



**Wei Xie, Ph.D.**

**Principal Investigator**

**Tsinghua University**

**HHMI International Research Scholar**

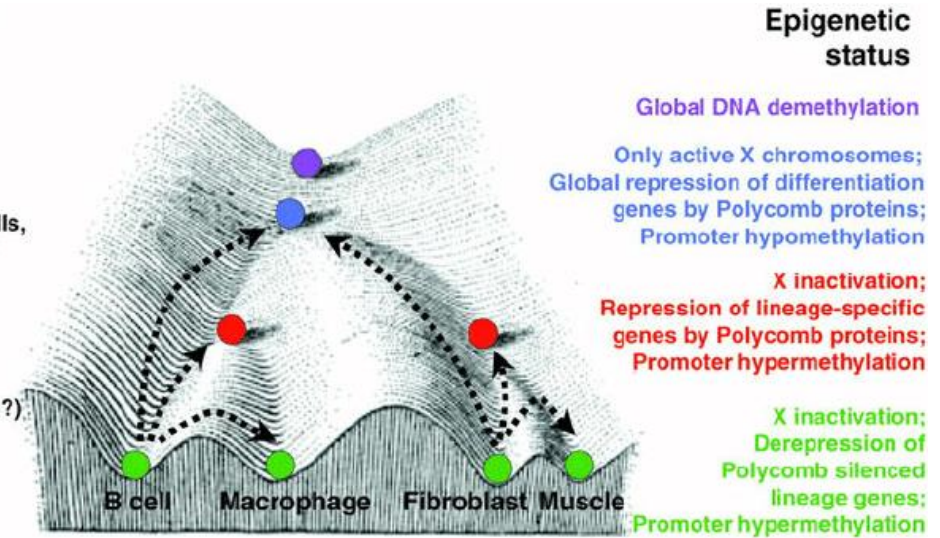
**Developmental potential**

**Totipotent**  
Zygote

**Pluripotent**  
ICM/ES cells, EG cells,  
EC cells, mGS cells  
iPS cells

**Multipotent**  
Adult stem cells  
(partially  
reprogrammed cells?)

**Unipotent**  
Differentiated cell  
types



**Yunhao Wang Ph.D.**

December 18, 2020

Department of Biomedical Informatics, Ohio State University

# Research @ Xie Lab since 2013

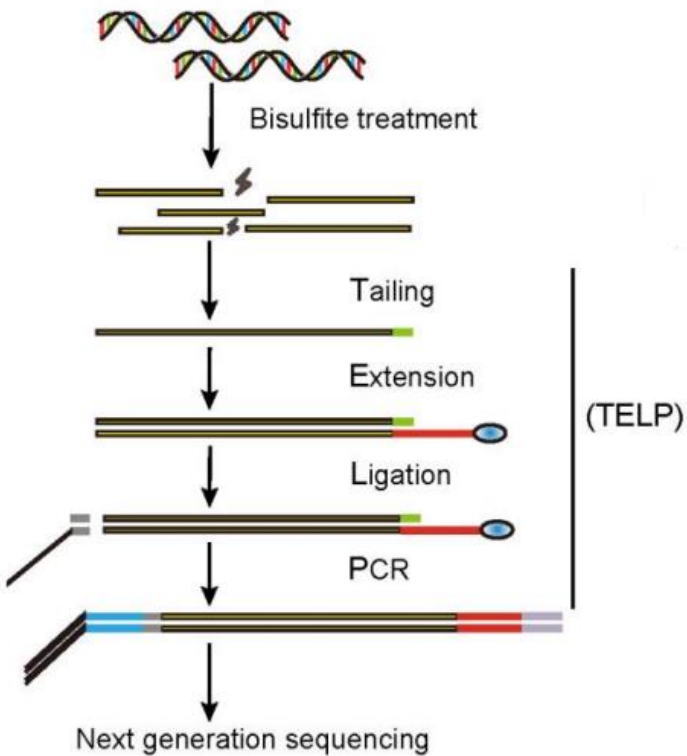
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- **Goal:** epigenetic regulation of mammalian early embryo development
- **Technology:** ultra-sensitive and low-input epigenetic analysis
  - **DNA methylation:** STEM-seq
  - **Histone modifications:** STAR ChIP-seq
  - **Chromatin accessibility:** miniATAC-seq
  - **3D chromatin architecture:** sisHi-C
  - **DNA binding:** Stacc-seq

