

Experimental evidences for nucleic acid (DNA & RNA) as carrier of genetic information

Introduction

- Life depends on the ability of the cells to **store** , **retrieve** & **translate** the **genetic information** required to make & maintain a living organism.
- This hereditary information is passed from one cell to its daughter cells at cell division , & from **one generation to the next** the organ's reproductive cells.
- The genetic information are stored within every living cell as its **genes**.
- A gene is a **basic unit of hereditary** & a **functional segment of DNA** (genetic material) that yield a product which is either a RNA or a protein.

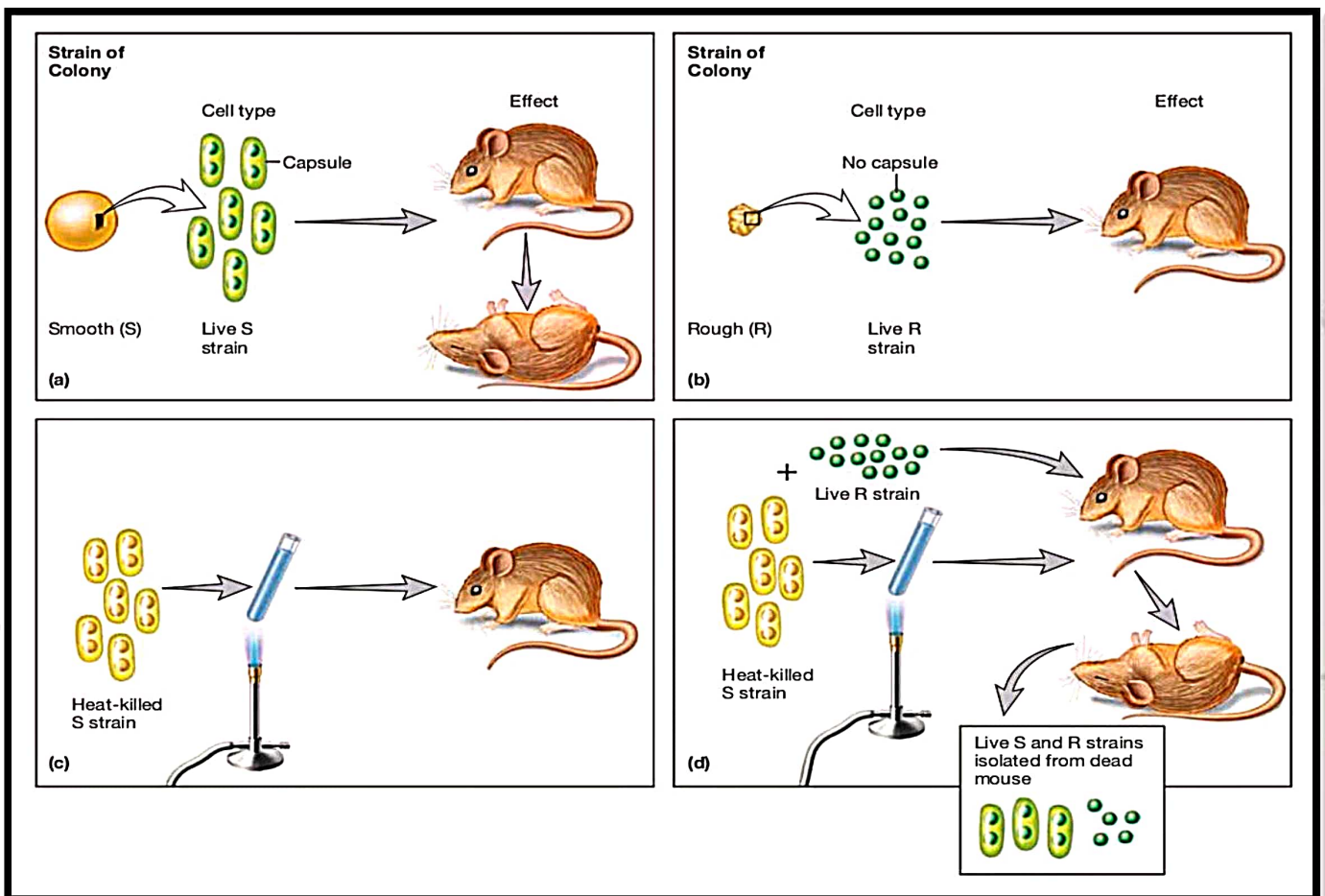
Nucleic acid as carrier of genetic information.

