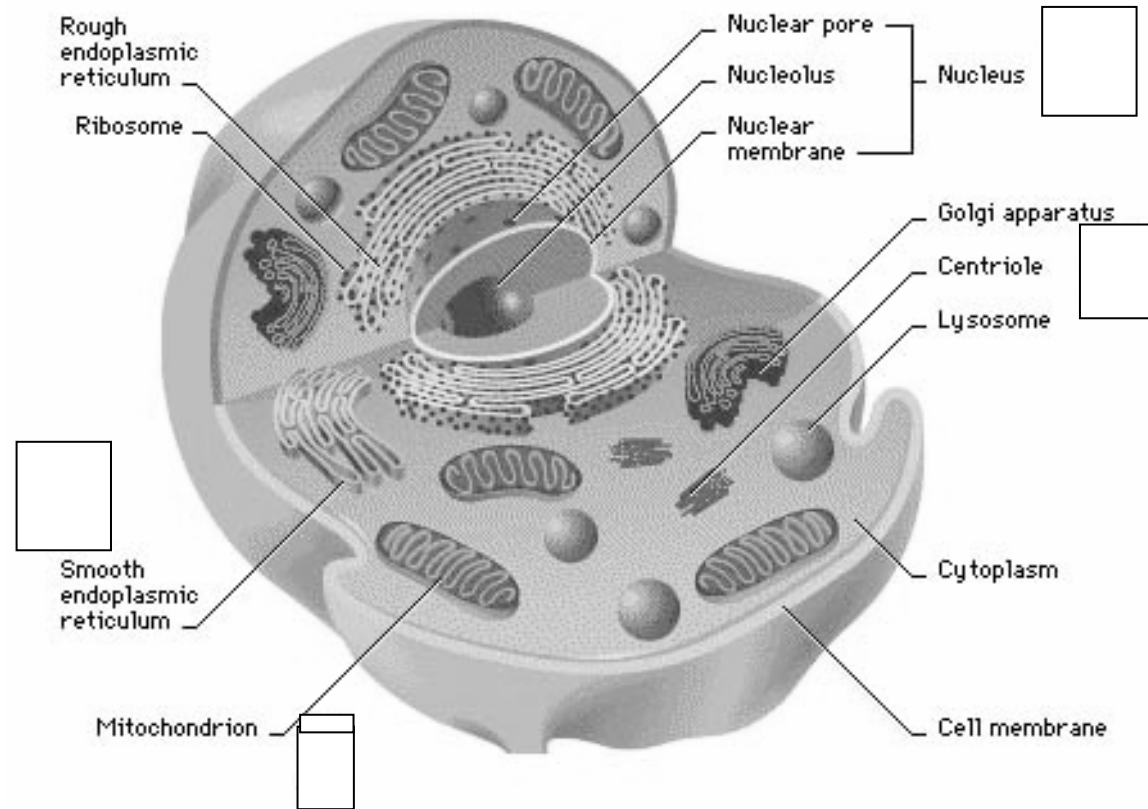


Life

# What is necessary for life?

Most life familiar to us: Eukaryotes

FREE LIVING  
Or Parasites



First appeared  $\sim 1.5 - 2 \times 10^9$  years ago

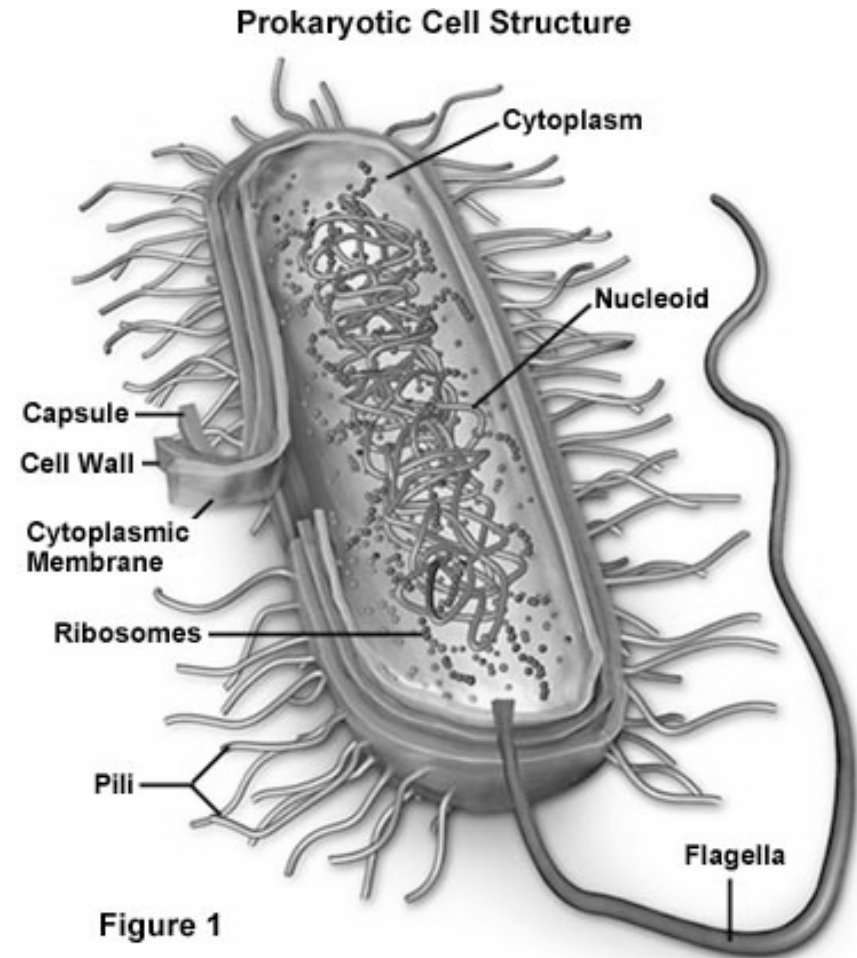
Requirements: DNA, proteins, lipids, carbohydrates,  
complex structure,  $\sim 10^4 - 10^5$  genes

# Prokaryotes (Bacteria and Archaea)

First appeared

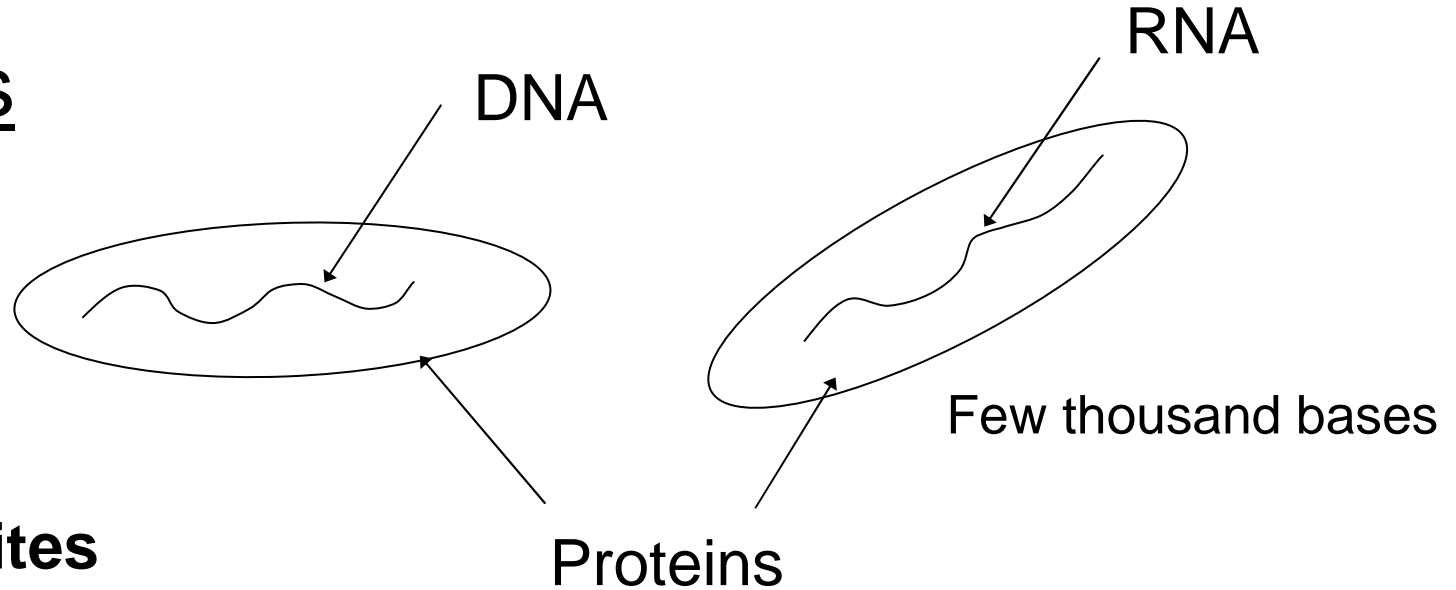
$\sim 3 - 4 \times 10^9$  years ago

FREE LIVING  
Or Parasites



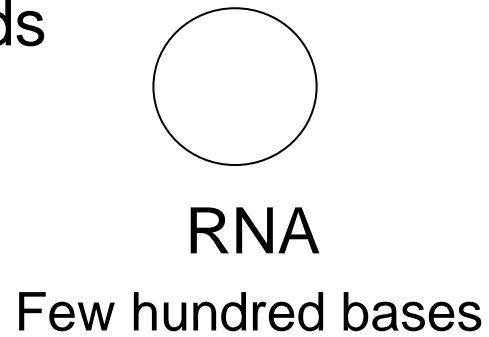
Requirements: DNA, protein, lipids, carbohydrates,  
simpler structure, few thousand genes

# Viruses

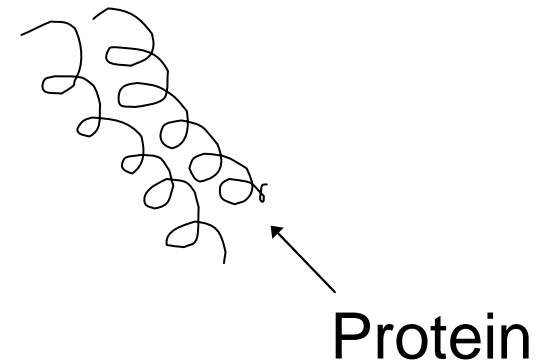


## Parasites

Viroids,  
Plasmids



Prions?



# Minimum Requirements for Life

Proteins and Nucleic Acids for simplest  
Or maybe only one.

Lipids and Carbohydrates for any thing  
more complex than a virus.

These are all macromolecules.

