scPloidy: Infer ploidy of single cells (or nuclei) based on single-cell (or single-nucleus) ATAC-seq data

New R package!

Available from

- CRAN
- https://github.com/fumi-github/scPloidy



Background, Aim

- Cells of different ploidy can constitute a tissue.
 - Hepatocyte: 2*x*, 4*x*, 8*x*
 - Cardiomyocyte: 2x, 4x
- Ploidy is related with cell function, damage, and regeneration.
- Standard experimental procedure is to stain DNA, separate by ploidy using a cell sorter, and perform single-cell analysis.
- If not sorted. Infer ploidy based on ATAC-seq data.

Methods

- ATAC-seq
 - Excise DNA fragments from open chromatin, and read by NGS.



- Idea
 - In a diploid cell, # of DNA fragments derived from one specific locus is 0, 1 or 2.
 - If tetraploid, 0, 1, 2, 3 or 4.
 - Model by binomial distribution and infer ploidy.



ATAC-seq fragment

Results

- Simulation
 - From human peripheral blood mononuclear cells (diploid), generate a mixture of 2x, 4x, 8x (1521 each).



- Actual example
 - Liver of a male rat (16 weeks old)
 - 4*x*, 8*x* should exist only in hepatocytes

| # of nuclei | Prediction | | |
|------------------|------------|------|-----|
| | 2x | 4x | 8x |
| Hepatocyte | 306 | 1427 | 593 |
| Endothelial cell | 457 | 125 | 0 |
| Stellate cell | 134 | 69 | 3 |
| Macrophage | 176 | 104 | 0 |
| T, NK cell | 81 | 40 | 0 |
| B cell | 38 | 18 | 1 |