

TOPIC : EPIGENETICS
SUBJECT : BOTANY
SEMESTER : M.Sc. Botany Semester IV
PAPER/COURSE : MBOTEC–1 (Unit III)
Cytogenetic & Crop Improvement



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EPIGENOMICS



Introduction

- An **epigenome** consists of a record of chemical changes to the DNA and the histone protein of an organism.
- These changes can be passed down to the offspring of an organism.
- **How are epigenetics and Epigenomics different?**
 - **Epigenetics** focuses on process that regulates gene expression.
 - **Epigenomics** focuses on the analysis of epigenetic changes across the entire genome

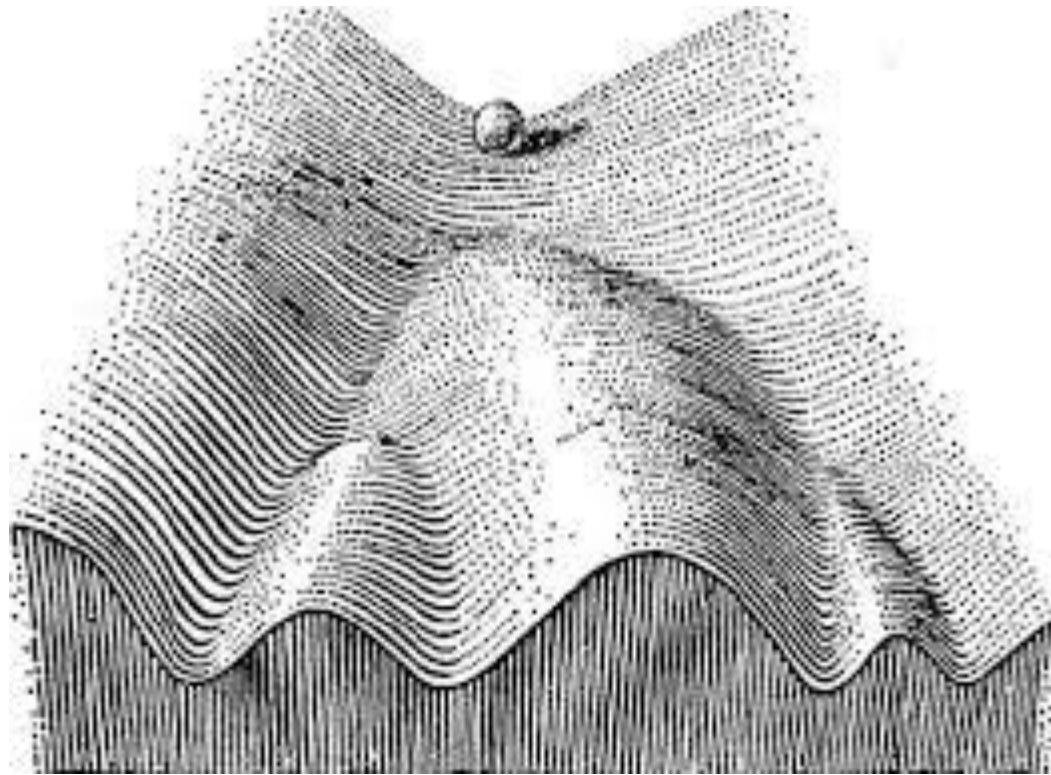
FATHER OF EPIGENETICS



Dr. V

WAL

Waddington(1942)



The image created by Waddington to represent the epigenetic landscape.

The Epigenetic Landscape

- Conrad Waddington was a hugely influential British polymath. He was born in 1903 in India.
- This image was created by Waddington to represent the epigenetic landscape. The position of the ball represents different cell fates. The image is incredibly powerful in helping to visualise what might be happening during cellular development.
- The ball at the top of the hill is the zygote, the single cell that results from the fusion of one egg and one sperm. As the various cells of the body begin to differentiate (become more specialised), each cell is like a ball that has rolled further down the hill and headed into one of the troughs.
- Once it has gone as far as it can go, it's going to stay there. Unless something extraordinarily dramatic happens, that cell is never going to turn into another cell type (jump across to another trough). Nor is it going to move back up to the top of the hill and then roll down again to give rise to all sorts of different cell types.

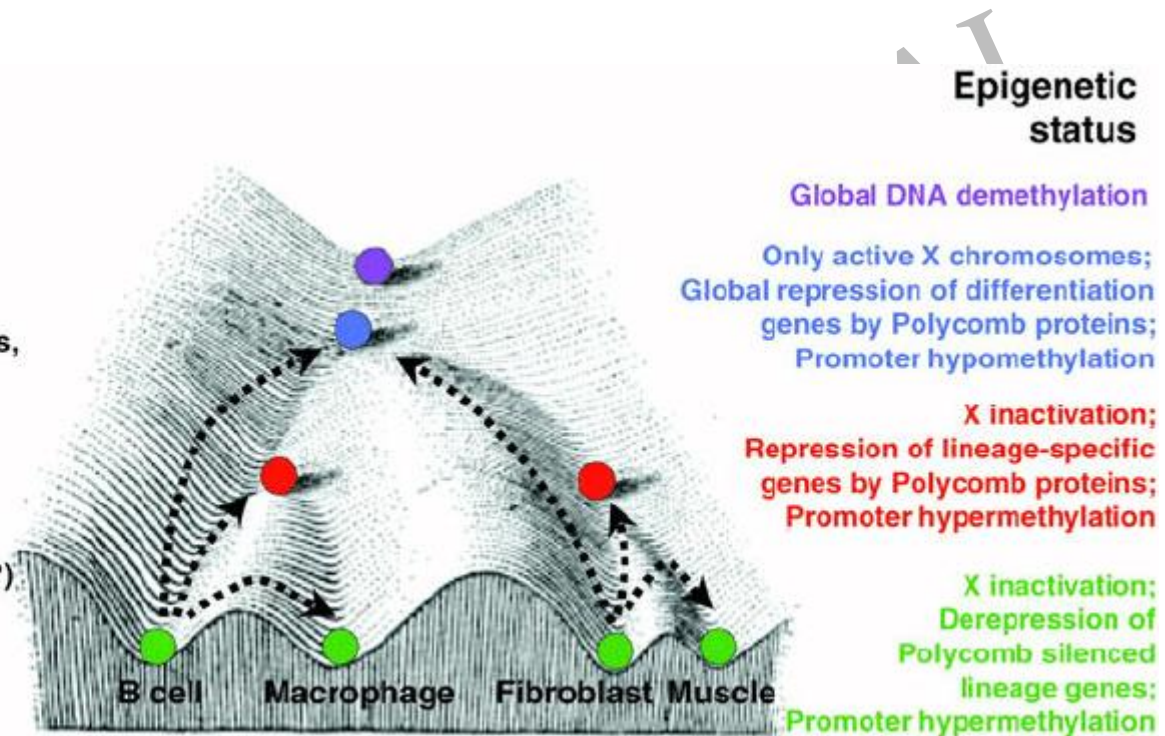
Developmental potential

Totipotent
Zygote

Pluripotent
ICM/ES cells, EG cells,
EC cells, mGS cells
iPS cells

Multipotent
Adult stem cells
(partially reprogrammed cells?)