# Macromolecules

H, C, N, O  
(S)

Proteins made of amino acids (20)  
Construction and catalysis (enzymes)

H, C, N, O  
(P)

Nucleic acids made of nucleotides

base    sugar    phosphate

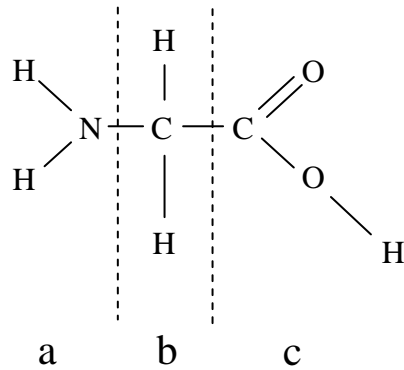
## Polymers and Monomers

# Proteins

Monomers are amino acids

20 kinds

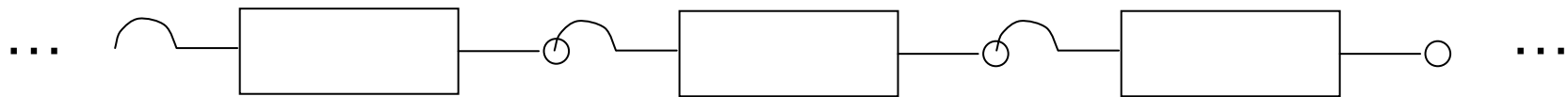
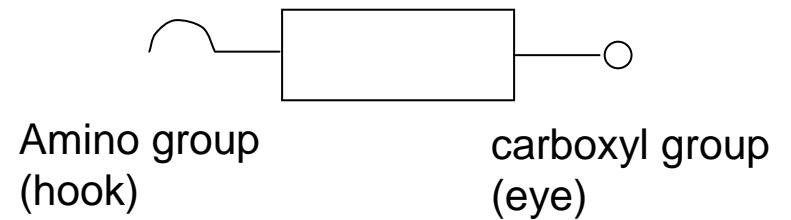
Glycine



Amino  
group

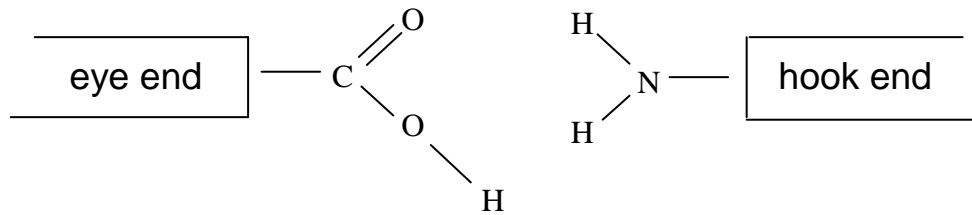
carboxyl  
group

Schematic

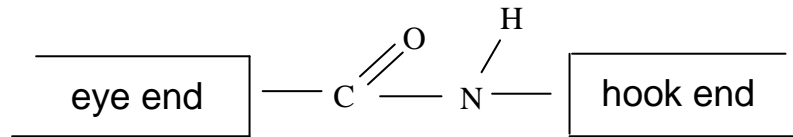


Section of Protein

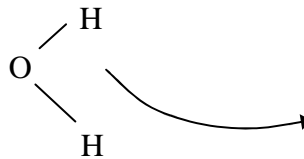
# A Peptide Bond at the Chemical Level



Before

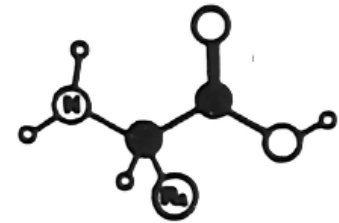


After

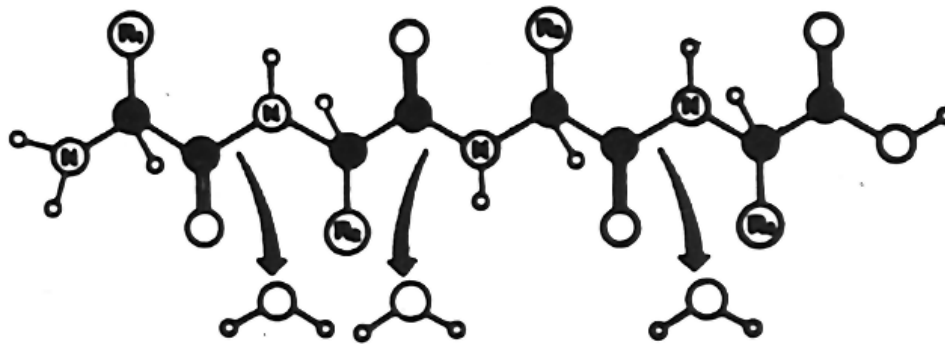


Note that a water molecule must be removed





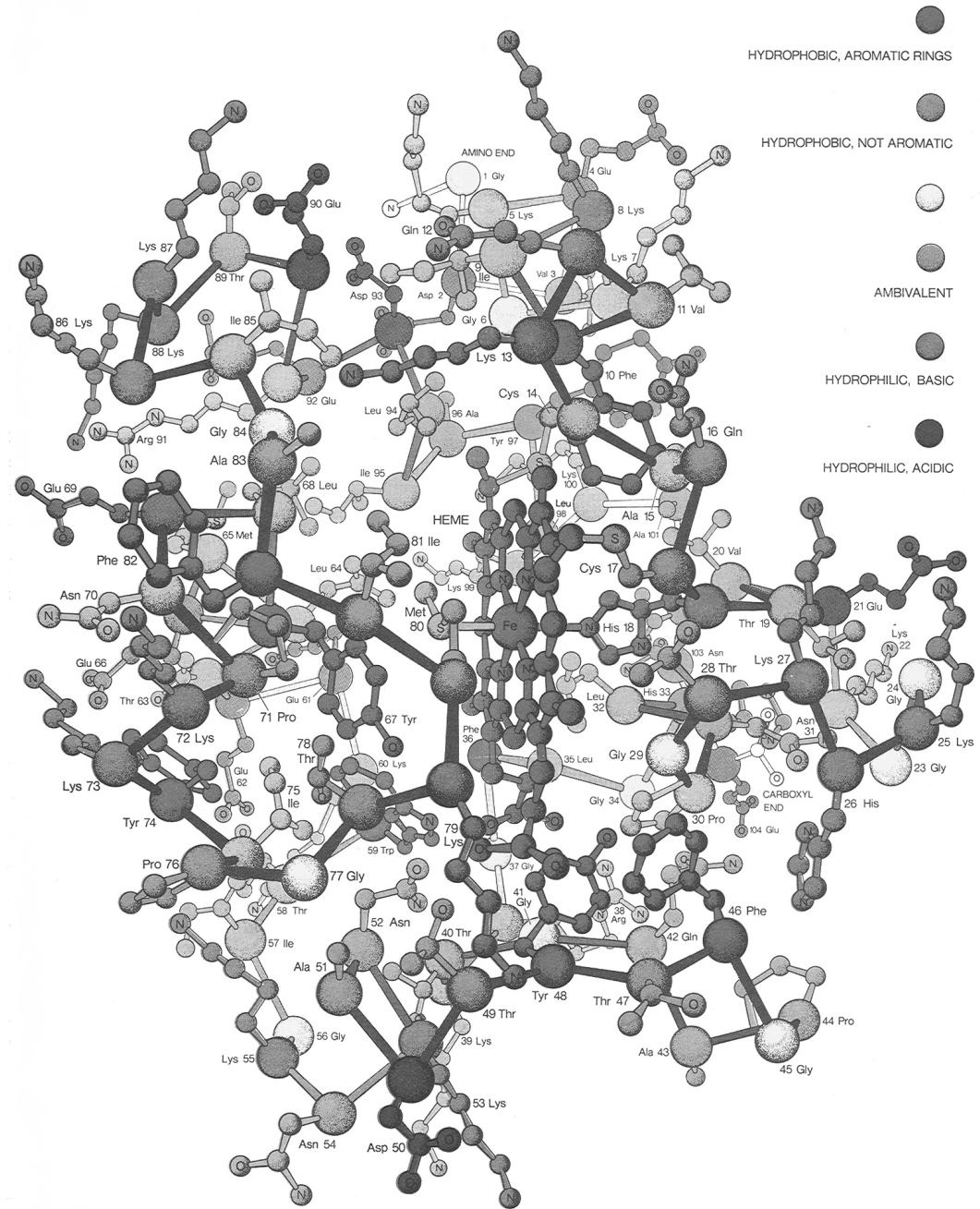
amino acids



protein

A complex protein:

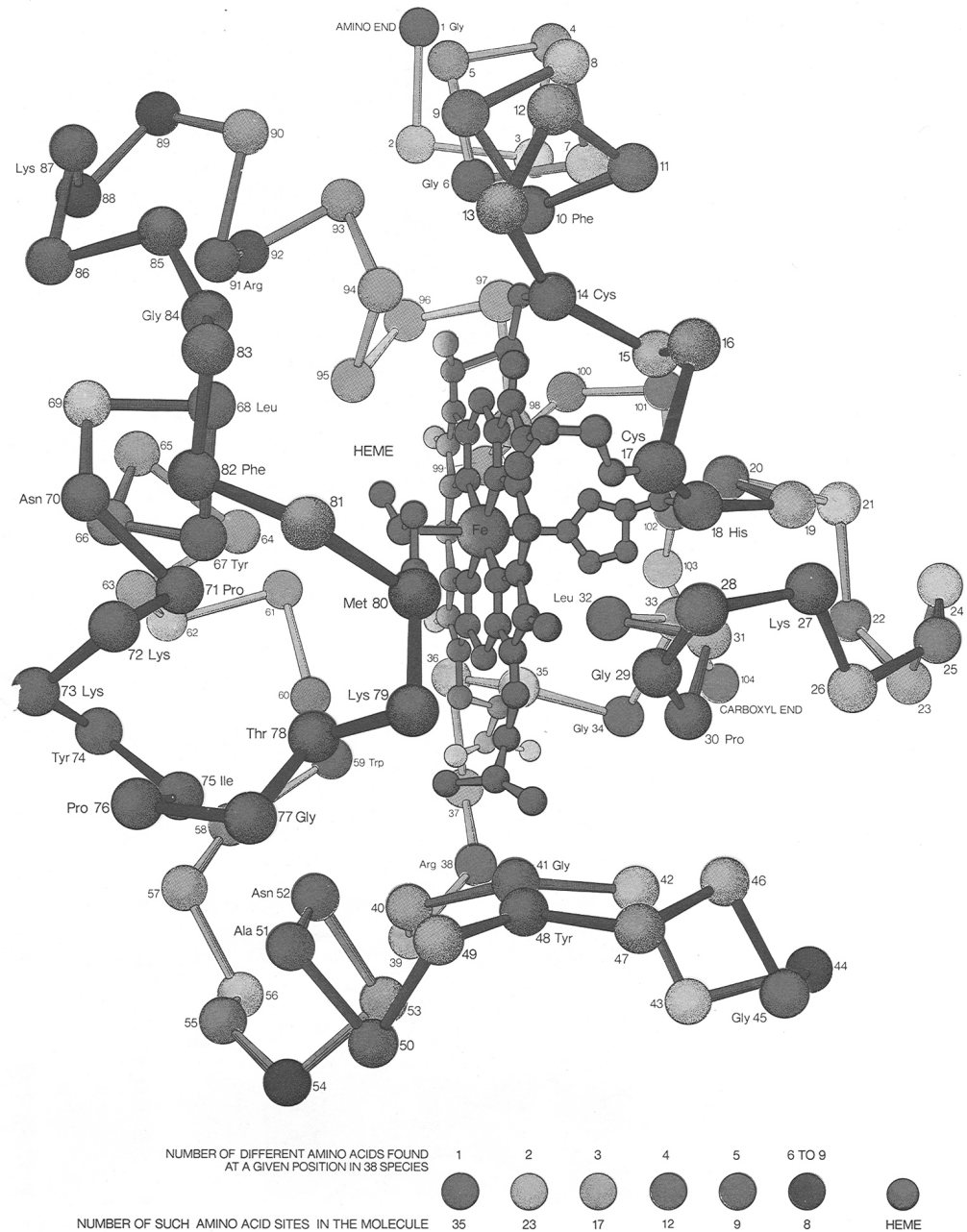
Involved in oxygen  
use Each circle is an  
amino acid



Stripped down view  
Can you find the  
amino end and the  
carboxyl end?

Note the “heme”,  
containing iron.

