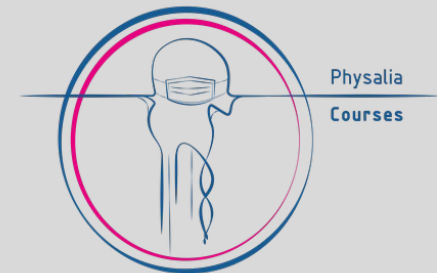


Hi-C analysis

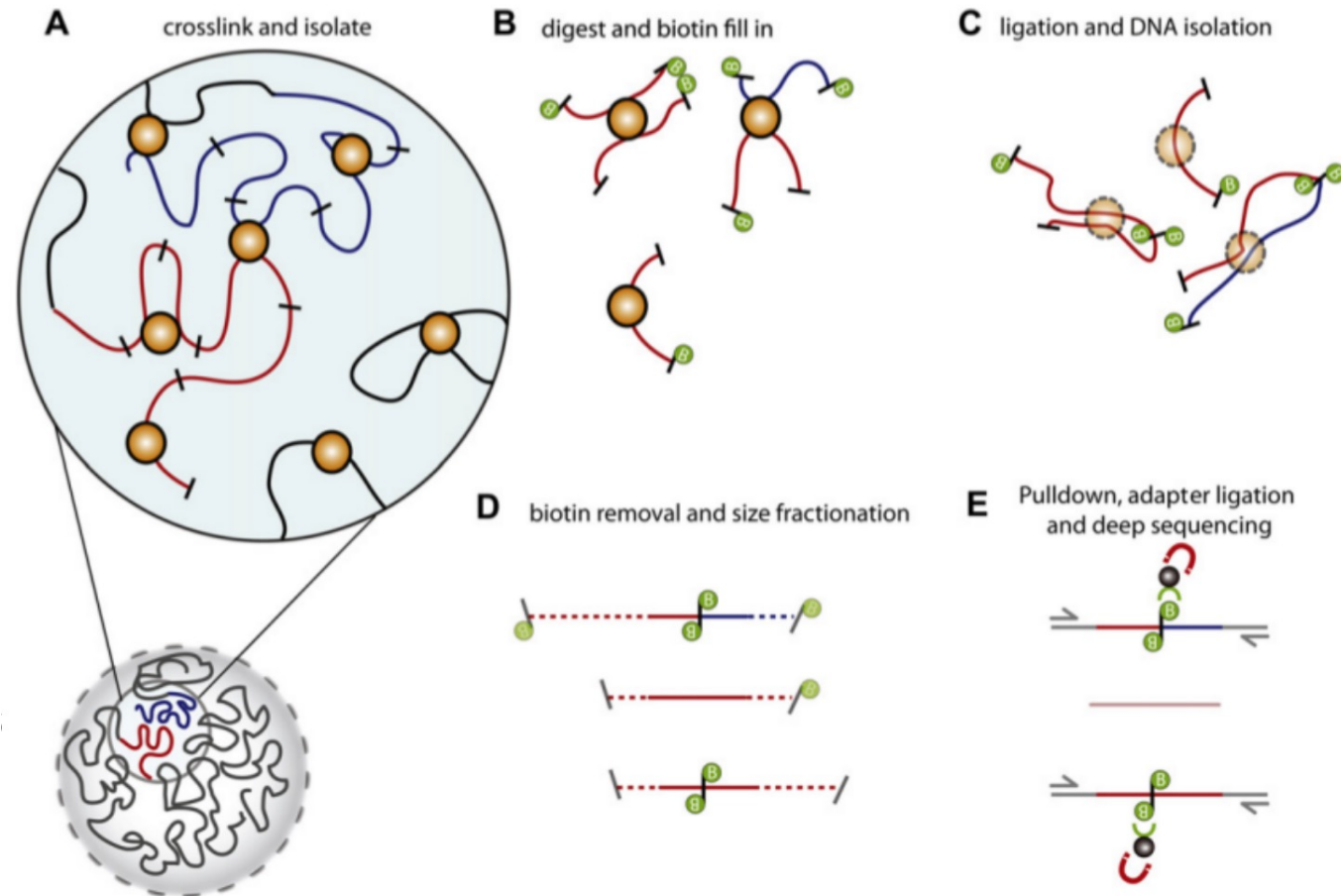
Epigenomics Data Analysis

Jacques Serizay

Physalia 2023

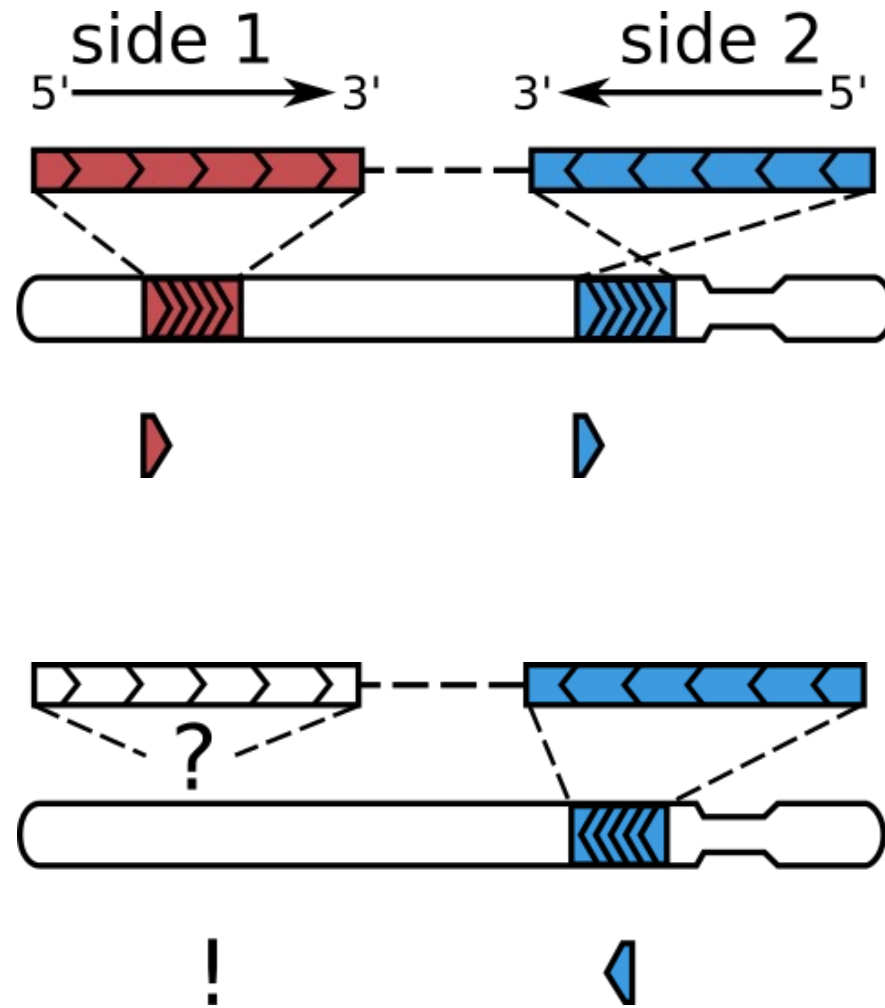


