

Gene - Its nature, regulation & manipulation

Definitions of gene :->

- (I) It is a segment of nucleic acid usually DNA, rare RNA.
- (II) It occupies a specific locus on chromosome.
- (III) It has a unique sequence of nucleotide base pair.
- (IV) It carries coded information for a specific polypeptide.
- (V) Can undergo crossing over & mutations.
- (VI) It may have continuous or split information.
- (VII) It is able to replicate itself and can produce phenotypic expression.

Cistron :-> It was introduced by Benzer in 1955. It is a segment of the DNA, which carry information for the production of one polypeptide chain.

★★ Types of gene :->

- (i) Housekeeping or Constitutive genes :->  
The genes which are constantly transcribed and expressed in all the cells, because the products they code are needed constantly for



cellular activities.

(ii) **Non-constitutive or luxury gene** :-> The genes which remain inactive and only express only in certain cells, where they need any product.

(iii) **Structural genes** :-> The genes which code for mRNAs which in turn synthesizes specific polypeptides on the ribosomes.

(iv) **Regulator gene** :-> The gene which code for repressor protein for regulating the transcription of cistron.

(v) **Operator gene** :-> The gene which acts as a switch to turn on and off the transcription of a structural gene as cell requires.

(vi) **Promoter gene** :-> The genes or sequences on DNA where RNA polymerase binds for the transcription of structural proteins.

(vii) **Split genes** :->

**Introns** :- The genes which are transcribed but not translated.

**Exons** :- The genes which are transcribed as well as translated.



**Split genes**: The genes which carry information in pieces rather than a continuous stretch are called split genes.

(viii) **Overlapping genes**: → Some genes code for two different polypeptides called overlapping genes. For eg: in some Bacteria & viruses.

(ix) **Transposon's (Jumping genes)**: → In repetitive DNA sequences change their position in DNA are called jumping genes.

★ **Regulation of gene expression**: The rate of protein synthesis is regulated by genetic apparatus and environmental factors. In accordance with the need of a cell. The mechanism which stimulate the expression of certain genes and inhibit the expression of others as required in the cells is called control or regulation of gene expression.

**Regulation of gene expression in Prokaryotes**: Bacteria can regulate the synthesis of enzymes in a way that the enzymes are produced only when the substrates are there to act upon.

**Lactase or Lac operon Hypothesis**: This hypothesis was given by Francois Jacob and



Jacques Monod in 1961, for the regulation of gene action as a result of their studies on the metabolism in *E. coli*.

★ Operon: It is a segment of DNA which is composed of one or more adjacent structural gene, an operator gene, a promoter gene and a regulated gene. Operon works in co-operation with Repressor protein and induces substances.

Operon are of two types:

(I) Inducible. →

(II) Repressible.

A regulator gene code for a protein is Repressor.

Repressor combine with operator gene to repress or stop its action. So it is called Regulator protein. It stops transcription of structural genes by turning off the operator gene.

★ Inducer: → It is a chemical substance which inactivate the repressor by modifying its structure. Inducer join the repressor form Repressor - Inducer complex, this complex prevents the repressor to operator gene of operators.



I Inducible Operon :-> This operon is switched on when a chemical, called Inducer is present. Lactose or Lac-operon of E. coli is an example of Inducible operon. It has following components.

(I) Structural gene :-> When Lactose is added to the E. coli culture. The structural genes produce specific polypeptides which acts as enzyme to catalyse the lactose. In the cell, the operon has 3 structural genes, Lac Z, Lac Y, Lac A, which lie adjacent to each other. which further transcribe mRNA and translate into enzymes i.e.  $\beta$ -galactosidase, Lactose permease, Transacetylase.

(II) Operator gene :-> The gene which controls the functioning of structural genes. It lies adjacent to the structural gene. This gene turned on by an Inducer and not expressed when operator gene is turned off by a Repressor. The Repressor blocks the binding of RNA polymerase to the promoter, so gene turned off.

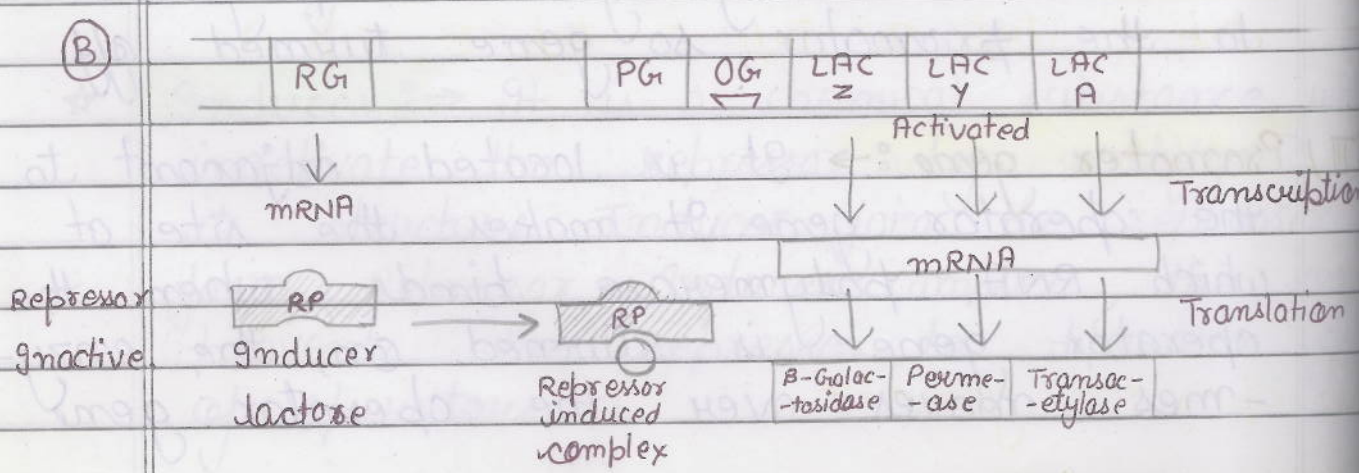
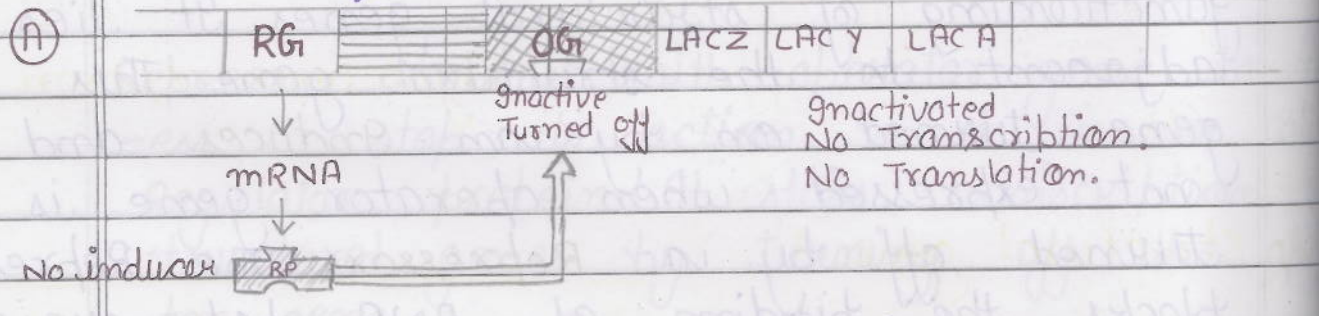
(III) Promoter gene :-> It is located adjacent to the operator gene. It makes the site at which RNA polymerase binds, when the operator gene is turned on, the enzymes moves over the operator gene.



to reach the structural genes and transcription starts. Promoter gene determine DNA strand act as a template for transcription.

(IV) **Regulator gene** :-> This gene control the operator gene in co-operation with chemical compound induces present in the cytoplasm.