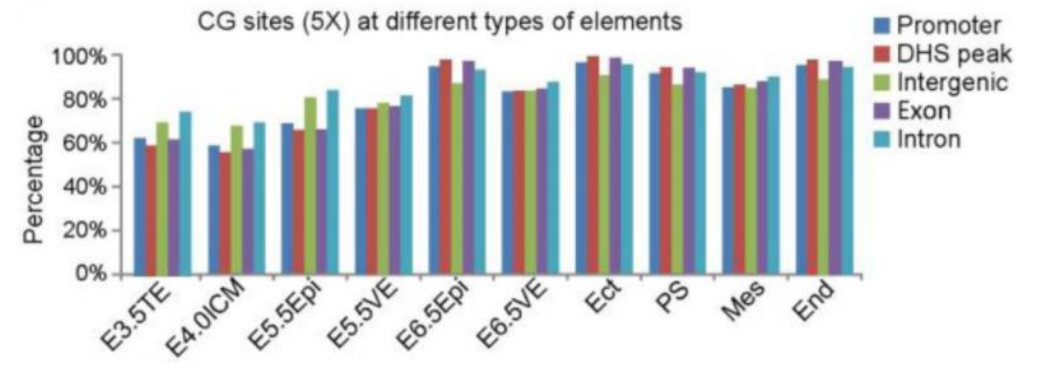
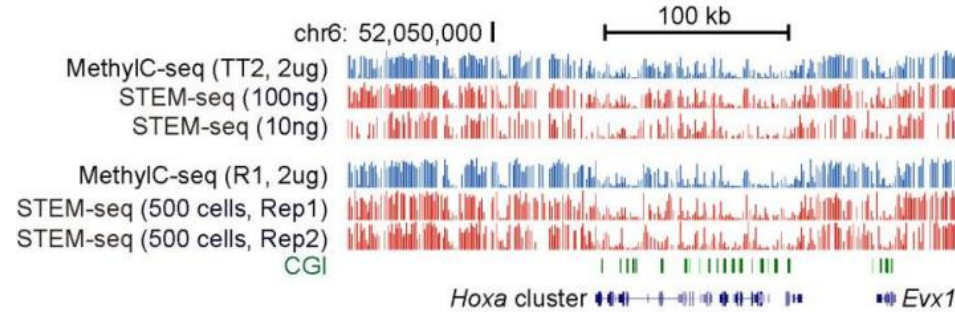
# DNA methylation: STEM-seq

## ➤ STEM-seq: small-scale TELP-enabled methylome sequencing (500 cells)

### The workflow of STEM-seq



### The comparison of STEM-seq and MethylC-seq (mESC)



### nature genetics

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nature > nature genetics > articles > article

Article | Published: 04 December 2017

### Dynamic epigenomic landscapes during early lineage specification in mouse embryos

Yu Zhang, Yunlong Xiang, Qiangzong Yin, Zhenhai Du, Xu Peng, Qijun Wang, Miguel Fidalgo, Weikun Xia, Yuanyuan Li, Zhen-ao Zhao, Wenhao Zhang, Jing Ma, Feng Xu, Jianlong Wang, Lei Li & Wei Xie