Hi-C experimental workflow



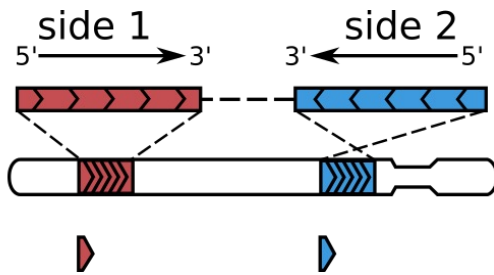
Belton et al., Methods 2012

Hi-C computational workflow



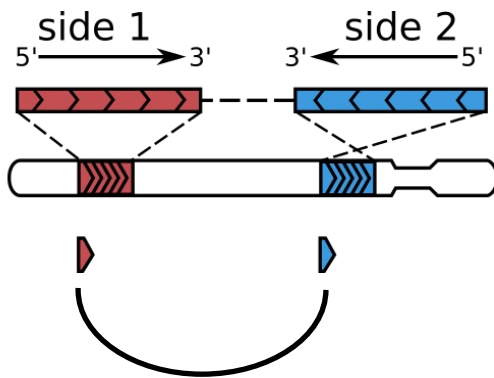
Open2C et al., Biorxiv 2023

Hi-C computational workflow



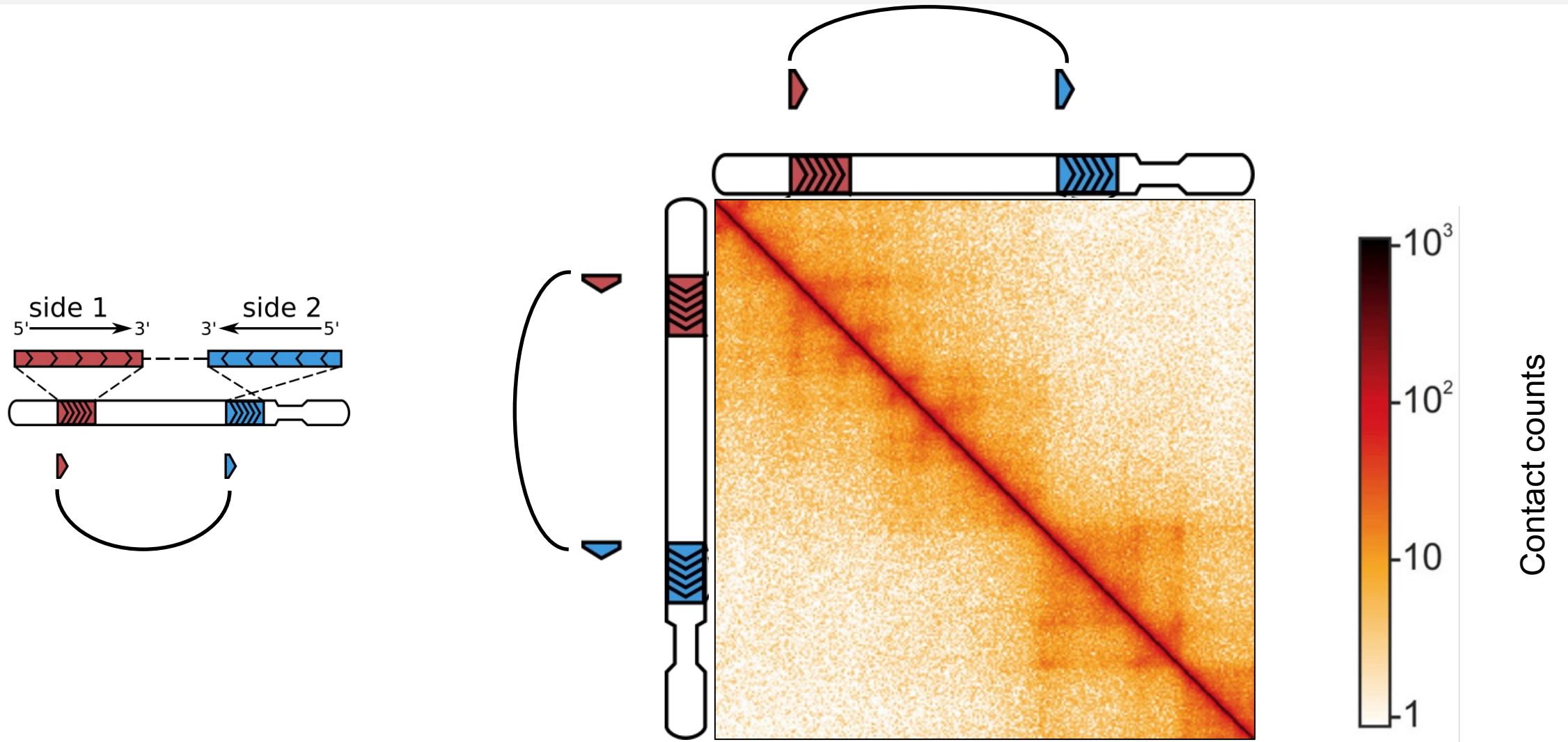
Open2C et al., Biorxiv 2023

Hi-C computational workflow



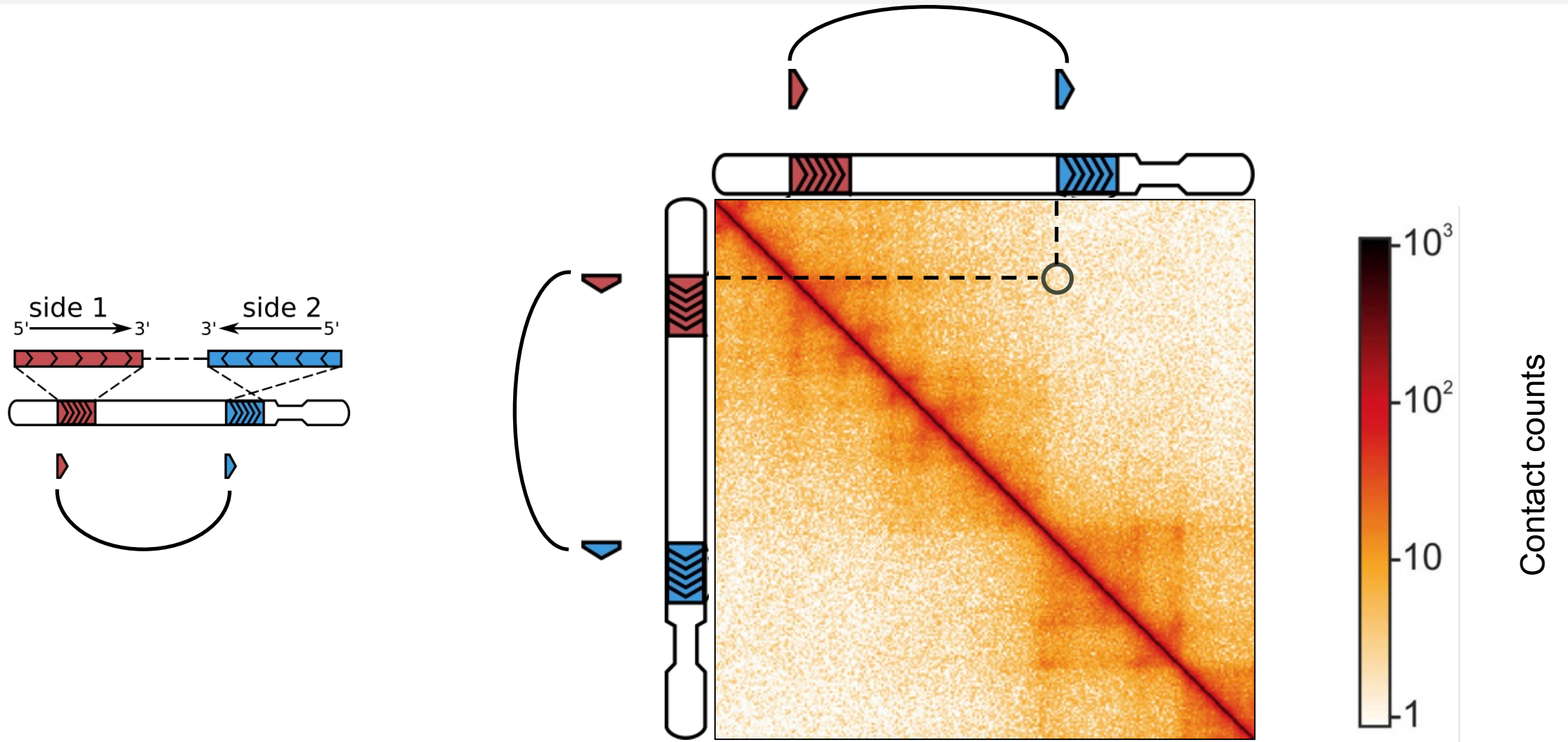