Repressor - Induces complex this complex prevents the repressor to bind with operator gene of operon. This frees the operator gene so that RNA polymerase can move from promoter to structural gene, then they transcribe mRNA and further change to enzyme, which are required.





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Result of lactose addition to *E. coli* culture:  
 Some lactose molecules are carried into the cell by the enzymes Lactose Permease molecules. As a small amount of this enzyme is present in the cell even when the operon is not working. These few lactose molecules are converted into active form which act as Inducers and bind the repressor - Inducer complex which fails to join with operator gene, which in turned on. The three structural genes are expressed as three enzymes to metabolise lactose to galactose & glucose.

Inducible operon function in catabolic pathway

(II) Repressible operon: → The operon is switched off when a chemical substance called co-repressor is present. Tryptophan operon of *E. coli* is example of Repressible operon.

(i) Structural genes: The tryptophan operon comprises 5 structural genes denoted as tup E, tup D, tup C, tup B, tup A they lie close to each other on chromosome. These genes form mRNA and then enzymes, which convert raw material into tryptophan.

(ii) Operator gene: → This gene is lying adja-



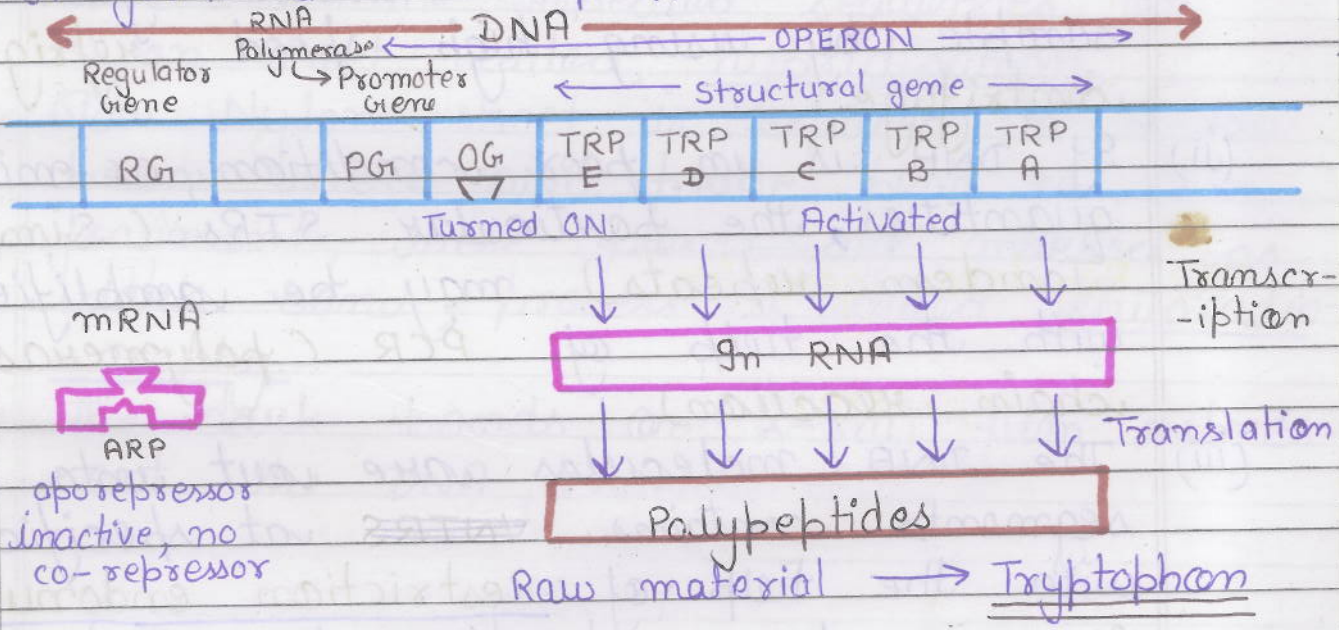
- adjacent to structural gene. It remains turned on, because it is not joined by the corepressor protein produced by the Regulator gene. So the structural gene produces enzymes. They lack catalyse the synthesis of Tryptophan from raw material.

(III) Promoter gene  $\Rightarrow$  This gene lies to the operator gene. It marks the site where RNA polymerase enzymes bind. When operator gene is turned on, this enzyme moves over the operator to structural gene & transcription starts.

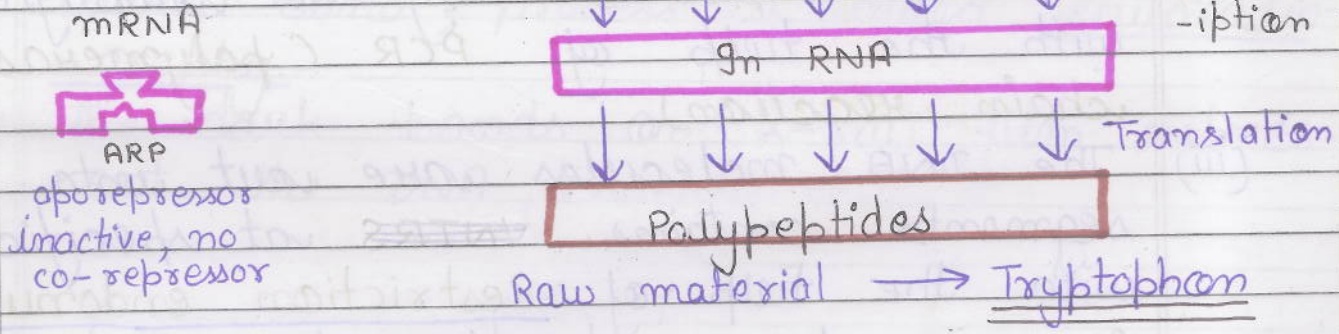
(IV) Regulator gene  $\Rightarrow$  It controls the operator gene in co-operation with a substance called co-repressor. The co-repressor may be end product of metabolism or received by the cell from outside. The regulator gene codes for a protein called corepressor and form complex and later join the operator gene, which is turned off. The structural genes stop transcription when organism gets tryptophan in excess from outside. It conserves raw material and energy by suspending the synthesis of this compound, in its own turn. This phenomenon is called feedback repression.



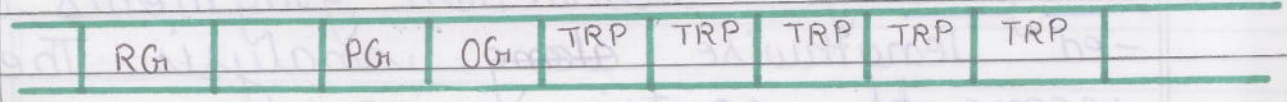
Co-repressor :- It is a non-protein compound which may come from outside or from metabolism within the cell. Its concentration controls transcription in the cell, if conc. of co-repressor is low, aporepressor-co-repressor complex is formed, the complex joins the operator gene and stop the production. If the co-repressor level falls, aporepressor fails to get co-repressor and does not join operator gene, it form the Co-repressor.



(I)



(II)





## "DNA Finger Printing"

In 1977, it was invented by Sir Alec Jeffreys.

\* Sources of DNA finger printing: → The source of DNA may be obtained from the tissue of blood, semen, urine, skin, saliva, cigarette buds, vaginal swab, single hair etc.

Procedure for DNA finger printing: →

- (i) The DNA molecules are isolated from the sample by using high speed refrigerated centrifuge.
- (ii) If DNA is in poor condition, or minute quantities, the particular STRs (Simple Tandem repeats) may be amplified with the help of PCR (polymerase chain reaction).
- (iii) The DNA molecules are cut into small segments containing VNTRs at specific sites with the help of Restriction endonucleases for restriction fragments analysis. These segments contain VNTRs.
- (iv) Now DNA fragments are arranged on agarose gel slab by electrophoresis. Fragments are arranged lengthwise along gel and electric charges.
- (v) The double stranded DNA now split into single stranded by using alkaline chemical.



