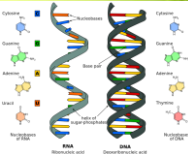


Francis Crick James Watson Maurice Wilkins Rosalind Franklin



## Molecular Biology

### 2.6- Structure of DNA and RNA

The Hierarchical Structure of DNA through to the Chromosome

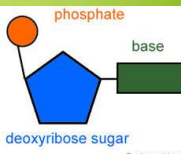
DNA (genes and other nucleotides) reside in 46 chromosomes

Genes are nucleotides that get expressed in the real world

Nucleotides are multiple segments of DNA base pairs

Human DNA stretched out measures some 6 feet / 1.8 meters

DNA is a combination of 4 possible amino acids, bound in pairs, in a double helix structure




phosphate base deoxyribose sugar

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## Essential idea:

- The structure of DNA allows efficient storage of genetic information.
  - 32 Billion base pairs, or sets of genetic "letters", make up the human genome.
  - Each uncoiled DNA strand is about 6 feet long
  - Do this for all your DNA, and the resulting strand would be 67 billion miles long—the same as about 150,000 round trips to the Moon.



The Hierarchical Structure of DNA through to the Chromosome

DNA (genes and other nucleotides) reside in 46 chromosomes

Genes are nucleotides that get expressed in the real world

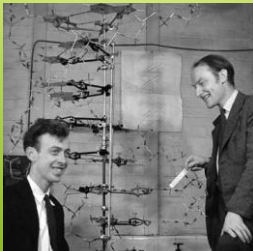
Nucleotides are multiple segments of DNA base pairs

Human DNA stretched out measures some 6 feet / 1.8 meters

DNA is a combination of 4 possible amino acids, bound in pairs, in a double helix structure


## Nature of science:

- Using models as representation of the real world
  - Crick and Watson used model making to discover the structure of DNA. (1.10)



## Theory of knowledge:

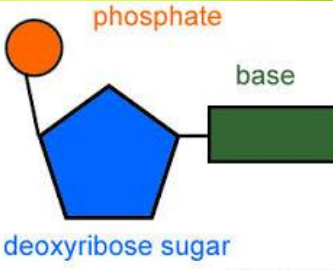
- The story of the elucidation of the structure of DNA illustrates that cooperation and collaboration among scientists exists alongside competition between research groups.
- To what extent is research in secret 'anti-scientific'?
- What is the relationship between shared and personal knowledge in the natural sciences?



Francis Crick James Watson Maurice Wilkins Rosalind Franklin

## Nucleic Acids

- The nucleic acids DNA and RNA are polymers of **nucleotides**.
  - Nitrogenous base
  - Pentose sugar
  - Phosphate group



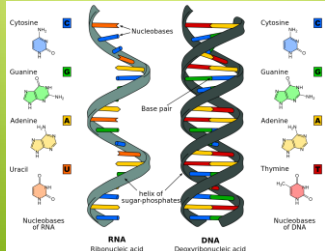
phosphate base deoxyribose sugar

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Use circles, pentagons and rectangles to represent phosphates, pentoses and bases

## Nucleic Acids

- DNA differs from RNA
  - in the number of strands present (1 strand in RNA)
  - the nitrogen base composition (Uracil not Thymine in RNA)
  - the type of pentose sugar (Ribose in RNA)



Nucleobases of RNA: Cytosine, Guanine, Adenine, Uracil

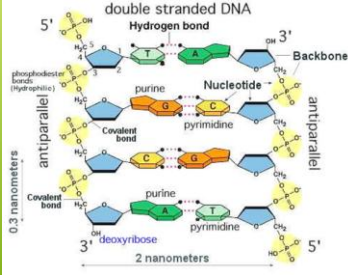
Nucleobases of DNA: Cytosine, Guanine, Adenine, Thymine