- Nucleic acid was first discovered by **Friedrich Miescher** from the nuclei of **pus cells** (Leukocytes) and called it **nuclein**.
- Later **Richard Altman** coined the term **nucleic acid**.
- Nucleic acid are the **polymer of nucleotides** (therefore are also called polynucleotides).
- There are two types of nucleic acids : **RNA** (ribonucleic acid) & **DNA** (deoxyribonucleic acid).
- **DNA** acts as genetic material & used to store the genetic information of **all cellular** forms of life & **many viruses**.
- **Some viruses** uses **RNA** as their genetic material.

Experimental evidences

1. Griffith's transformation experiment

- Fred Griffith experimented on the transfer of virulence in the pathogen *streptococcus pneumoniae* commonly called **pneumococcus**.
- He injected mice with **pathogenic strains** of pneumococcus which have a **capsule** & forms **smooth** colonies (**S – Strain**). As a result the **mice died**.

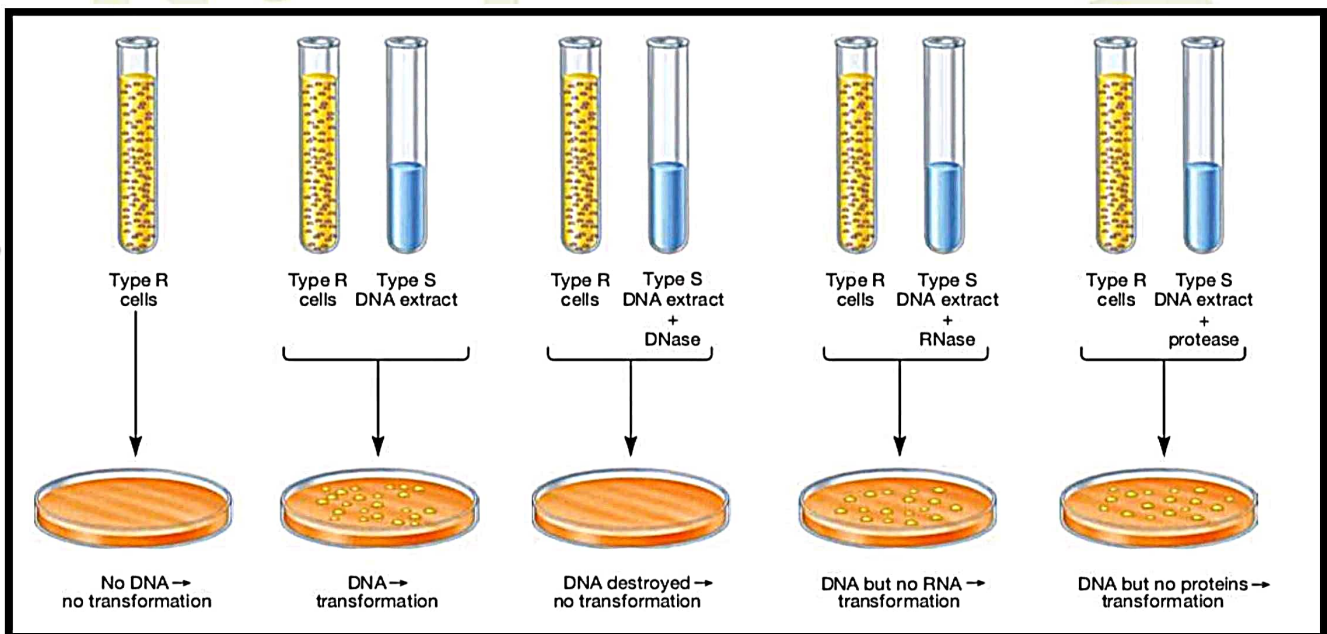


- **Mice survived** when injected with a **non pathogenic strain** of pneumococcus which **lacks a capsule** & forms **rough** colonies (**R – strain**).