- (vi) Now place the Nylon nitrocellulose sheet on agarose gel slab. So DNA fragments copied on it with the help of southern blotting technique (Invented by E.M southern)
- (vii) Now DNA probes are prepared in lab, which contains repeated sequences of nucleotides complementary to those on VNTRS.
- (viii) These probes are made radioactive which bind to the repeated sequences on Nylon sheet called hybridisation.
- (ix) Now Nylon sheet is exposed to R-ray beam where DNA probe bind to DNA fragment, these places are marked as dark band, process is called autoradiography.
- (x) The dark bands on x-ray film represent the DNA fingerprint.



66 DNA : The genetic material 99

Evidences or Proof that DNA is Genetic Material

★ Experiment 1 :->

Griffith's and Avery's transformation experiment :->

In 1928 S.F. Griffith experimented with Streptococcus pneumoniae (Diplococcus) a bacterium that causes Pneumonia.

There are two strains of bacterium:

- (a) A smooth virulent strain (S-strain), protected by polysaccharides capsule and cause Pneumonia.
- (b) A rough harmless strain (R-strain) without capsule not cause pneumonia.
- (I) When rough bacteria injected with live smooth bacteria, mice died due to pneumonia.
- (II) When rough bacteria injected to mice, did not die.
- (III) Mice injected heat-killed smooth bacteria mice survived.
- (IV) Now mice injected with mixture of live rough bacteria and heat-killed smooth bacteria. It cause Pneumonia and killed the mice.



(V) Griffith found smooth bacteria in the blood of dead mice from this he concluded that a chemical substance transform from the death, smooth bacteria, into rough bacteria and later transform into smooth bacteria.

(VI) These transformed smooth bacteria recovered from the dead mice, when grown in culture produced smooth bacteria.

(2) Bacteriophage experiment :->

Bacteriophage :-> A virus which infect bacteria and kill them is called Bacteriophage.

In 1952 Alfred D. Hershey and Martha Chase did experiment with bacteria Escherichia coli and bacteriophage T<sub>2</sub>, both inhabiting in human intestine.

(i) They are different radio active isotopes to label DNA and Protein. Ist they label Protein coat of Bacteriophage T<sub>2</sub> with radioactive Sulphur S<sup>35</sup>. Because protein contain sulphur and DNA does not contain sulphur.

(ii) Now T<sub>2</sub> bacteriophage used to infect normal bacteria.

(iii) Now protein coats were separated from bacterial cell walls by shaking and centrifugation.

(iv) It is found that radioactivity was associated with protein coats & none was inside the bacterial cell, this shows



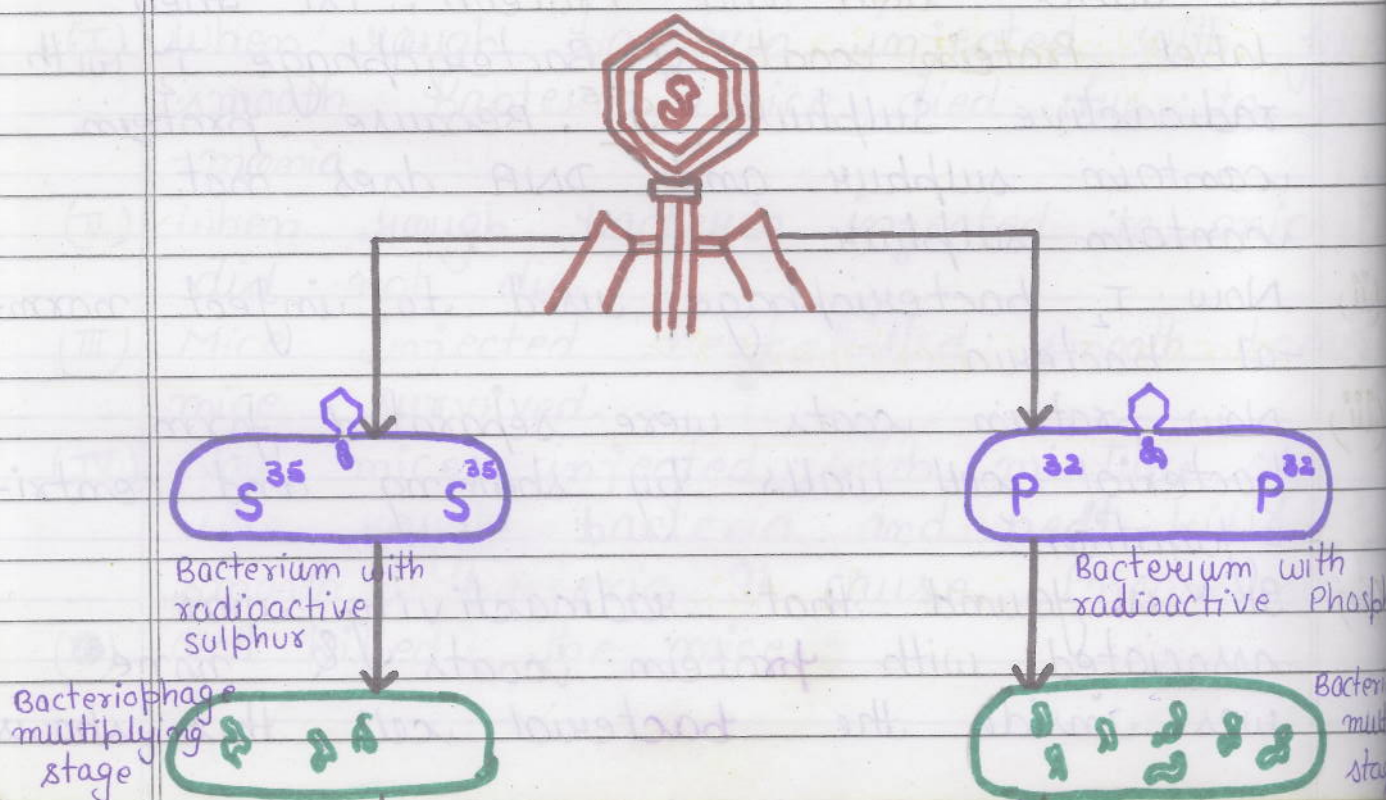
that protein coat of Bacteriophage did not enter inside bacterial cell.