RNA: Ribonucleic acid

DNA: Deoxyribonucleic acid

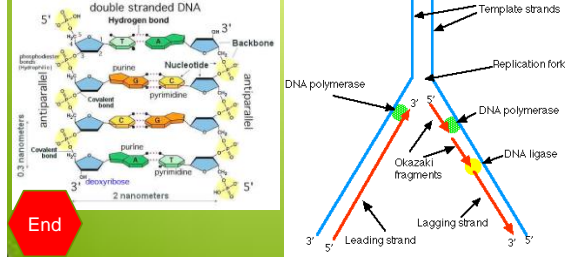
## Nucleic Acids

- DNA antiparallel strands are linked by hydrogen bonding between complementary nitrogen base pairs.
- Nucleotides link by covalent phosphodiester linkages



## Nucleic Acid Directionality

- Nucleic acids have ends, defined by the 3' and 5' carbons of the sugar in the nucleotide
- Can only add nucleotides at the 3' end.
- Build from 5' to 3'.



## Historical Experiments on DNA

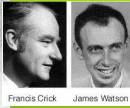
- [Frederick Griffith](#)
- [Avery-MacLeod-McCarty](#)
- [Erwin Chargaff](#)
- [Hershey-Chase](#)
- [Franklin-Wilkins-Watson-Crick](#)



← Back



Avery



MacLeod



## Frederick Griffith (1928)

- Something Can Transform Bacteria
- Conducted experiments with *Streptococcus pneumoniae*
  - Injected mice with two strains: (S) strain and a (R) strain.
  - The S strain is virulent (mice died); it has a mucous capsule
  - The R strain is not virulent (mice lived); it has no capsule.



**EXPERIMENT** Bacteria of the "S" (smooth) strain of *Streptococcus pneumoniae* are pathogenic because they have a capsule that protects them from an animal's defense system. Bacteria of the "R" (rough) strain lack a capsule and are nonpathogenic. Frederick Griffith injected mice with the two strains as shown below.

**RESULTS** Mouse dies, Mouse healthy, Mouse healthy, Mouse dies

**CONCLUSION** Griffith concluded that the living R bacteria had been transformed into pathogenic S bacteria by an unknown, heritable substance from the dead S cells.

## Frederick Griffith (1928)

- He injected mice with heat-killed S strain bacteria; the mice lived.
- He injected mice with a mixture of heat-killed S strain and live R strain bacteria; the mice died and living S strain pneumococcus were recovered from their bodies.
- Griffith concluded, some substance transformed the R strain.

**EXPERIMENT** Bacteria of the "S" (smooth) strain of *Streptococcus pneumoniae* are pathogenic because they have a capsule that protects them from an animal's defense system. Bacteria of the "R" (rough) strain lack a capsule and are nonpathogenic. Frederick Griffith injected mice with the two strains as shown below.

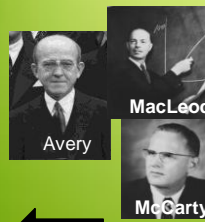
**RESULTS** Mouse dies, Mouse healthy, Mouse healthy, Mouse dies

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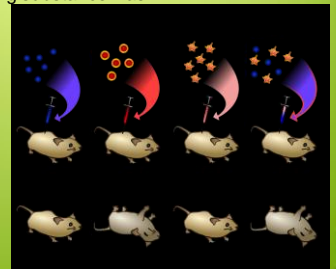
← Back

## Oswald Avery, Colin MacLeod, Maclyn McCarty (1944)

- Used deoxyribonucleopolymerase to make the Griffith extract non transformable
- Shows the transforming substance was DNA.



← Back



### Erwin Chargaff (1947)

- Performed detailed analysis of base content of DNA.
- Purine bases
  - double-ring structure
  - adenine (A) and guanine (G).
- Pyrimidine bases
  - single-ring structure
  - Thymine (T) and cytosine (C)
- Chargaff's Rules:**
  - The amount of A, T, G, and C in DNA varies from species to species.
  - In each species, the amount of A=T and G=C.

