

Human Fibroblast IMR90 Hi-C Data (Dixon et al.)

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May 7, 2020

1 Introduction

The Hi-C technic was first introduced by [Lieberman-Aiden et al. \[2009\]](#). In the continuity with 3C, 4C and 5C technics, the goal of the Hi-C is to simultaneously detect all chromosomal contacts in a single experiment. All these technics aim at measuring the population-averaged frequency at which two genomic loci physically interact in three-dimensional space. In Hi-C, after a first crosslink and digestion, all genomic fragments are labeled with a biotinylated nucleotide before ligation. These junctions can then be purified efficiently by streptavidin-coated magnetic beads, and finally sequenced using a standard Illumina paired-end protocol.

The data available in this package were published by [Dixon et al. \[2012\]](#) and downloaded from the GEO website (GSE35156, sample GSM862724). This publication is one of the key papers in the field for two main reasons: i) it was the first time that Hi-C data were generated at such resolution (up to 20kb), ii) this resolution highlighted a new short range structure defined as topological domains (TADs), with high frequencies of intra-domain chromatin interactions but infrequent inter-domain chromatin interactions ([Nora et al. \[2012\]](#)).

If you use *HiCDataHumanIMR90*, please cite:

- Servant N (2014). HiCDataHumanIMR90: Human Fibroblast IMR90 HiC data from Dixon et al. 2012. R package version 1.1.0.
- Dixon JR, Selvaraj S, Yue F, Kim A et al. (2012) Topological domains in mammalian genomes identified by analysis of chromatin interactions. *Nature* 485(7398):376-80.

2 Hi-C Data

The `hic_imr90_40` object is a *HTClist* object (see the *HiTC* package for more information ([Servant et al. \[2012\]](#))). It contains the complete genome-wide HiC data, with all inter and intrachromosomal contact maps at a resolution of 40kb.

Human Fibroblast IMR90 Hi-C Data (Dixon et al.)

```
> require(HiCDataHumanIMR90)
> require(HiTC)
> data(Dixon2012_IMR90)
> ## Show data
> show(hic_imr90_40)

HTClist object of length 325
25 intra / 300 inter-chromosomal maps

> ## Is my data complete (i.e. composed of intra + inter chromosomal maps)
> isComplete(hic_imr90_40)

[1] TRUE

> ## Note that a complete object is not necessarily pairwise
> ## (is both chr1-chr2 and chr2-chr1 stored ?)
> isPairwise(hic_imr90_40)

[1] FALSE

> ## Which chromosomes ?
> seqlevels(hic_imr90_40)

[1] "chr1" "chr2" "chr3" "chr4" "chr5" "chr6" "chr7" "chr8" "chr9"
[10] "chr10" "chr11" "chr12" "chr13" "chr14" "chr15" "chr16" "chr17" "chr18"
[19] "chr19" "chr20" "chr21" "chr22" "chrX" "chrY" "chrM"

> ## Details about a given map
> detail(hic_imr90_40$chrXchrX)

HTC object
Focus on genomic region [chrX:1-155270560]
CIS Interaction Map
Matrix of Interaction data: [3882-3882]
Binned data - window size = 40000
3882 genome intervals
Total Reads = 15349610
Number of Interactions = 3362484
Median Frequency = 1
Sparsity = 0.112

> ## Descriptive statistics
> head(summary(hic_imr90_40))

      seq1 seq2  nbreads nbinteraction averagefreq medfreq sparsity
chr1chr1 chr1 chr1 25914788      4524734      5.7274      1  0.8835
chr1chr2 chr1 chr2  504332      497291      1.0142      1  0.9869
chr1chr3 chr1 chr3  440865      434917      1.0137      1  0.9859
chr1chr4 chr1 chr4  456924      450005      1.0154      1  0.9849
```

Human Fibroblast IMR90 Hi-C Data (Dixon et al.)

```
chr1chr5 chr1 chr5 399067 393926 1.0131 1 0.986
chr1chr6 chr1 chr6 382580 377654 1.013 1 0.9858
```