Open2C et al., Biorxiv 2023

Hi-C computational workflow



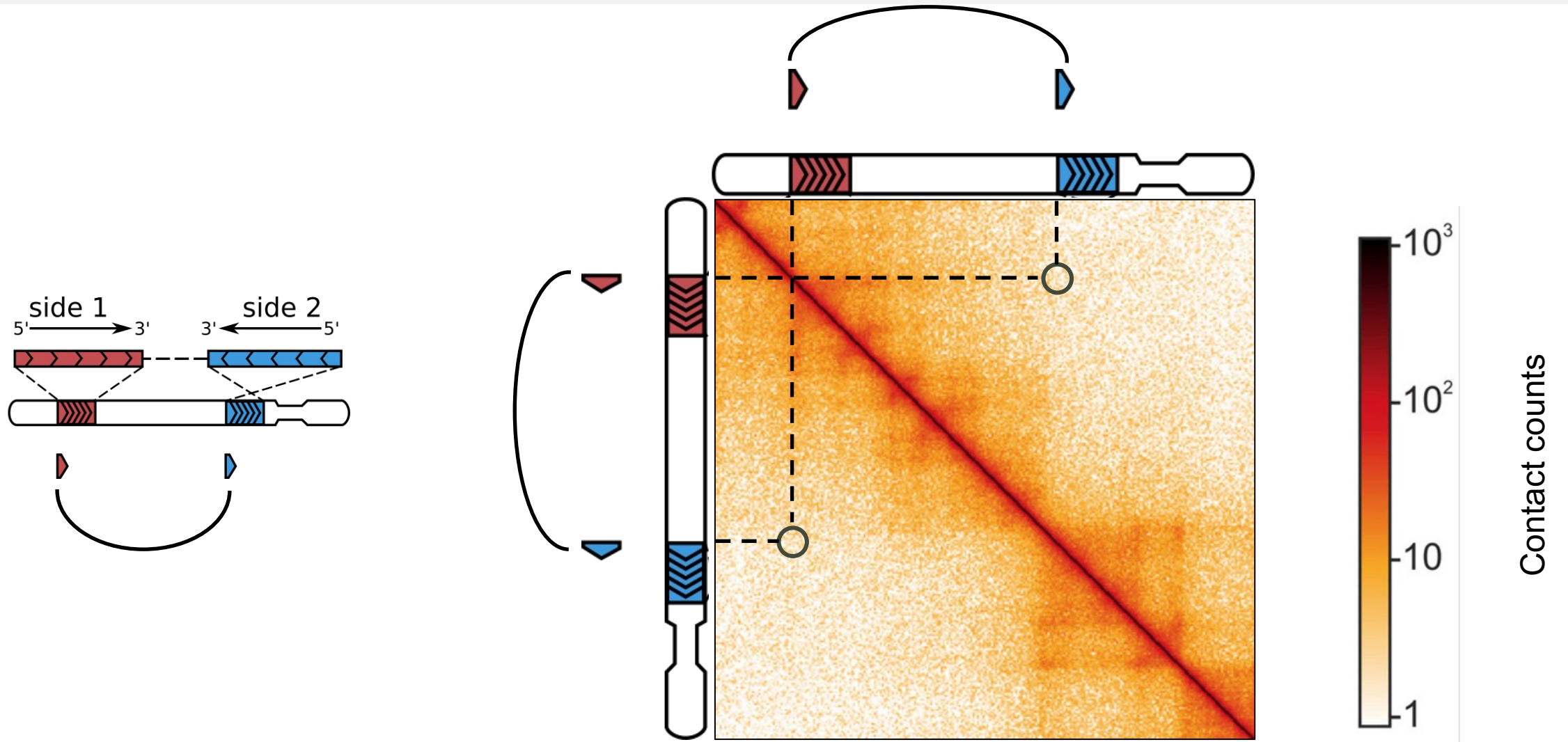
Open2C et al., Biorxiv 2023

Hi-C computational workflow



Open2C et al., Biorxiv 2023

Hi-C computational workflow



Open2C et al., Biorxiv 2023

Hi-C pipeline



Get .bcl files



Create fastq files



QC: remove/trim low quality reads



Align fastq to BAM



Filter duplicates, artifacts, ...