★★ In second case :->

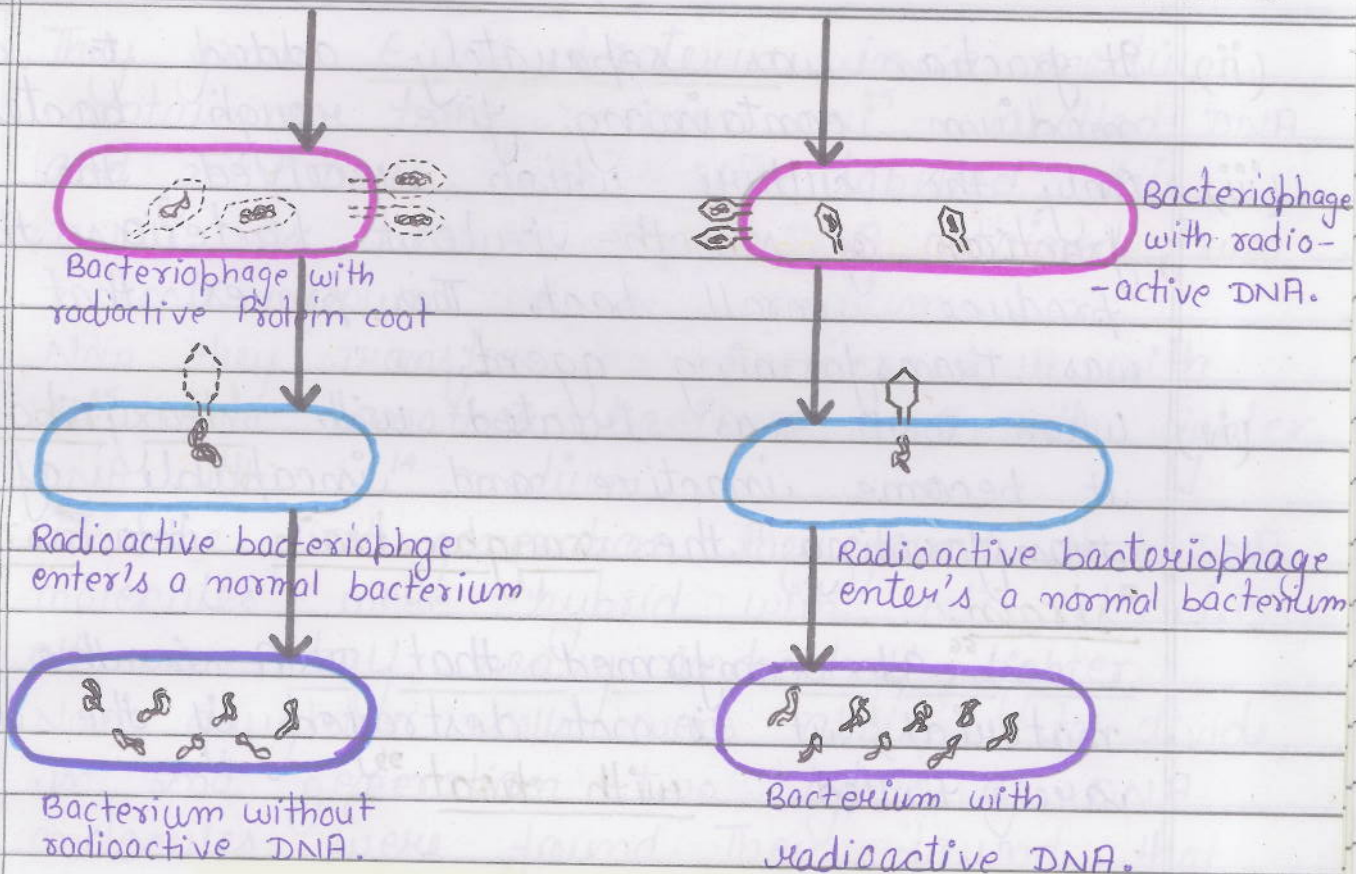
(i) Now Hershey & Chase did experiment with DNA core. Now DNA core of bacteriophage is labelled with radioactive  $P^{32}$ , Phosphorus, because DNA contain Phosphorus and Protein does not contain it.

(ii) Now bacteriophage used to infect the bacteria.

(iii) After centrifugation, it was confirmed that radioactivity was observed inside the bacterial cell, it proves that DNA is the material which pass to bacterial cell not protein. Hence it is proved that DNA is the genetic material.







Experiment of Hershey and Chase to show that DNA is the Hereditary or genetic material in the Bacteriophage  $T_2$ .

© Transformation  $\Rightarrow$  It is a phenomenon in which DNA is isolated from R-strain of bacteria, introduced from S-strain of bacteria is called transformation.

AVERY'S Experiment  $\Rightarrow$  In 1944, O.T. Avery, C. Macleod and M. McCarty.

(i) They separate the extract of smooth, virulent bacteria, into protein, DNA and carbohydrates fraction.



- (ii) It fraction was separately added to culture medium containing five rough bacteria.
- (iii) Only the culture which received the DNA fraction of smooth-virulent bacteria that produce small bact. This proves that DNA was transforming agent.
- (iv) When DNA was treated with deoxyribonuclease it became inactive and incapable of transforming the rough strain into smooth strain.
- It confirmed that DNA is the genetic material, it is not destroyed if the cells are killed with heat.

### ★★ Replication and Synthesis of DNA

It is the process of synthesis of another daughter DNA from its parent DNA is called replication.

Mode of Replication :-> Mode of replication is said to be "Semi-conservative"

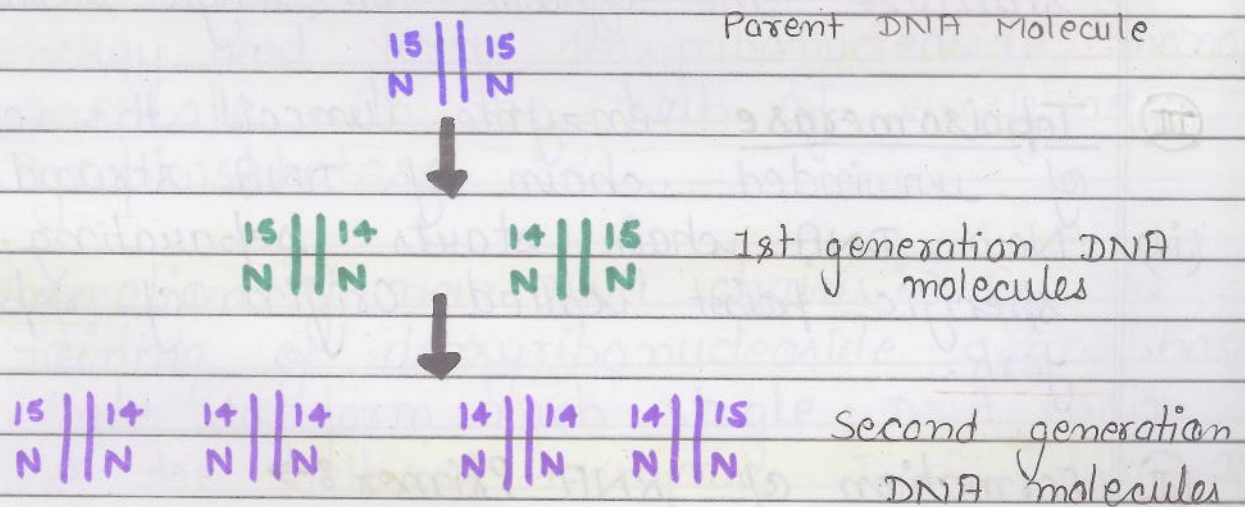
Replication :-> Occurs in nucleus in eukaryotes. In prokaryotes it occurs in cytoplasm.

Discovery :- Semiconservative mode of DNA was demonstrated by Taylor in 1957.

★★ Experimental evidences was given by Meselson and Stahl in 1958.



1. They grow E. coli bacterio in vo medium containing heavy nitrogen  $N^{15}$  - labelled DNA, and get DNA with  $N^{15}$  - labelled, this DNA was heavier than DNA obtained from E. coli grown in  $N^{14}$  - medium.
2. Now they transfer the heavier DNA with  $N^{15}$  into another bacterial cell with lighter DNA with  $N^{14}$  medium.
3. In the first generation they found that DNA molecules were hybrid with  $N^{15} - N^{14}$ . i.e. all were half heavy and half lighter.
4. Now daughter cells were allowed to divide, in 2nd generation two types of DNA molecules were found. They found that 50% half heavy with  $N^{15} - N^{14}$  hybrid and 50% light with  $N^{14} - N^{14}$  density. This contains the semiconservative mode of replication



★ Mechanism for DNA replication  $\Rightarrow$  Mechanism of replication involves complex process which require many enzymes, Protein



factors and metal ions.

### (I) Activation of Deoxyribonucleotides :->

There are four types of deoxyribonucleoside monophosphate are found, they are, de dGMP, de CMP, de TMP. (Adenosine, Guanosine, Cytidine, thymidine, monophosphates). are activated by enzymes Phosphorylase producing phosphate & form de ATP, de GTP, de CTP & de TTP.

