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UNIVERSITY

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FACULTY OF ENGINEERING & TECHNOLOGY  
DEPARTMENT OF BIOTECHNOLOGY

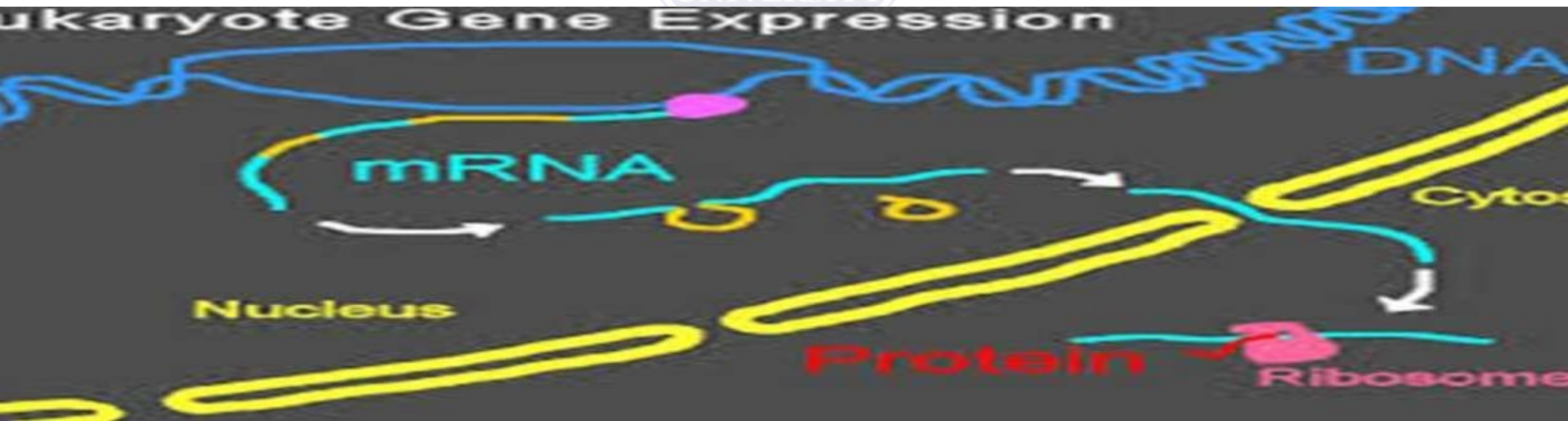
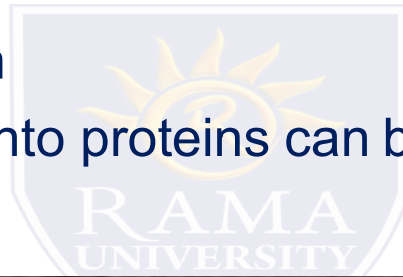
## Eukaryotic Gene Regulation

Eukaryotic cells have a much larger genome Eukaryotes have much greater cell specialization

Thus eukaryotic cells contain an enormous amount of DNA that does not program the synthesis of RNA or protein

This requires complex organization

In eukaryotes expression of gene into proteins can be controlled at various locations



