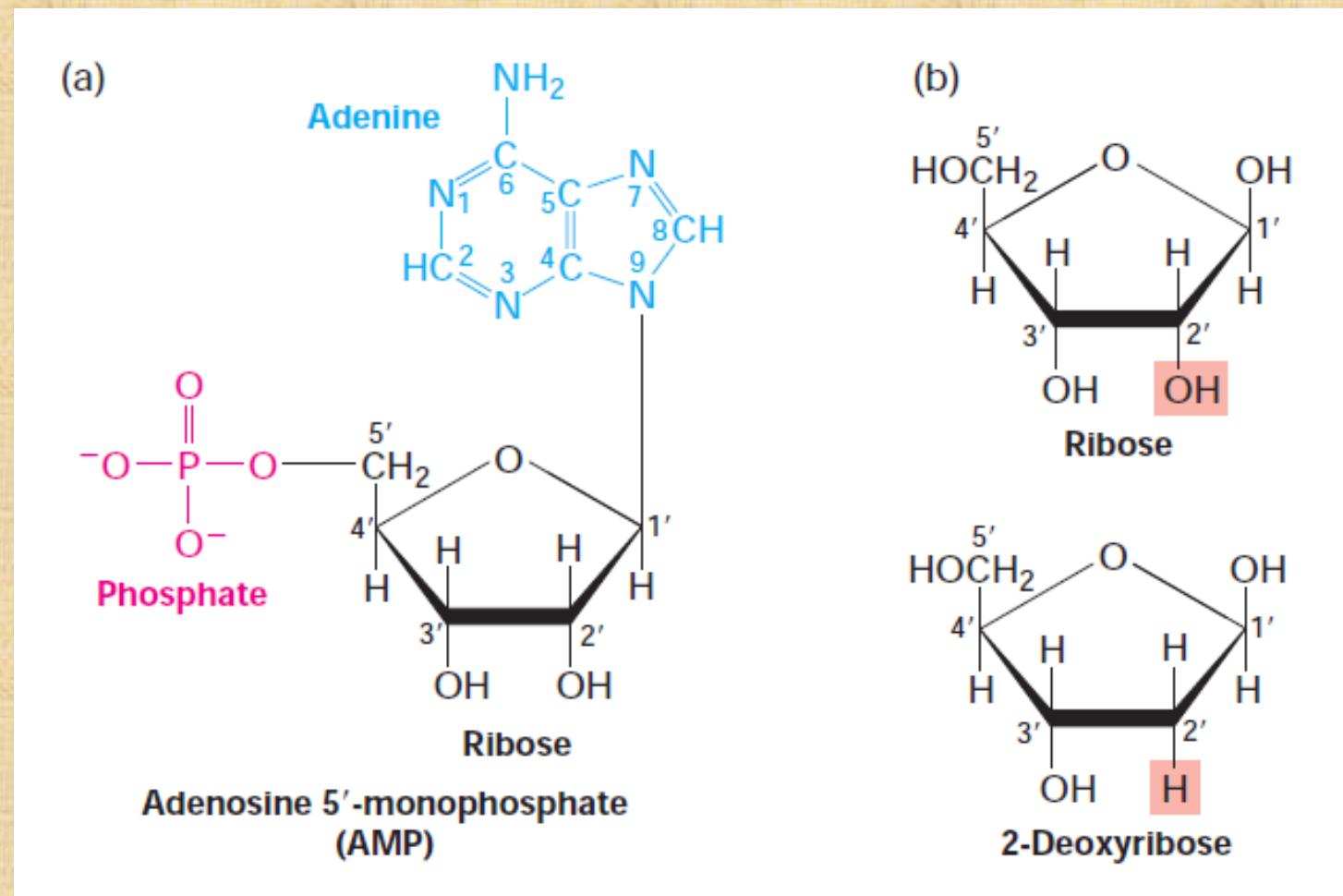


Thymine (T)