### (II) Exposure of parent DNA bases :->

(i) The parent DNA double strand uncoils & splits into single DNA strand by breaking of Hydrogen bonds.

(ii) The unwinding is done by enzyme Helicase by using ATP and Non-enzymatic single stranded DNA-binding Protein (SSB) to stabilize the chain in single strand.

(iii) Topoisomerase enzyme uncoil the coiling of unwinded chain of DNA strand.

(iv) Now DNA chain starts separating at a specific point called origin of replication fork.

### (III) Formation of RNA Primer :->

(i) A short chain of RNA is formed on the DNA template of the 5<sup>th</sup> end is called RNA Primer.



Ligase

Pyrophosphatase

Helicase

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- (ii) RNA Primer is formed with the help of enzyme RNA Primase. Because enzyme DNA Polymerase cannot initiate the synthesis of new DNA strand.
- (iii) DNA Primer is formed at free ends of one DNA strand and at the fork end of other strand, later primer removed and gaps filled to form DNA strand.
- (iv) **Base Pairing**  $\Rightarrow$  All the bases like Deoxyribo-nucleoside triphosphates joined by the pairing rule of Watson & Crick i.e. A-T, G-C, C-G and T-A.
- (v) **Conversion to Deoxyribonucleoside monophosphate**  $\Rightarrow$   
Now the deoxyribonucleoside triphosphates join to each other at each single DNA chain, breaks off their inner high energy and form deoxyribonucleoside monophosphate with the help of enzyme Pyrophosphatase.
- (vi) **Formation of new DNA chains**  $\Rightarrow$  By the joining of deoxyribonucleoside monophosphate to form each single DNA chain with the help of enzyme DNA Polymerase  $\alpha$  and  $\delta$  and with metal ions  $Mn^{+2}$  and  $Mg^{+2}$ . They produce the double chains which are identical to each other.



## ★ Leading and Lagging strands :

(1) **Leading strand**  $\Rightarrow$  The DNA Polymerase polymerise the deoxyribonucleotides in the 5'-3' direction. Because the two strands of DNA are anti-parallel, so new strand formed on old strand in opposite direction from 5'-3' is called leading strand.

★★ **Okazaki Fragments** : On the other parent strand, short DNA segments are formed in 5'-3' direction, starting from RNA Primers. These DNA segments are called Okazaki fragments.

(2) **Lagging strand**  $\Rightarrow$  A separate RNA primer formed for the synthesis of each Okazaki fragment. These fragments joined together and form continuous strand that is lagging strand.

## (VII) Editing (Proof-reading) and DNA repairs :

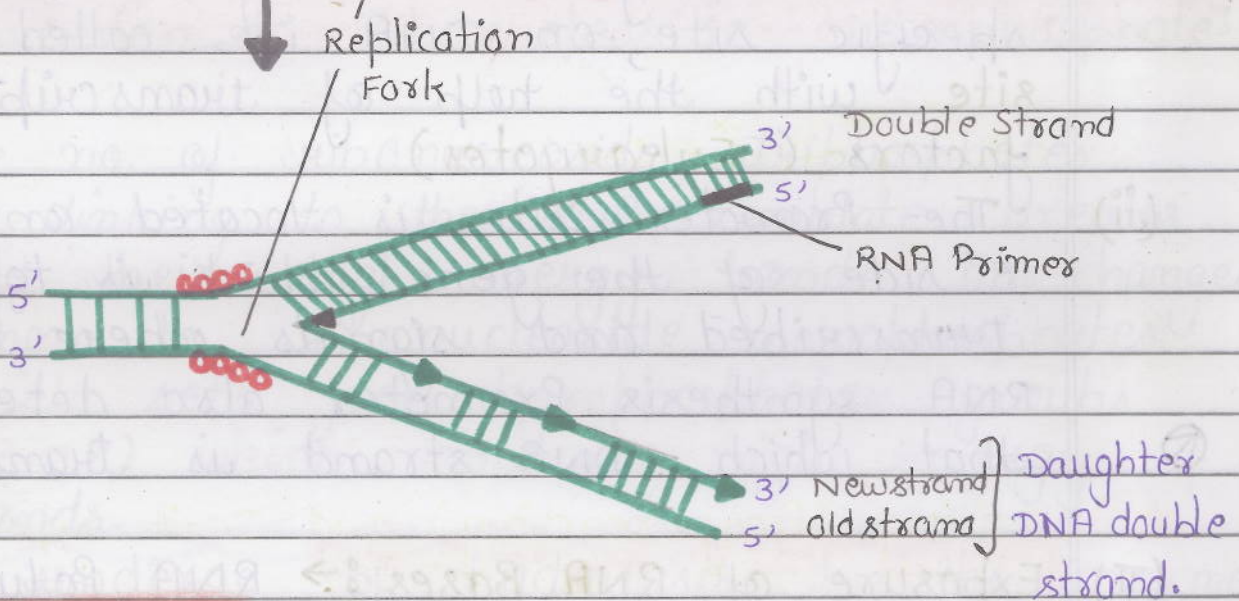
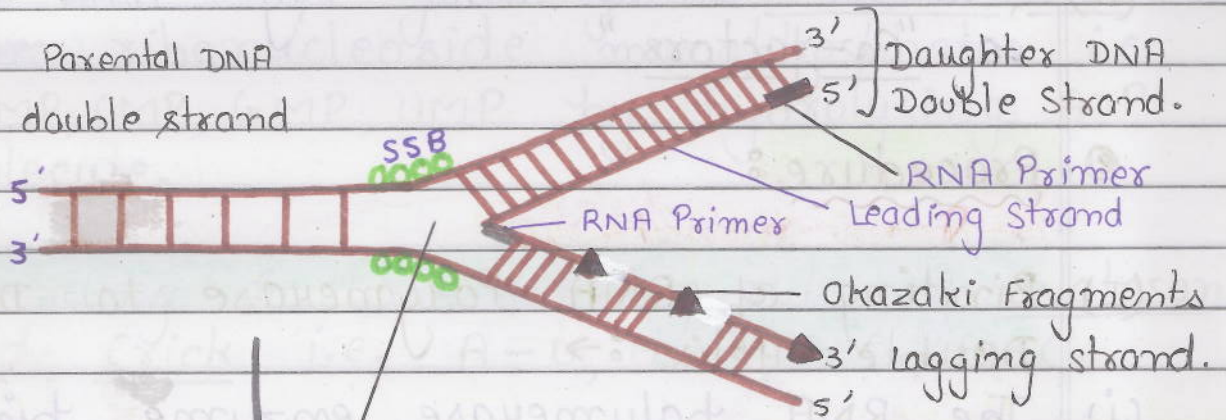
(i) To know the accurate base pairing of replication, sometime wrong bases do get inserted, they are noted and removed by DNA Polymerase.

(ii) Abnormal region of DNA are enclosed by enzymes Nucleases and missing segments of DNA strand resynthesized by DNA Polymerase.



(VIII) Helix Formation : Each daughter's double DNA strand become spirally coiled to form Double Helix.

★★ Diagrammatic representation :->



★★ Transcription :-> It is the process of synthesis of RNA from any one of DNA template is called Transcription.

Discovery : It was discovered by Jeraxel Hurwitz, Samuel B. Weiss and Audrey Stevens in 1950 in vitro experiment.



## Mechanism of Transcription :->

### Material required =>

- (i) Enzyme "RNA Polymerase."
- (ii) DNA template.
- (iii) Four types of ribonucleoside triphosphates.
- (iv) Divalent metal ions  $Mg^{+2}$  and  $Mn^{+2}$  as "Co-factors".

### ⊙ Procedure :

#### (I) Binding of RNA Polymerase to DNA Double Helix :->

(i) The RNA polymerase enzyme binds to a specific site on DNA i.e. called Promoter site with the help of transcription factors (Eukaryotes).