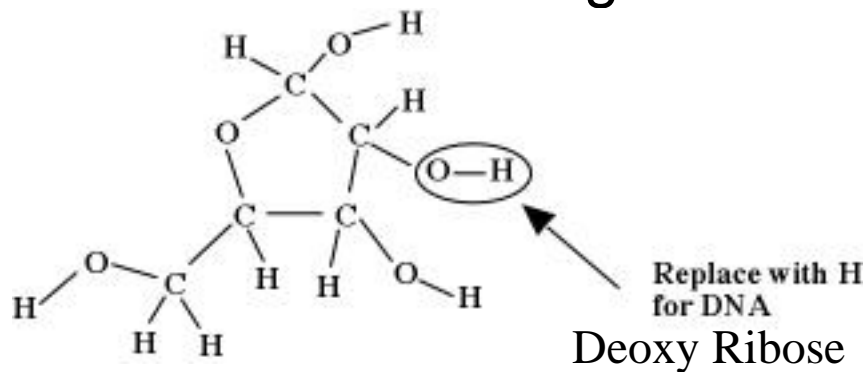
Function depends  
on structure, which  
depends on folding,  
which depends on  
order of amino acids



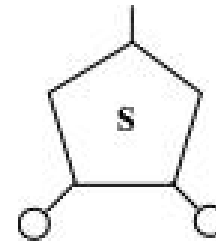
# Nucleic Acids (DNA, RNA)

Made of sugars, phosphates, bases

Sugar



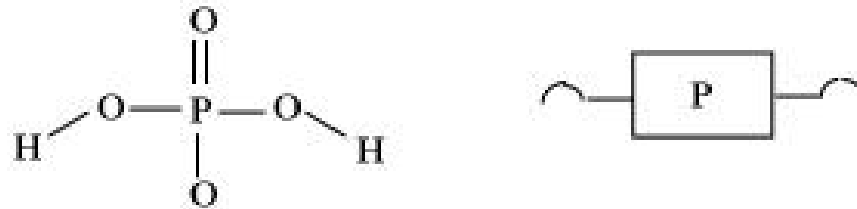
Schematic



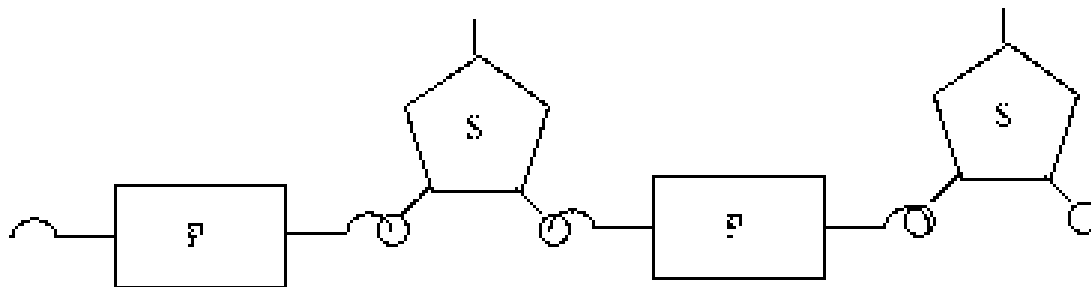
Ribose Sugar  
5 C, 5 O, 10 H

Ribonucleic acid (RNA) uses ribose sugar;  
Deoxyribonucleic acid (DNA) uses deoxyribose sugar

phosphate



sugars & phosphates linked  
phosphodiester bonds

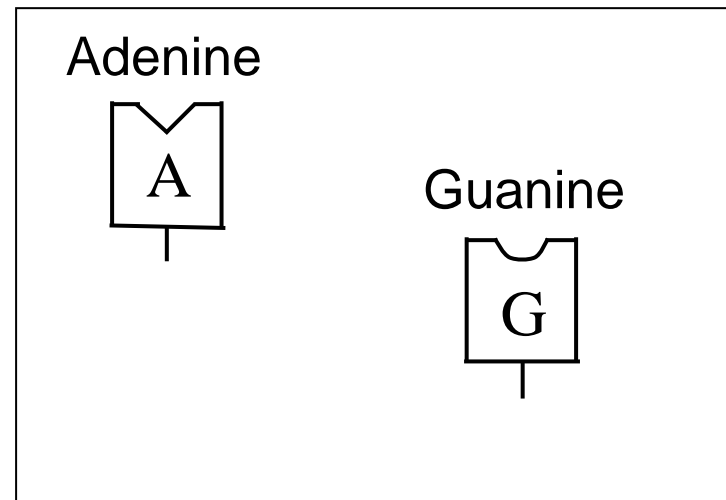
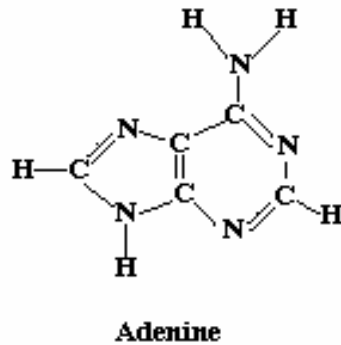


Segment of side of ladder structure

# Nucleic Acids (cont.)

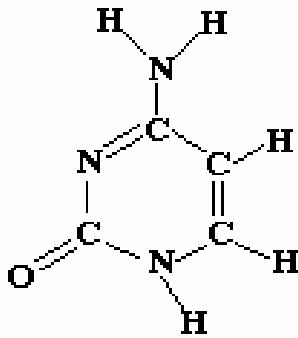
Bases: Carry Genetic Code

Purines



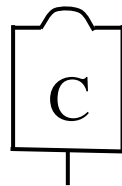
Equal numbers of C and N

# Pyrimidines

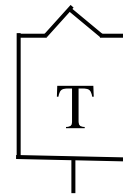
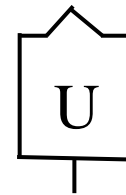


Cytosine

More C than N



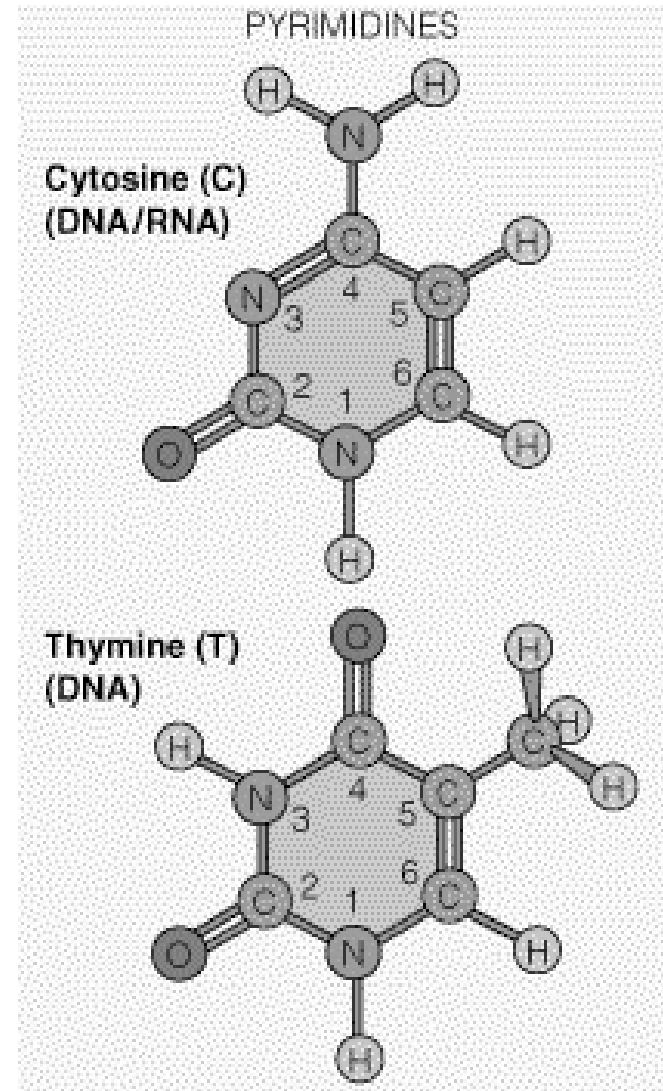
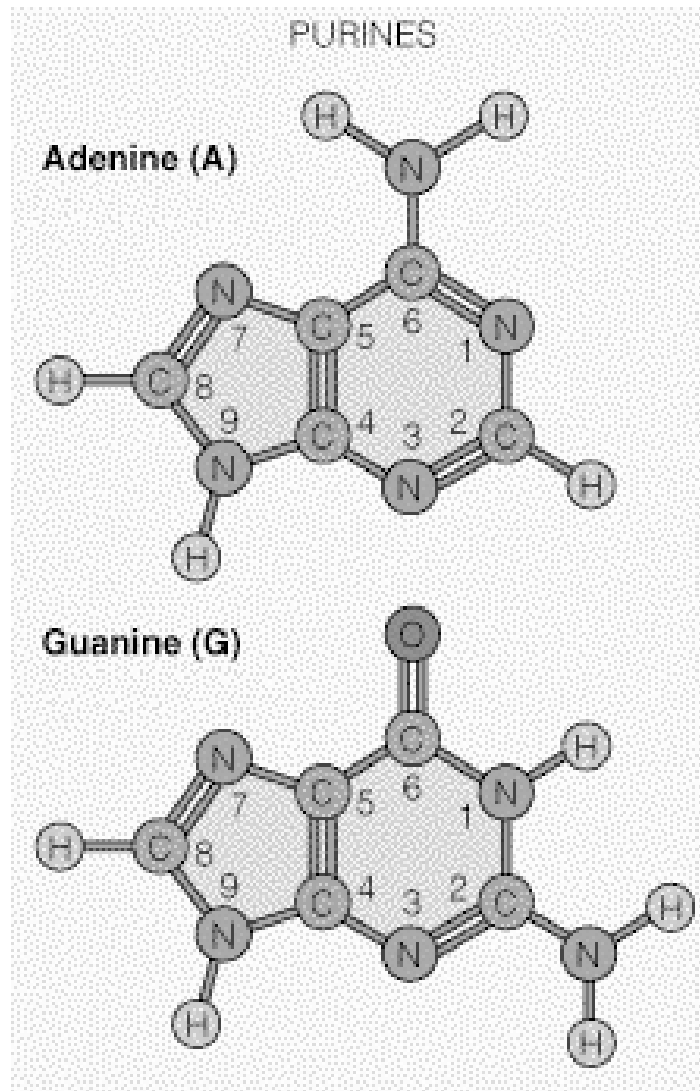
Cytosine



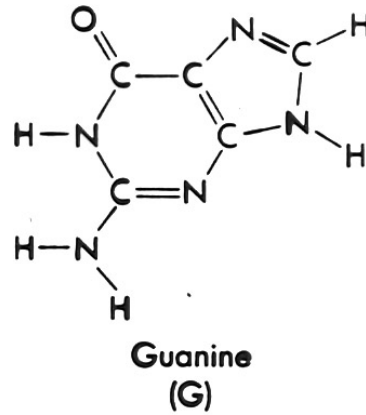
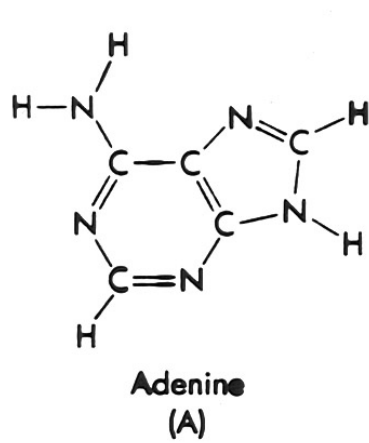
Uracil / Thymine

RNA / DNA

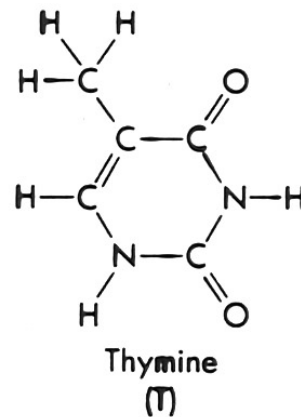
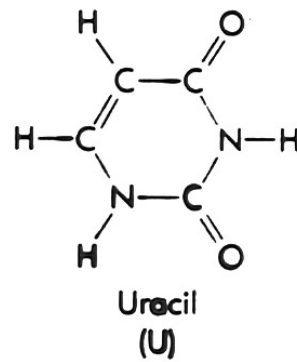
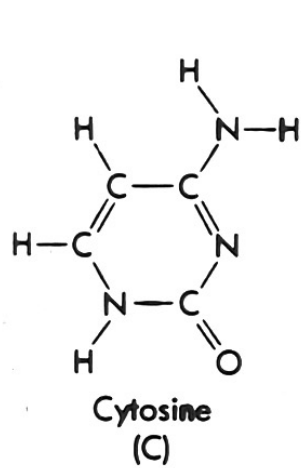
## Bases in Nucleic acids: Purines and Pyrimidines