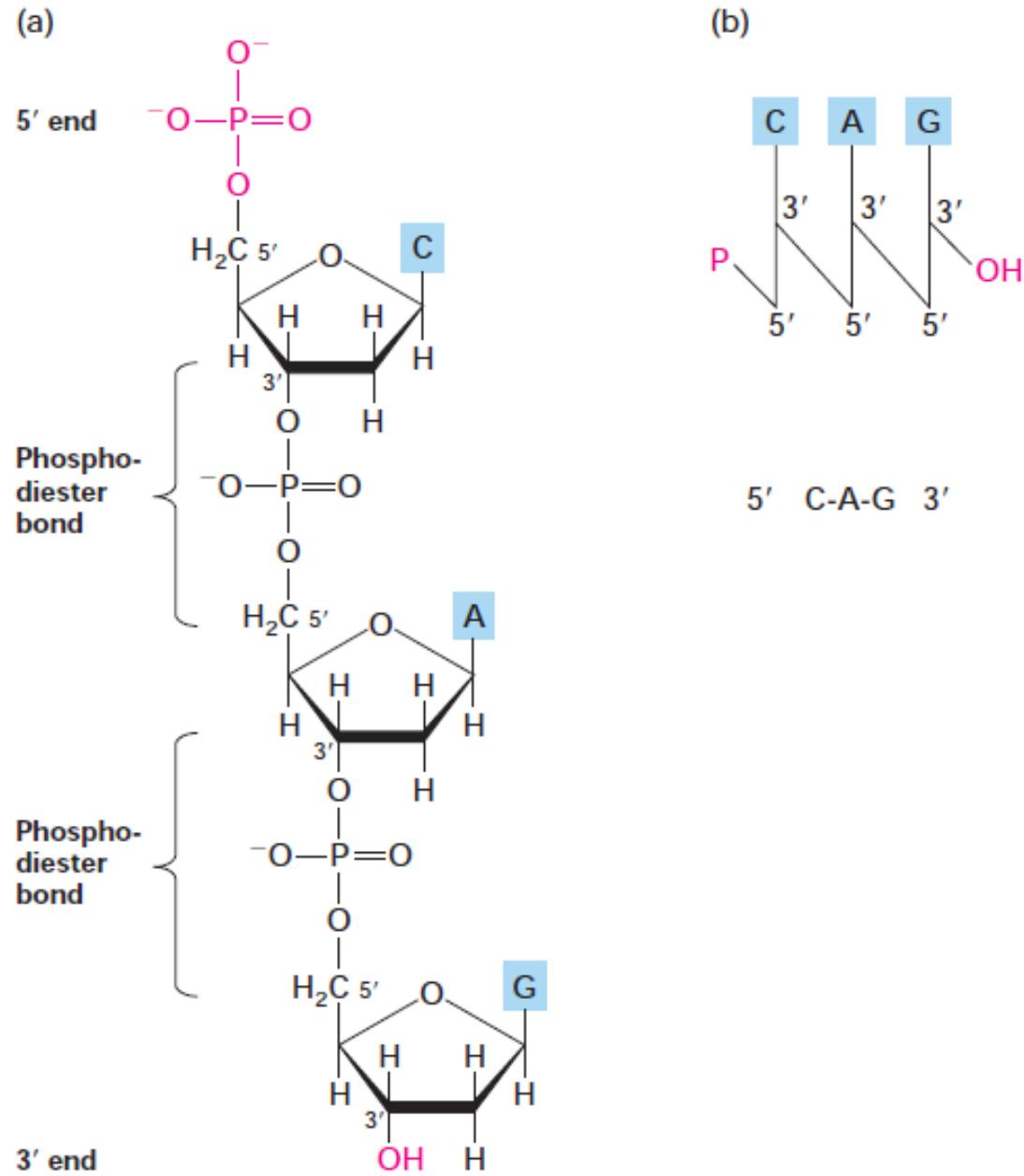


Cytosine (C)

Chemical structures of the principal bases in nucleic acids. In nucleic acids and nucleotides, nitrogen 9 of purines and nitrogen 1 of pyrimidines (red) are bonded to the 1' carbon of ribose or deoxyribose. U is only in RNA, and T is only in DNA. Both RNA and DNA contain A, G, and C.



Common structure of nucleotides. (a) Adenosine 5'-monophosphate (AMP), a nucleotide present in RNA. By convention, the carbon atoms of the pentose sugar in nucleotides are numbered with primes. In natural nucleotides, the 1' carbon is joined by a β -linkage to the base (in this case adenine); both the base (blue) and the phosphate on the 5'hydroxyl (red) extend above the plane of the furanose ring. (b) Ribose and deoxyribose, the pentoses in RNA and DNA, respectively.



Alternative representations of a nucleic acid strand illustrating its chemical directionality. Shown here is a single strand of DNA containing only three bases: cytosine (C), adenine (A), and guanine (G). (a) The chemical structure shows a hydroxyl group at the 3' end and a phosphate group at the 5' end. Note also that two phosphodiester bonds link adjacent nucleotides; this two-bond linkage commonly is referred to as a *phosphodiester bond*. (b) In the “stick” diagram (top), the sugars are indicated as vertical lines and the phosphodiester bonds as slanting lines; the bases are denoted by their single-letter abbreviations. In the simplest representation (bottom), only the bases are indicated. By convention, a polynucleotide sequence is always written in the 5' → 3' direction (left to right) unless otherwise indicated.