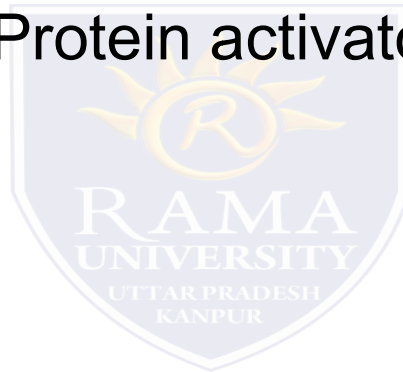
## of Eukaryotic gene Regulation

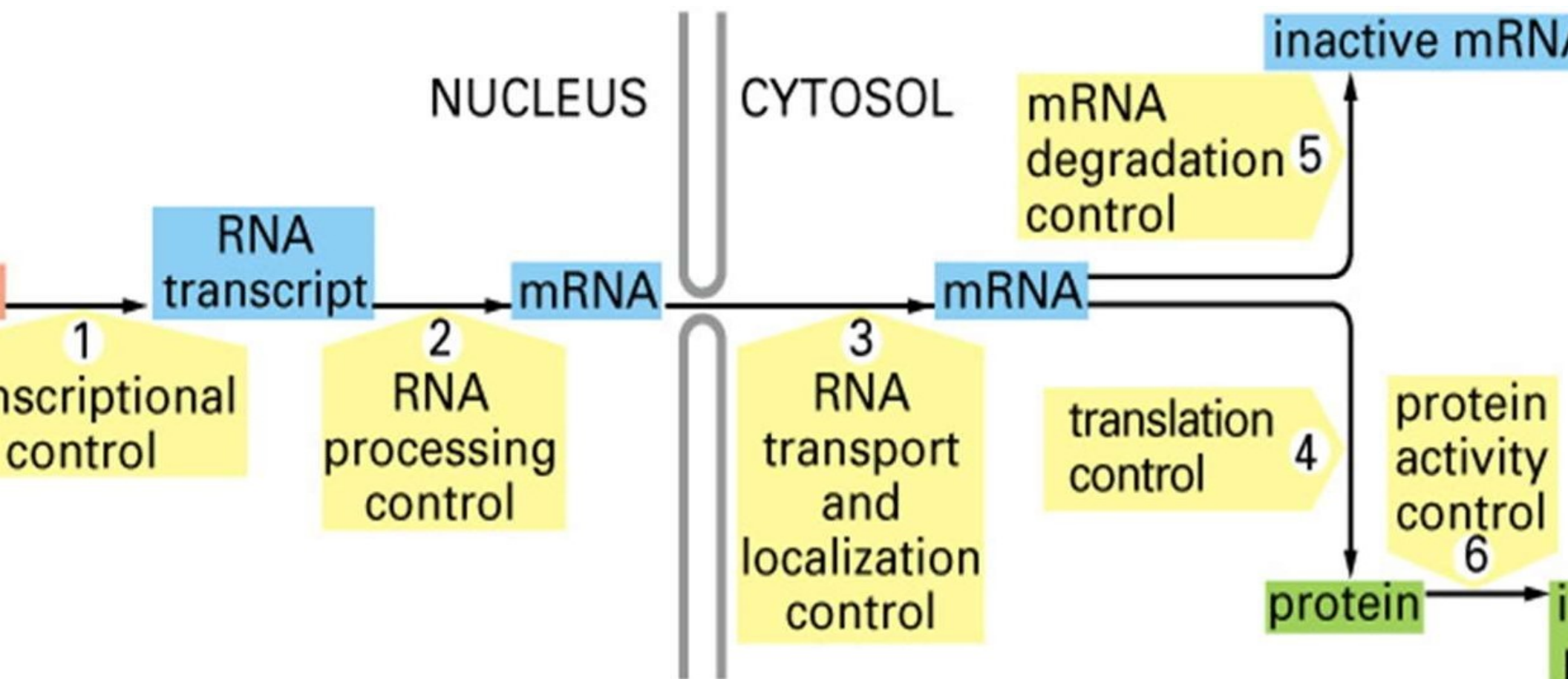
Transcriptional control. 2. RNA processing control.

RNA transport & localisation control

Translation control.

mRNA degradation control. 6. Protein activator control.





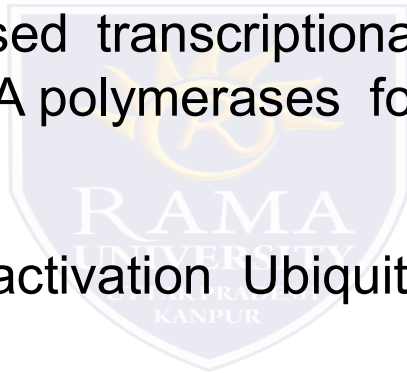
### Two forms of chromatin

Euchromatin – A less coiled transcriptionally active region which can be easily accessed by the RNA polymerases.

Heterochromatin – A highly condensed transcriptionally inactive region. The genes in this region cannot be accessed by the RNA polymerases for active transcription.

### Ubiquitination:

Ubiquitination of H2A – Transcriptional inactivation  
Ubiquitination of H2B -  
Transcriptional activation



## Mechanisms affects the chromatin Structure:

- Histone modifications – These modifications make a region of gene either transcriptionally active or inactive.