Purines

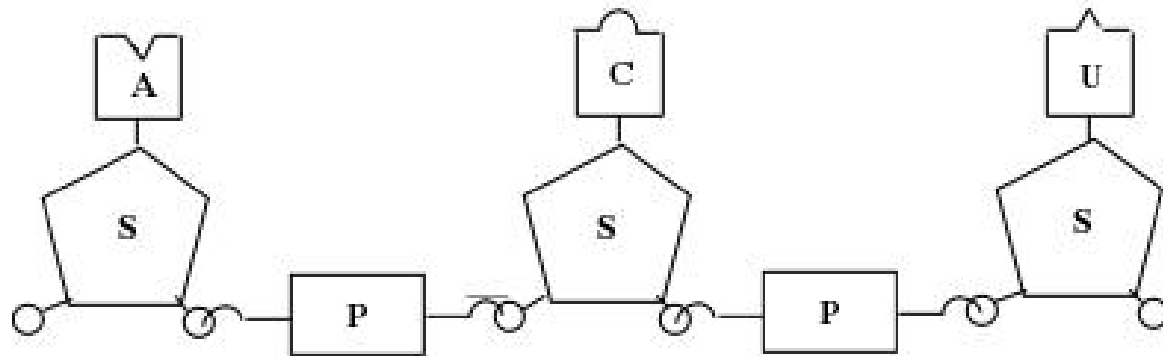


Pyrimidines

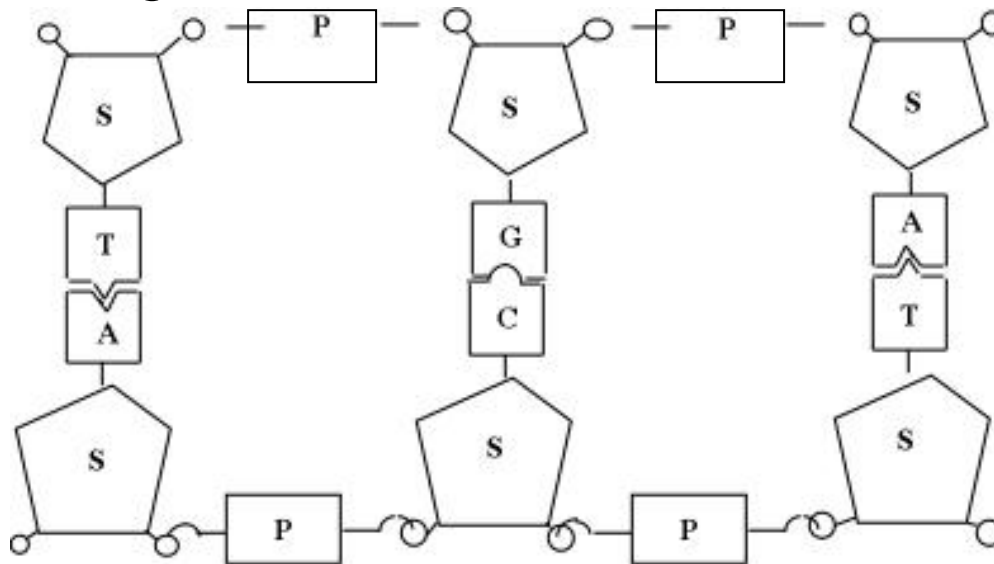
Note Uracil

# Nucleic Acids (cont.)

## Segment of RNA

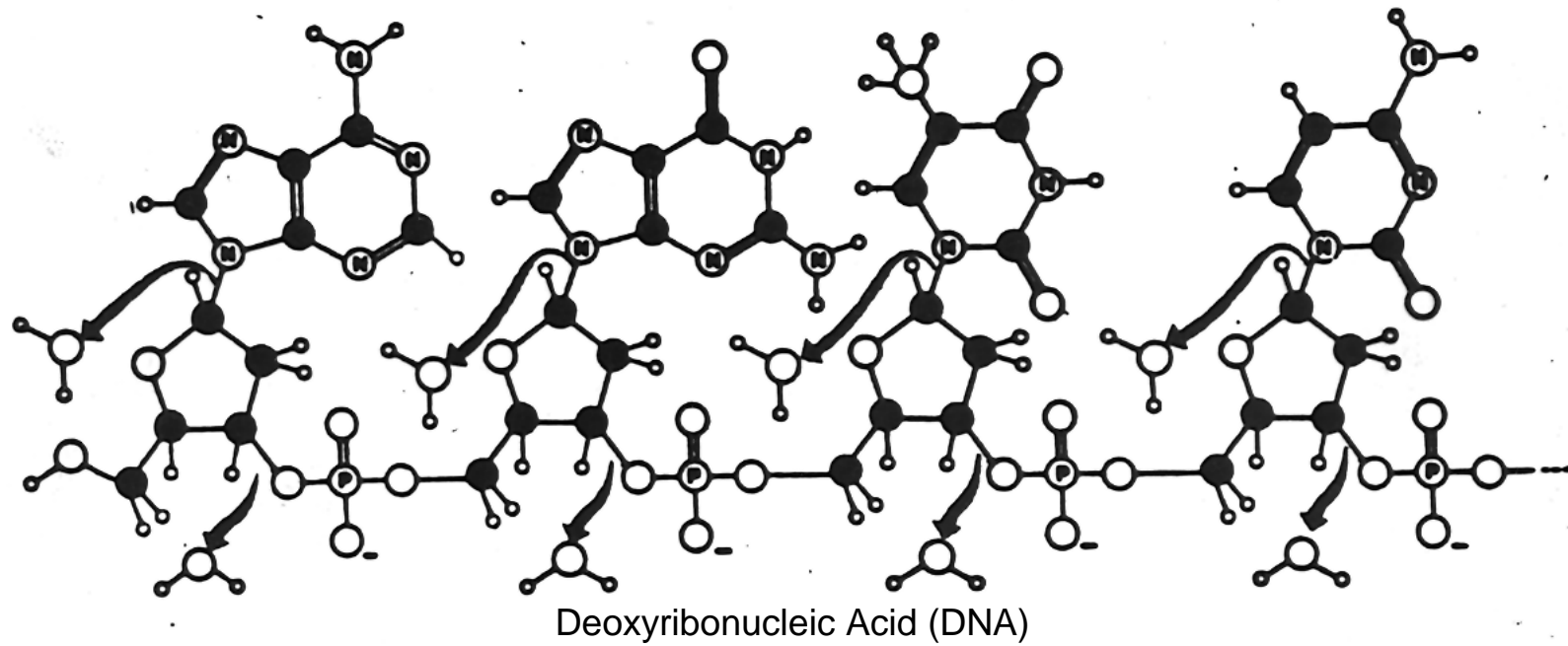
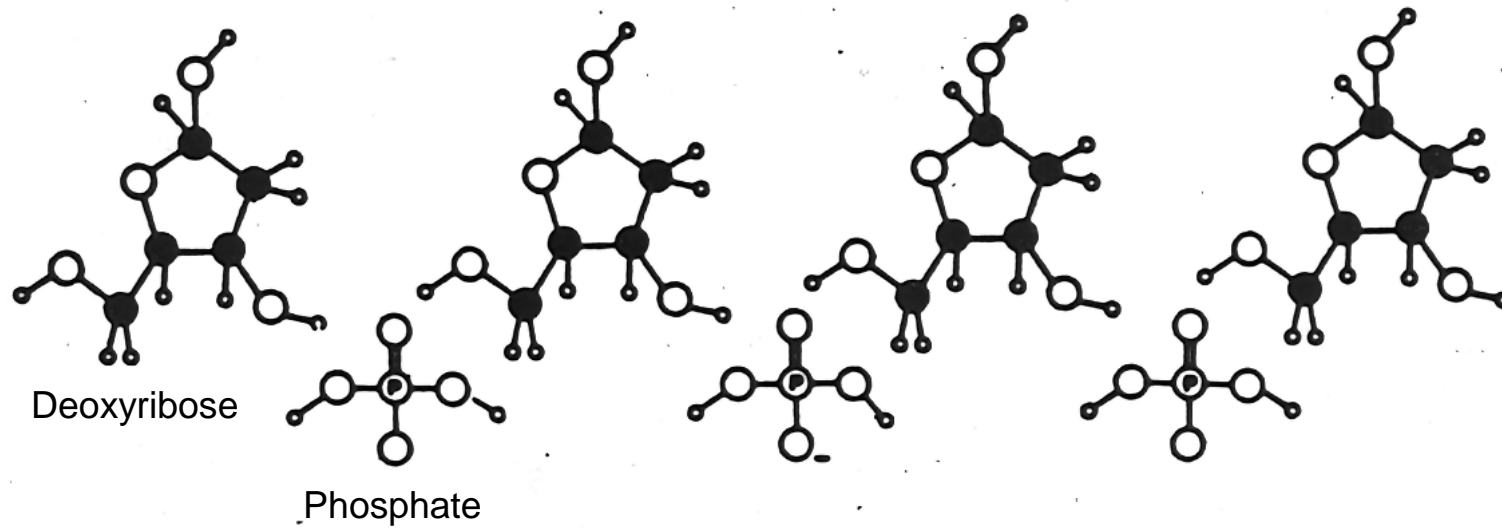


