The ChIA-PET data and analysis pipeline



Sheng Li

The Jackson Laboratory for Genomic Medicine

ENCODE NA call, May 26th, 2017

The goals and scopes of the ChIA-PET mapping center in ENCODE4

Our major goal is to produce high-quality, high-resolution, and comprehensive maps of long-range chromatin interactions between the structural and functional elements in human and mouse genomes.

Our scopes of data production:

- CTCF ChIA-PET for structural genome organization
- RNAPII ChIA-PET for transcriptional chromatin architecture
- 125 different biological samples focusing on
 - ENCODE cell lines
 - Subtypes of immune cells from human blood donors
 - Patient-derived iPSCs and iPSC-derived neurons
 - Mouse cell lines and primary tissues

Mapping and visualization



ChIA-PET process pipeline



PET clusters



PRE-PROCESSING

INTERACTION DISCOVERY



Signal vs noise assessment (1)



Li et al Genome Biology 2010

Li et al Cell 2012

6

Signal vs noise assessment (2)



Lib ID	Seq method	Bead dilution	Uniquely mapped PETs	Inter-species PETs	Noise rate	
LDS0221	Miseq	Low	5,746,593	608,618	10.59%	low dilution library
LDS0224	Miseq	Low	8,667,508	721,158	8.32%	
LDS0225	Miseq	Low	8,337,772	833,831	10.00%	
LDS0226	Miseq	Low	9,137,320	852,414	9.33%	
Total			31,889,193	3,016,021	9.46%	
LDS0227	Miseq	High	8,128,874	387,685	4.77%	high dilution library
LDS0228	Miseq	High	5,768,369	259,150	4.49%	
LDS0229	Miseq	High	5,745,104	226,540	3.94%	
LDS0230	Miseq	High	8,302,149	400,521	4.82%	
LDS0231	Miseq	High	5,192,920	83,834	1.61%	
LDS0232	Miseq	High	5,148,641	135,162	2.63%	
LDS0230N	Nextseq	High	141,314,800	6,801,514	4.81%	
LDS0231N	Nextseq	High	190,200,755	3,194,298	1.68%	
LDS0232N	Nextseq	High	157,849,191	4,167,384	2.64%	
Total			527,650,803	15,656,088	2.97%	

Signal vs noise assessment (3)



Hi-C

CTCF ChIA-PET



Tang et al., Cell 2015



Haplotype-resolved chromatin interactions



Single nt resolution of chromatin interactions linked to genetics



SNP-based validation of CTCF binding and looping