Unipotent
Differentiated cell types



Explained form of Waddington's landscape

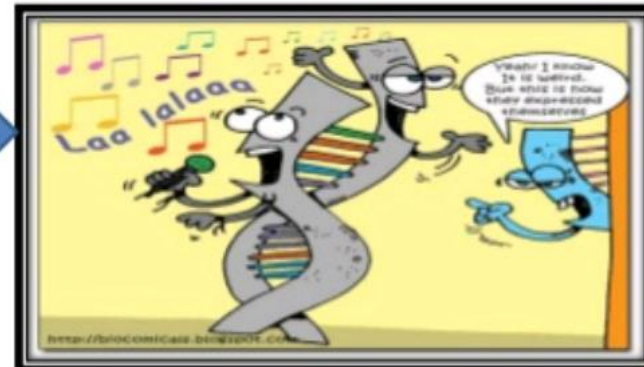
What is EPIGENETICS ?

Epigenetics literally means “above” Or “ on top of” genetics: It refers to external modification to DNA that turns **genes 'on' or 'off'**.



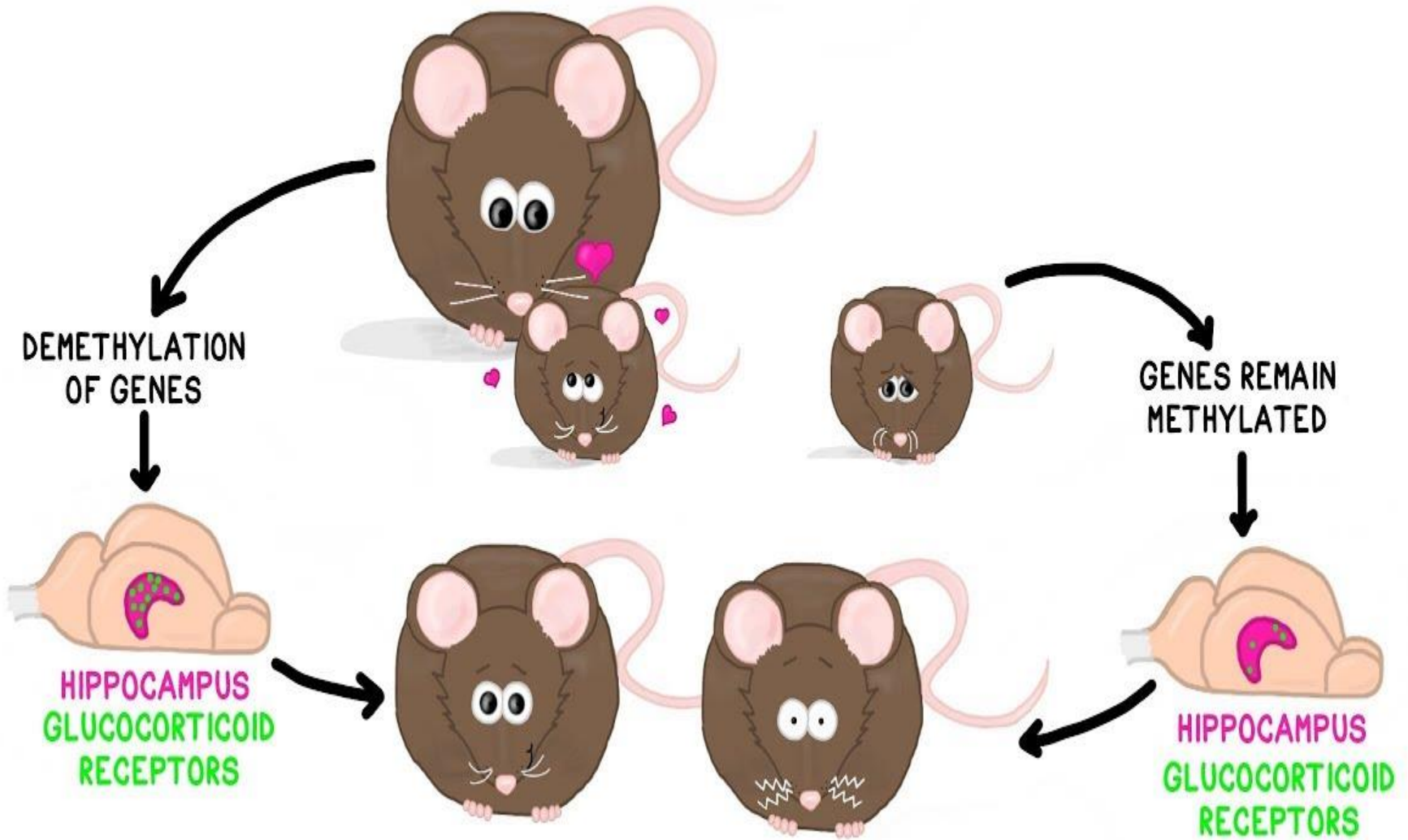
“The study of changes **in gene function** that are heritable and donot entail a change in DNA sequence”

- Corrent wu and morris(2001)



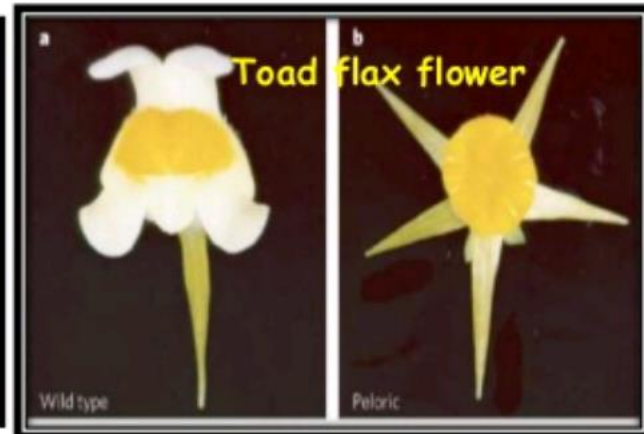
Epigenomics : Study of the complete set of epigenetic modifications on the genetic material of a cell .

EPIGENETICS:



NATURE VS. NURTURE?

EPIGENETICS...