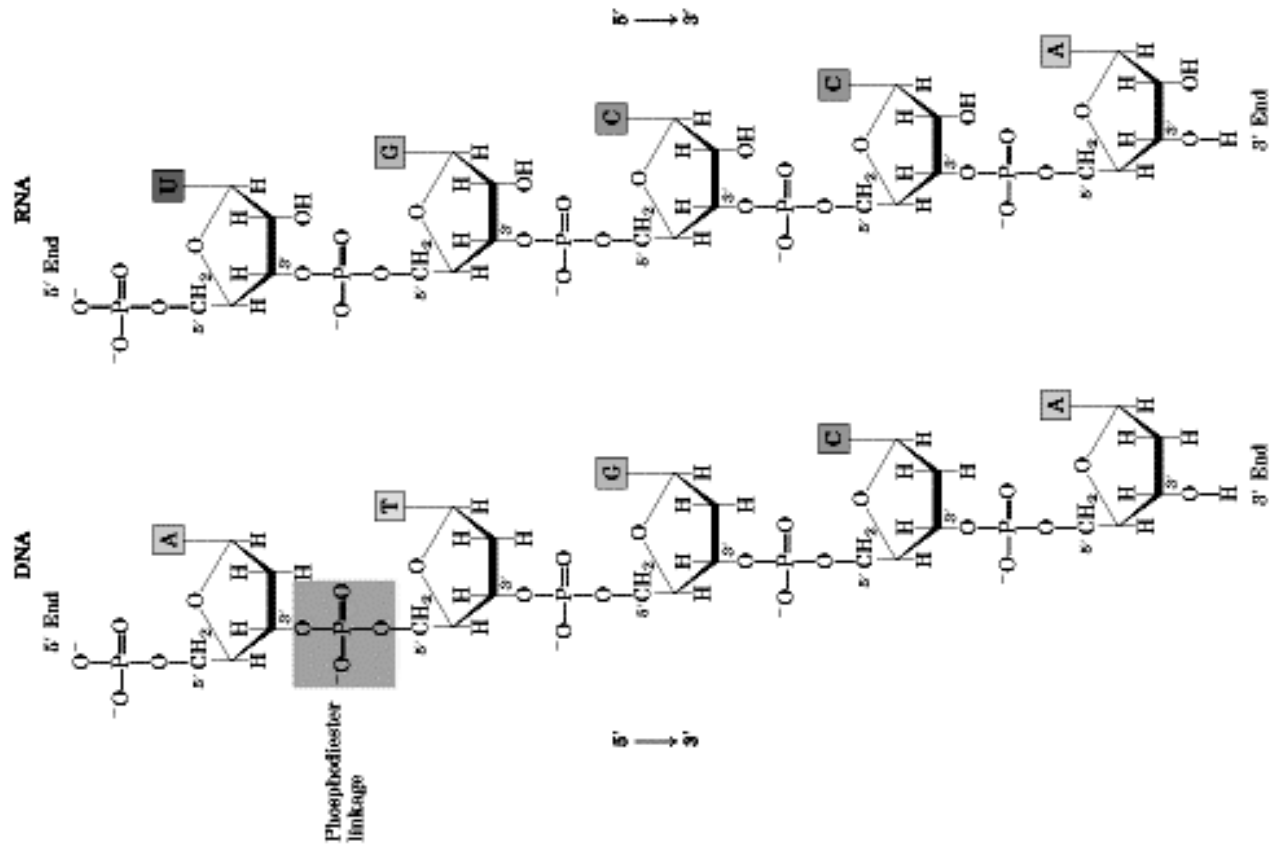
## Segment of DNA



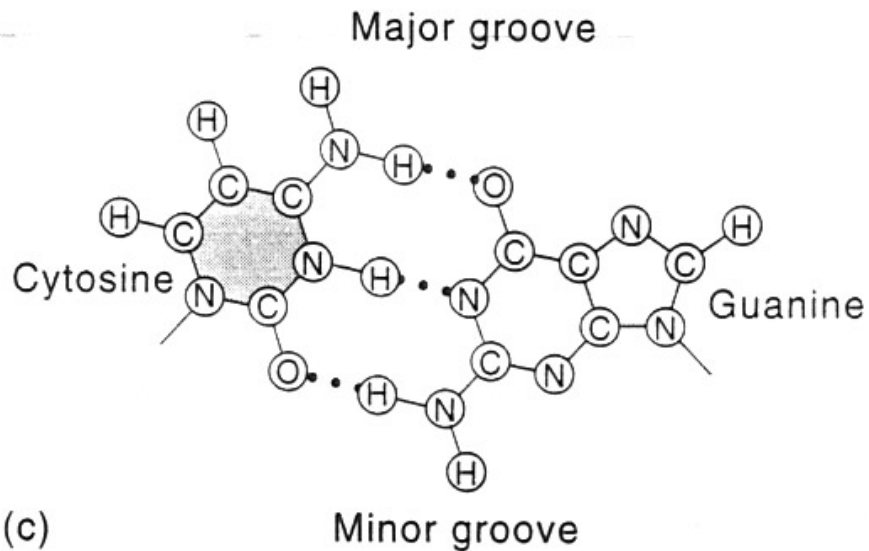
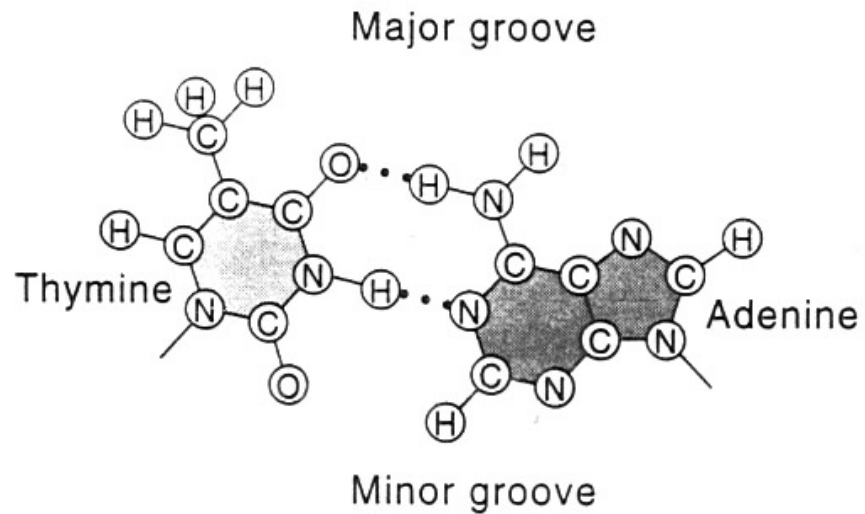
Note that T  
replaces U in DNA

## At the Chemical Level

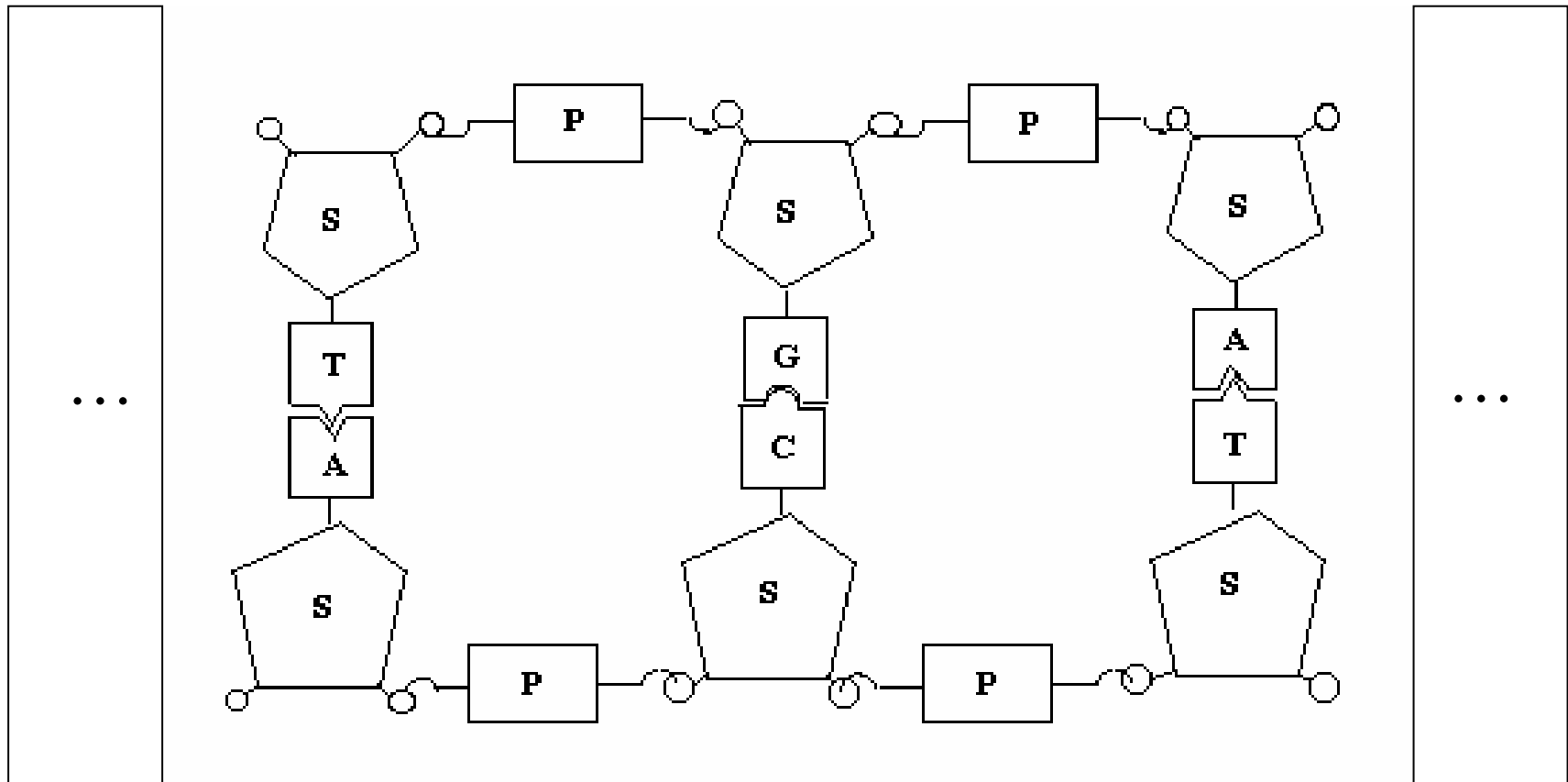




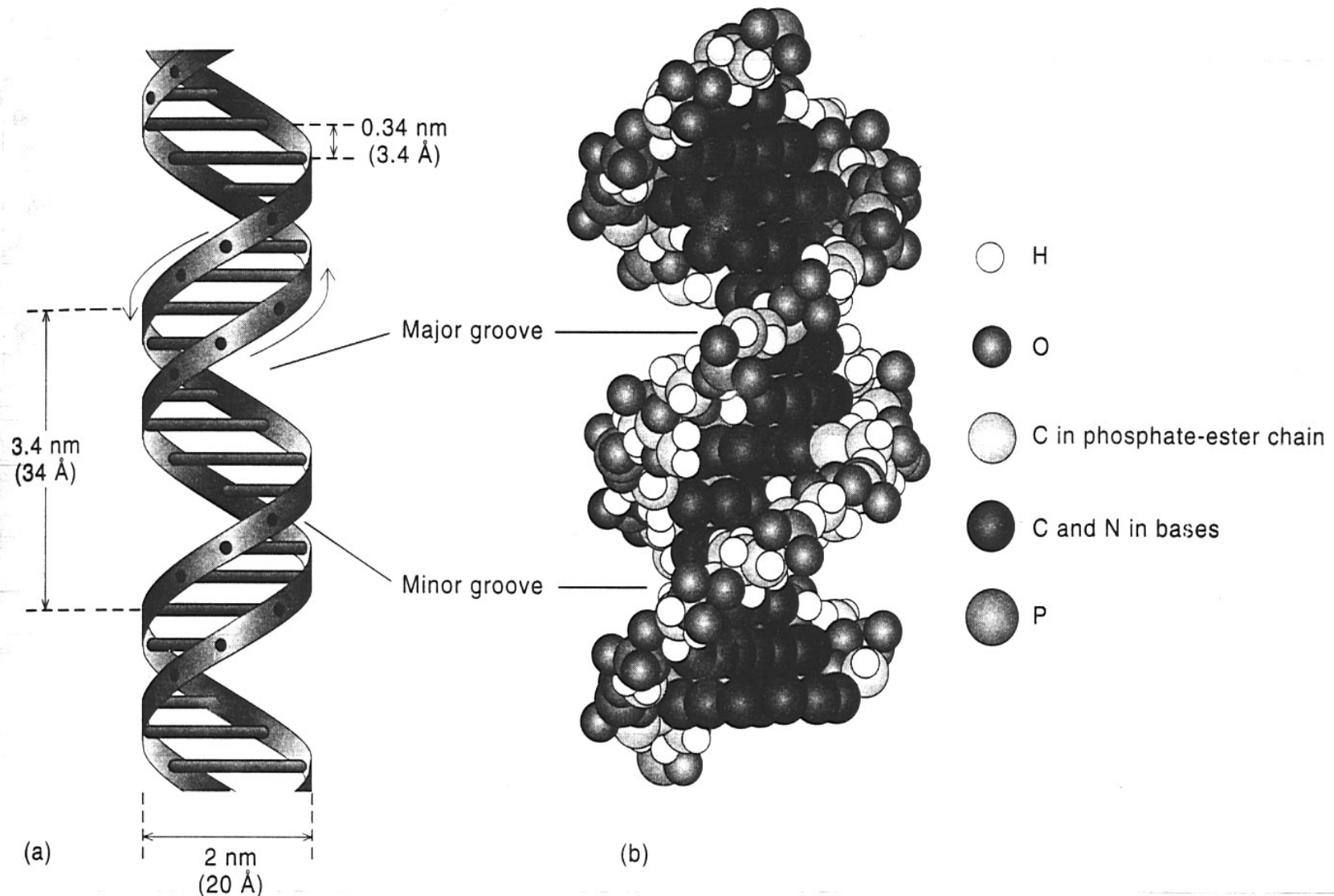
# Hydrogen Bonds (weak) connect the bases across the two sides of DNA