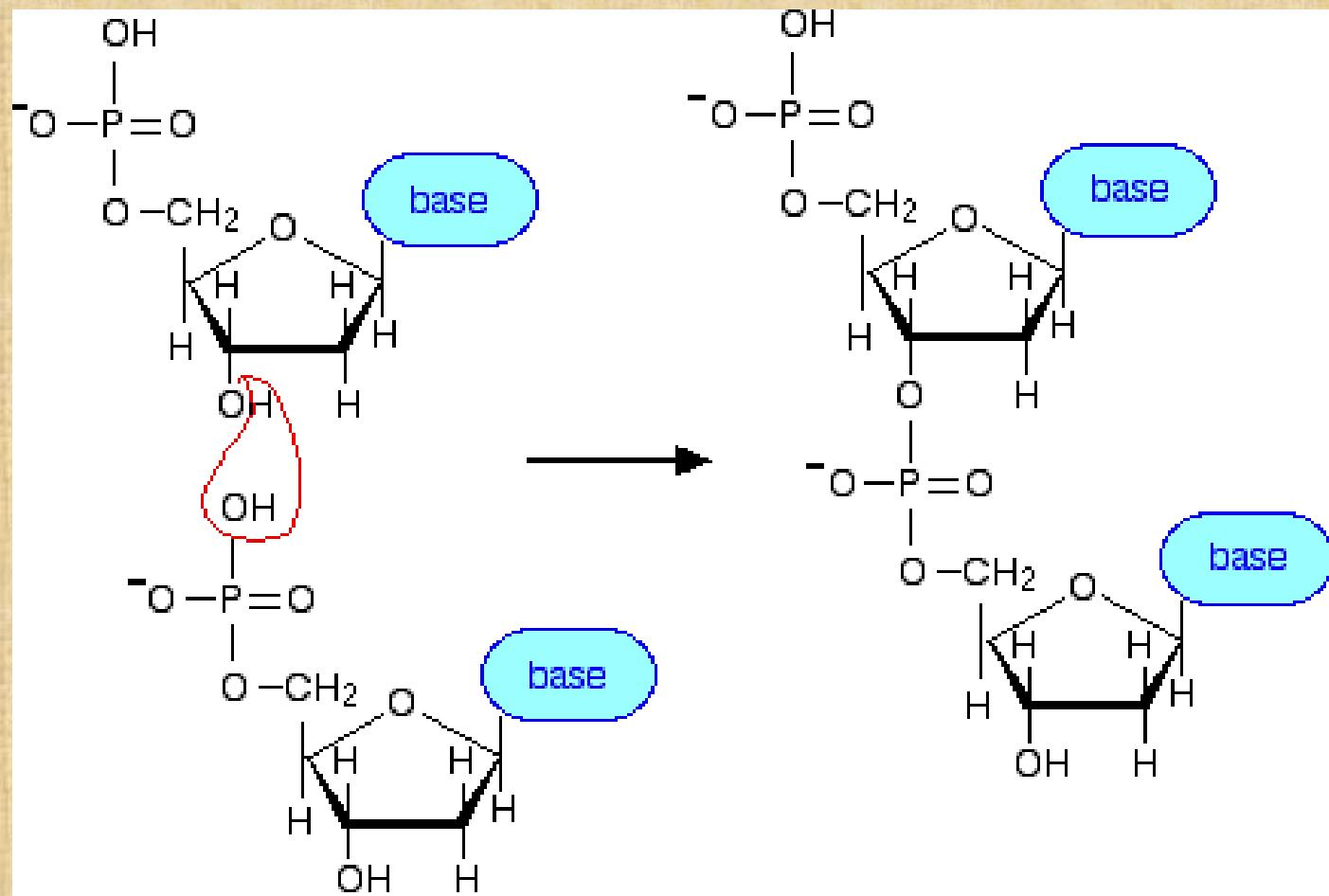


TABLE 2-2

Terminology of Nucleosides and Nucleotides

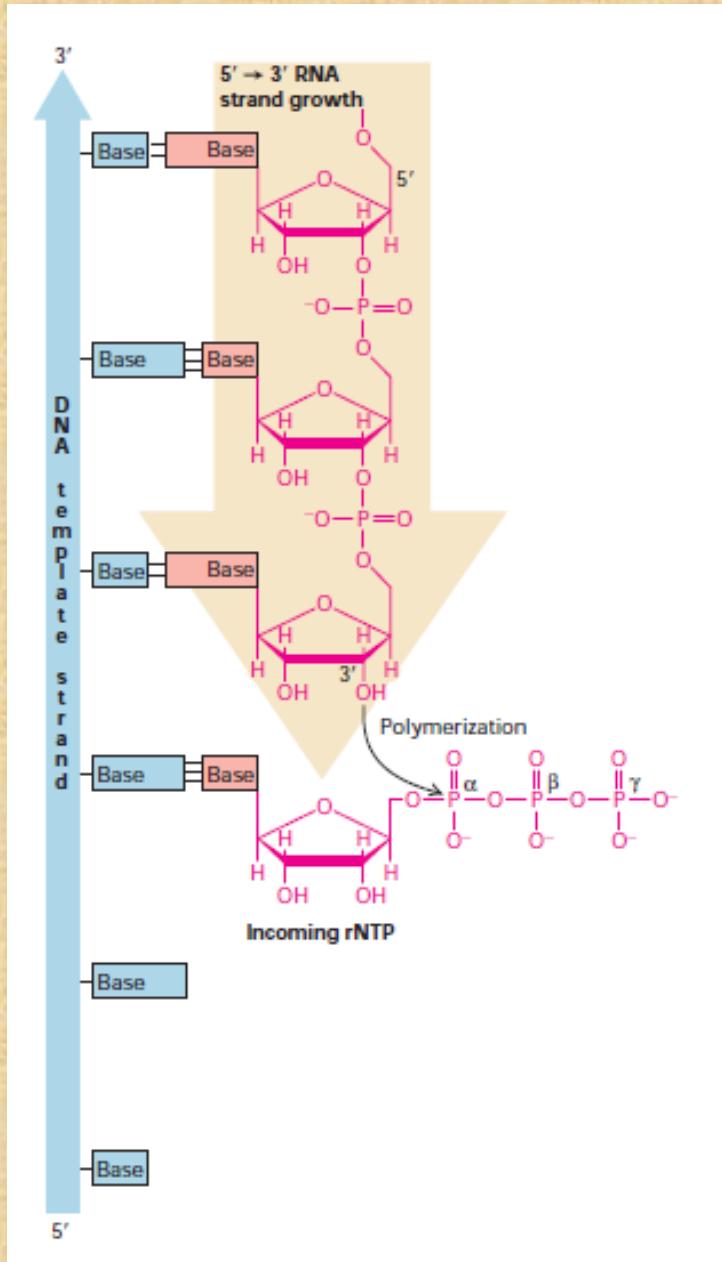
		Bases			
		Purines		Pyrimidines	
		Adenine (A)	Guanine (G)	Cytosine (C)	Uracil (U) Thymine [T]
Nucleosides	{ in RNA	Adenosine	Guanosine	Cytidine	Uridine
	{ in DNA	Deoxyadenosine	Deoxyguanosine	Deoxycytidine	Deoxythymidine
Nucleotides	{ in RNA	Adenylate	Guanylate	Cytidylate	Uridylate
	{ in DNA	Deoxyadenylate	Deoxyguanylate	Deoxycytidylate	Deoxythymidylate
Nucleoside monophosphates		AMP	GMP	CMP	UMP
Nucleoside diphosphates		ADP	GDP	CDP	UDP
Nucleoside triphosphates		ATP	GTP	CTP	UTP
Deoxynucleoside mono-, di-, and triphosphates		dAMP, etc.			

TABLE 4-2 Known Deviations from the Universal Genetic Code

Codon	Universal Code	Unusual Code*	Occurrence
UGA	Stop	Trp	<i>Mycoplasma, Spiroplasma</i> , mitochondria of many species
CUG	Leu	Thr	Mitochondria in yeasts
UAA, UAG	Stop	Gln	<i>Acetabularia, Tetrahymena, Paramecium</i> , etc.
UGA	Stop	Cys	<i>Euplotes</i>

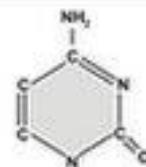
* “Unusual code” is used in nuclear genes of the listed organisms and in mitochondrial genes as indicated.

SOURCE: S. Osawa et al., 1992, *Microbiol. Rev.* 56:229.



Polymerization of ribonucleotides by RNA polymerase during transcription. The ribonucleotide to be added at the 3' end of a growing RNA strand is specified by base pairing between the next base in the template DNA strand and the complementary incoming ribonucleoside triphosphate (rNTP). A phosphodiester bond is formed when RNA polymerase catalyzes a reaction between the 3' O of the growing strand and the phosphate of a correctly base-paired rNTP. RNA strands always are synthesized in the 5'→3' direction and are opposite in polarity to their template DNA strands.