(ii) The Promoter site is located on the 5' side of the gene which is to be transcribed and signals where to start RNA synthesis. Promoter also determines that which DNA strand is transcribed.

#### (II) Exposure of RNA Bases :-> RNA Polymerase moves along the DNA & unwinds the DNA complex into two chains. In the

⊙ region of gene which is to be transcribed. This exposes the A, T, C & G. Only one strand is called sense strand of DNA and other strand acts as anti-sense and non-coding strand.



(III) **Base pairing**  $\Rightarrow$  The Ribonucleoside triphosphates, i.e. Adenosine triphosphate, Guanosine triphosphate, Cytidine triphosphate & Uridine triphosphate (ATP, GTP, CTP, UTP). act as raw material are activated with the help of enzyme Phosphorylase ~~from~~ ribonucleoside monophosphates i.e. AMP, CMP, GMP, UMP by hydrolysing ATP molecule.

According to base pairing rule of Watson and Crick i.e. A-U, U-A, C-G, G-C.

(IV) **Conversion to Ribonucleoside monophosphate:**

- (i) The no. of ribonucleoside triphosphates on linking to the DNA template breaks off their high energy bonds. This changes them to ribonucleoside monophosphates which sets free pyrophosphates groups (P<sub>2</sub>P) which contain high energy bonds.
- (ii) It undergoes by hydrolysis by the enzyme Pyrophosphatase.

(V) **Formation of RNA chain**  $\Rightarrow$

- (i) Each ribonucleoside monophosphate joined to DNA template the ribonucleotide arrived earlier, make the RNA chain longer.



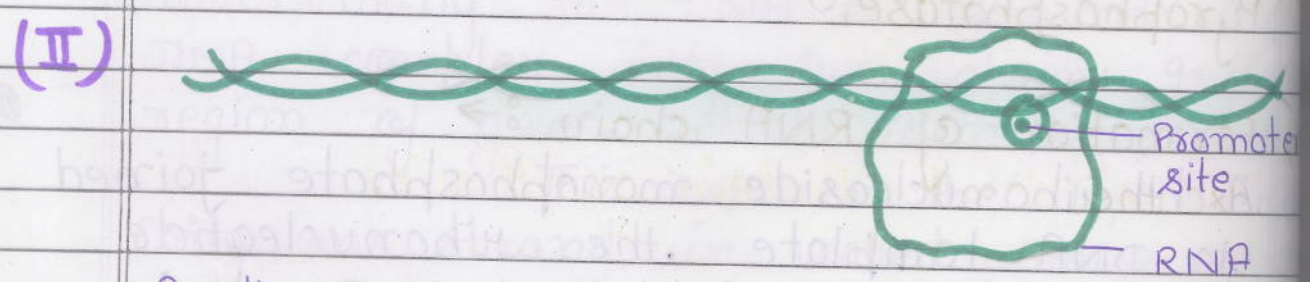
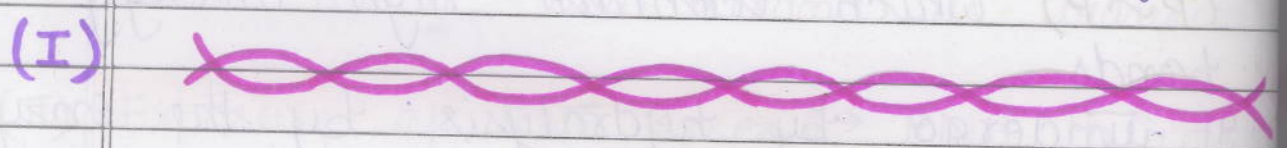
(ii) The process is catalysed by enzyme Polymerase and divalent ions  $Mg^{2+}$  or  $Mn^{2+}$ .

(VI) Separation of RNA chain :->

(i) As the transcription proceeds, RNA molecule dissociates from DNA.

(ii) When the polymerase reaches to terminator signal on the DNA, it leaves DNA, one gene forms several molecules of RNA, which are released from the DNA.

(VII) Return of DNA segment to original  
As the chain grows and separated from DNA, the DNA molecule gets hydrogen bonded to the mis-sense strand, and two are spirally coiled to assume the double helix form.

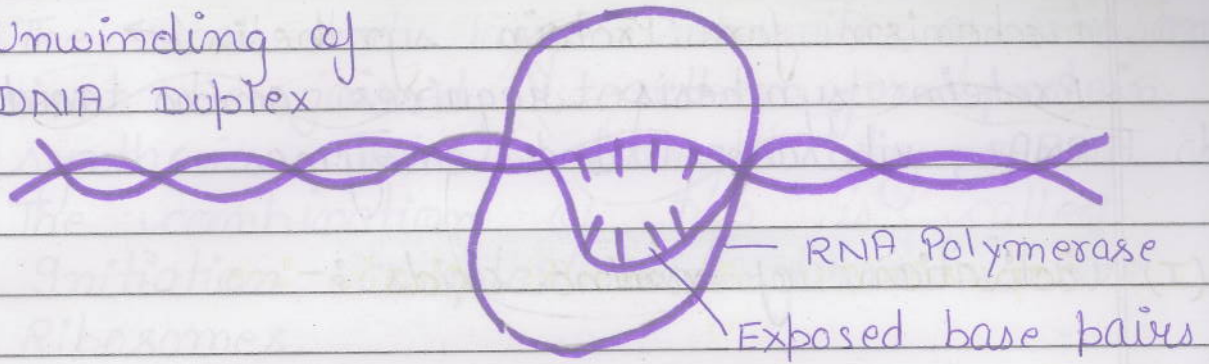


Binding and RNA with DNA duplex

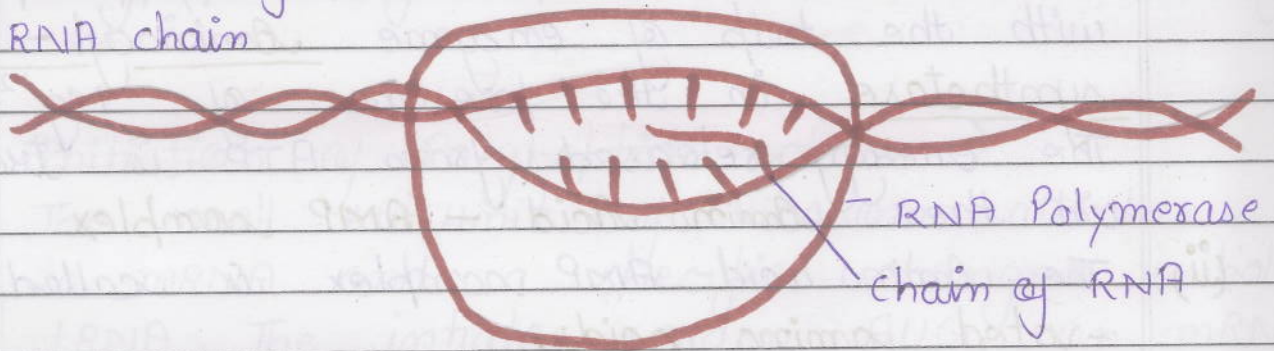
Polymerase Transcription factors



(III) Unwinding of DNA Duplex



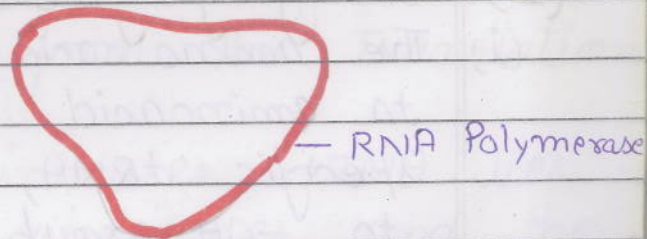
(IV) Initiation of RNA chain



(V)

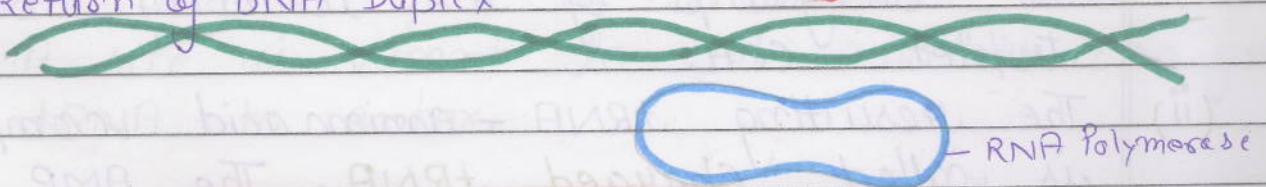


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(VI)

Return of DNA Duplex



Translation: (Protein synthesis)

It is the process by which triplet base sequences of mRNA guides the linking of a specific amino acid to form a polypeptide.



