

# (Epi)genomic environmental response in Drosophila

Cristina Vieira

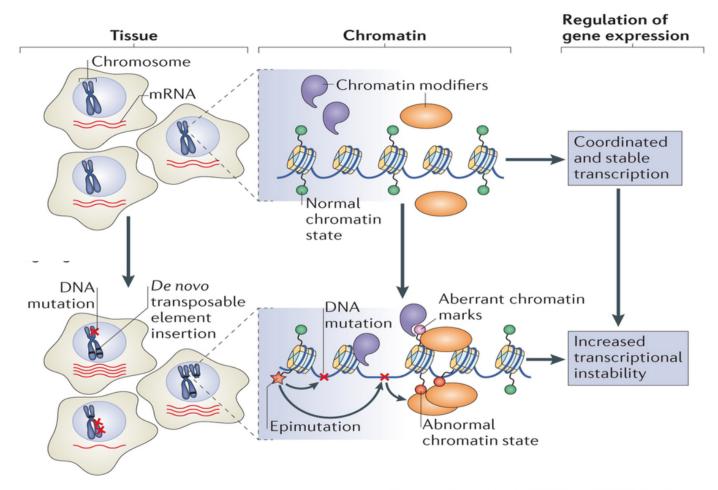








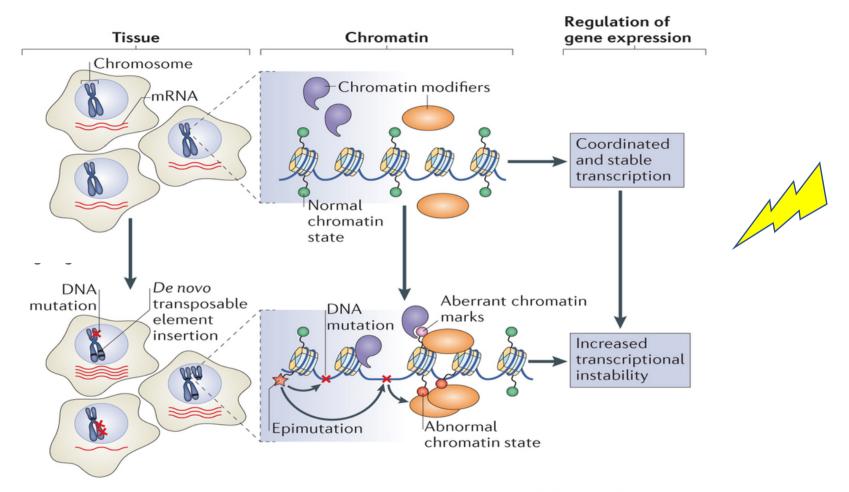
#### Genome stability



Nature Reviews | Molecular Cell Biology

Benayoun et al. 2015

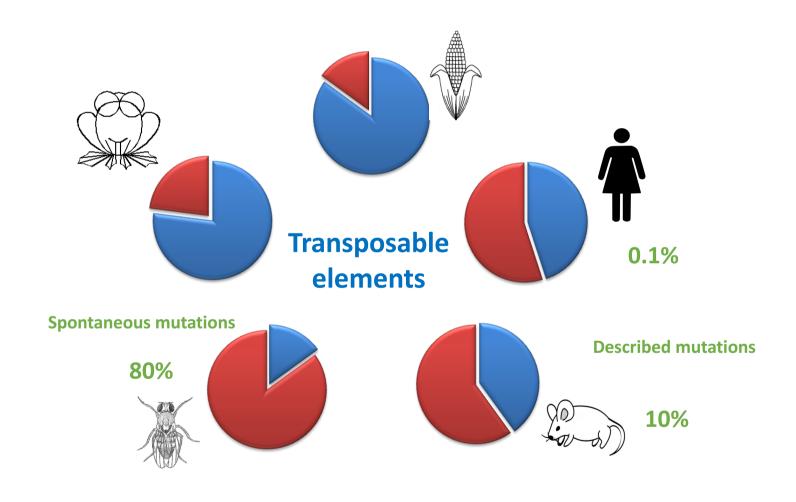
#### Genome stability



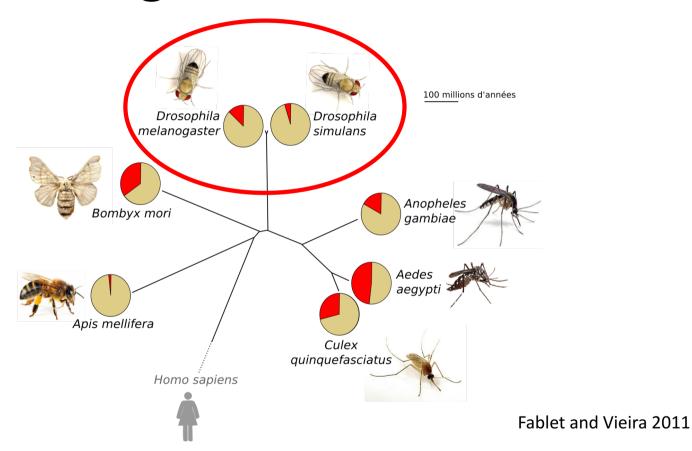
