

DNA The Genetic Material





# Scientific History

- The march to understanding that DNA is the genetic material
  - T.H. Morgan (1908)
  - Frederick Griffith (1928)
  - Avery, McCarty & MacLeod (1944)
  - Erwin Chargaff (1947)
  - Hershey & Chase (1952)
  - Watson & Crick (1953)
  - Meselson & Stahl (1958)

### 1908 | 1933

# Chromosomes related to phenotype

- T.H. Morgan's Experiment
  - working with Drosophila
  - <u>associated phenotype with</u> <u>specific chromosome</u>
    - white-eyed male had specific X
      chromosome

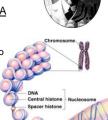




### 1908 | 1933

### Genes are on chromosomes

- · Morgan's conclusions
  - genes are on chromosomes
  - but is it the <u>protein</u> or the <u>DNA</u> of the chromosomes that are the genes?
    - initially proteins were thought to be genetic material... Why?

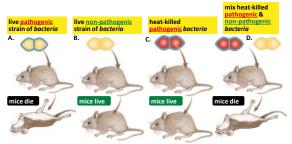


### 1928 The "Transforming Principle"

- · Frederick Griffith's Experiment
  - Streptococcus pneumonia bacteria
    was working to find cure for pneumonia
  - <u>harmless live bacteria ("rough") mixed</u> with heat-killed pathogenic bacteria ("smooth") causes fatal disease in mice
  - a substance passed from dead bacteria to live bacteria to change their phenotype
    - "Transforming Principle"



# The "Transforming Principle"



<u>Transformation</u> = change in phenotype something in heat-killed bacteria could still transmit disease-causing properties

### 1928

# The "Transforming Principle"

· Frederick Griffith's Conclusions

- a substance passed from dead bacteria to live bacteria to change their phenotype, which he called the "Transforming Principle"



### 1944 DNA is the "Transforming Principle"

- · Avery, McCarty & MacLeod's Experiments
  - purified both DNA & proteins separately from Streptococcus pneumonia bacteria • which will transform non-pathogenic bacteria?
  - injected protein into bacteria
  - no effect
  - injected DNA into bacteria transformed harmless bacteria into virulent bacteria



### 1944 | ??!!

# Avery, McCarty & MacLeod

- Conclusion
  - first experimental evidence that DNA was the genetic material







**Oswald Avery** 

# 1952 | 1969 Confirmation of DNA

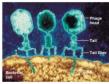
· Hershey & Chase's Experiment

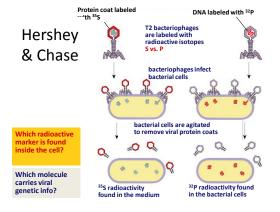
- grew bacteroiophage viruses in 2 media, radioactively labeled with either:



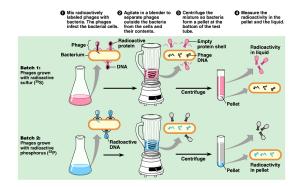
Hershey

- <u>35S in their proteins</u>
- <sup>32</sup>P in their DNA
- infected bacteria with labeled phages











# **Blender experiment Conclusions**

- · Radioactive phage & bacteria in blender
  - <sup>35</sup>S phage
    - · radioactive proteins stayed in supernatant
    - therefore viral protein did NOT enter bacteria
  - <sup>32</sup>P phage
    - radioactive DNA stayed in pellet
    - therefore viral DNA did enter bacteria
  - Confirmed DNA is "transforming factor"





Martha Chase

Alfred Hershey

# Chargaff's Conclusions

#### DNA composition:

- varies from species to species
- all 4 bases not in equal quantity
- bases present in characteristic ratio, where A=T, C=G
  - humans:
- A = 30.9%
- T = 29.4% G = 19.9%
- C = 19.8%



1947

Erwin Chargaff

### 1953 | 1962

# Structure of DNA

- Watson & Crick's Conclusions
  - developed double helix model of DNA
    - other leading scientists working on question:
    - Rosalind Franklin
      - Maurice Wilkins
      - Linus Pauling



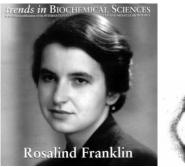




### Watson and Crick



# Rosalind Franklin (1920-1958)

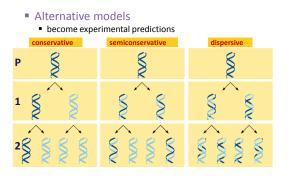




# But how is DNA copies Replication of DNA base pairing suggests that it will allow each side to serve as a template for a new strand

"It has not escaped our notice that the specific pairing we have postulated immediately suggests a possible copying mechanism for the genetic material." — Watson & Crick

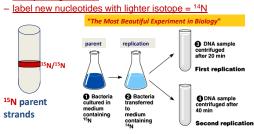
# Models of DNA Replication



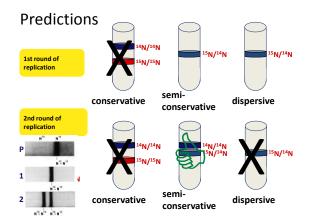
1958

# Semiconservative replication

- Meselson & Stahl's Experiment
  - <u>label "parent" nucleotides in DNA strands with</u> <u>heavy nitrogen = <sup>15</sup>N</u>









### Meselson & Stahl

Conclusions

DNA replication is semi-conservative





# Scientific History

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   T.H. Morgan (1908)
  - genes are on chromosomes
  - Frederick Griffith (1928)
  - <u>a transforming factor can change phenotype</u>
  - Avery, McCarty & MacLeod (1944)
  - transforming factor is DNA
  - Erwin Chargaff (1947)
  - <u>Chargaff rules: A = T, C = G</u>
  - Hershey & Chase (1952)
    <u>confirmation that DNA is genetic material</u>
  - Watson & Crick (1953)
    - determined double helix structure of DNA
  - Meselson & Stahl (1958)
    <u>semi-conservative replication</u>