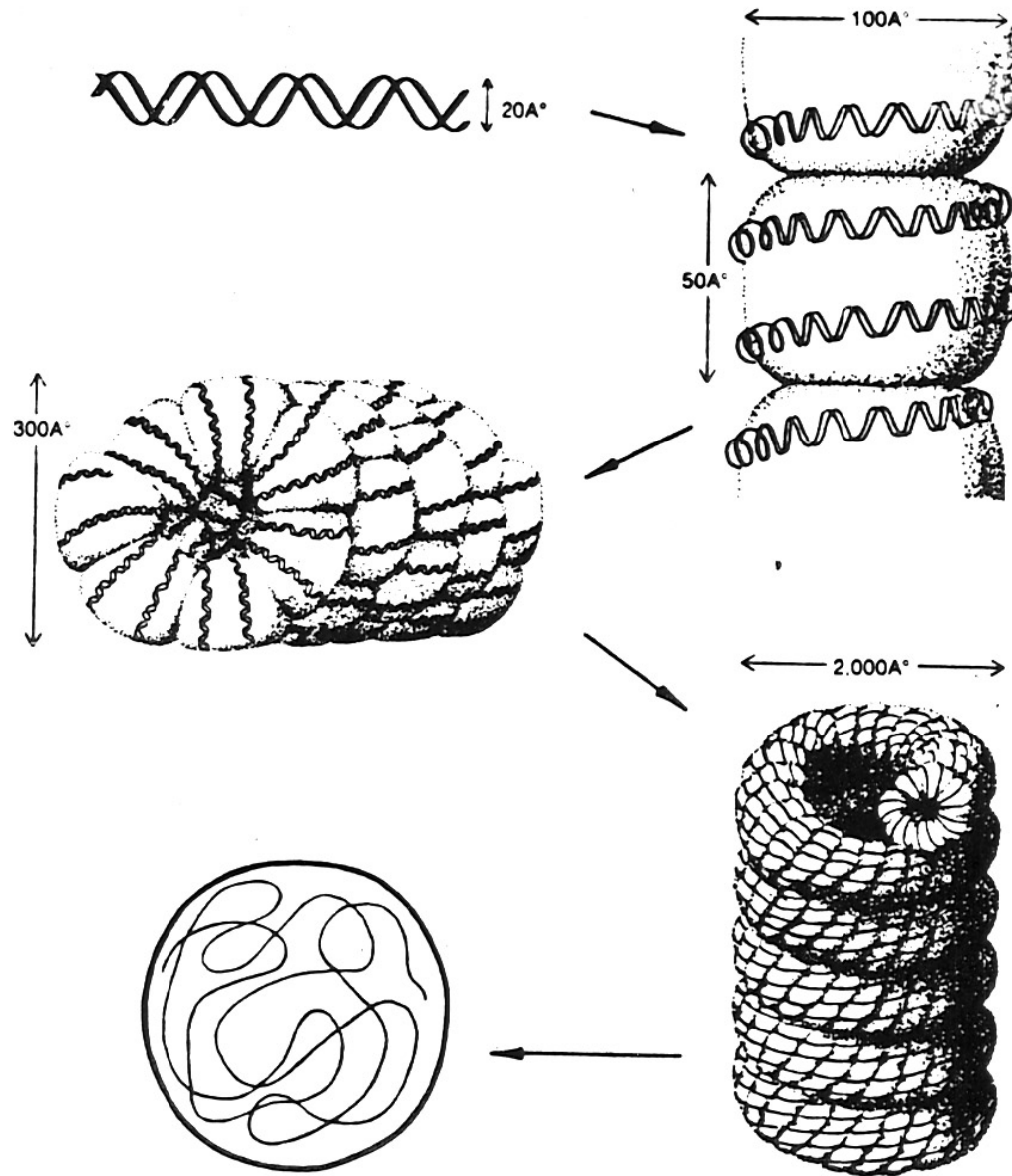
# Segment of DNA



The two strands of DNA form a double helix, connected between bases by hydrogen bonds



## Further wrapping to make compact chromosome





# Information Storage

- Nucleic acids store information
- The information specifies proteins
- The information can be replicated
- This allows inheritance

## Base pairing rules

A - T

G - C

- U

⇒ Replication of order  
(reproduction)

Nucleic Acid - Protein

↪ Genetic Code ↪

## Codon

3 base sequence       $\longrightarrow$       Amino Acid

## Gene

Sequence of codons       $\longrightarrow$       Protein

1 gene       $\longrightarrow$       1 protein

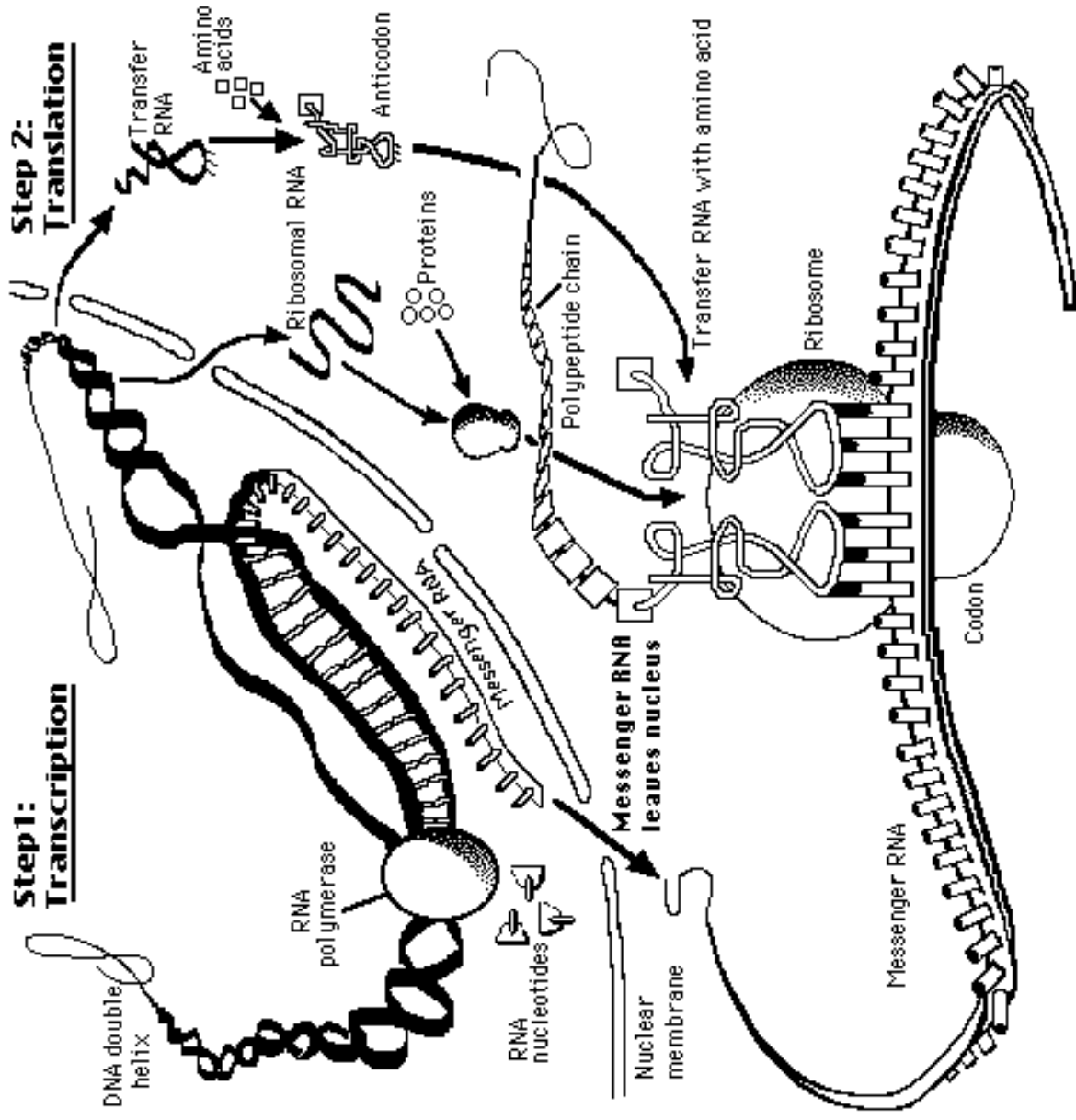
e.g.	tobacco mosaic virus	4 genes
	bacteria	$\sim 10^3$ genes
	human cell	$\sim 25,000$ genes (update)

# For mRNA Genetic Code

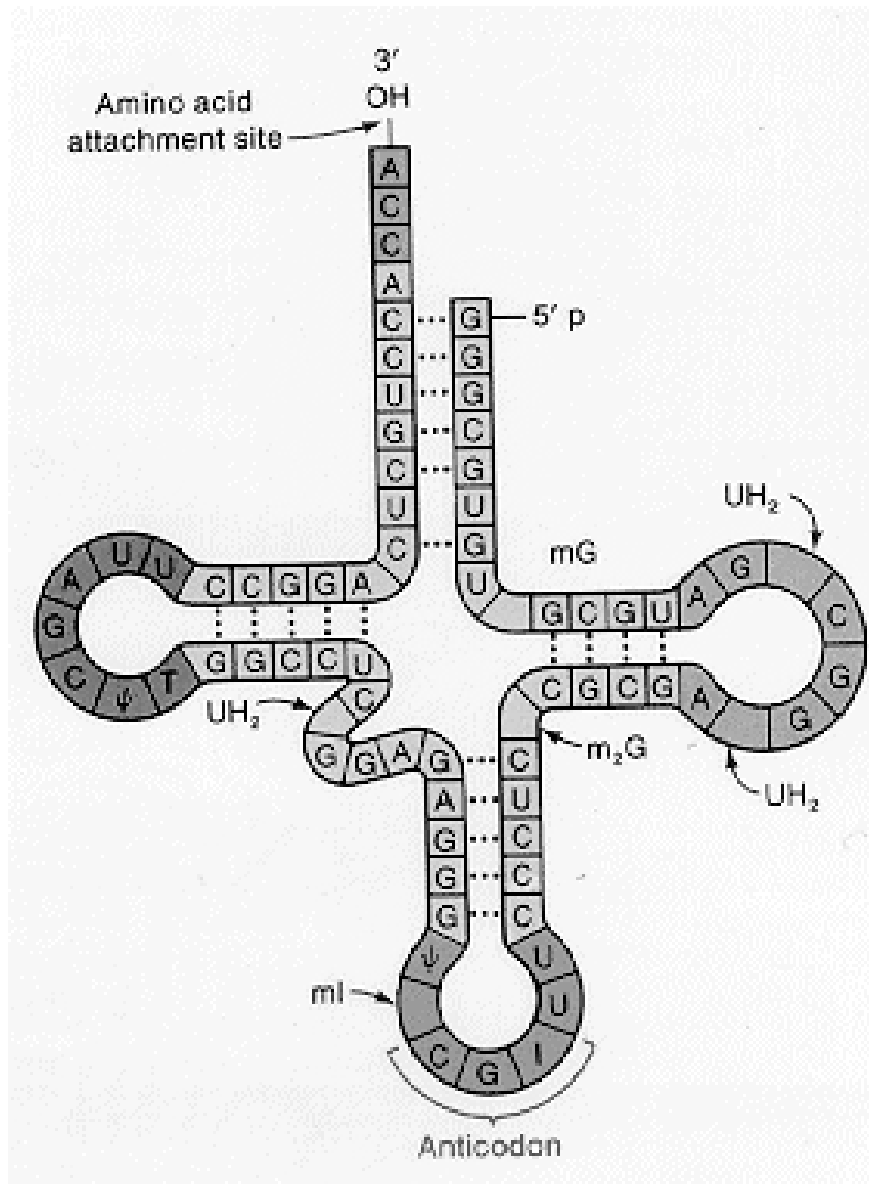
First RNA Base	U	C	A	G	Third RNA BASE
U	Phenylalanine	Serine	Tyrosine	Cysteine	U
	Phenylalanine	Serine	Tyrosine	Cysteine	C
	Leucine	Serine	Stop	Stop	A
	Leucine	Serine	Stop	Tryptophan	G
C	Leucine	Proline	Histidine	Arginine	U
	Leucine	Proline	Histidine	Arginine	C
	Leucine	Proline	Glutamine	Arginine	A
	Leucine	Proline	Glutamine	Arginine	G
A	Isoleucine	Threonine	Asparagine	Serine	U
	Isoleucine	Threonine	Asparagine	Serine	C
	Isoleucine	Threonine	Lysine	Arginine	A
	Start/Methionine	Threonine	Lysine	Arginine	G
G	Valine	Alanine	Aspartic Acid	Glycine	U
	Valine	Alanine	Aspartic Acid	Glycine	C
	Valine	Alanine	Glutamic Acid	Glycine	A
	Valine	Alanine	Glutamic Acid	Glycine	G