### Molecular Cell

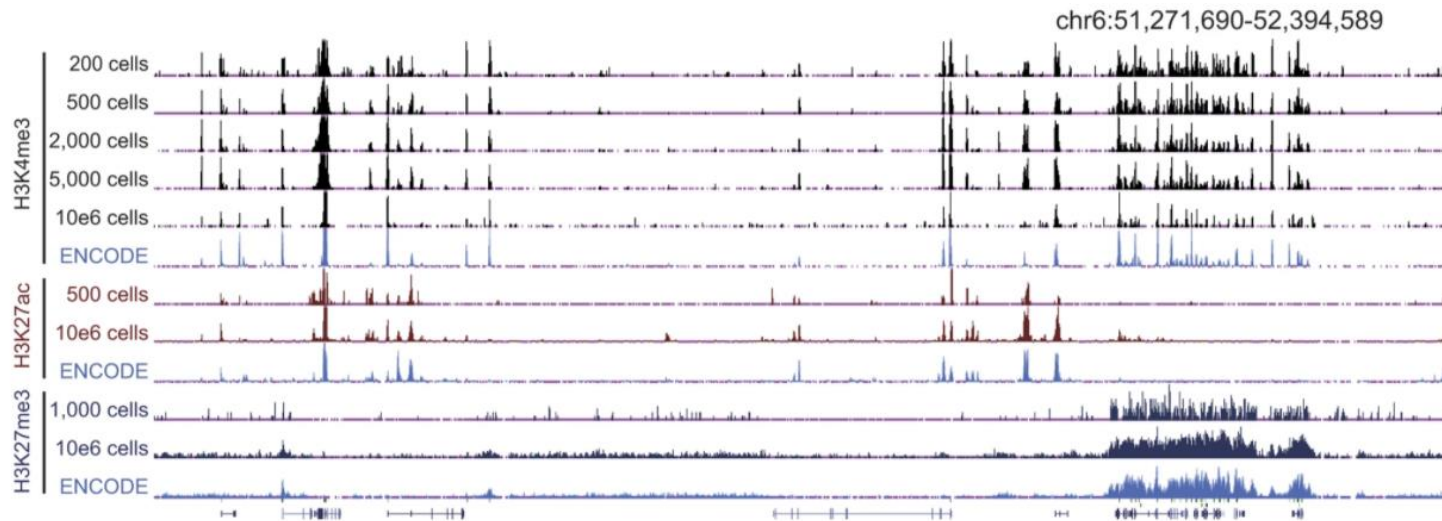
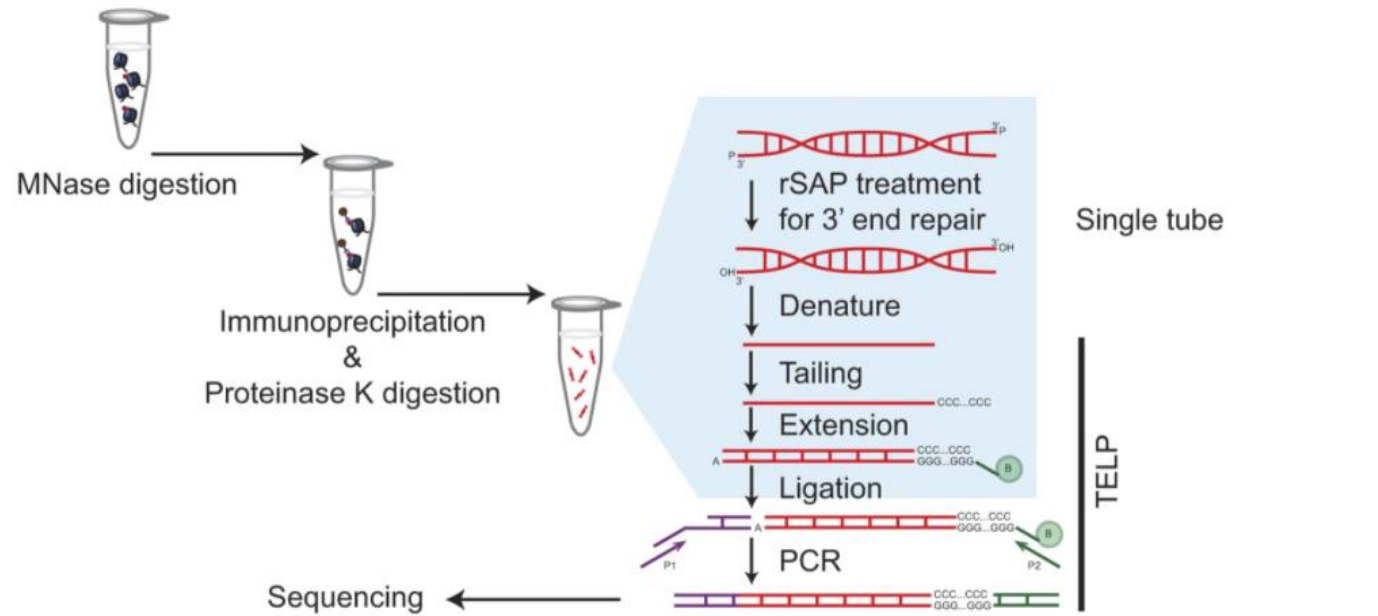
ARTICLE | VOLUME 64, ISSUE 6, P1062-1073, DECEMBER 15, 2016

### Isoform Switch of TET1 Regulates DNA Demethylation and Mouse Development

Wenhao Zhang<sup>10</sup>, Weikun Xia<sup>10</sup>, Qijun Wang, ... Shaorong Gao, Yong-hui Jiang, Wei Xie, 11