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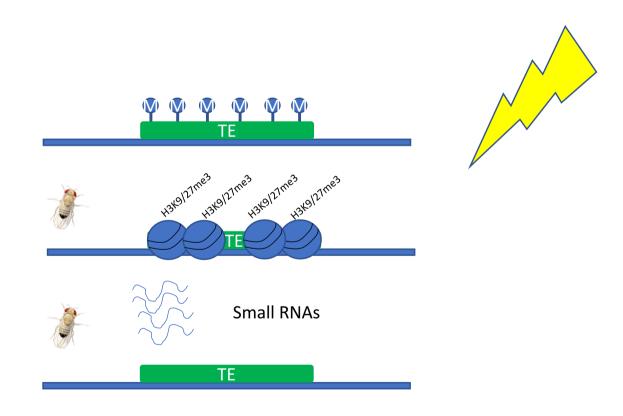
### TE proportions in some sequenced genomes



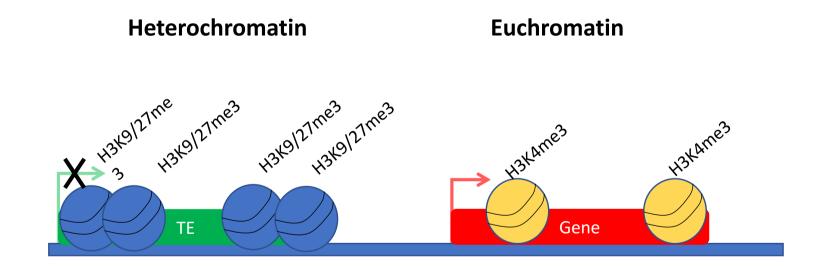
### Proportion of TE in some sequenced genomes



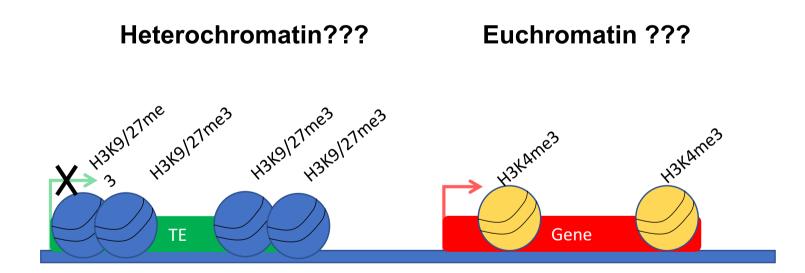
Does environmental changes affect the stability of the genome mediated by TEs?