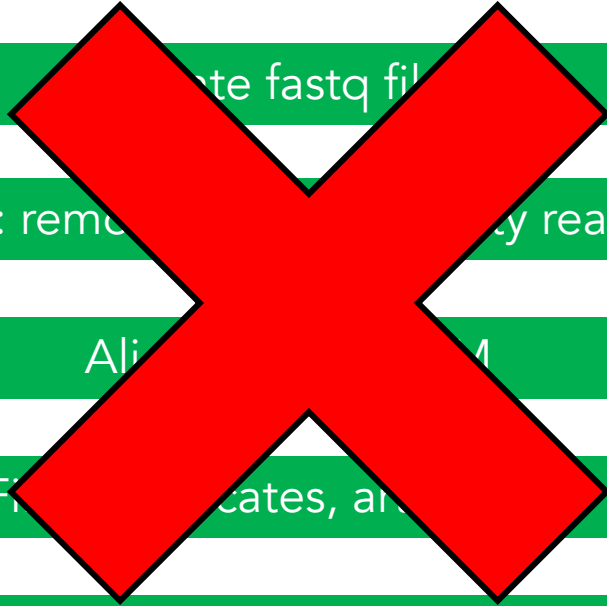
Generate tracks



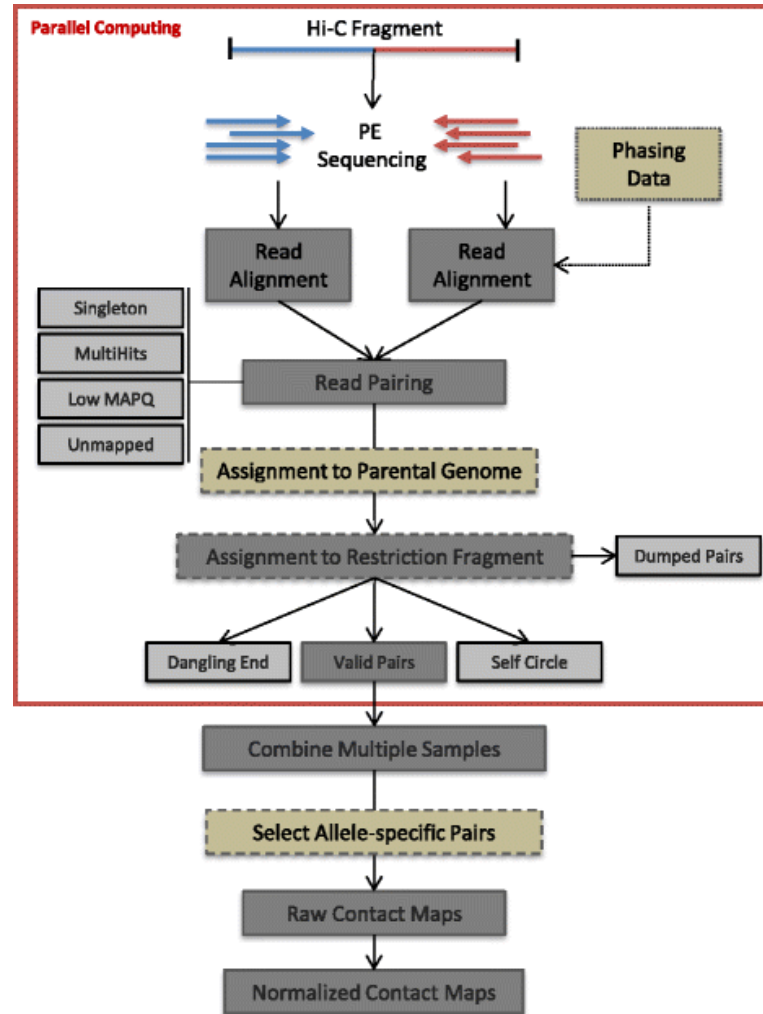
Assay-specific downstream analysis

Hi-C pipeline

- ✓ Get .bcl files
- ✓ Generate fastq files
- ✓ QC: remove low quality reads
- ✓ Align to reference genome
- ✓ Filter duplicates, and
- ✓ Generate tracks
- Assay-specific downstream analysis