# Histone modifications: STAR ChIP-seq

## ➤ STAR ChIP-seq: small-scale TELP-assisted rapid ChIP-seq (200 cells)



nature

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Journal information

nature > letters > article

Published: 14 September 2016

### Allelic reprogramming of the histone modification H3K4me3 in early mammalian development

Bingjie Zhang, Hui Zheng, [...] Wei Xie

Nature 537, 553–557(2016) | Cite this article

nature genetics

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nature > nature genetics > articles > article

Article | Published: 16 December 2019

### Epigenomic analysis of gastrulation identifies a unique chromatin state for primed pluripotency

Yunlong Xiang, Yu Zhang, Qianhua Xu, Chen Zhou, Bofeng Liu, Zhenhai Du, Ke Zhang, Bingjie Zhang, Xiaoxiao Wang, Srimita Gayen, Ling Liu, Yao Wang, Yuanyuan Li, Qiujun Wang, Sundee Kalantry, Lei Li & Wei Xie

nature genetics

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Journal information

nature > nature genetics > articles > article

Article | Published: 29 April 2019

### SETD2 regulates the maternal epigenome, genomic imprinting and embryonic development

Qianhua Xu, Yunlong Xiang, Qiujun Wang, Leyun Wang, Julie Brind'Amour, Aaron Blair Bogutz, Yu Zhang, Bingjie Zhang, Guang Yu, Weikun Xia, Zhenhai Du, Chunyi Huang, Jing Ma, Hui Zheng, Yuanyuan Li, Chao Liu, Cheryl Lyn Walker, Eric Jonasch, Louis Lefebvre, Min Wu, Matthew C. Lorincz, Wei Li, Li Li & Wei Xie

Science

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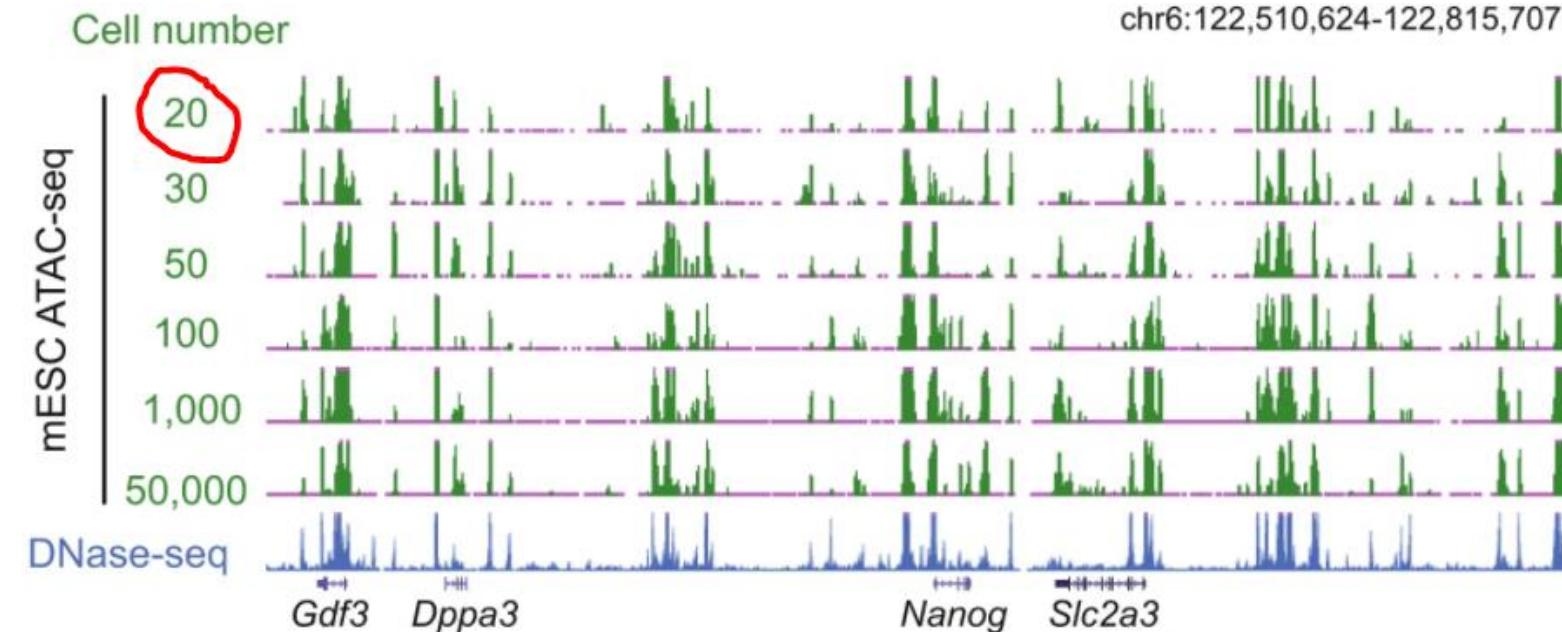
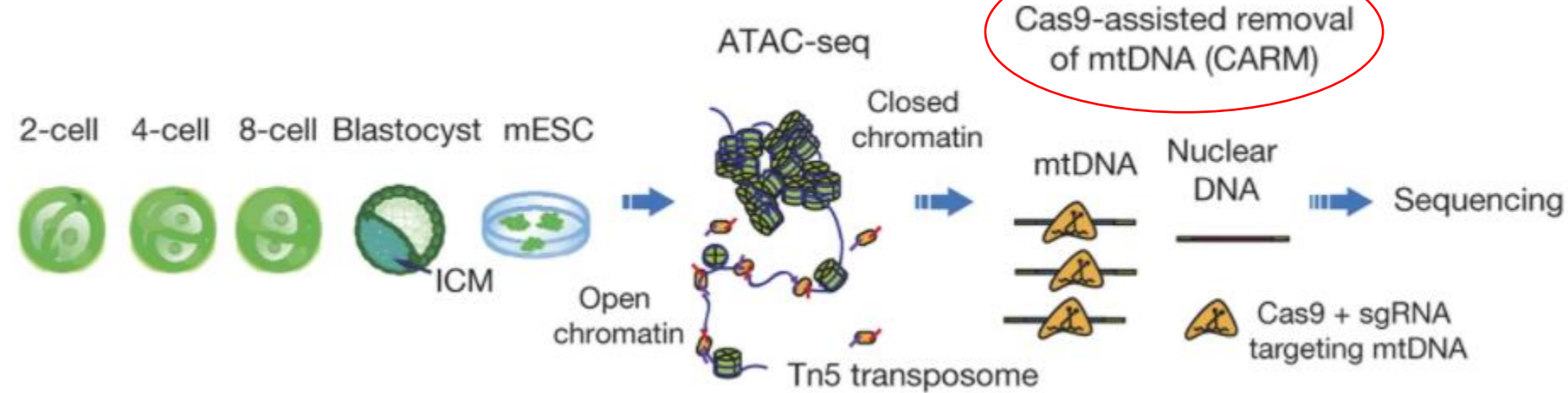
### Resetting histone modifications during human parental-to-zygotic transition