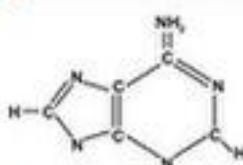
Cytosine



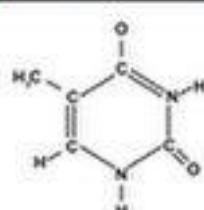
Guanine



Adenine

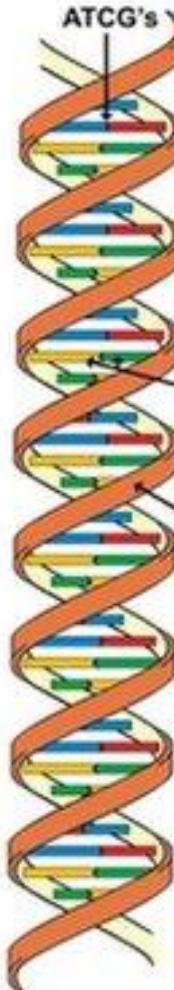


Thymine



Nitrogenous
Bases

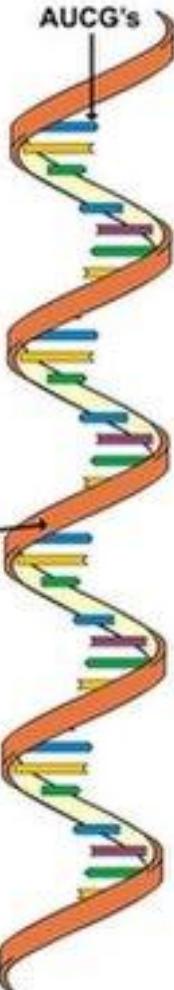
ATCG's



DNA

Deoxyribonucleic Acid

AUCG's



RNA

Ribonucleic Acid

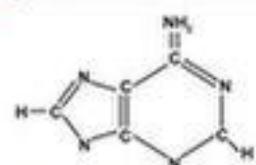
Cytosine



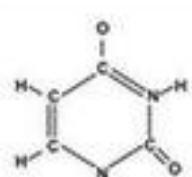
Guanine



Adenine



Uracil