Amino Acids

# PROTEIN SYNTHESIS



# Structure of a tRNA



# Video

- <http://www.teachersdomain.org/9-12/sci/life/cell/proteinsynth/index.html>

## Variations in the Code

### 1. “Wobble” Bases

The third base in a codon can sometimes vary.

tRNA

U

G

mRNA

A or G

C or U

Comparison to genetic code  $\Rightarrow$  no change  
in amino acids



# For mRNA Genetic Code

First RNA Base	U	C	A	G	Third RNA BASE
U	Phenylalanine	Serine	Tyrosine	Cysteine	U
	Phenylalanine	Serine	Tyrosine	Cysteine	C
	Leucine	Serine	Stop	Stop	A
	Leucine	Serine	Stop	Tryptophan	G
C	Leucine	Proline	Histidine	Arginine	U
	Leucine	Proline	Histidine	Arginine	C
	Leucine	Proline	Glutamine	Arginine	A
	Leucine	Proline	Glutamine	Arginine	G
A	Isoleucine	Threonine	Asparagine	Serine	U
	Isoleucine	Threonine	Asparagine	Serine	C
	Isoleucine	Threonine	Lysine	Arginine	A
	Start/Methionine	Threonine	Lysine	Arginine	G
G	Valine	Alanine	Aspartic Acid	Glycine	U
	Valine	Alanine	Aspartic Acid	Glycine	C
	Valine	Alanine	Glutamic Acid	Glycine	A
	Valine	Alanine	Glutamic Acid	Glycine	G

Amino Acids

2. Some organisms use slightly different codes, with one or more changes in codon translation.

First seen in mitochondrial DNA.

Now known in some nuclear DNA

The code has evolved since the last common ancestor (But not much).

# Summary

1. Atoms needed: H, C, O, N, small amounts of P (phosphorus), S (sulfur)
2. Two basic molecules needed for life: proteins, nucleic acids
3. Both are polymers - made of simpler monomers. The monomers function as words or letters of alphabet. Information is the key.

## Summary (cont.)

4. Proteins and nucleic acids closely linked at fundamental level. Communicate through genetic code. All organisms have almost the same genetic code. It must have originated very early in evolution of life.
5. In present day organisms, protein synthesis must be directed by nucleic acids, but nucleic acid reading or replication requires enzymes (proteins). Chicken-Egg problem

# Origin of Life: I

## Monomers to Polymers

# Synthesis of Monomers

Life arose early on Earth (within  $0.7 \times 10^9$  y)

## 1. Conditions

1. Liquid Water
2. Reducing or Neutral Atmosphere
3. Energy Sources

2. Originally thought atmosphere was  
 $\text{NH}_3$ ,  $\text{CH}_4$ ,  $\text{H}_2\text{O}$ ,  $\text{H}_2$

Miller -Urey Experiment

Now Believe       $\text{CO}_2$ ,  $\text{H}_2\text{O}$ ,  $\text{N}_2$

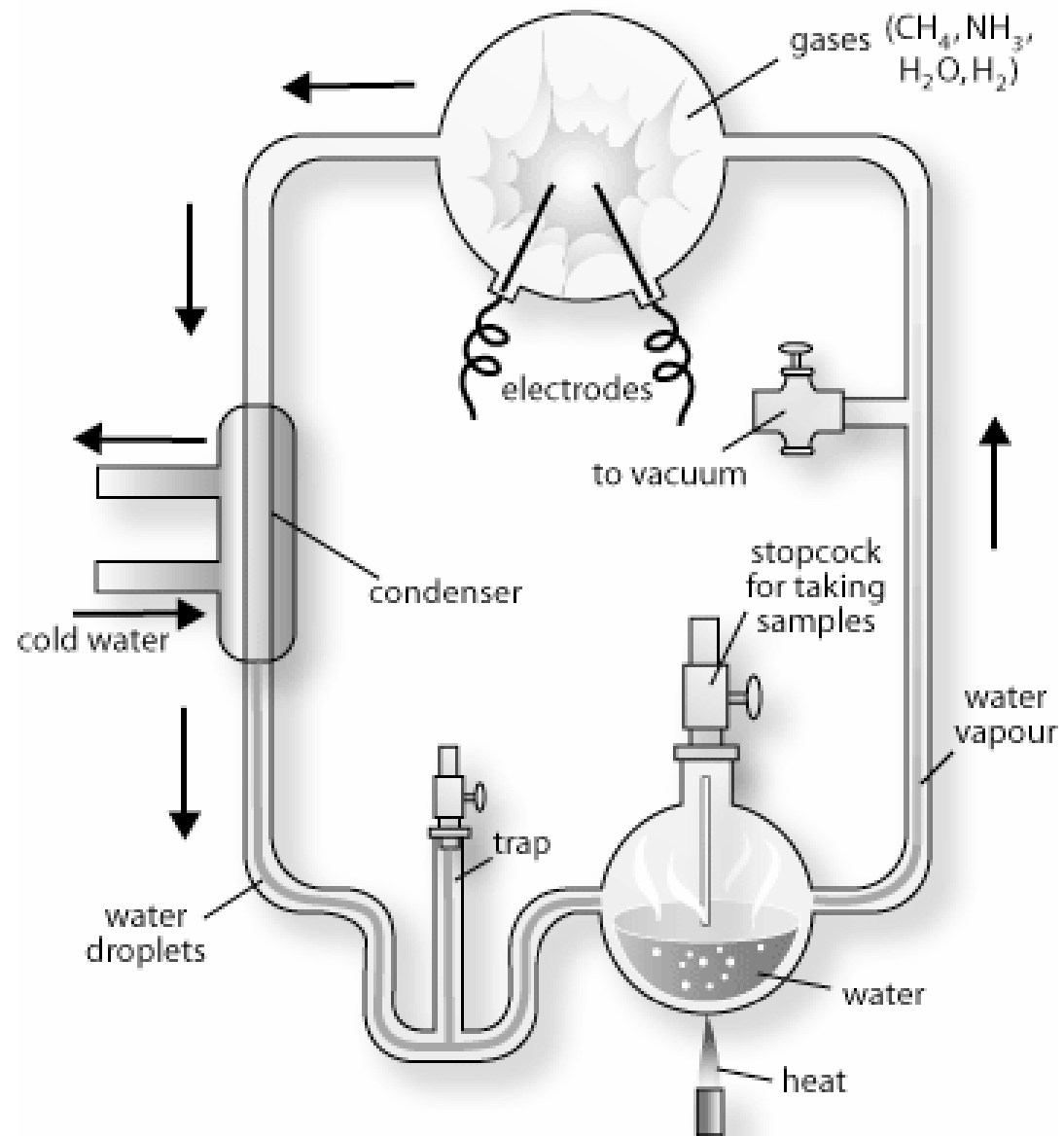
3. Energy Sources

Ultraviolet Light (No Ozone)

Lightning

Geothermal (Lava, Hot Springs, Vents, ...)

# Miller -Urey Experiment

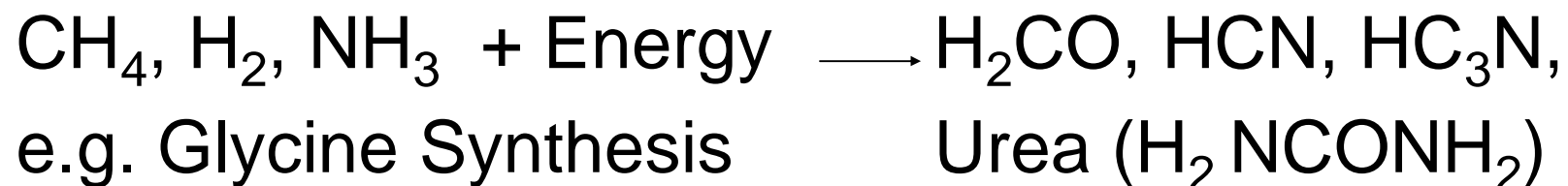




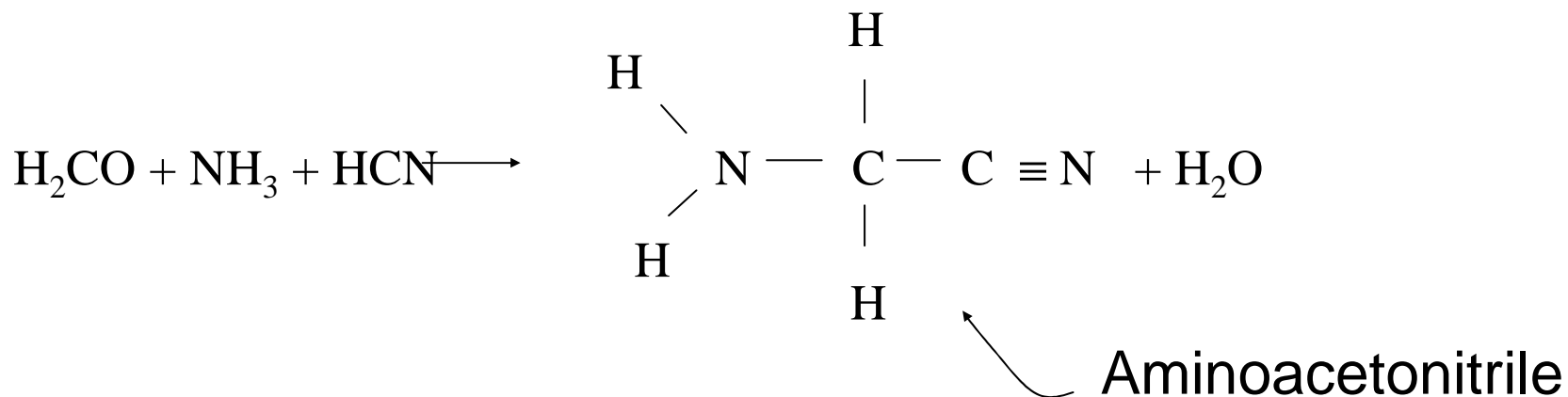
<u>COMPOUND</u>	<u>Relative Yield</u>
<b>Glycine</b>	270
Sarcosine	21
<b>Alanine</b>	145
N-methylalanine	4
Beta-alanine	64
Alpha-amino-n-butyric acid	21
Alpha-aminoisobutyric acid	0.4
<b>Aspartic acid</b>	2
<b>Glutamic acid</b>	2
Iminodiacetic acid	66
IminoaceticOpropionic acid	6
Lactic acid	133
Formic acid	1000
Acetic acid	64
Propionic acid	56
Alpha-hydroxybutyric acid	21
Succinic acid	17
Urea	8
N-methyl urea	6

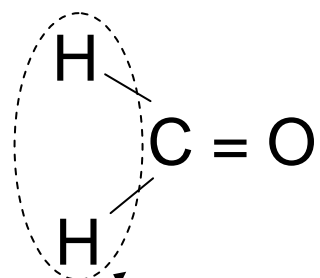
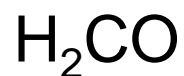
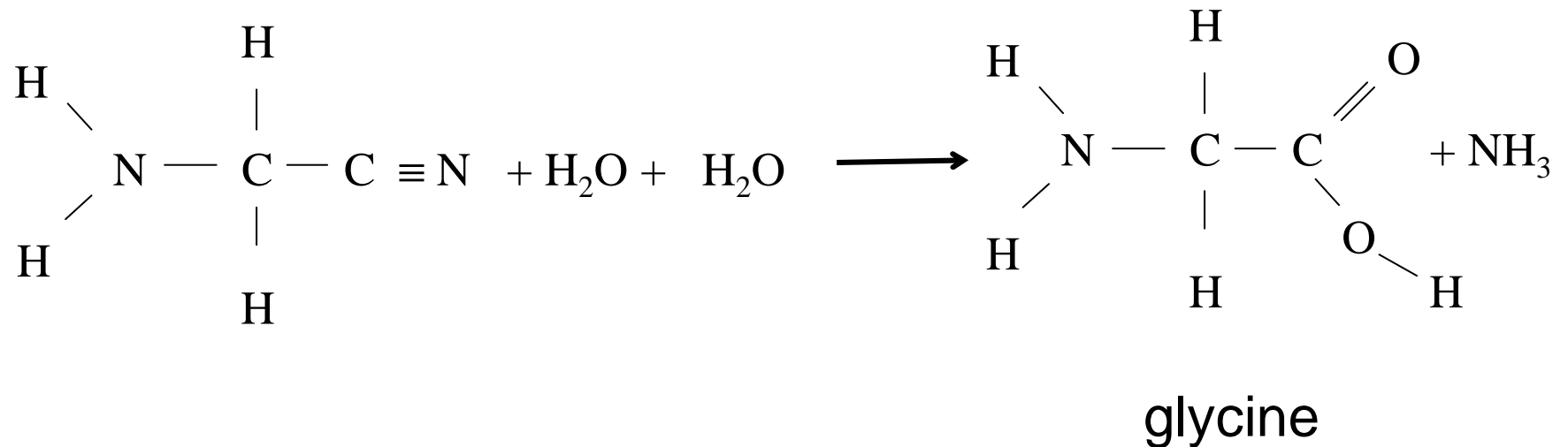
# How did Amino Acids form in Miller -Urey Experiment?

