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Transcriptional Interactome in Mouse Genome

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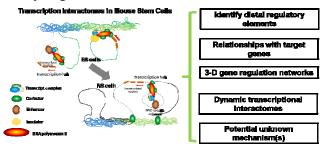
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Abstract

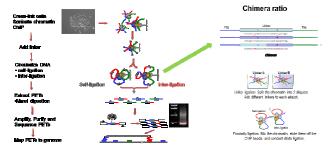
Distant regulatory elements have been shown to influence gene function *via* long-range chromatin interactions. In this study, we depict the dynamic transcriptional interactomes mediated by RNA polymerase II (RNAPII) with the newly developed Chromatin Interaction Analysis by Pair End diTag (ChIA-PET) to define the chromatin organizations pertinent to transcription regulation and epigenetic control in pluripotent embryonic stem cells (mES), neural stem cells (mNS) and neurosphere cells (mNP). With over 150 million ChIA-PET sequencing, thousands of *cis*- and *trans*- chromatin interactions tethered by RNAPII are defined. We plan to integrate the transcription interactome maps with global transcription expression changes, promoter activities, gene activities and major epigenomic features to define spatial distribution and temporal regulation of transcription networks.

I. Project goal



2. approaches used

CHIA-PET Procedure

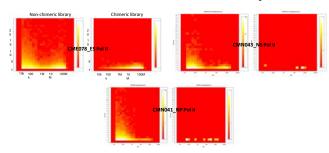


Result

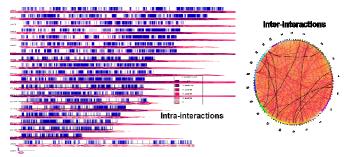
I. CHIA-PET sequencing and mapping summary

Library ID		ES_CME078	NS_CMN043	NP_CMN041
Total PETs		160.9M	152.6M	166.7M
Unique mapped PETs		94.6M (58.8%)	72.1M (47.3%)	79.8M (47.9%)
Merged Unique PETs (±2bp)		68.9M (41.8%)	38.7M (25.4%)	8.1M (4.9%)
Self-ligation PETs	PETs Binding Sites	26.9M (40.3%) 24,201	5.6M (14.3) 14,867	721K (8.9%) 11,819
intra-chromsoroal	PETs Interactions (2+)	3.004 (4.5%) 72,427	2.1M (5.5%) 9,671	765K (9.4%) 6,581
Inter-chromsomal	PETs Interactions (2+)	36.M (54.6%) 278,656	30.1M (79.7%) 59,152	6.5M (79.8%) 1,700

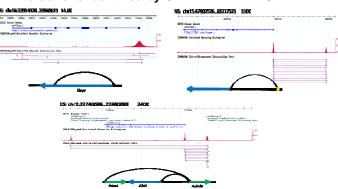
2. Evaluation noise level via chimera library



3. Overview of the mES Interactome across Genome



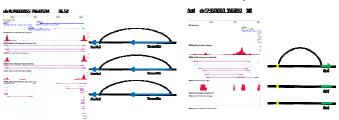
4. Interactions identified by CHIA-PET in mES



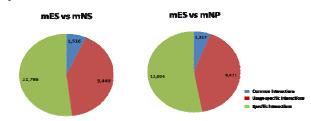
5. Specific Interactions

a. Common Interactions identified

b. Tissue specific interactions



6. Dynamic interactome across three cell lines



Ongoing works

- 1. Validation our dataset via various experiments, such as 3C, FISH
- 2. integrating with other genomic datasets, CHIP-Seq, RNA-Seq, RIP-Seq, methylome
- 3. Dynamic interactome analysis across the mouse genome
- 4. Modeling the 3-D gene regulation networks

References

Fullwood. M et al., Nature 462, 58-64, 2009 Li. G et. al., Genome Biology, 11(2):R22, 2010 Handoko. L et. al., Nature Genetics 43, 630-638, 2011