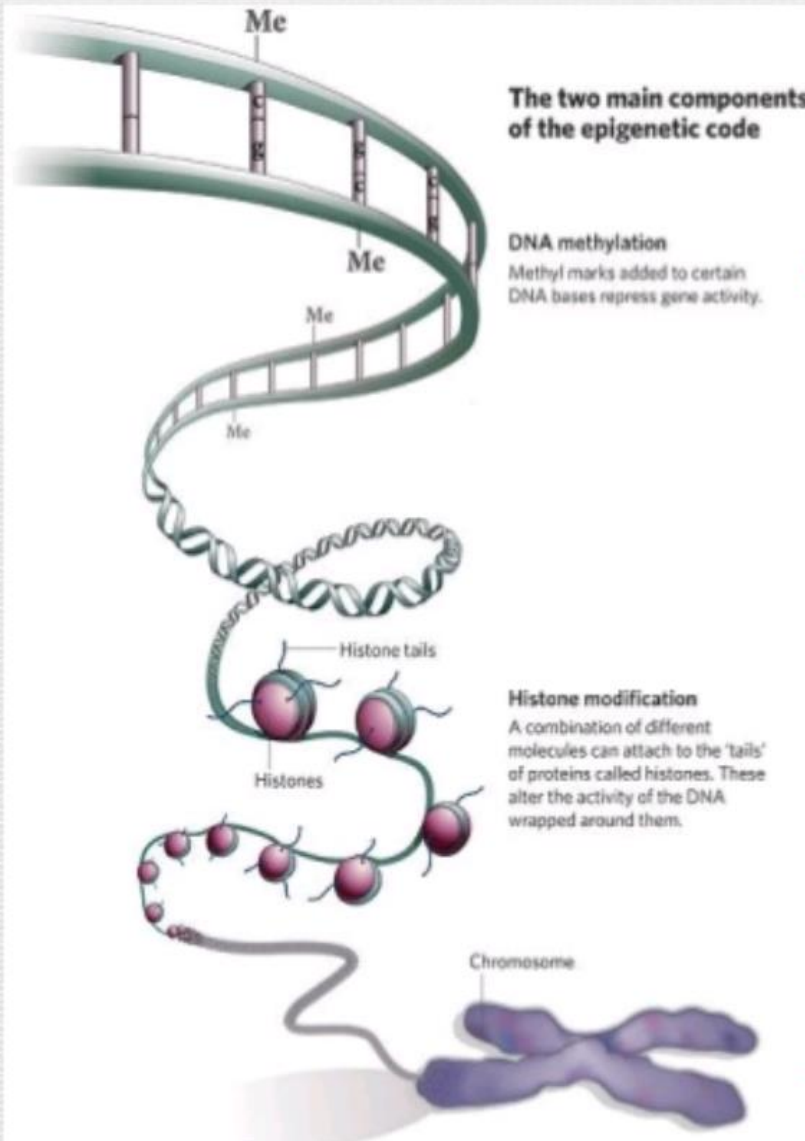
Genome is identical, Yet looks different –Epigenetics

Epigenetics

Studies show associations between epigenetic changes and:

- Environmental chemicals:
 - Air pollution
 - Arsenic
 - BPA
 - Flame retardants
 - Heavy metals
 - Organotins
 - PFCs
 - POPs
 - Pesticides
 - Phthalates
 - Radiation
 - Selenium
 - Solvents
 - Diet/nutrition
 - Overweight/obesity
 - Stress
 - Viruses
 - Vitamin D levels
-pretty much everything

RELATION OF EPIGENETIC CHANGES WITH DIFFERENT FACTORS

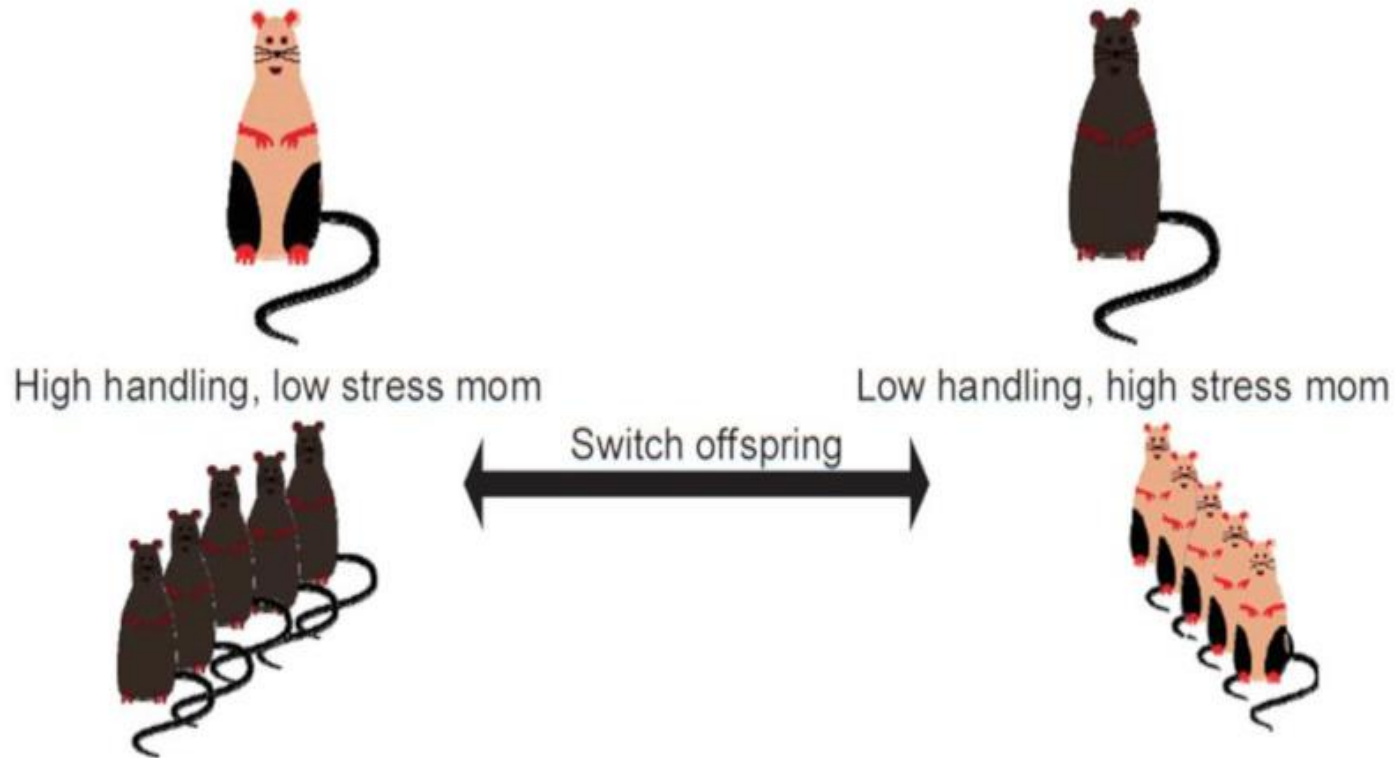


Epigenomics

Epi (Greek for above) -**genetics** refers to changes in the phenotype or gene expression caused by mechanisms other than changes in the underlying DNA sequence.