- Infection with **heat killed S-strain** had **no effect**.
- Injection with a **live R-strain** & a **heat killed S-strain** gave the mice pneumonia & live S-strains could be isolated from the **dead mice**.
- Based on this information, Griffith concluded that Heat killed S-strains bacteria were responsible for converting live non virulent R-strain bacteria into virulent S-strain bacteria & called the phenomenon **transformation**.

2. Avery – MacLeod – McCarty experiment

- **Oswald Avery, Colin MacLeod & Maclyn McCarty** set out to discover which constituent in the heat killed virulent pneumococcus was responsible for Griffith's transformation.
- They discovered **DNA is the transforming principle** which resulted in transformation of Rough strain cells to S-strain cells.

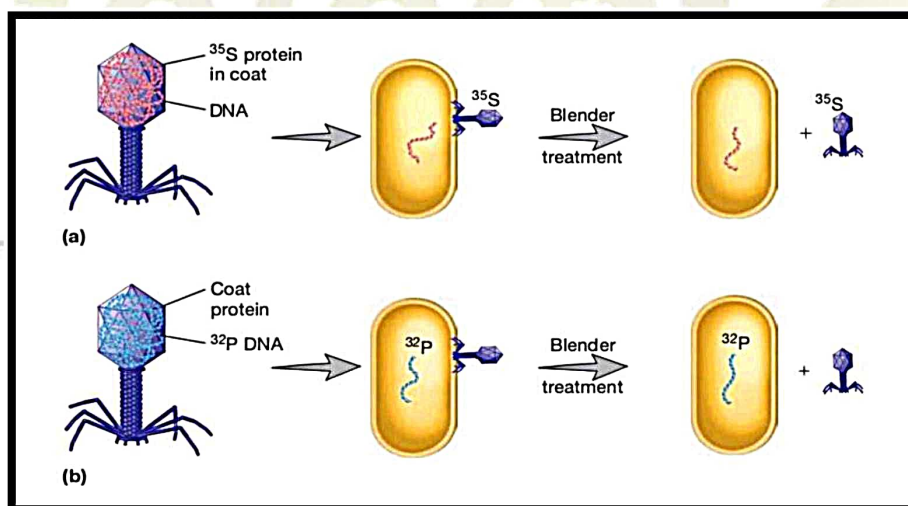


- Extract from S-strain cells was treated with **RNase**, **DNase** & **protease**. They are then mixed with R-strain cells.

- Capsulated *S*-strain cell appeared in all cultures , except those in which heat killed *S*-strain cell extract has been treated with DNase , an enzyme that destroys DNA. These results suggested that **DNA was the molecule responsible for transformation.**

3. Hershey – Chase experiment

- This experiment was conducted by **Alfred Hershey & Martha Chase** that identified **DNA as genetic material of phages**
- They designed an experiment using **radioisotopes of sulfur & phosphorus** to keep separate track of **viral proteins & Nucleic acids** during the infection process.
- **Proteins contain sulfur** but DNA doesn't. Conversely **DNA contain phosphate** but Proteins do not. Thus radioactive forms of Phosphate (^{35}P) or Sulphur (^{35}S) can be selectively incorporated into either DNA or protein.
- They used the **T2 phage** (genetic material is **linear dsDNA**) & the bacterium **E.coli** .