## Strecker Synthesis



Reactive





form Aldehyde

More complex group - other aldehydes

more complex amino acids

Lower yield if atmosphere was  $\text{N}_2$ ,  $\text{CO}_2$ ,  $\text{H}_2\text{O}$

(If  $\text{H}_2/\text{CO}_2 > 2$ , get good yield)

## Problems with Miller - Urey

Atmosphere was  $\text{N}_2$ ,  $\text{CO}_2$ ,  $\text{H}_2\text{O}$

$\text{NH}_3$ ,  $\text{CH}_4$  would react  $\longrightarrow$   $\text{N}_2$ ,  $\text{CO}_2$

Try  $\text{N}_2$ ,  $\text{CO}_2$ ,  $\text{H}_2\text{O}$  in Miller - Urey simulation

Only get trace amounts of glycine

Need  $\text{CH}_4$  to get more complex amino acids

Need  $\text{H}_2/\text{CO}_2 > 2$  to get much of any amino acid

## Miller - Urey with Cosmic Rays

A group in Japan has obtained good yields of amino acids from slightly reducing gases  
(CO<sub>2</sub>, CO, N<sub>2</sub>, H<sub>2</sub>O)

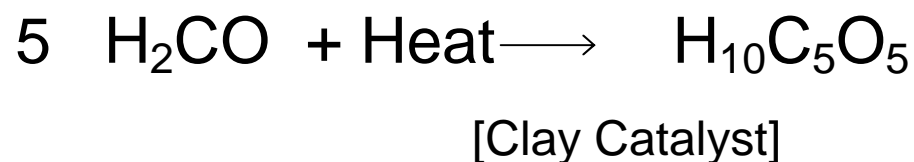
When they used high energy protons  
(simulate cosmic rays)

Apparently not Strecker Synthesis  
(Low abundance of aminoacetonitrile)

# Building Blocks of Nucleic Acids

Not formed in Miller - Urey But some intermediates were

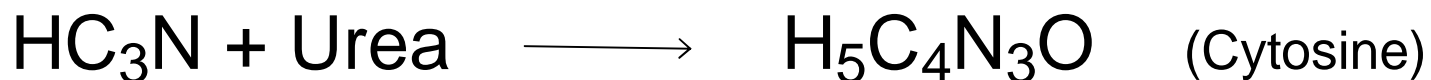
## 1. Ribose Sugar:



## 2. Bases



b) Pyrimidines



### 3. Phosphate

Rock Erosion

Less understood than amino acids

Other Possibilities:

Seafloor Vents

Interstellar Molecules

Comets

## Alternative Delivery

Molecular clouds - strongly reducing, contain many molecules used in Miller-Urey ( $\text{H}_2$ ,  $\text{NH}_3$ ,  $\text{H}_2\text{O}$ ,  $\text{CH}_4$ ) and intermediates ( $\text{HCN}$ ,  $\text{H}_2\text{CO}$ ,  $\text{HC}_3\text{N}$ ) and possibly glycine

Problem: These would not have survived in part of disk where Earth formed

But interstellar ices  $\longrightarrow$  comets

Evidence from similar molecules

(e.g.  $\text{C}_2\text{H}_2$ ,  $\text{CH}_4$ ,  $\text{HNC}$ , ...)

Clearly indicates interstellar chemistry



Cratering record on moon, ...

⇒ heavy bombardment early in history

Comets and their debris could have brought  
large amounts of “organic” matter to Earth  
(and maybe oceans)

Some evidence for non-biological amino acids  
in layer deposited after asteroid impact 65  
million years ago

# Sources of Organic Molecules

Quantitative comparison by Chyba & Sagan, Nature  
1992, Vol. 355, p. 125

Currently, Earth accretes  $\sim 3.2 \times 10^6 \text{ kg y}^{-1}$  from  
interplanetary dust particles (IDP)

$\sim 10\%$  organic carbon  $\Rightarrow 3.2 \times 10^5 \text{ kg y}^{-1}$

$\sim 10^3 \text{ kg y}^{-1}$  comets

$\sim 10 \text{ kg y}^{-1}$  meteorites

$\sim 10^3 \times$  more at  $4.5 \times 10^9 \text{ yr ago}$  (?)

(cratering record)

UV + reducing atmosphere  $2 \times 10^{11} \text{ kg y}^{-1}$

But if  $\text{H}_2/\text{CO} \lesssim 0.1$  IDP's dominant source

# Alternative Sites

## Locally reducing environments

### 1. Ocean vents

Sources of  $\text{CH}_4$  and  $\text{H}_2\text{S}$

Current Vents have ecosystems based on energy from chemicals - not photosynthesis

$\text{H}_2\text{S} \rightarrow \text{Bacteria} \rightarrow \text{Clams, Tube Worms}$

Pre-biotic amino acid synthesis?

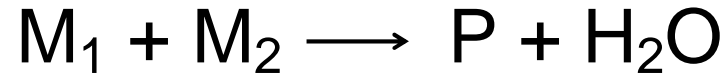
## 2. Inside Earth

Many bacteria now known to live deep (~ 2 miles) in Earth. Again, on chemicals, adapted to high temperature genetic makeup very ancient

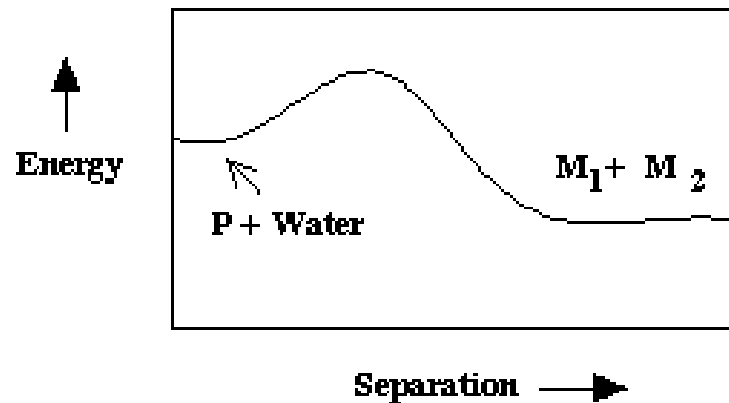
## 3. Hot Springs

Bacteria may be important in precipitating minerals. again adapted to high T and ancient

# Synthesis of Polymers



← more likely in liquid  $H_2O$



Solutions

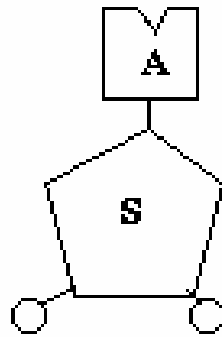
Remove  $H_2O$  (Drying, Heat)

Sydney Fox  $\longrightarrow$  Proteinoids

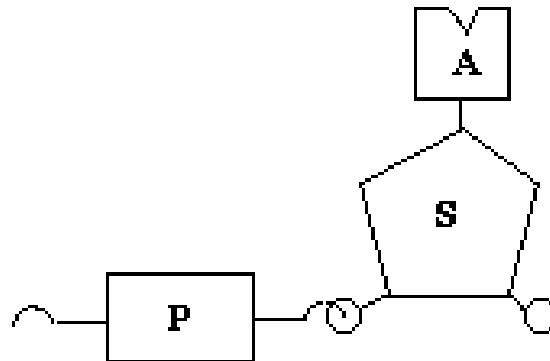
Energy Releasing Reactions ( $H_2NCN$  or  $HC_3N$ )

Catalysts: Clays

Problem is worse for Nucleic acids because more complex



nucleoside



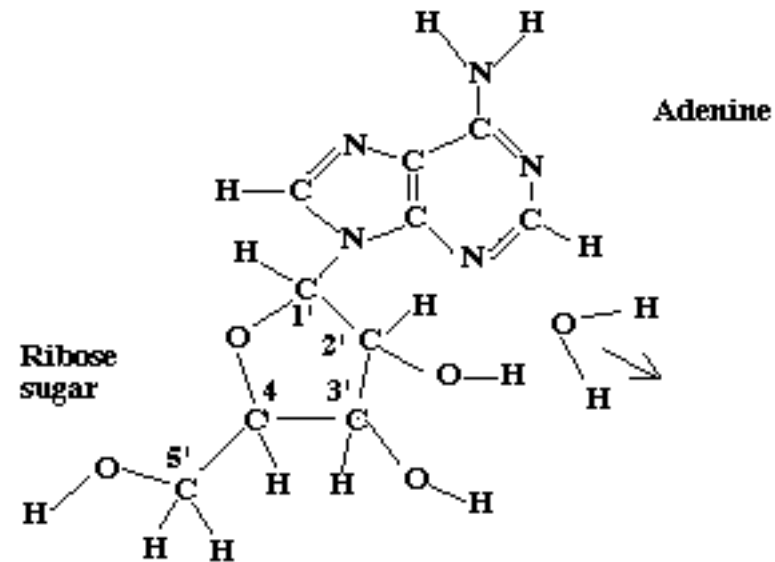
nucleotide

Monomers of nucleic acids



# Synthesis of Adenosine

Base on 1' Carbon (Why?)

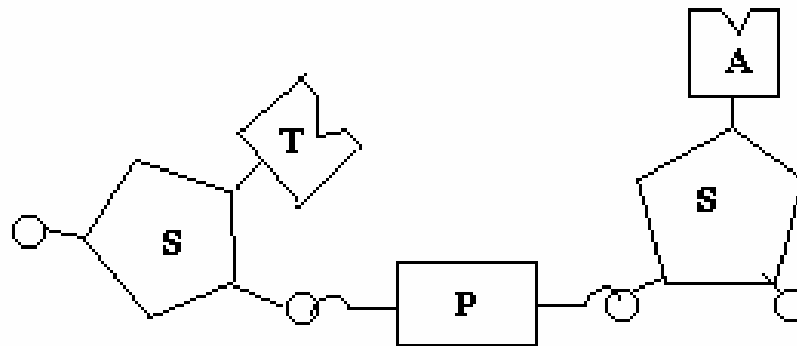


Also phosphates 3' & 5' carbons

Otherwise



Misalignment



Leslie Orgel has had some success in getting high percentage of correct linkages, in presence of Zinc ions.



## Experimental Results

Sugar + base + heat yield some nucleosides

Activated nucleosides + phosphoric acid +  $\text{Zn}^{+2}$

Get polymers up to 50 nucleotides in length

linkages (mostly) correct