- DNA methylation
- Histone modifications

Epigenomics and stress relation



— If stress is **genetic**, offspring will have same stress level as **biological parent** —
If stress is **environmental**, offspring will have same stress level as **foster parent**

Stress Gene Experiment



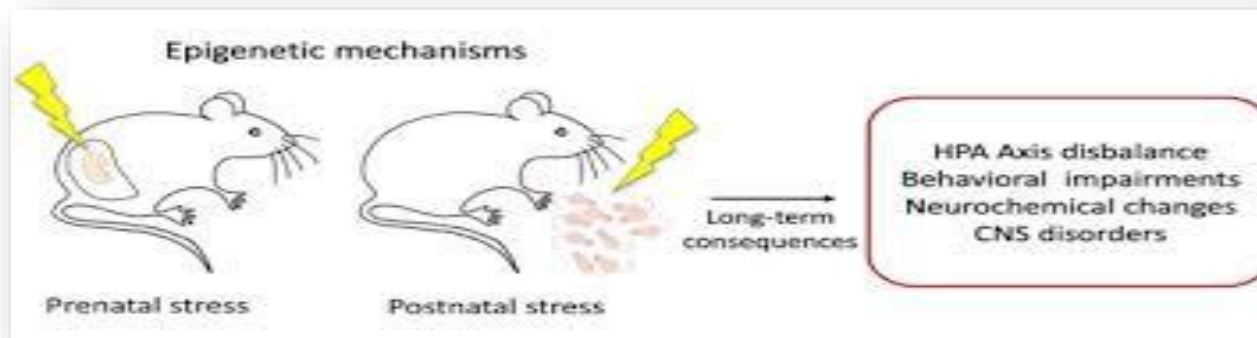
Group 1:

The babies' DNA is tested and epigenetic tags are visible on the stress response gene, turning it "on." These rats respond very well in stressful situations for the rest of their lives.



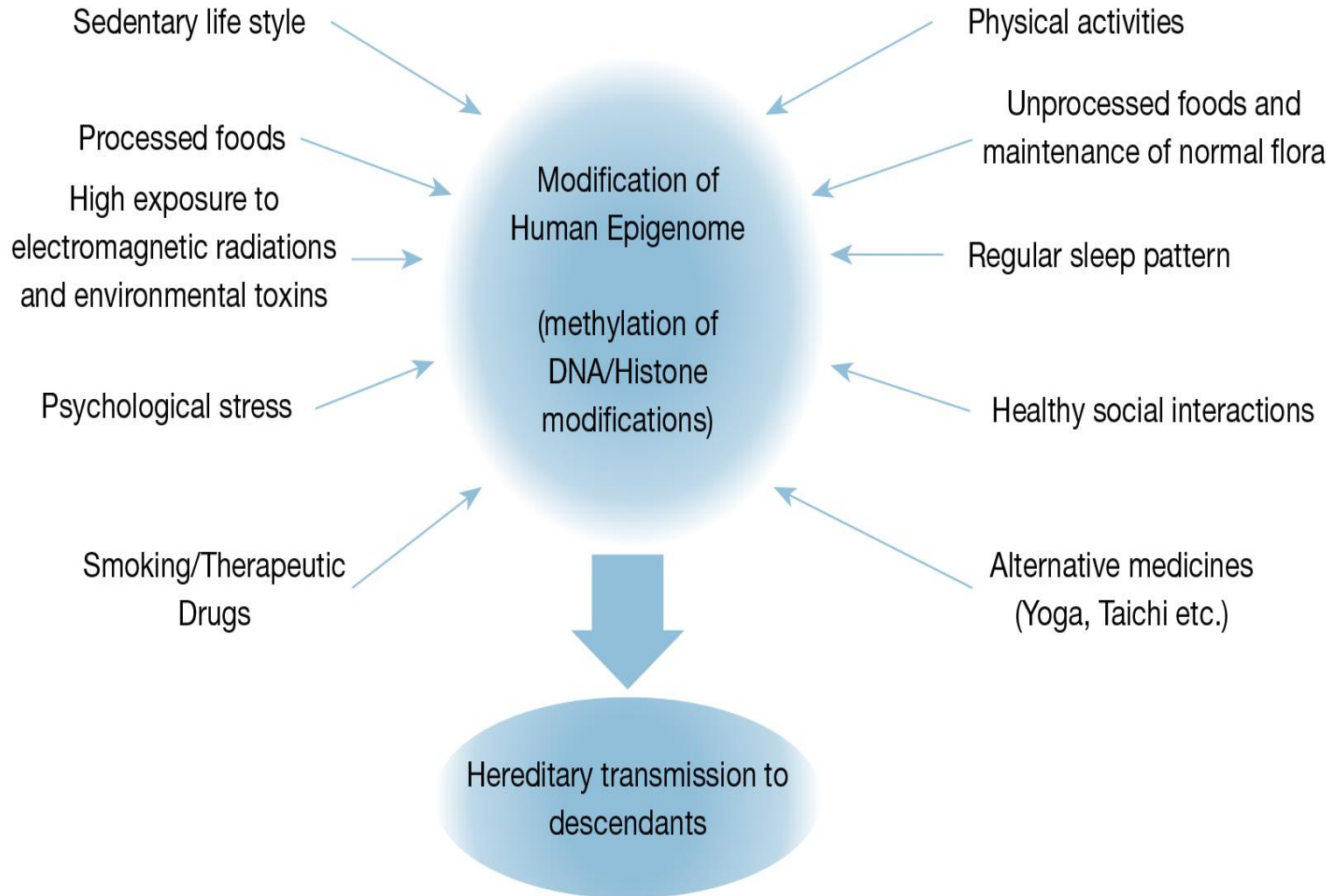
Group 2:

The babies' DNA is tested and epigenetic tags are NOT present on the stress response gene, keeping it "off." These rats do NOT respond well in stressful situations for the rest of their lives.