Weikun Xia<sup>1,\*</sup>, Jiawei Xu<sup>2,\*,†</sup>, Guang Yu<sup>1,\*</sup>, Guidong Yao<sup>2,†</sup>, Kai Xu<sup>1,†</sup>, Xueshan Ma<sup>2,†</sup>, Nan Zhang<sup>2,†</sup>, Bofen...



# Chromatin accessibility: miniATAC-seq

## ➤ miniATAC-seq: an improved ATAC-seq (20 cells)



nature

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[Journal information](#)

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Published: 15 June 2016

### The landscape of accessible chromatin in mammalian preimplantation embryos

Jingyi Wu, Bo Huang, He Chen, Qiangzong Yin, Yang Liu, Yunlong Xiang, Bingjie Zhang, Bofeng Liu, Qiujuan Wang, Weikun Xia, Wenzhi Li, Yuanyuan Li, Jing Ma, Xu Peng, Hui Zheng, Jia Ming, Wenhao Zhang, Jing Zhang, Geng Tian, Feng Xu, Zai Chang, Jie Na, Xuerui Yang & Wei Xie

nature

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Letter | Published: 02 May 2018

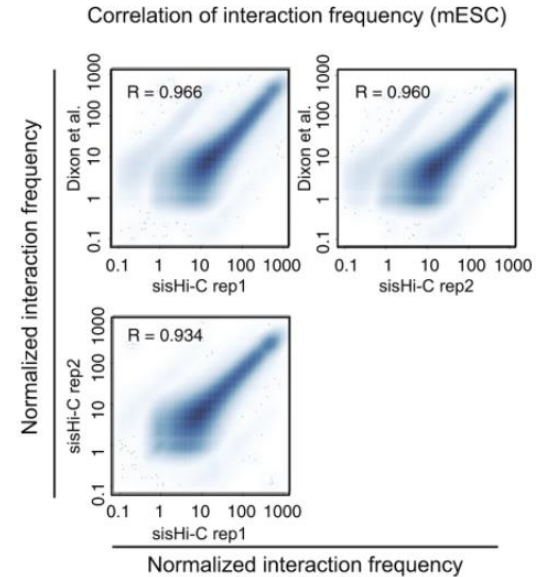
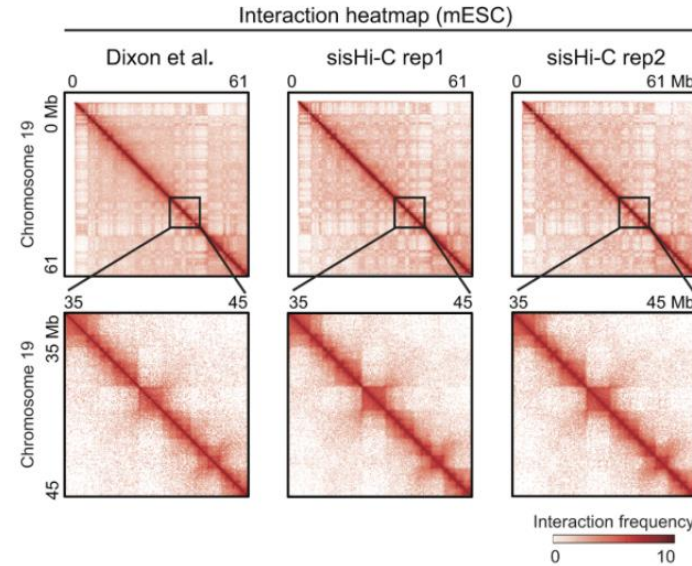
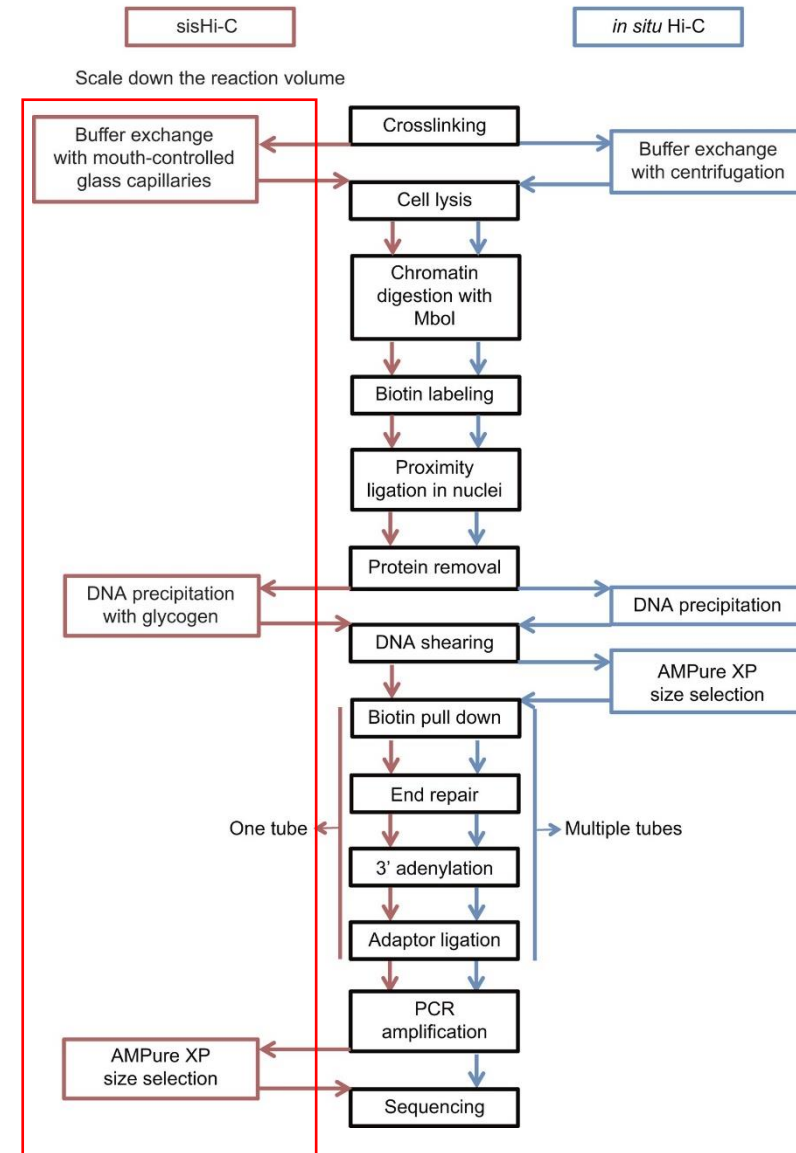
### Chromatin analysis in human early development reveals epigenetic transition during ZGA