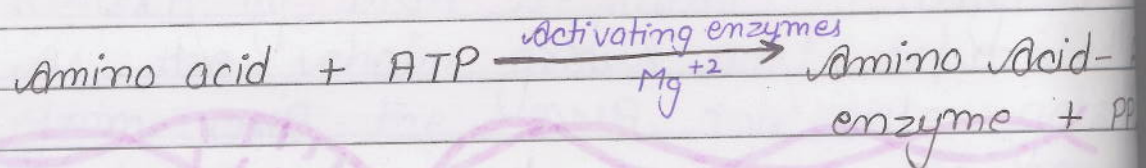
## Mechanism for Protein synthesis :->

Protein synthesis requires amino acids, RNAs, ribosomes and enzymes.

### (I) Activation of amino acids :-

(i) The amino acids react with ATP to form amino acid-AMP complex and Pyrophosphate with the help of enzyme Aminocyl-synthetase, in the presence of  $Mg^{+2}$ . The energy released from ATP is trapped in the amino acid-AMP complex.

(ii) The amino acid-AMP complex is called activated amino acids.



### (II) Charging of tRNA :-

(i) The amino acid-AMP-enzyme complex binds to amino acid binding site of the specific tRNA, where its  $-COOH$  group reacts to  $-OH$  group of the terminal base triplet CCA.

(ii) The resulting tRNA-amino acid complex is called charged tRNA. The AMP and enzymes are freed, these freed enzymes can activate and attach another amino acids.

### (III) Activation of Ribosomes :-



The small and large subunits of ribosome must be joined together for protein synthesis. This is brought by mRNA chain. The combination of two is called initiation complex and form active Ribosomes.

(IV) Polypeptide formation :- It involves 3 stages :-

(a) Initiation of Polypeptide chain  $\Rightarrow$

(i) The small subunit of ribosome attaches to mRNA and a specific changes initiates tRNA. The initiates codon AUG on mRNA signals the start of Translation.

(ii) The initiates tRNA join the initiation codon by its anticodon and carries the amino acid methionine. Now the large subunit of ribosomes join the small subunit. This form the translation initiation complex.

(iii) At this stage the initiates tRNA lies at the P site of ribosomes and the A site is vacant to let another charged tRNA to enter.

(b) Elongation of Polypeptide chain :-

(i) In the elongation stage of translation, amino acids are added one by one to the 1st amino acid. It is



done with certain proteins called Elongation factors.

(ii) A charged tRNA anticodon sequence and specific amino acid attached to its carrier end. This tRNA shift to site of ribosomes and A site lies vacant for another charged tRNA.

(iii) With the help of Peptidyl Transferase shifts amino acid on another tRNA at A-site of ribosomes and form bond called Peptide bond. Bond is formed b/w  $-COOH$  grp. and free  $-NH_2$  group another amino acids.

(iv) Now tRNA at A site carrying amino acid move to P-site, the process is called Translocation. It requires energy which is provided by Hydrolysis of GTP to GDP.

(v) Now the mRNA move with its codon to the A site. The uncharged tRNA shift from P site to E site and leaves the ribosomes.

(vi) Now the next cycle starts, a third tRNA with its specific amino acid enters at the A site of the ribosomes. It binds by anticodon and further elongates the chain and further elongation form Polypeptide by repeating the process.

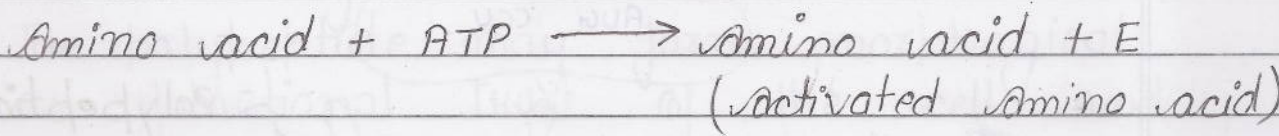
and release of Polypeptide



(i) At the end of mRNA chain there is a stop or terminator codon (UAA, UAG or UGA) reaches at A site and is not coded by anticodon of tRNA, no further addition of amino acids occur.

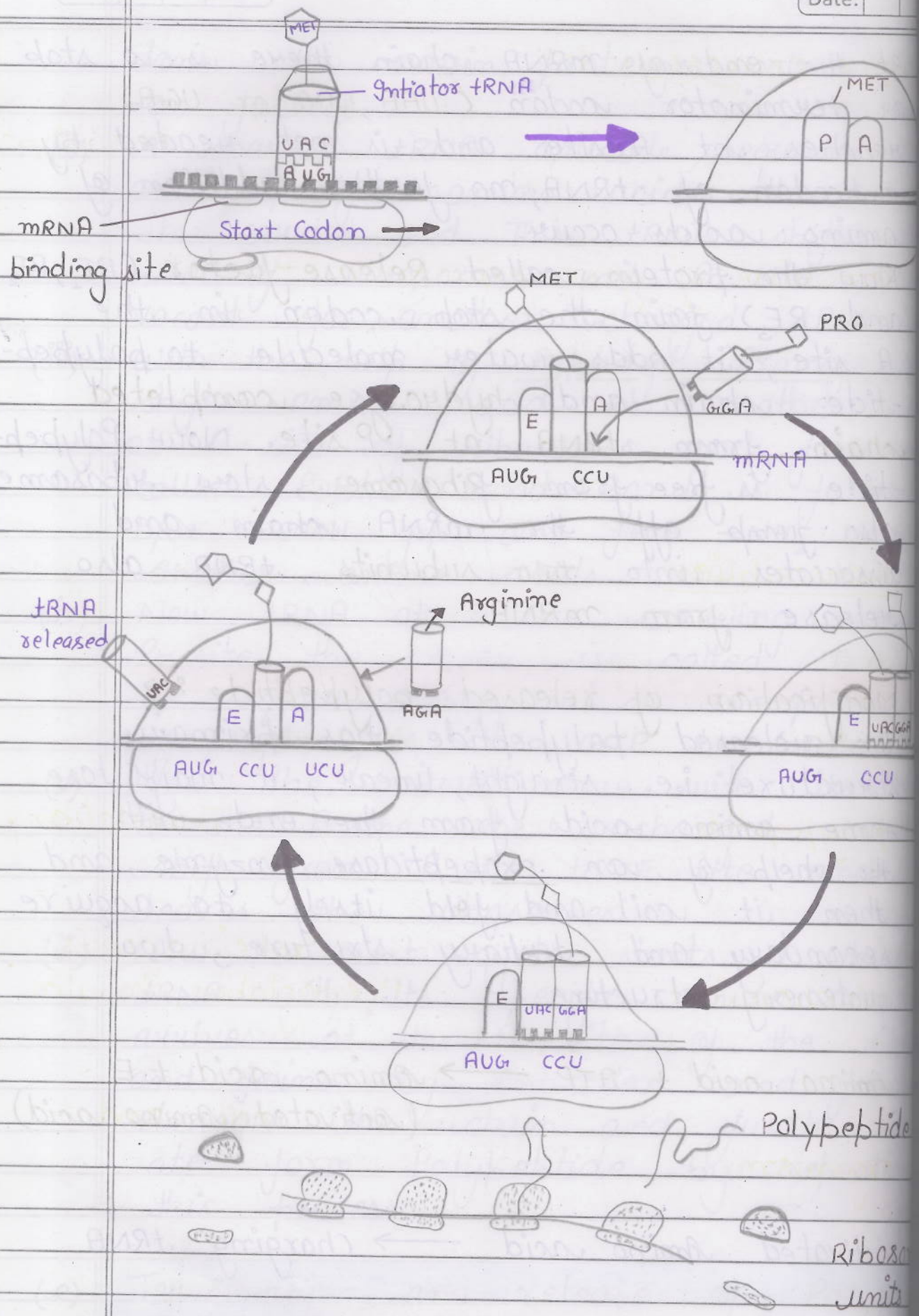
(ii) Now the protein called Release factor ( $RF_1$ ,  $RF_2$  and  $RF_3$ ) join the stop codon in the A site, it adds water molecule to polypeptide chain and hydrolyse completed chain from tRNA at P site. Now Polypeptide is free from ribosomes, small ribosomes also jump off the mRNA chain and dissociates into two subunits, tRNA also release from mRNA.