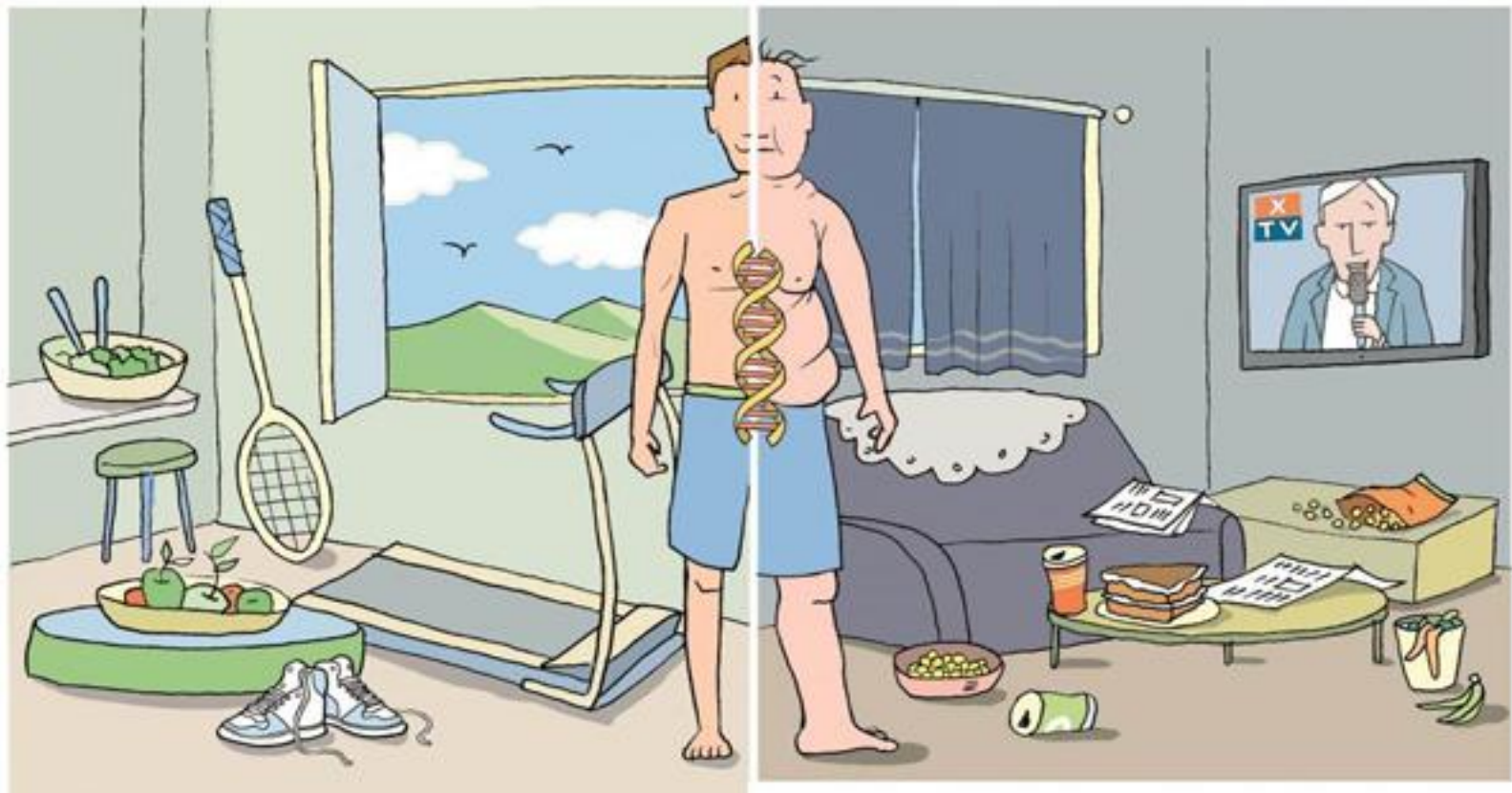


Factors predispose epigenetic maladaptation

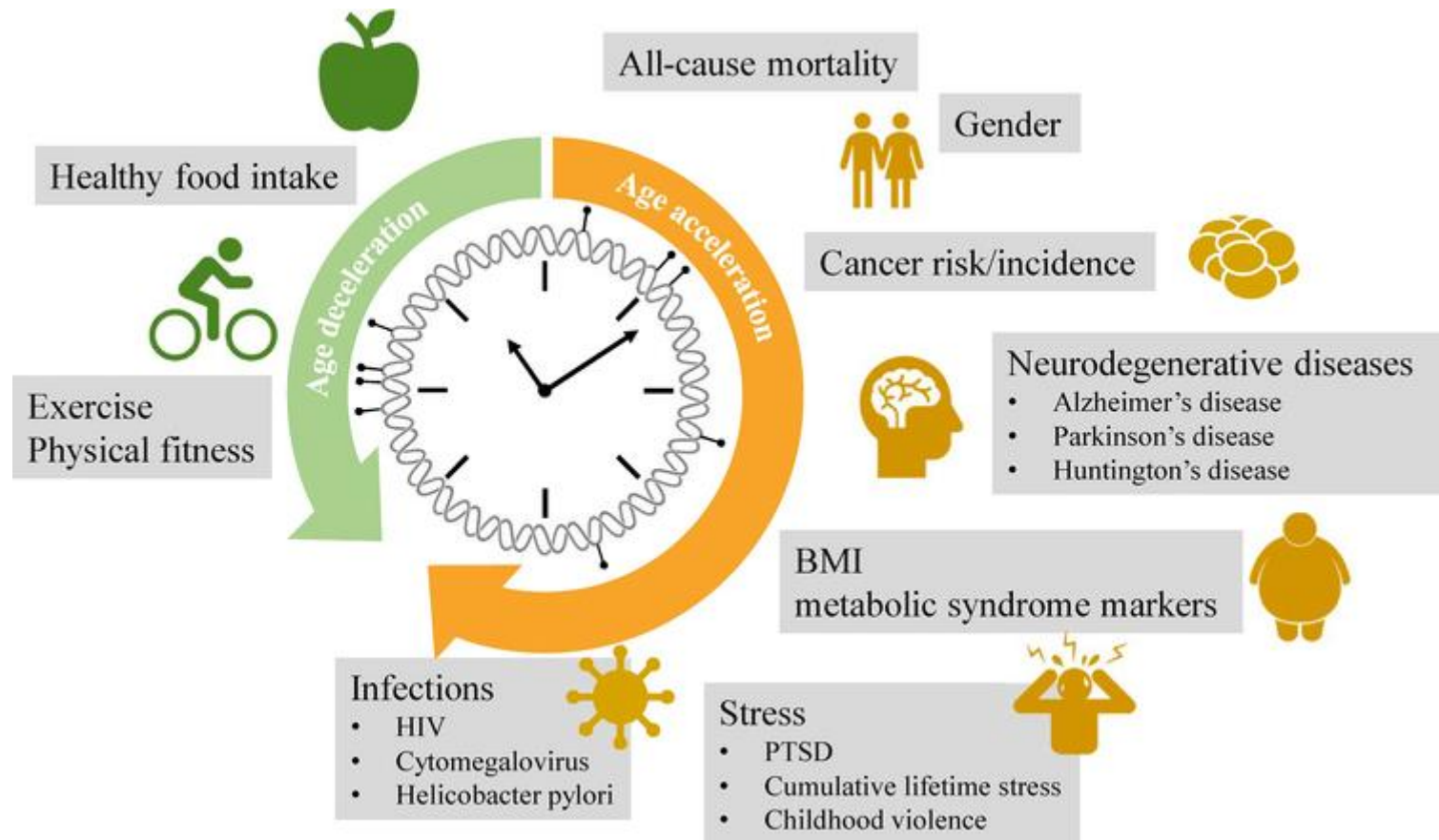
Factors bring about beneficial epigenetic adaptation