Jingyi Wu, Jiawei Xu, [...] Yingpu Sun

# 3D chromatin architecture: sisHi-C

## ➤ sisHi-C: small-scale *in situ* Hi-C (500 cells)

Comparison of Hi-C protocols



nature

Explore our content ▼ Journal information ▼

nature > letters > article

Published: 13 July 2017

## Allelic reprogramming of 3D chromatin architecture during early mammalian development

Zhenhai Du, Hui Zheng, Bo Huang, Rui Ma, Jingyi Wu, Xianglin Zhang, Jing He, Yunlong Xiang, Qiujun Wang, Yuanyuan Li, Jing Ma, Xu Zhang, Ke Zhang, Yang Wang, Michael Q. Zhang, Juntao Gao, Jesse R. Dixon, Xiaowo Wang, Jianyang Zeng & Wei Xie

## Molecular Cell

ARTICLE | VOLUME 77, ISSUE 4, P825-839.E7, FEBRUARY 20, 2020

## Polycomb Group Proteins Regulate Chromatin Architecture in Mouse Oocytes and Early Embryos

Zhenhai Du <sup>11</sup> • Hui Zheng <sup>11</sup> • Yumiko K. Kawamura <sup>11</sup> • ... Kikue Tachibana • Antoine H.F.M. Peters • Wei Xie <sup>12</sup> • Show all authors • Show footnotes

## Molecular Cell

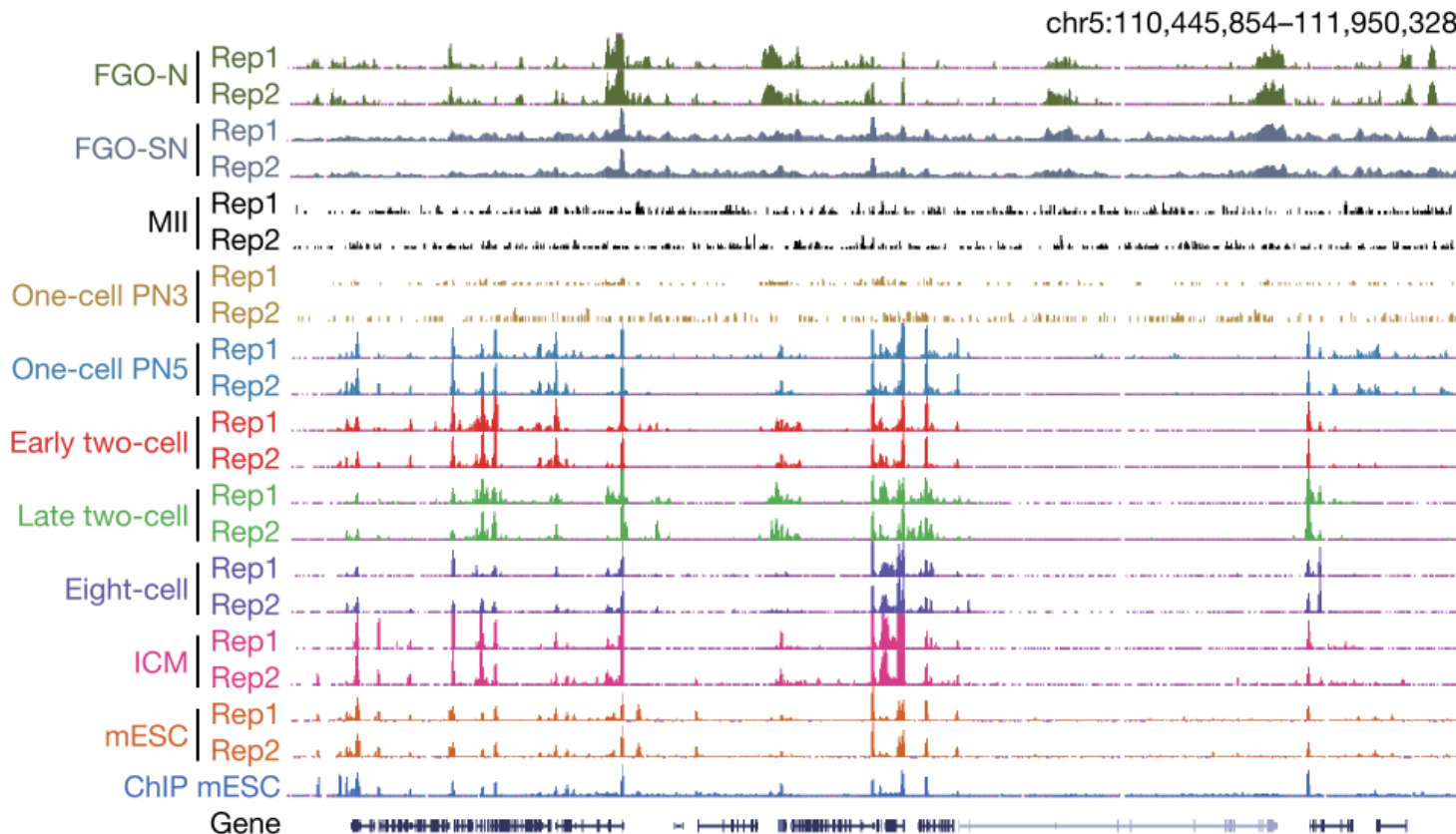
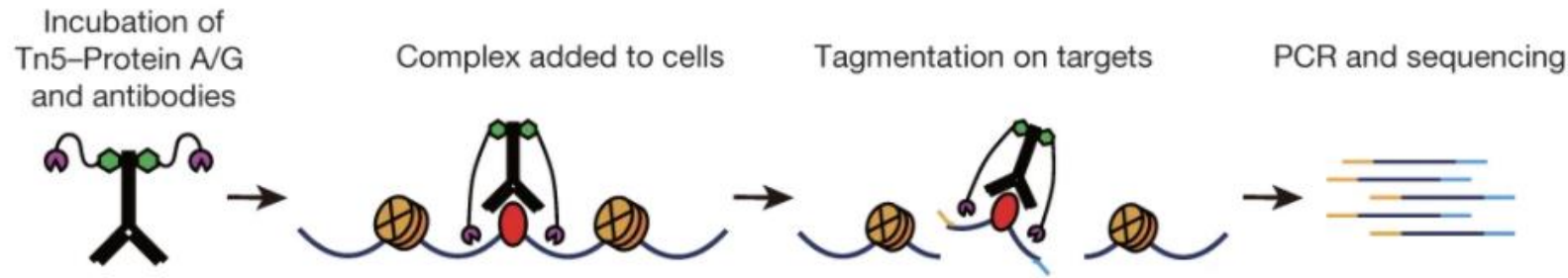
ARTICLE | VOLUME 73, ISSUE 3, P547-561.E6, FEBRUARY 07, 2019

