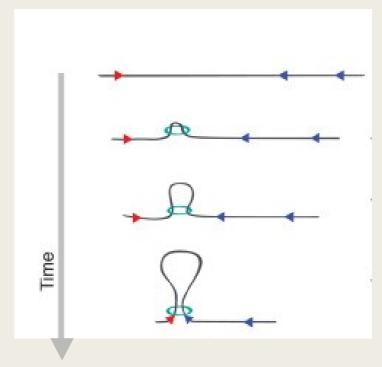
### LOOP CALLERS

Relating chromatin loops to GWAS studies in CLL
CARLOS MADARIAGA ARAMENDI
François Serra – Alba Lepore – Jon Sánchez

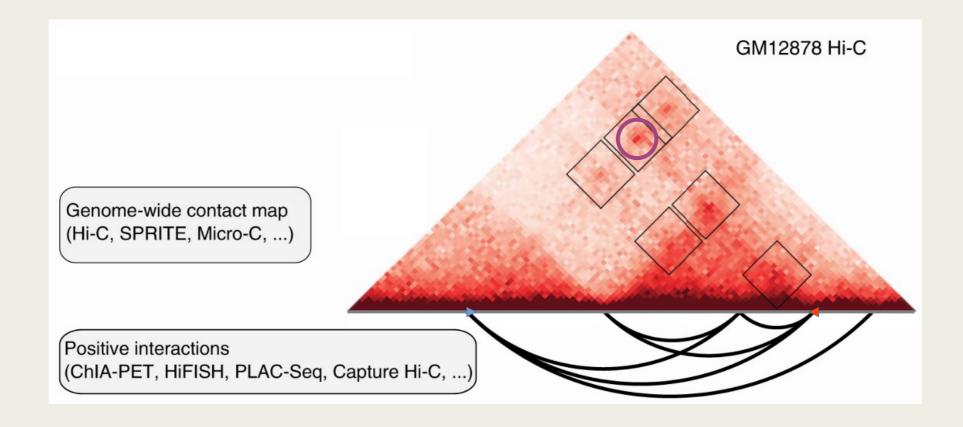
# Chromatin loops

- What are they?
- How they occur?
- What is it function?
- When they happen?
- Types of looped structures



The Cohesin Release Factor WAPL Restricts Chromatin Loop Extension

# Chromatin loops

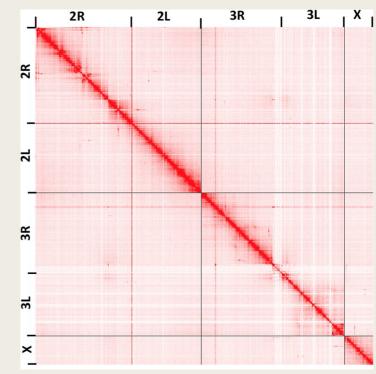


# Definition and purpose

Tools that implement an algorithm which detects chromatin loops

This is made using Hi-C data; protocol derived from Chromosome Conformation Capture (3C) and its successors (4C & 5C), that allows to study the 3D structure of genomes

Using Hi-C data we can create an interaction map of the chromatin ,so we can infer the chromatin loops reflecting interactions of promoters & enhancers, gene loops, etc.



Hi-C interaction map for assembled A. Funestus scaffolds generated using the Juicebox Hi-C visualization program

#### Different Tools Available

- The different tools detect different loops
- Also the number of loops detected is different
- More accuracy analyzing the intersecting loops



**HOMER** 

Detected Loops by different Software tools

#### **HOMER**

- Hypergeometric Optimization of Motif EnRichment
- Set of Motif Discovery and next-gen sequencing analysis too
- Analyze ChIP-Seq, GRO-Seq, RNA-Seq, DNase-Seq and Hi-C
- Installation: Using perl by command line
- Procedure: Trim FASTQ files, align them to the reference genome and assembly them into a HOMER-style tag directory. After that, the analysis is carried out to find the loops and regions of our interest.
- We can also use Hi-C Summary files (tab-delimired text files)

#### **HiCCUPS**

- Juicer tool that implements an algorithm for finding chromatin loops
- Juicer: one-click pipeline for processing Hi-C datasets
- Installation: It requires Juicer, which also requires some dependencies, as GNU, BWA or CUDA and NVIDIA GPU
- Usage: You need to specify the matrix size, the chromosomethie resolution(s), the Hi-C file to analyze and the output dire
- It outputs several files; the 'merged\_loops' file is what we

#### MUSTACHE

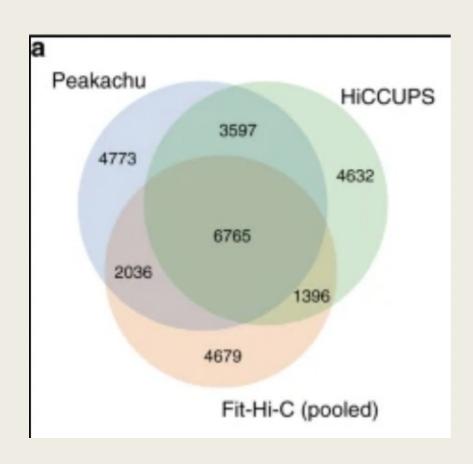


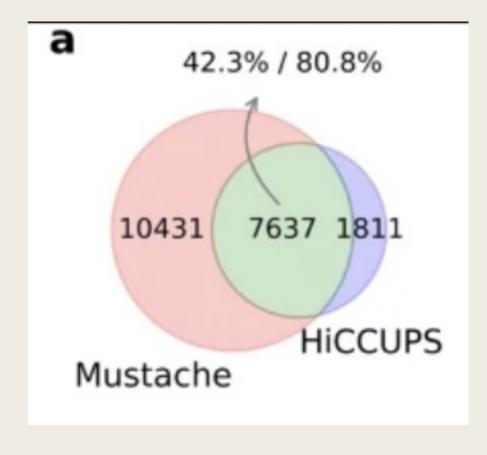
- Multi-scale Detection of Chromatin Loops from Hi-C and Micro-C Maps using Scale-Space Representation
- Tool for multi-scale detection of chromatin loops from Hi-C and Micro-C contact maps in high resolutions
- Several ways to install it: Conda (recommended), Docker, PIP or Github.
- Input: Text format, Juicer .hic files, Cooler .cool and .mcool files. Output: TSV format
- Subject chromosome and resultion also need it to run the tool.

#### PEAKACHU

- Unveil Hi-C Anchors and Peaks
- Takes genome-wide contact data as input and returns coordinates of likely interactions such as chromatin loops.
- ML framework based on sklearn to generate random forest models trained on example interactions predicted by an arbitrary experiment
- Designed to accept any genome-wide contact map (Micro-C...)
- Requires several packages. Conda recommended to install
- Input: .cool or .hic files. Also .bedpe text file with training data.
- It can work as a regular loop caller with the 'score\_genome' function

## Traditional vs Modern Loop Callers





- Stephan Kadauke and Gerd A. Blobel. *Chromatin loops in gene regulation* Biochim Biophys Acta. 17–25 (2008)
- Joachim Wolff, Rolf Backofen, Björn Grüning. Loop detection using Hi-C data with HiCExplorer. bioRxiv. (2020)
- Salameh, T.J., Wang, X., Song, F. et al. A supervised learning framework for chromatin loop detection in genome-wide contact maps. Nat Commun 11, 3428 (2020)

#### References



# THANK YOU FOR YOUR ATTENTION! ANY QUESTIONS?