### Acetylation

- $\uparrow$ Acetylation ----  $\downarrow$  Condensation of DNA -----  $\uparrow$  Transcription of genes in that region Ubiquitination

Ubiquitination of H2A – Transcriptional inactivation Ubiquitination of H2B - Transcriptional activation

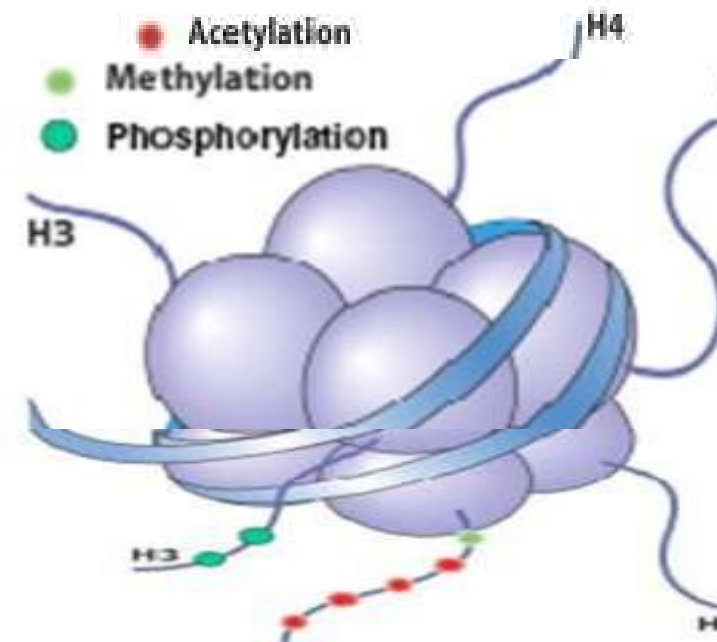
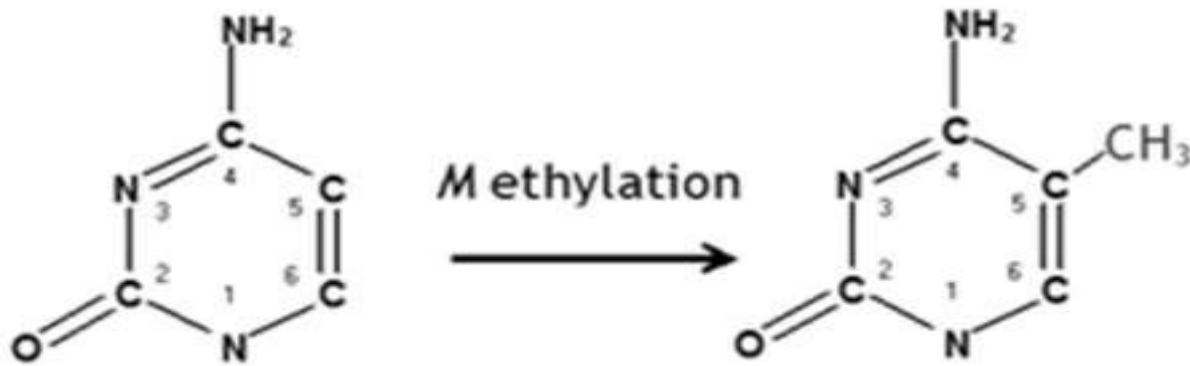
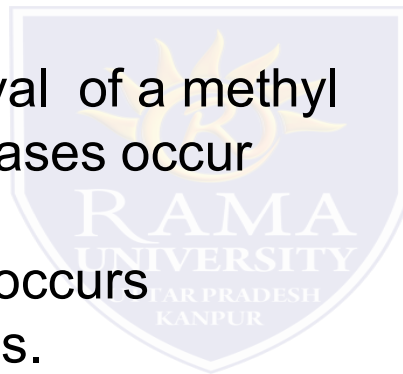
# Methylation

5-methylcytosine: is the addition or removal of a methyl group to cytosine bases occur consecutively.

5-methylcytosines occur consecutively.

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5-methylcytosines occur consecutively. Methylation occurs often in symmetrical CG sequences.





# ACETYLATION :

HATs and coactivators leads to euchromatin formation

# METHYLATION:

HDACs and corepressors leads to heterochromatin formation

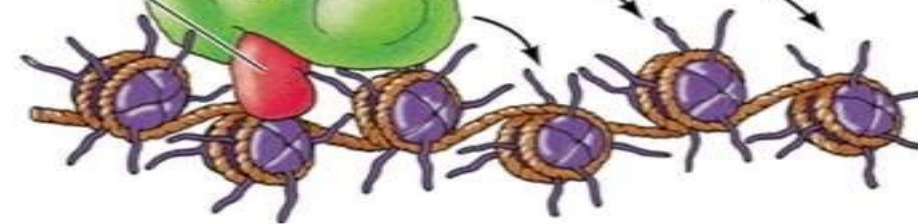


(B)