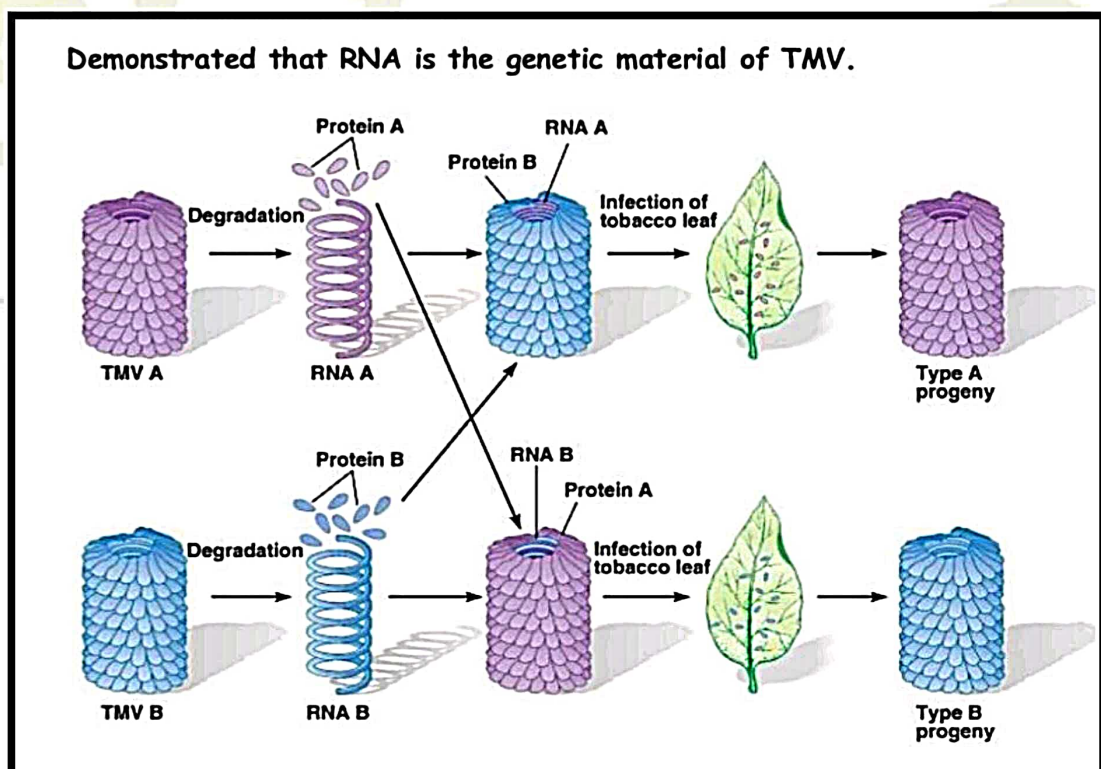


- When **^{32}P labeled phages** are used to infect *E.coli*. They found that ^{32}P DNA entered the cell & the **next generation** of phages carried **^{32}P labeled DNA.**
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- When **³⁵S labeled phages** are used to infect *E. coli*. They found that **³⁵S labelled protein** stayed outside the bacterial cell and is not **³⁵S is absent in next generation** of phages.
- Since the DNA entered the bacterial cell & is responsible for the production of new phages during the infection process, the **DNA must be genetic material. not the protein**

4. Fraenkel-Conratt experiment

- The genome of viruses may be DNA or RNA. **Most of the plant viruses** have **RNA** as their hereditary material.
- **Fraenkel-Conrat (1957)** conducted experiments on **tobacco mosaic virus (TMV)** to demonstrate that in some viruses **RNA acts as genetic material**.
- TMV is a small virus composed of a single molecule of **helical RNA** encapsulated in a **cylindrical protein coat**.



- *Fraenkel-Conrat and Singer took **two different strains of TMV** and **separated the RNAs from protein coats**, reconstituted **hybrid viruses** by mixing the proteins of one strain with the RNA of the second strain, and vice versa.*
 - *When the **hybrid viruses** were **rubbed** into **live tobacco leaves**, the **progeny viruses** produced were always found to be phenotypically and genotypically **identical to the parental type** from where the **RNA** had been isolated.*
 - *Thus the **genetic information** of TMV is **stored in the RNA** and not in the protein.*
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