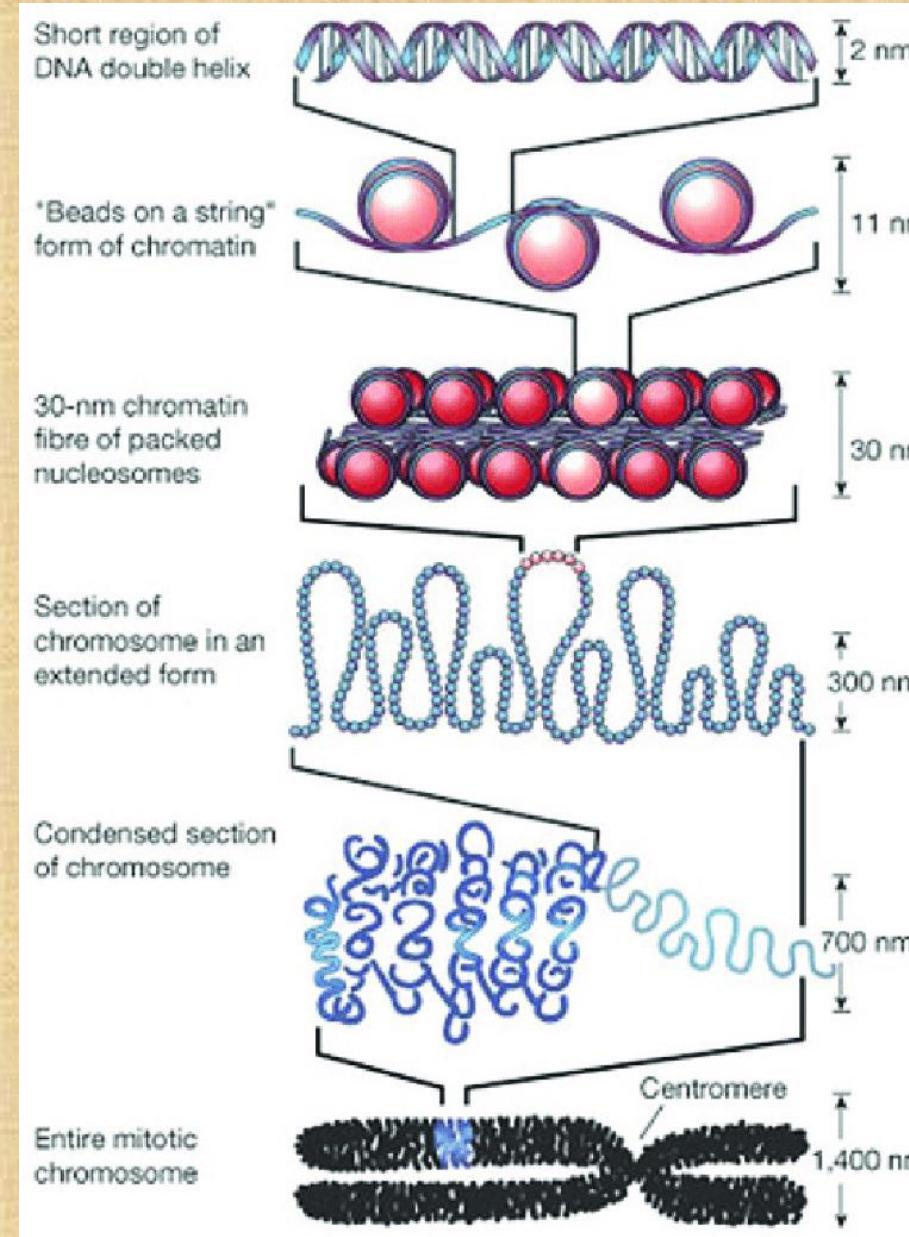


Replaces Thymine in RNA.

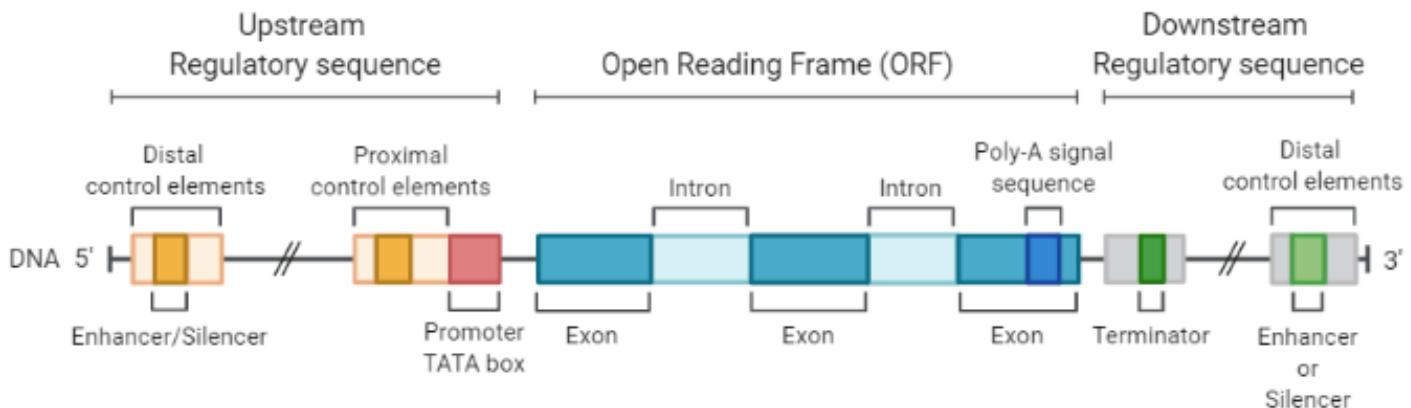
Nitrogenous
Bases

Sl.N o.	Organism	Chromosome Number	Genome Size (MB)	Number of genes
1	<i>Escherichia coli</i>	1	4.6	4288
2	<i>Saccharomyces cerevisiae</i>	16	13	6034
3	<i>Drosophila melanogaster</i>	8	180	13600
4	<i>Arabidopsis thaliana</i>	10	125	26000
5	<i>Oryza sativa</i>	24	390	38000
6	<i>Homo sapiens</i>	46	2900	25000