(d) Modification of released polypeptide :->  
The released polypeptide has primary structure i.e. straight, linear, it may lose some amino acids from the end with the help of an exopeptidase enzyme, and then it coil and fold itself to acquire secondary and tertiary structure, two quaternary structure



Activated amino acid  $\longrightarrow$  Charging tRNA







## ★ Gene expression :->

It is a molecular mechanism by which gene can show its potential in the phenotype of an organism.

## Mechanism of gene expression :->

- (i) A gene contains code for a particular polypeptide in the form of a specific sequence of its base pairs. It transfers its code to mRNA transcribe from it. Only the sense strand of DNA is capable of transcription. Other genes transcribe mRNA and tRNAs.
- (ii) The ribosomes bind with mRNA, and with the help of tRNAs select the required amino acid and links them in proper sequence received from gene to form polypeptide process is called Translation.
- (iii) The polypeptide chain formed may act as a structural protein and form some organelles as form Proteinaceous biochemical such as Haemoglobin, Insulin etc. or enzyme.
- (iv) The polypeptide may form morphological as functional trait of the cell and organism. So the molecular structure of a gene is expressed in a phenotype.



eg: → One gene  $\xrightarrow{\text{Transcription}}$   $\left\{ \begin{array}{l} \text{rRNA} \\ \text{mRNA} \\ \text{tRNA} \end{array} \right. \xrightarrow{\text{Trans-lation}} \left\{ \begin{array}{l} \text{One} \\ \text{Polypeptide} \end{array} \right.$

Polypeptide  $\xrightarrow{\text{EXPRESSION}}$   $\left\{ \begin{array}{l} \text{Organelle} \\ \text{Enzyme} \\ \text{Proteinaceous} \end{array} \right.$

A change in the gene world Biochemical change the code & give rise to different Polypeptide.

### ★ Central Dogma or Flow of Information

"It is the flow of information from DNA to mRNA and hence to a polypeptide is called central Dogma."

- (i) The chromosomal DNA is the 1<sup>o</sup> - information centre. It contains complete information about all the specific proteins, to be synthesized in the cells for their use.
- (ii) The information coded in the DNA the particular sequence of bases will genetic code.
- (iii) The one of the two strands of DNA the sense strand, contain information to guide the synthesis of Polypeptide and a single stranded nucleic acid molecule can carry information from DNA to the site of Protein synthesis.



white

<sup>66</sup> mould - Fungus<sup>99</sup>  
Fungus - Rhizopus

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(iv) The information passed on by the DNA to mRNA and then it direct the synthesis of required polypeptide chain.



Relationship b/w Genes and enzymes :->

The relationship was found in 1902 by Archibald Garrod. He told that the genes operate by producing enzymes. He suggest that each of the disease inherited by offspring from the parents is caused by the lack of an enzyme which change one metabolic substance to another. He called such diseases Inborn errors of metabolism.

Lack of enzyme is due to the absence of normal form of gene which control the synthesis of that enzyme.

★ One gene - One enzyme Hypothesis :->  
(Beadle and Tatum's Hypothesis)

This hypothesis was given by Beadle & Tatum in 1948, with the help of their experiment on the pink bread mould, "Neurospora crassa."

(i) They grow this mould in simple medium containing salt, sugar, amino acids, Purines and Pyrimidines with the help of enzymes. Such a normal or wild type of individual are called Prototroph.



## Melanocyte cells - Melanin

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(ii) They expose this mould to X-ray, which cause the mutation by changing the sequence of DNA. They found that mutants were unable to grow in minimal medium unless additional -ients were added to the medium.

Such mutants are called Auxotrophs.  
(iii) The different mutants need the amino ornithine to citruline to arginine for growth.

(iv) It is known that a cell can sequentially change ornithine to citruline and then arginine, for their growth, they lack enzyme due to mutation in gene, which code this enzyme. Similarly another gene was lack of enzyme which change ornithine to citruline. This mutant need citruline for growth.

(v) This led them to thought that each gene had the information for producing single enzyme, so they proposed their famous One-gene One-enzyme hypothesis in 1948.

This hypothesis states that each gene controls the synthesis of specific enzyme or particular protein.

This hypothesis got them Nobel Prize in 1958.



## \* Modification of Beadle and Tatum's

Hypothesis :-> It has been now found that

- (i) All genes do not produce enzymes, certain proteins contain more than one polypeptide chain. For eg: Haemoglobin molecule consists of two  $\alpha$  and  $\beta$  polypeptide chain. This shows that more than one gene may control the synthesis of a protein.

Now it is stated that one gene is responsible for the formation of one polypeptide chain.

Now one gene - One enzyme hypothesis has been replaced by one gene - one enzyme polypeptide chain principles.

\* Genetic code :-> The information exist as the particular sequences of bases in the DNA strand is called genetic code.

\* 2nd definition: The sequence of base triplets in DNA molecule, each triplet representing one amino acid of a Polypeptide also called genetic code.

\* Characteristics of genetic code :->

- (i) The genetic code is a triplet code. Three adjacent bases termed as codon.



(ii) The adjacent codon's do not overlap, they do not share any base. Each single base is part of only one codon.

AUG UAA  
 III III

C C U, C A U

AUG  
 III

CCU CUC UCA

(iii) The genetic code is commaless. There is punctuation marks b/w coding triplets.

(iv) The genetic code is universal i.e. a codon in the DNA and mRNA specifies the same amino acids in the protein synthesizing systems of all organisms from bacteria to man.

(v) The genetic code is "degenerate" it lacks specificity and one amino acid often has more than one code triplet. Methionine and tryptophan have single triplet codons. All other amino acids are specified by 2-6 base triplets.

eg: Phenylalanine has two codons UUU & UUC  
 Arginine has six codons - CGU, CGC, CGA, CGG, AGA and AGG.

(vi) Each codon codes for only one amino acid, none for more than one.

(vii) Nonsense or Terminator codons. Three of the 64 codons are terminator codons UAA, UAG and UGA. (i) UAA is called the



- (ii) UAG is called Amber (iii) UGA is called OPAL.
- (viii) The codon's AUG and GUG are k/o start codon or Initiation codon.
- (ix) DNA is a linear polynucleotide chain & a protein is a linear polypeptide chain.
- (x) A specific gene transcribes a specific mRNA which produces a specific polypeptide.