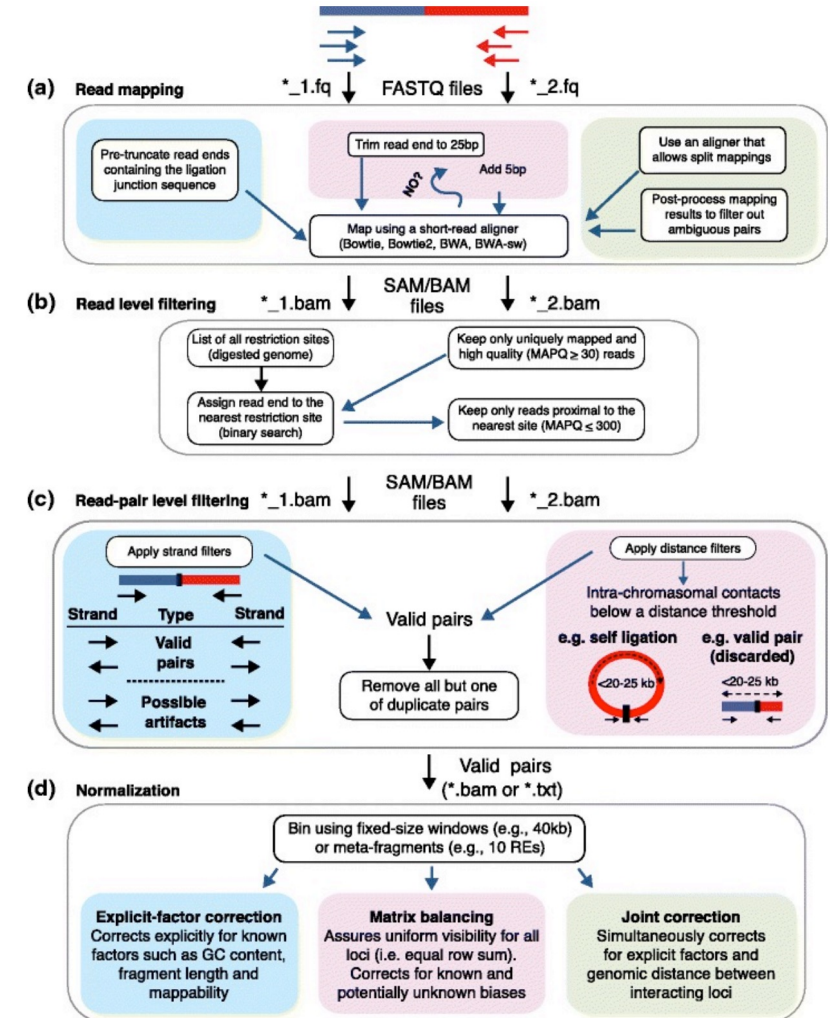


Hi-C pipeline(s)



Servant et al., Genome Biol. 2015

Ay & Noble Genome Biol 2015



Hi-C pipeline(s) – 2015

| Tool | Short-read aligner(s) | Mapping improvement | Read filtering | Read-pair filtering | Normalization | Visualization | Confidence estimation | Implementation language(s) |
|---------------------|-----------------------|---------------------|----------------|---------------------|----------------------------|---------------|-----------------------|----------------------------|
| HiCUP [46] | Bowtie/Bowtie2 | Pre-truncation | ✓ | ✓ | — | — | — | Perl, R |
| Hiclib [47] | Bowtie2 | Iterative | ✓ ^a | ✓ | Matrix balancing | ✓ | — | Python |
| HiC-inspector [131] | Bowtie | — | ✓ | ✓ | — | ✓ | — | Perl, R |
| HIPPIE [132] | STAR | ✓ ^b | ✓ | ✓ | — | — | — | Python, Perl, R |
| HiC-Box [133] | Bowtie2 | — | ✓ | ✓ | Matrix balancing | ✓ | — | Python |
| HiCdat [122] | Subread | — ^c | ✓ | ✓ | Three options ^d | ✓ | — | C++, R |
| HiC-Pro [134] | Bowtie2 | Trimming | ✓ | ✓ | Matrix balancing | — | — | Python, R |
| TADbit [120] | GEM | Iterative | ✓ | ✓ | Matrix balancing | ✓ | — | Python |
| HOMER [62] | — | — | ✓ | ✓ | Two options ^e | ✓ | ✓ | Perl, R, Java |
| Hicpipe [54] | — | — | — | — | Explicit-factor | — | — | Perl, R, C++ |
| HiBrowse [69] | — | — | — | — | — | ✓ | ✓ | Web-based |
| Hi-Corrector [57] | — | — | — | — | Matrix balancing | — | — | ANSI C |
| GOTHIC [135] | — | — | ✓ | ✓ | — | — | ✓ | R |
| HiTC [121] | — | — | — | — | Two options ^f | ✓ | ✓ | R |
| chromoR [59] | — | — | — | — | Variance stabilization | — | — | R |
| HiFive [136] | — | — | ✓ | ✓ | Three options ^g | ✓ | — | Python |
| Fit-Hi-C [20] | — | — | — | — | — | ✓ | ✓ | Python |

^aHiclib keeps the reads with only one mapped end (single-sided reads) for use in coverage computations

^bHIPPIE states that it rescues chimeric reads. No details are given

^cHiCdat reports no substantial improvement in successfully aligned read pairs when iterative mapping in Hiclib is used for *Arabidopsis thaliana* Hi-C data

^dHiCdat provides three options for normalization: coverage and distance correction, HiCNorm and ICE

^eHOMER provides two options for normalization: simpleNorm corrects for sequencing coverage only and norm corrects for coverage plus the genomic distance between loci

^fHiTC provides two options for normalization: normLGF implements HiCNorm and normICE implements ICE algorithm from Hiclib

^gHiFive provides three options - Probability, Express, and Binning - for normalization. The Express and Binning algorithms correspond to matrix balancing and explicit-factor correction schemes, respectively

Servant et al., *Genome Biol.* 2015

Ay & Noble *Genome Biol* 2015