Coactivator

Activator

HAT

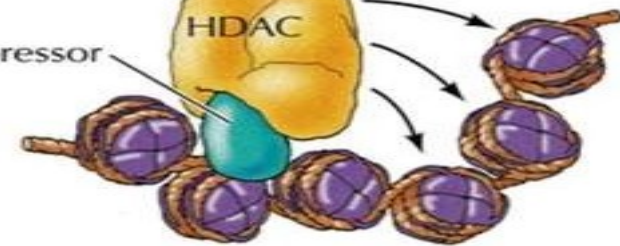


Acetylated histones  
Active chromatin

Corepressor

Repressor

HDAC



Deacetylated histones  
Inactive chromatin



## Regulation of Transcription:

Eukaryotes – There are two types of promoters which are:

**Basal promoters**

**Upstream promoters**

**Basal promoter or core promoter** - These promoters reside within 40bp upstream of the start site. These promoters are seen in all protein coding genes.

**Upstream promoters** - These promoters may lie up to 200bp upstream of the transcriptional initiation site. The structure of this promoter and the associated binding factors keeps varying from gene to gene.

## Transcriptional control:

controlling when and how often a given gene is transcribed

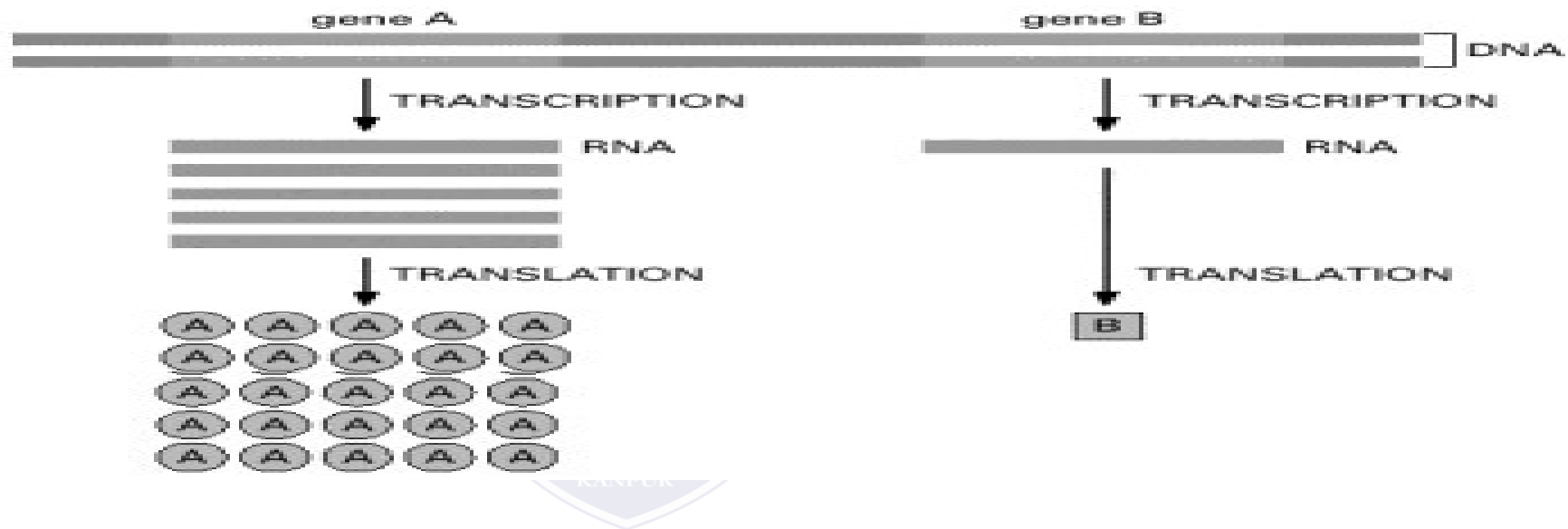
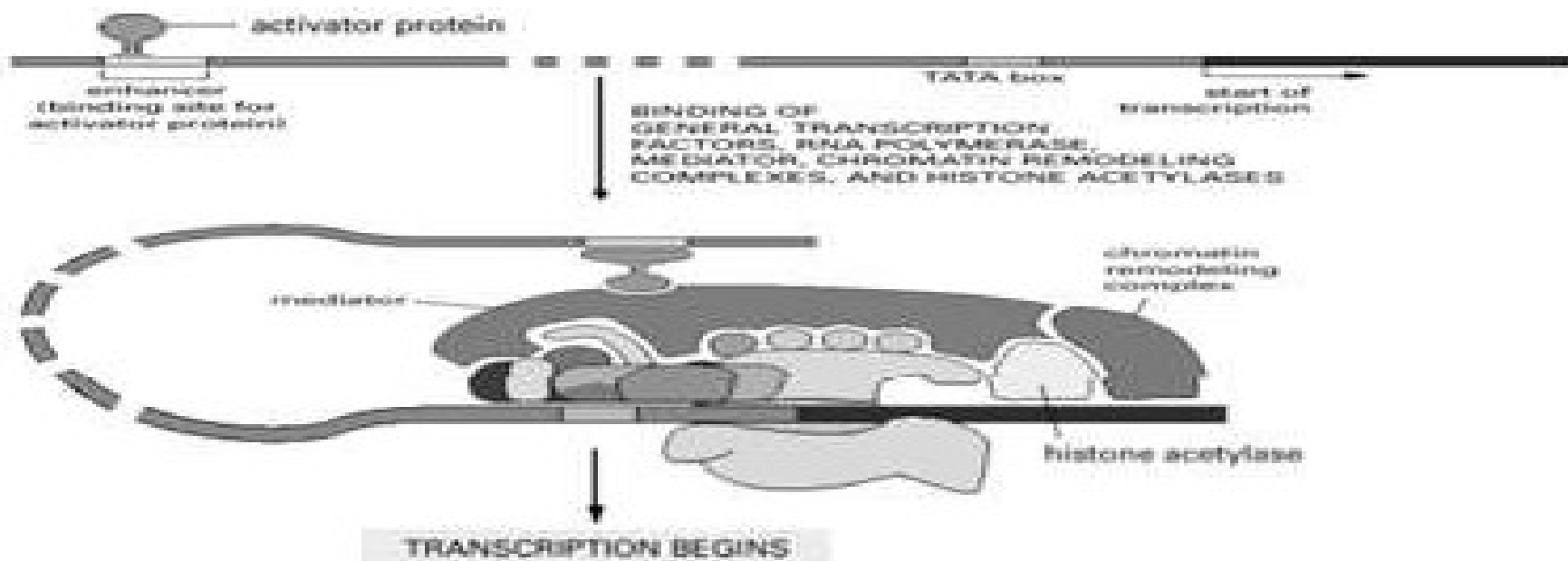