#### Genome Homeostasis



## Genome Homeostasis >> Genome instability? With or without environemental external changes?



#### "controlling elements"



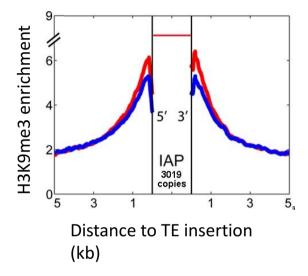
Barbara McClintock (1902-1992)





#### Retrotransposon-Induced Heterochromatin Spreading in the Mouse Revealed by Insertional Polymorphisms

Rita Rebollo<sup>1,2</sup>, Mohammad M. Karimi<sup>2</sup>, Misha Bilenky<sup>3</sup>, Liane Gagnier<sup>1,2</sup>, Katharine Miceli-Royer<sup>1,2</sup>, Ying Zhang<sup>1,2</sup>, Preeti Goyal<sup>2</sup>, Thomas M. Keane<sup>4</sup>, Steven Jones<sup>2,3</sup>, Martin Hirst<sup>3</sup>, Matthew C. Lorincz<sup>2\*</sup>, Dixie L. Mager<sup>1,2\*</sup>



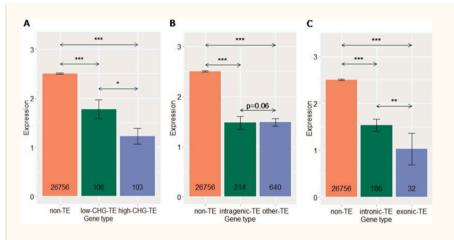


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#### Epigenetic regulation of intragenic transposable elements impacts gene transcription in *Arabidopsis thaliana*

Tu N. Le<sup>1</sup>, Yuji Miyazaki<sup>1</sup>, Shohei Takuno<sup>2</sup> and Hidetoshi Saze<sup>1,\*</sup>



#### Figure 4.

Negative impact of intragenic TEs and their methylation on gene expression ( $log_2(RPKM+1)$ ) in A. thaliana. (A) Expression of genes containing high-CHG-methylated TEs, low-CHG-methylated TEs and genes without TE insertions. (B) Expression of genes containing intragenic TEs, overlapped with other TEs and genes without TE insertions. (C) Expression of genes harboring exonic TEs and intronic TEs, and genes without TE insertions. (\*), (\*\*\*), (\*\*\*) correspond to P < 0.05, 0.005 and 0.0005, the MWU test, respectively.

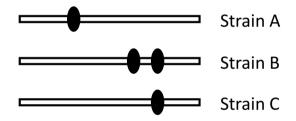


### Are TE genome expression modifiers with environmental changes?

- We need:
  - TE insertion polymorphism and gene expression
  - Stress disturbing genome homestasis
- Different approaches
  - Candidate genes
  - Genome wide

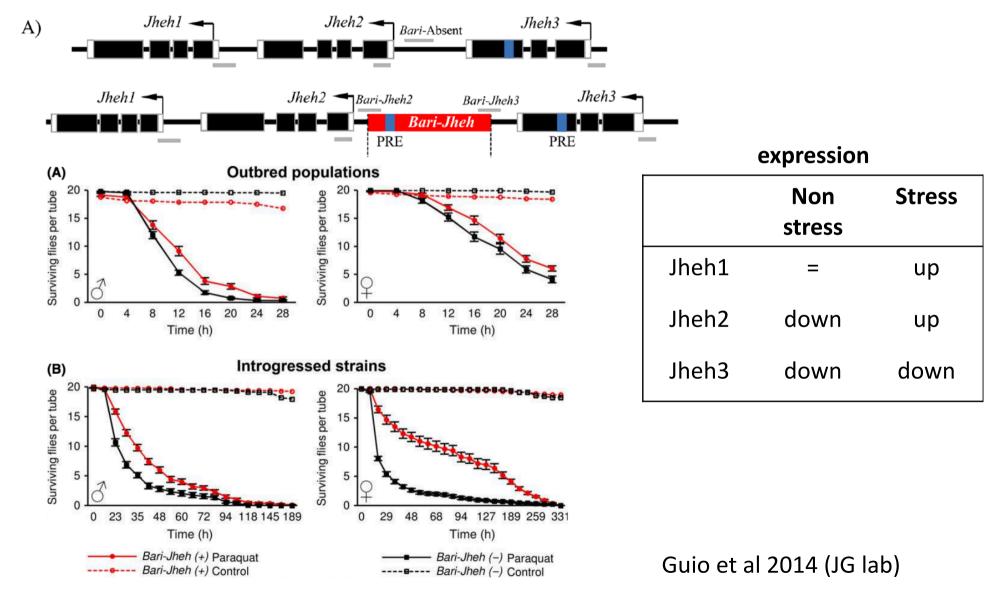
#### Genome expression modifiers

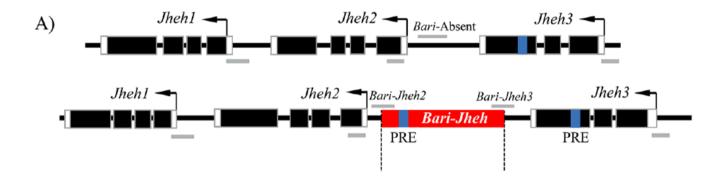
• TE insertion polymorphism and gene expression