3 Topological Domains

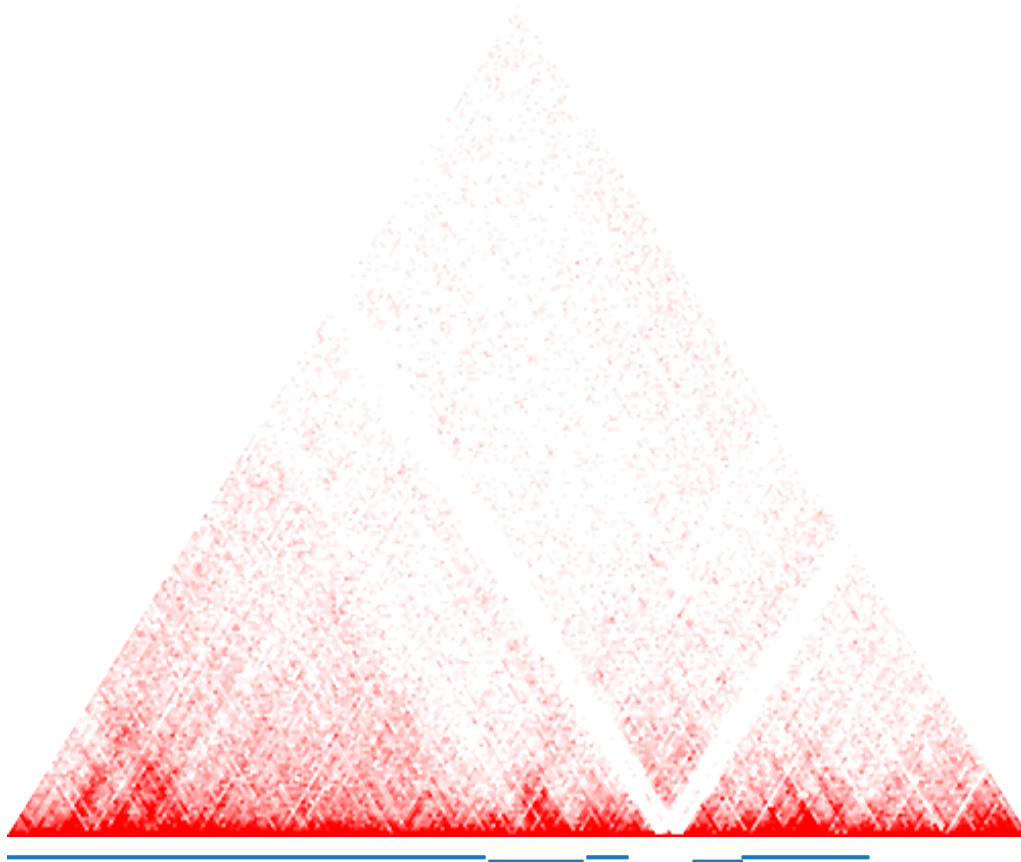
The `tads_imr90` object is a `GRanges` object with all TADs detected from this Hi-C data.

```
> show(tads_imr90)

GRanges object with 2338 ranges and 0 metadata columns:
      seqnames          ranges strand
      <Rle>            <IRanges> <Rle>
TAD-1   chr1      770138-1290137   *
TAD-2   chr1     1290138-1850140   *
TAD-3   chr1     1850141-2330140   *
TAD-4   chr1     2330141-3610140   *
TAD-5   chr1     3770141-6077413   *
...     ...           ...         ...
TAD-2334 chrX 146992309-148552096   *
TAD-2335 chrX 148592096-149929342   *
TAD-2336 chrX 149929343-151969344   *
TAD-2337 chrX 152089345-152746806   *
TAD-2338 chrX 152786807-154946806   *
-----
seqinfo: 23 sequences from an unspecified genome; no seqlengths
```

```
> ## Extract region
> regx <- extractRegion(hic_imr90_40$chrXchrX,
+                       chr="chrX", from=95000000, to=105000000)
> ## Plot Hi-C data with TADs
> plot(regx, tracks=list(tads_imr90), maxrange=20)
```

Human Fibroblast IMR90 Hi-C Data (Dixon et al.)



Package versions

This vignette was generated using the following package versions:

- R version 4.0.0 (2020-04-24), x86_64-pc-linux-gnu
- Running under: Ubuntu 18.04.4 LTS
- Matrix products: default
- BLAS: /home/biocbuild/bbs-3.11-bioc/R/lib/libRblas.so
- LAPACK: /home/biocbuild/bbs-3.11-bioc/R/lib/libRlapack.so
- Base packages: base, datasets, grDevices, graphics, methods, parallel, stats, stats4, utils
- Other packages: BiocGenerics 0.34.0, GenomInfoDb 1.24.0, GenomicRanges 1.40.0, HiCDataHumanIMR90 1.8.0, HiTC 1.32.0, IRanges 2.22.1, S4Vectors 0.26.0
- Loaded via a namespace (and not attached): Biobase 2.48.0, BiocManager 1.30.10, BiocParallel 1.22.0, BiocStyle 2.16.0, Biostrings 2.56.0, DelayedArray 0.14.0, GenomInfoDbData 1.2.3, GenomicAlignments 1.24.0, Matrix 1.2-18, RColorBrewer 1.1-2, RCurl 1.98-1.2, Rcpp 1.0.4.6, Rsamtools 2.4.0,

Human Fibroblast IMR90 Hi-C Data (Dixon et al.)

SummarizedExperiment 1.18.1, XML 3.99-0.3, XVector 0.28.0, bitops 1.0-6,
compiler 4.0.0, crayon 1.3.4, digest 0.6.25, evaluate 0.14, grid 4.0.0, htmltools 0.4.0,
knitr 1.28, lattice 0.20-41, matrixStats 0.56.0, rlang 0.4.6, rmarkdown 2.1,
rtracklayer 1.48.0, tools 4.0.0, xfun 0.13, yaml 2.2.1, zlibbioc 1.34.0

Human Fibroblast IMR90 Hi-C Data (Dixon et al.)

References

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- E. Lieberman-Aiden, N. L. van Berkum, L. Williams, M. Imakaev, T. Ragooczy, A. Telling, I. Amit, B. R. Lajoie, P. J. Sabo, M. O. Dorschner, R. Sandstrom, B. Bernstein, M. A. Bender, M. Groudine, A. Gnirke, J. Stamatoyannopoulos, L. A. Mirny, E. S. Lander, and J. Dekker. Comprehensive mapping of long-range interactions reveals folding principles of the human genome. *Science*, 326(5950):289–293, Oct 2009. doi: 10.1126/science.1181369. URL <http://dx.doi.org/10.1126/science.1181369>.
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- N. Servant, B. R. Lajoie, E. P. Nora, L. Giorgetti, C. Chen, E. Heard, J. Dekker, and E. Barillot. Hitc : Exploration of high-throughput 'c' experiments. *Bioinformatics*, Aug 2012. doi: 10.1093/bioinformatics/bts521. URL <http://dx.doi.org/10.1093/bioinformatics/bts521>.