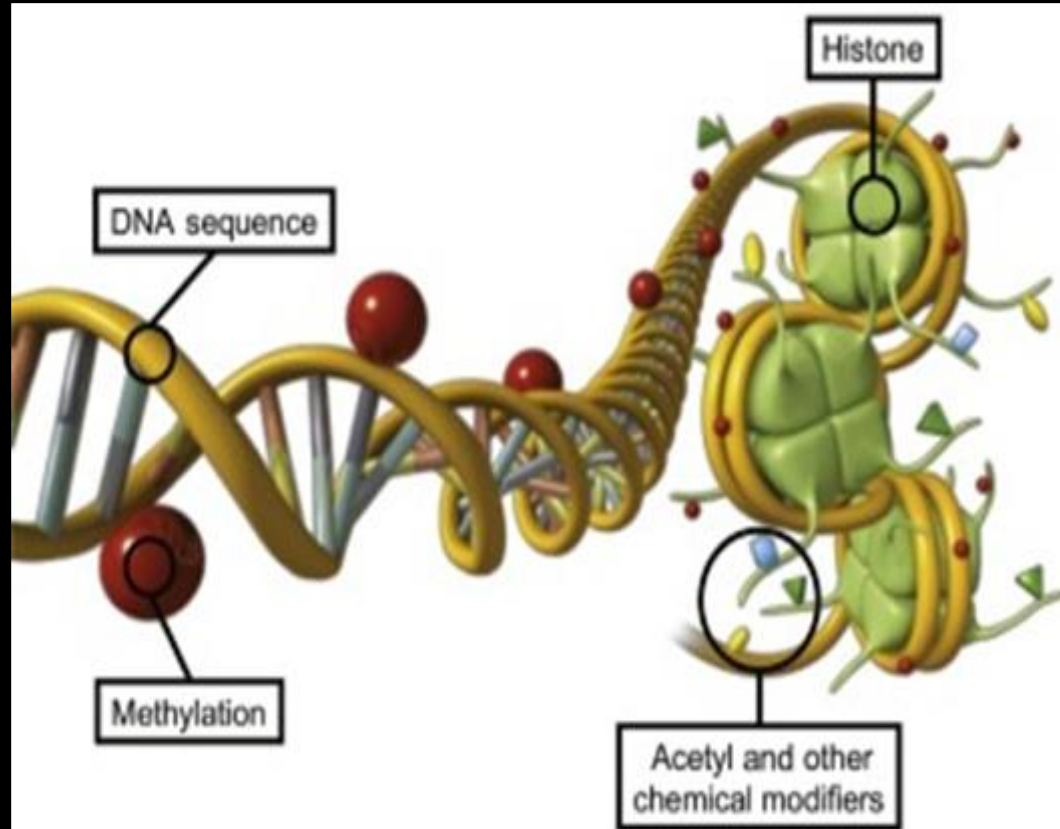
MODIFICATION OF HUMAN EPIGENOME



HUMAN EPIGENOME...



Epigenetics mechanism



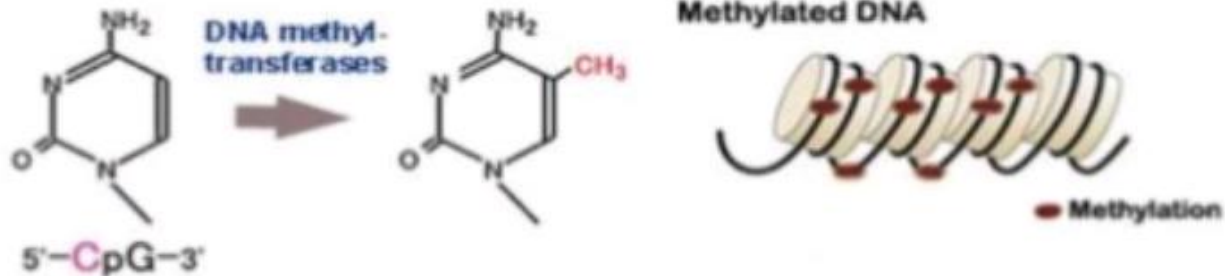
Epigenetic mechanisms and marks

- DNA methylation
- Histone modifications
- Chromatin structure
- Noncoding RNA
- RNA methylation

Trends in Endocrinology & Metabolism

DNA METHYLATION

- The methylation reaction is catalyzed by the enzyme *DNA methyltransferases* (DNMTs).
- The most frequent positions for methylation in eukaryotic DNA are C residues that are present next to G (CpG) islands Murrell et al, 2005



METHYLATION



Stimulates binding of some proteins on chromosome



Binding of Histone Deacetylases(HDACs)



Genes become non-accessible to RNA polymerase

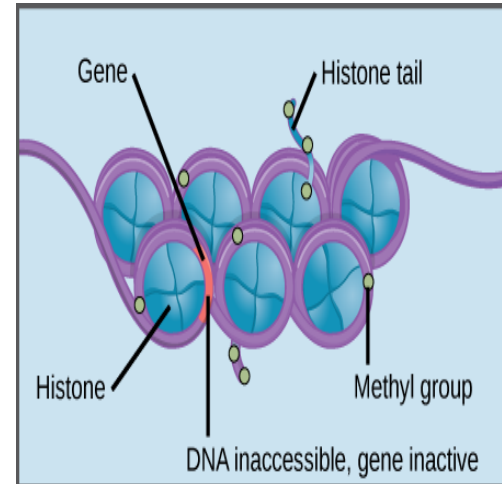
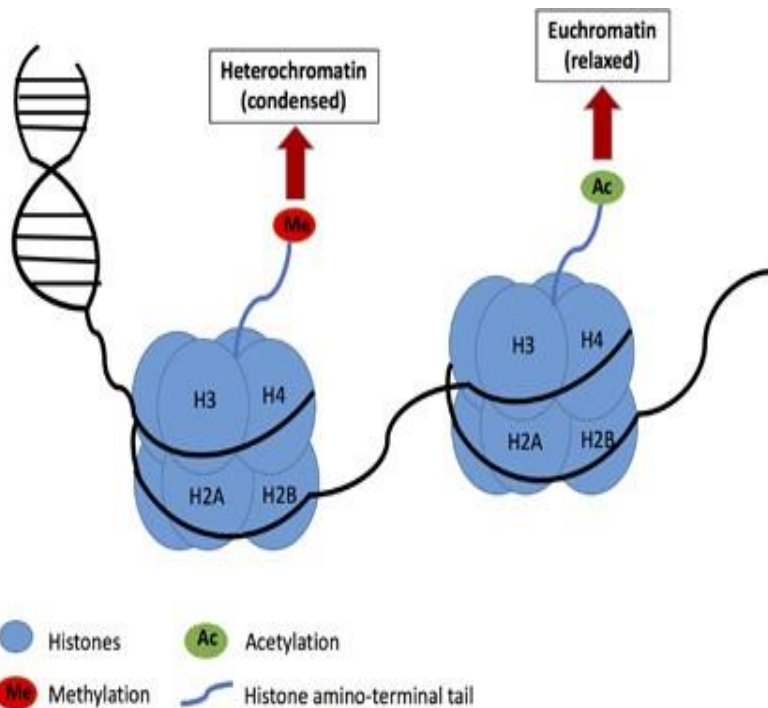
HISTONE MODIFICATIONS