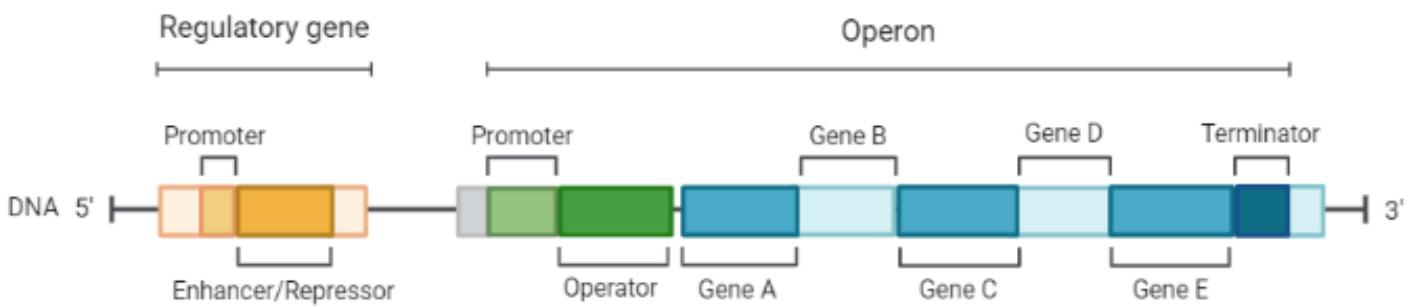
Eukaryotic chromosome organization



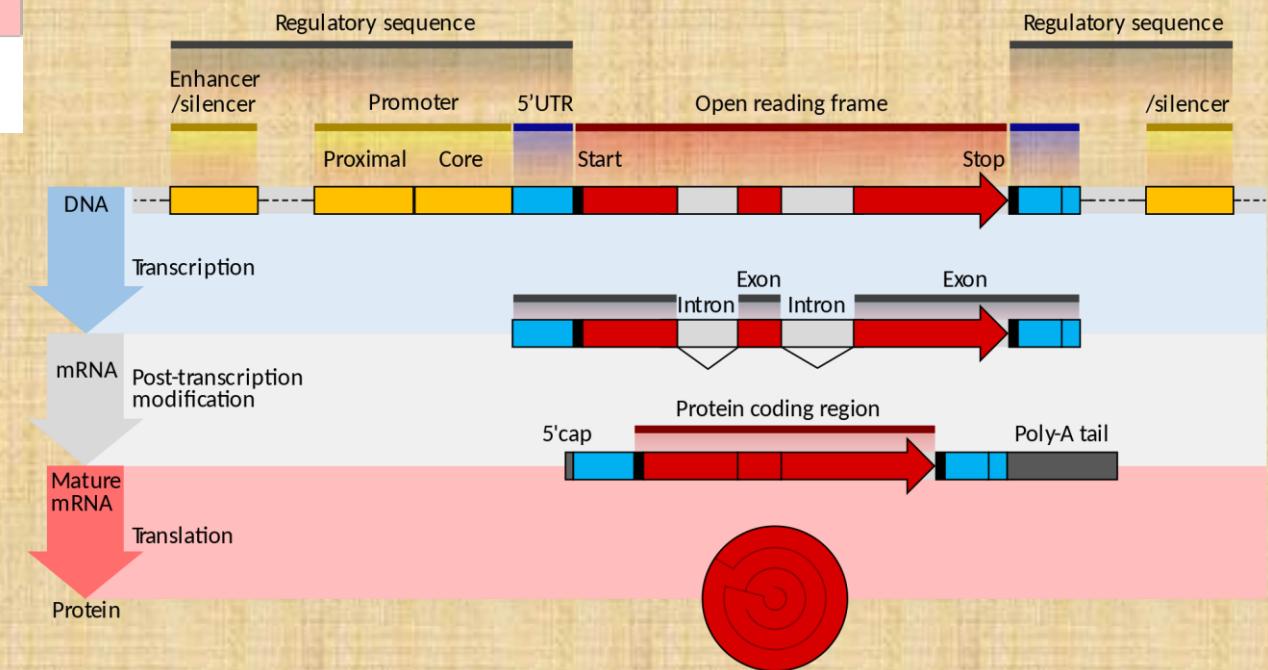
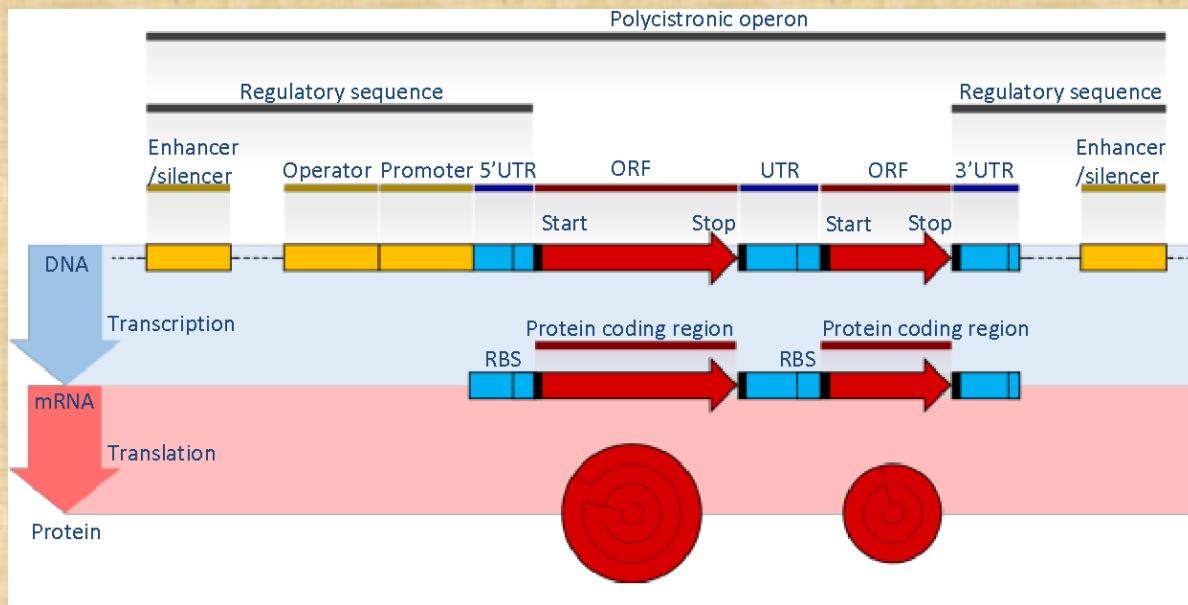
Eukaryotic Gene Structure