### TE insertion polymorphism and gene expression

### The case of Bari-Jheh (Juvenile hormone epoxy hydrolase)





#### Is Bari affecting the chromatin structure?

#### **Predictions:**

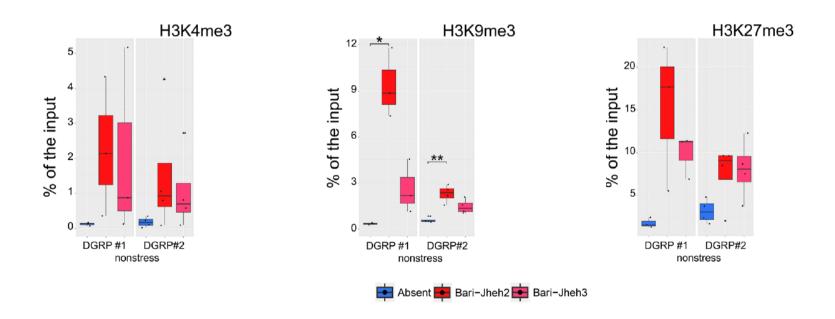
Trithorax group Response Elements (TREs) that recruit H3K4 methyltransferases,

Polycomb group Response Elements (PREs) that recruit H3K27 methyltransferases

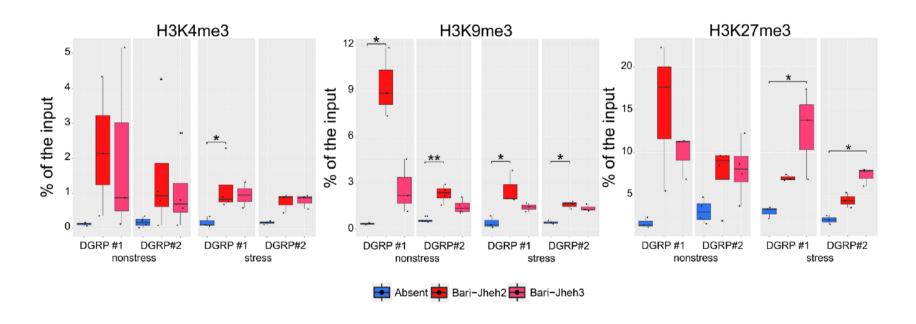
no TRE, one PRE in the Bari-Jheh sequence, and one PRE in the coding region of Jheh3

- Two genetic backgrounds
- Three histone marks:
  - H3K4m3 associated with active chromatine
  - H3K9m3 repressive mark associated with TEs
  - H3K27m3 associated with facultative heterochromatin (silenced or repressed promoters and enhancers)

 Bari-Jheh is associated with an enrichment of H3K9me3 histone mark in nonstress conditions.

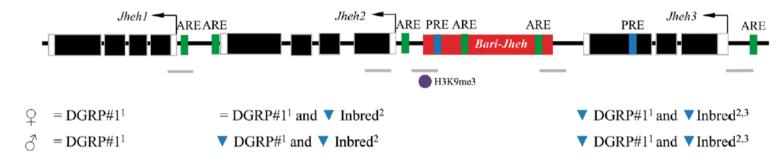


 Bari-Jheh is also associated with an enrichment for H3K4me3 and H3K27me3 chromatin marks in oxidative stress conditions.