The unwrapping of DNA from histones is necessary for expression of genes. Hence, change in the structure of chromatin leads to variation in the gene expression. This is called as **chromatin-remodelling**.

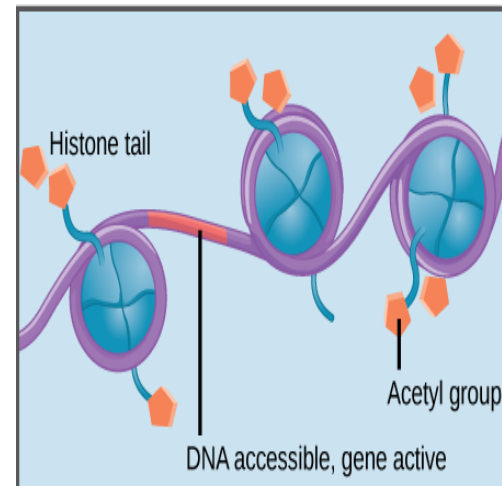
TYPES OF HISTONE MODIFICATION

- Methylation
- Acetylation and Deacetylation
- Phosphorylation
- Ubiquitylation

METHYLATION OF DNA & ACETYLATION OF HISTONE



Methylation of DNA and histones causes nucleosomes to pack tightly together. Transcription factors cannot bind the DNA, and genes are not expressed.



Histone acetylation results in loose packing of nucleosomes. Transcription factors can bind the DNA and genes are expressed.

HISTONE PHOSPHORYLATION

- Occurs at S, T & Y residues.
- Unlike acetylation and methylation, histone phosphorylation seems to function by establishing interactions between other histone modifications and serving as a platform for effector proteins.

HISTONE UBIQUITYLATION

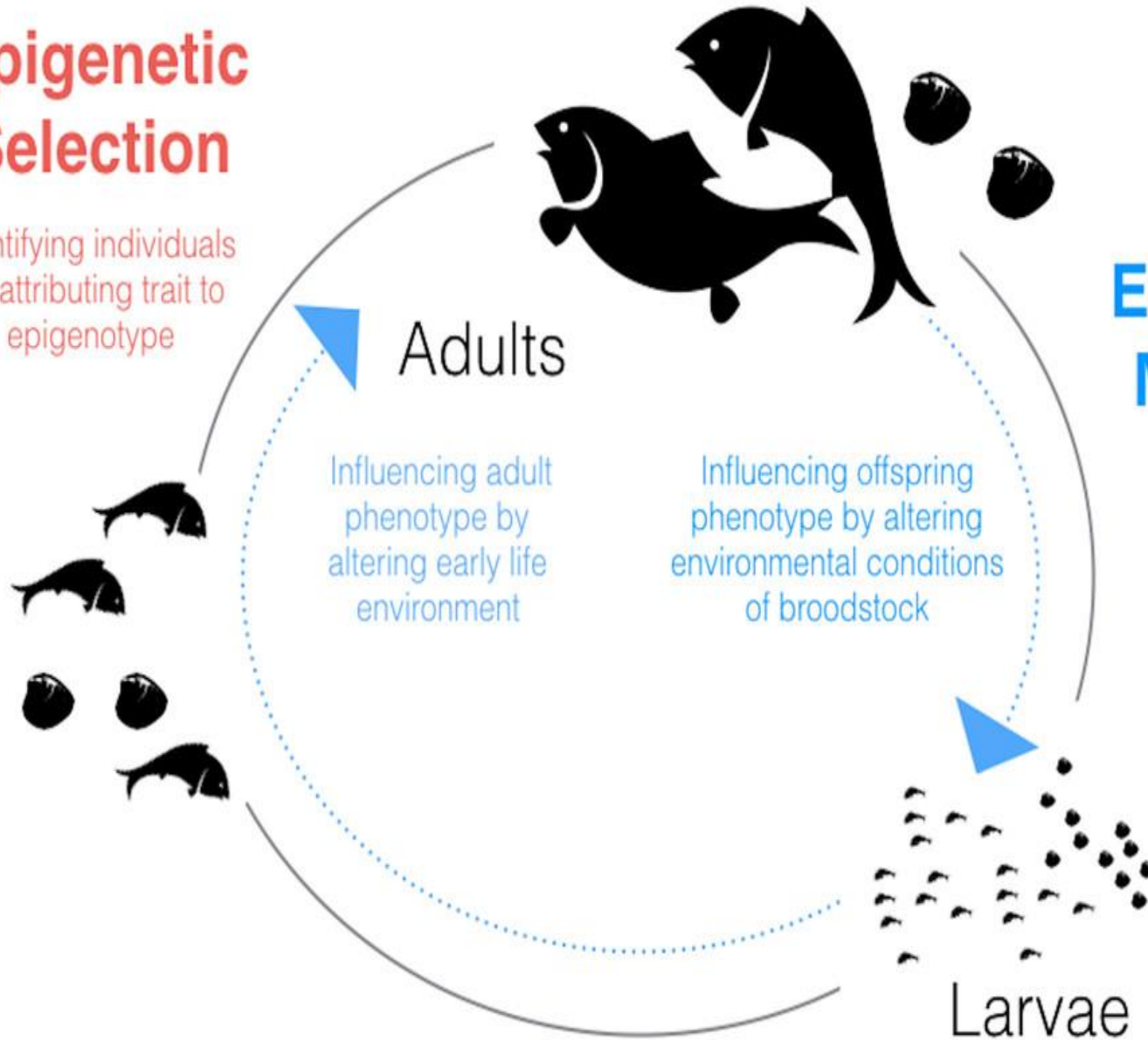
- Histone H2A and H2B are two of the most highly ubiquitylated proteins found in the nucleus.
- The most abundant forms are monoubiquitylated H2A on K119 and monoubiquitylated H2B on K123
- Ubiquitylated H2B is associated with transcription activation.
- Ubiquitylation of H2A and H2B is reversible, and is tightly regulated by histone ubiquitin ligases and deubiquitylating enzymes.