## Reprogramming of Meiotic Chromatin Architecture during Spermatogenesis

Yao Wang <sup>5</sup> • Hanben Wang <sup>5</sup> • Yu Zhang <sup>5</sup> • ... Qinghua Shi • Xin Wu • Wei Xie <sup>6</sup> • Show all authors • Show footnotes

# DNA binding: Stacc-seq

➤ **Stacc-seq: small-scale Tn5-assisted chromatin cleavage with sequencing (200 cells, within 3.5 hour)**



nature

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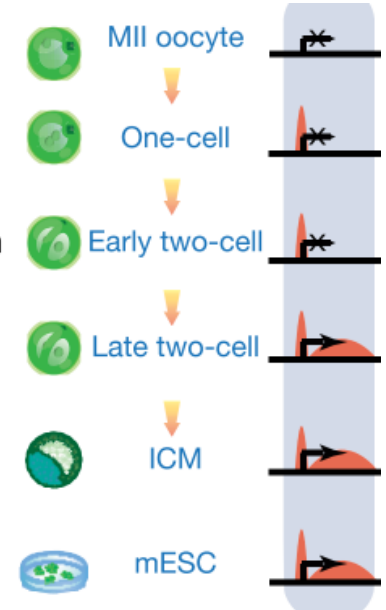
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Article | Published: 28 October 2020

## The landscape of RNA Pol II binding reveals a stepwise transition during ZGA

Bofeng Liu, Qianhua Xu, Qiujun Wang, Su Feng, Fangnong Lai, Peizhe Wang, Fangyuan Zheng, Yunlong Xiang, Jingyi Wu, Junwei Nie, Cui Qiu, Weikun Xia, Lijia Li, Guang Yu, Zili Lin, Kai Xu, Zhuqing Xiong, Feng Kong, Ling Liu, Chunyi Huang, Yang Yu, Jie Na & Wei Xie

Unloading



Reloading

Pre-configuration

Weak poising

Production

Strong poising

**Thanks**