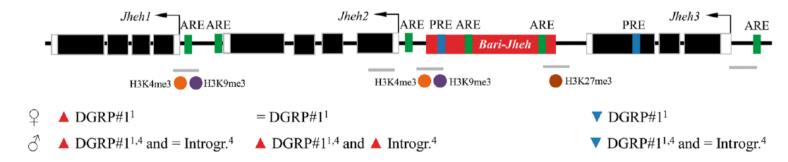


## Summary of the histone enrichments found in the Bari-Jheh genomic region.

#### A) nonstress conditions



#### B) stress conditions



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### Is stress disturbing genome homestasis? (McClintock)

- Response to environment with different recently wild-type strains of Drosophila
  - D. melanogaster and D. simulans





Judit Salces-Ortiz





- Freshly collected strains
- Phenotypic data (cold, starvation, oxidative stress)
- Genomic data (TE content)
- RNAseq data
- (ChIPseq+piRNA)











Sonia Martinez

Marie Fablet Nelly Burlet Judit Salces-Ortiz

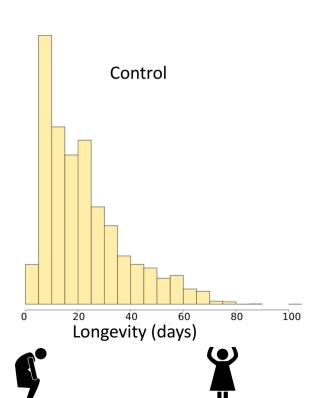
Bianca Menezes

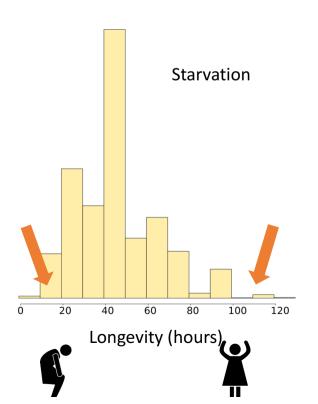
#### **Environment effects**

• Phenotypic measures in different environments (30 inbred lines/condition): longevity, starvation, cold, oxidative stress

Selection of a total of 8 lines

- -2 species
- -2 populations
- -2lines





#### Environment effects: gene expression

 How to measure the expression of TEs or piRNAs without loosing information due to the repetitive nature of TE sequences?

• Development of specific software: TE tools (Lerat et al 2017)

#### TE tools



- TE sequences (fasta file)
  - Annotated or non annotated genomes
- "rosette" file (text file allowing different levels of analyses)

chromosome	insertion	type	family	class	expression	etc
2L_1 (2134612221346196) LTR/G	297_Dsi_1	297_Dsi	Gypsy	LTR	somatic	
2L_1 (2168187721682629) LTR/G	297_Dsi_2	297_Dsi	Gypsy	LTR	somatic	
2L_1 (2169967521700427) LTR/Gy	297_Dsi_3	297_Dsi	Gypsy	LTR	somatic	
2R_1 (120098120414) LTR/Gypsy	297_Dsi_31	297_Dsi	Gypsy	LTR	somatic	
2R_1 (12369021237012) LTR/Gyp	297_Dsi_32	297_Dsi	Gypsy	LTR	somatic	
2R_1 (412736413170,413270413	ACCORD2_Dmel_12	ACCORD2_Dmel	Gypsy	LTR	germline	
2R_1 (427330428124) LTR/Gypsy	ACCORD2_Dmel_13	ACCORD2_Dmel	Gypsy	LTR	germline	
2R_1 (996594996745) LTR/Gypsy	ACCORD2_Dmel_14	ACCORD2_Dmel	Gypsy	LTR	germline	
2R_1 (10779131080665,1082958	BAGGINS1_69	BAGGINS1	LOA	LINE	germline	
2R_1 (10809871081517) LINE/LOA	BAGGINS1_70	BAGGINS1	LOA	LINE	germline	
2R_1 (10947231094905) LINE/LOA	BAGGINS1_71	BAGGINS1	LOA	LINE	germline	

#### TE tools



- Takes into account of the multicopy status and thus the diversity of TEs at the copy level
- => we can work on TE families not present in the reference genome
- => better mapping of reads on copy rather than on consensus

- Differential expression analysis with DESeq2
- mRNA and small RNA

#### TE tools



#### Available

- https://github.com/l-modolo/TEtools
- Or by the web site lbbe.univ-lyon1.fr