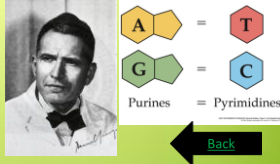


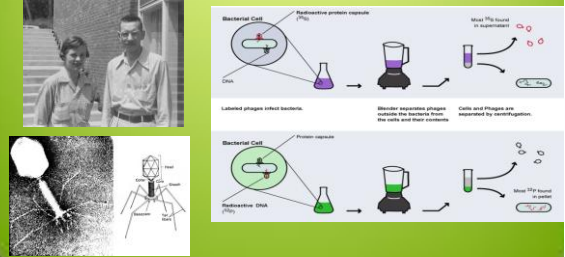
Table 1-1. Molar Proportions of Bases\* in DNAs from Various Sources

Organism	Tissue	Adenine	Thymine	Guanine	Cytosine
<i>Escherichia coli</i> (K12)	—	26.0	23.9	24.9	25.2
<i>Diphtheria</i> (poison)	—	29.8	31.6	30.5	18.0
<i>Mycobacterium tuberculosis</i>	—	15.1	14.6	34.9	35.4
Yeast	—	31.3	32.9	18.7	17.1
<i>Phaenocarpa</i> (larva)	Sperm	32.8	32.1	17.7	18.4
Human	Sperm	27.8	27.5	22.2	22.6
Sheep	Brain tissue	30.6	28.4	21.4	21.5
Human	Thymus	30.3	29.4	21.5	19.0
Human	Liver	30.5	30.3	19.5	19.7
Human	Spleen	30.7	31.2	19.5	18.8

\* Calculated as ratios of nitrogen contents per 100 g amino phosphorus in hydrolyzates.  
Source: E. Chargaff and J. Davidson, eds. *The Nucleic Acid*. Academic Press, 1955.

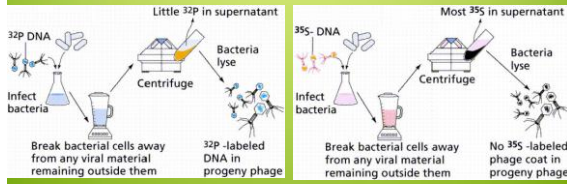
### Alfred Hershey and Martha Chase (1952)

- Used bacteriophage T2 in their experiments.
- See if protein coat or DNA directed reproduction of virus.
- In two separate experiments, they labeled the protein coat with radioactive <sup>35</sup>S and the DNA with radioactive <sup>32</sup>P.



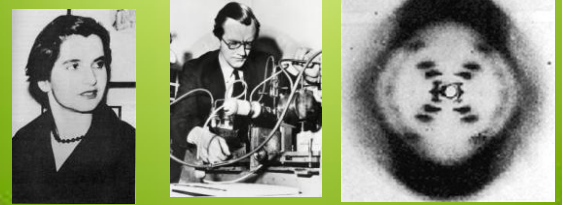
### Alfred Hershey and Martha Chase (1952)

- Viral coats are sheared away from bacterial cells and are separated by centrifugation.
- Results: radioactive <sup>32</sup>P alone is taken up by bacterial host and incorporated in virus reproduction.
- Their results reinforced the notion that DNA is the genetic material.



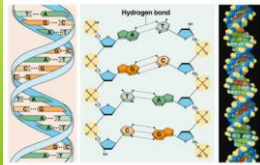
### Rosalind Franklin and Maurice Wilkins (1953)

- Franklin produced X-ray diffraction photograph of DNA.
- Wilkins gave Watson/Crick photo without Franklin's knowledge.
- Photo provided evidence that DNA had the following features:
  - DNA is a helix.
  - One part of the helix is repeated.



### James Watson and Francis Crick (1953)

- Used information generated by Franklin
- Built a model of DNA as double helix
- Sugar-phosphate molecules on outside
- Paired bases on inside.
- Sugar-phosphate backbones are antiparallel.



### James Watson and Francis Crick (1953)

- Using information generated by Chargaff.
  - Width is 2 nm.
  - Width of DNA due to purines paired to pyrimidines.
  - Chargaff's rules are consistent
    - A hydrogen-bonded to T
    - G hydrogen-bonded to C.
- Watson, Crick and Wilkins received the Nobel Prize in 1954 for their model of DNA.