Figure 6. Genes can be expressed with different efficiencies.

Gene A is transcribed and

translated much more efficiently than gene B. This allows the amount of protein A in the cell to be much greater than that of protein B.

# Transcriptional control – regulation by RNA polymerase:



# Enhancers

Enhancers can be located upstream, downstream or within the gene that is transcribed

The binding of these enhancers with enhancer binding proteins (transcription factors) increases the rate of transcription of that gene to a greater extent.

Promoters are capable of initiating lower levels of transcription.

Enhancers are responsible for the cell or tissue specific transcription.

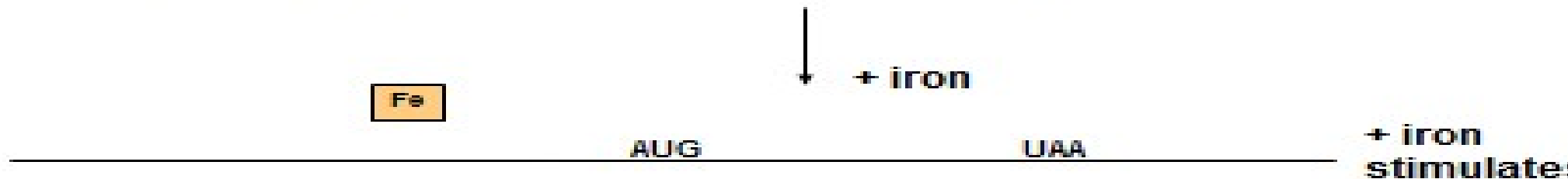
Each enhancer has its own transcription factor that it binds to.

## *Control at mRNA stability*

A stem loop is stabilised by the 90 kDa protein in the absence of iron. This time, the stem loop is at the 5' end of the mRNA.



The presence of the stem loop prevents translation of this mRNA by blocking the progress of the ribosomes along the mRNA.



In the presence of iron, the hairpin is lost, the ribosomes can translate the mRNA and ferritin protein synthesis is increased.