Prokaryotic Gene Structure



Prokaryotic and Eukaryotic gene structure

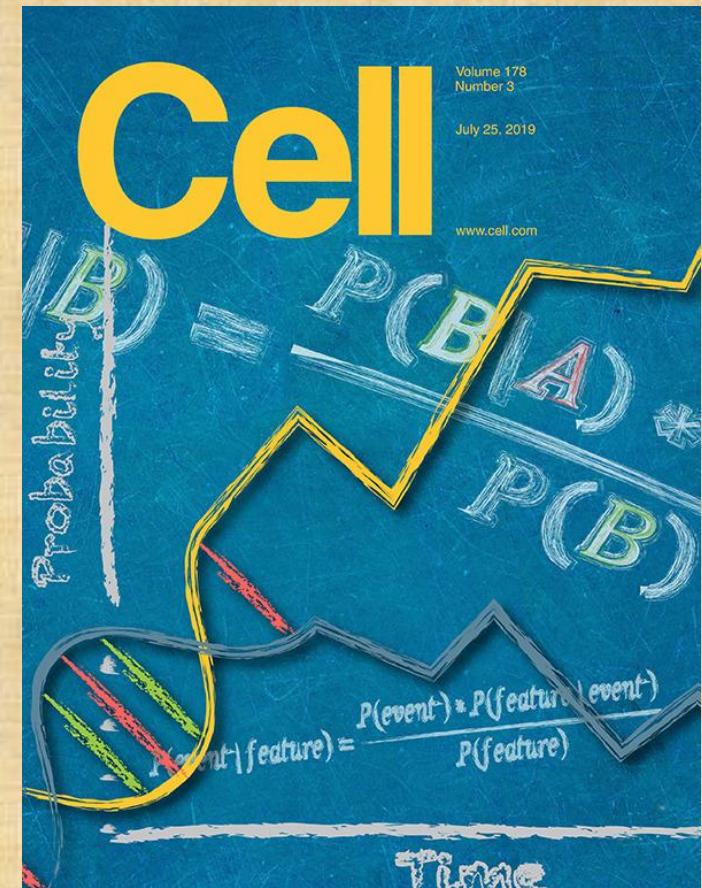




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Impact factor: 42 (2019)



Publisher: Cell Press
Impact factor: 36.637 (2019)



Herbert Boyer (pictured) and Stanley Cohen created the first genetically modified organism, a bacteria resistant to the antibiotic kanamycin in 1973.



Rudolf Jaenisch created the first genetically modified animal, a mouse in 1974

Simian Virus 40 DNA Sequences in DNA of Healthy Adult Mice Derived from Preimplantation Blastocysts Injected with Viral DNA

Rudolf Jaenisch and Beatrice Mintz

PNAS April 1, 1974 71 (4) 1250-1254; <https://doi.org/10.1073/pnas.71.4.1250>



Mary-Dell Chilton: produced the first genetically modified plants using Agrobacterium carrying the disarmed Ti plasmid (**1983**)

A chimaeric antibiotic resistance gene as a selectable marker for plant cell transformation

Michael W. Bevan, Richard B. Flavell & Mary-Dell Chilton

Nature 304, 184–187 (1983) | Download Citation ↴