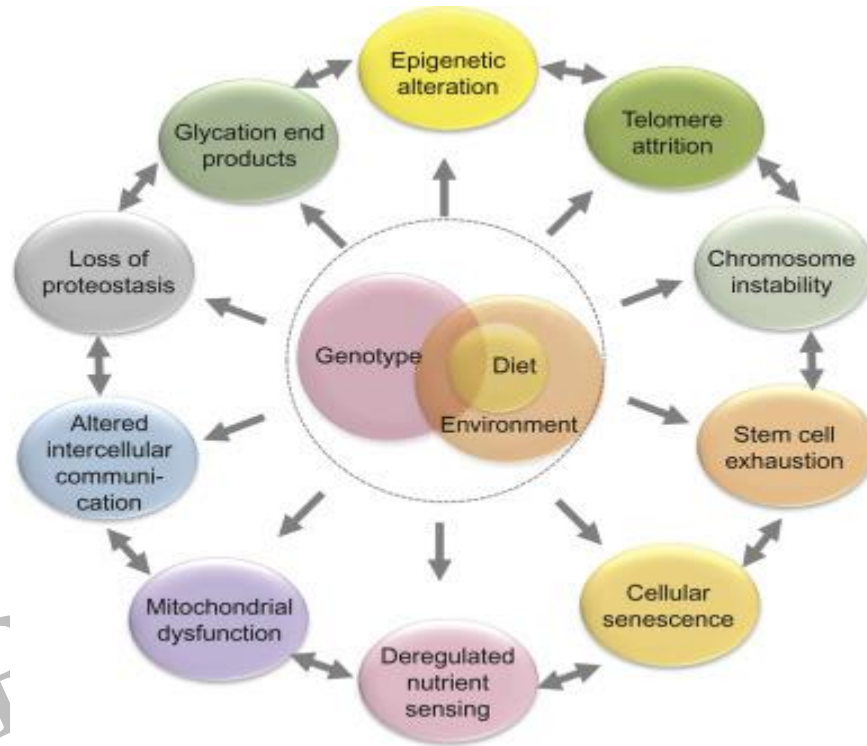
NON-CODING RNAs

- About 90% of the eukaryotic genome is transcribed. Interestingly, only 1 – 2% of these transcripts encode for proteins; the majority are transcribed as non-coding RNAs (ncRNAs).
 1. MicroRNA (miRNA)
 2. Piwi-interacting RNA (piRNA)
 3. Small-interacting RNA (siRNA)
 4. Long non-coding RNA (lncRNA)

Epigenetic Selection

Identifying individuals by attributing trait to epigenotype

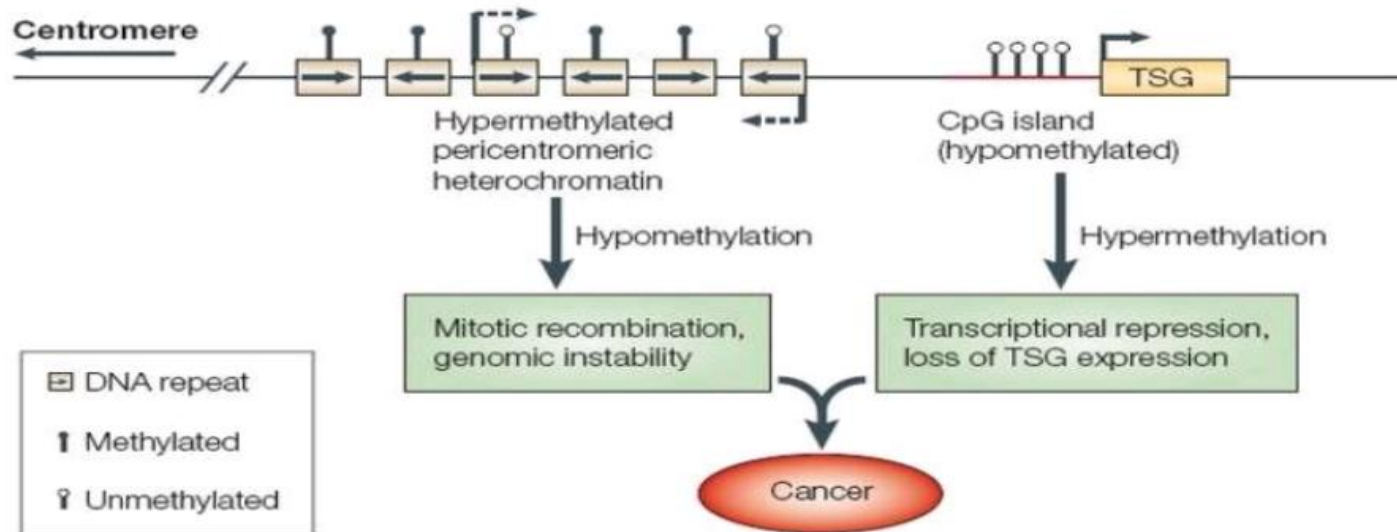




**RELATION OF EPIGENOMIC CHANGES
WITH DIFFERENT FACTORS**

DEVELOPMENT OF EPIGENOMICS IN CROP RESEARCH

- Initially, the epigenomics based researches were mostly contributed towards the understanding the phenomena of various human diseases, mainly cancer Murell *et al*, 2005; Thakur *et al*, 2013.



Small RNA and degradome deep sequencing reveals drought- and tissue-specific miRNAs and their important roles in drought-sensitive and drought-tolerant tomato genotypes

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Objective:

To know the role of “**Epigenetic regulation**” for drought tolerance in Tomato by miRNA analysis in drought tolerant and drought sensitive tomato root and up ground tissues.

Advantages of Epigenomics

- **Inheritable over many generations.**
- **New source of polymorphism.**
- **Basis for selection and plant breeding.**
- **Produce new source of variation and response to stressfull environment.**
- **It could emerge at high frequency in a single generation.**
- **DNA methylation is generator of epiallel and permanent mutant allele.**

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- ❑ Environmental epigenetics. Bollati V, Baccarelli A. Heredity (Edinb). 2010 Jul;105(1):105-12. Epub 2010 Feb 24.

THANKYOU

Dr. RASHMI KOMAL