BIOGRAPHICAL SKETCH

Provide the following information for the key personnel and other significant contributors in the order listed on Form Page 2. Follow this format for each person. **DO NOT EXCEED FOUR PAGES.**

NAME Chen, Xin	POSITION TITLE Associate Professor
eRA COMMONS USER NAME (credential, e.g., agency login) XIN.CHEN	

EDUCATION/TRAINING (Begin with baccalaureate or other initial professional education, such as nursing, and include postdoctoral training.)

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INSTITUTION AND LOCATION	DEGREE (if applicable)	YEAR(s)	FIELD OF STUDY
University of Science & Technology, Hefei, China	B.S.	9/91 - 6/96	Molecular Biology
University of Texas at Austin, TX	Ph.D	9/96 - 12/01	Mol. Cell & Dev. Biology
Stanford University School of Medicine	Postdoc	4/02 - 12/07	Developmental Biology

A. Personal Statement

B. Positions and Honors

Positions	and Em	ploy	/ment

1998 – 1999	Teaching Assistant, Department of Zoology, Univ. of Texas at Austin
1996 – 2001	Research Assistant, Molecular Cell and Developmental Biology, Univ. of Texas at Austin
	Advisor: Dr. Janice A. Fischer
2002-2007	Postdoctoral fellow, Dept. of Developmental Biology, Stanford University School of Medicine
	Advisor: Dr. Margaret T. Fuller.
2008-2014	Assistant Professor, Biology Department, The Johns Hopkins University
2015-	Associate Professor, Biology Department, The Johns Hopkins University

Other Experience and Professional Memberships

2001-	American Society of Genetics
2012	NIH DEV1 study section, ad hoc reviewer
2013	NIH Special Emphasis Panel/Scientific Review Group (Transgenerational Effects from
	Environmental Exposures) Study Section, member
2014	Faculty of 1000 member

Honors	
1991	The Fourth Zhang Zhong-Zhi Technology Scholarship in honor of First Place in the National-
	wide College Entrance Examination in Auhui province, USTC, China
1992	Excellent Student Fellowship, USTC, China
1993	Guang Hua Educational Fund Fellowship, USTC, China
1995	The Eighth Zhang Zhong-Zhi Technology Scholarship, USTC, China
1996 – 1998	Molecular Biology Institute Fellowship for Predoctoral Studies, UT, Austin
1998	Travel Funds from Department of Zoology, UT, Austin
2000	Travel Funds from Center for Developmental Biology, UT, Austin
2001	Dorothea Bennett Memorial Graduate Fellowship, UT, Austin
2001	Professional Development Award from the Office of Graduate Studies, UT, Austin
2002	Outstanding Dissertation award from Graduate School, University of Texas, Austin. One of the
	four awardees for academic years 2001 –2002, UT, Austin
2003	Katherine McCormick Travel Fund, Stanford, CA
2005	NIH NRSA Postdoctoral Fellowship, NICHD (declined)
2005 – 2006	The Leukemia and Lymphoma Society Special Fellow, Stanford, CA
2006 – 2010	The NIH Pathway to Independence (PI) Award (K99/R00)

2008- 2011	Edward Mallinckrodt, Jr. Foundation Young Investigator Award, the 49th Mallinckrodt Scholar
2009- 2011	Basil O'Connor Starter Scholar Research Award, March of Dimes
2009- 2014	David and Lucile Packard Fellowship for Science and Engineering
2011- 2012	Dean's Award for Excellence in Scholarship, Zanvyl Krieger School of Arts and Sciences, Johns Hopkins University
2015	Inaugural Catalyst Award, Zanvyl Krieger School of Arts and Sciences, Johns Hopkins University
2016	Finalist of the President's Frontier Award, Johns Hopkins University
2016	Howard Hughes Medical Institute, Bill & Melinda Gates Foundation, and the Simons Foundation Faculty Scholar
2016	Finalist for the 2016 Blavatnik National Awards for Young Scientists
2017	Finalist of the President's Frontier Award, Johns Hopkins University

C. Peer-reviewed Publications (in chronological order)

- 1. **Chen, X.**, Overstreet, E., Wood, S. A. and Fischer, J. A. (2000) On the conservation of function of the *Drosophila* Fat facets deubiquitinating enzyme and Fam, its mouse homolog. Dev. Genes Evol. 210: 603- 610.
- 2. **Chen, X**.*, Li, Q.* and Fischer, J. A. (2000) Genetic analysis of the *Drosophila* DNAprim gene: The function of the 60-kD primase subunit of DNA polymerase opposes the fat facets signaling pathway in the developing eye. <u>Genetics</u> 156: 1787-1795. (* equal contribution)
- 3. **Chen, X.** and Fischer, J. A. (2000) In vivo structure/function analysis of the *Drosophila* fat facets deubiquitinating enzyme gene. <u>Genetics</u> 156: 1829-1836.
- 4. **Chen, X.** and Fischer, J. A. (2002) A P element transformation vector for high levels of gene expression in germline cells of the ovary and undifferentiated cells in the developing eye of *Drosophila*. <u>Plasmid</u> 47: 61-65.
- 5. **Chen, X.**, Zhang, B. and Fischer, J. A. (2002) A specific protein substrate for deubiquitinating enzyme: Liquid facets is the substrate of Fat facets. <u>Genes and Development</u> 16: 289-294. One of the cover stories.
- 6. Overstreet, E., **Chen, X.**, Wendlend, B., and Fischer, J. A. (2003) Either part of a *Drosophila* Epsin protein, divided after the ENTH domain, functions in endocytosis of Delta in the developing eye. <u>Current Biology</u> 13: 854-860.
- 7. Hiller, M., Chen, X., Pringle, M.J., Suchorolski, M., Sancak, Y., Viswanathan, S., Bolival, B., Marino, S. and Fuller, M.T. (2004) Testis-specific TAF homologs collaborate to control a tissue-specific transcription program. <u>Development</u>, 131: 5297-5308.
- 8. **Chen, X.**, Hiller, M., Sancak, Y. and Fuller, M. T. (2005) Tissue specific TAFs counteract Polycomb to turn on terminal differentiation. <u>Science</u> 310: 869-872. Comment in BioEssays 28:330-334 (2006), by Ringrose, L. Featured in Faculty of 1000 Biology.
- 9. Krishnamoorthy, T., **Chen, X.**, Govin, J., Cheung, W.L., Dorsey, J., Schindler, K., Winter, E., Allis, C. D., Khochbin, S., Fuller, M. T., and Berger, S. L. (2006) Phosphorylation of histone H4 Ser1 regulates sporulation in yeast and is conserved in fly and mouse spermatogenesis. <u>Genes and Development</u>, 20: 2580–2592. PMCID: PMC1578680.

One of the cover stories. Comment in Genes and Development 20: 2487–2491(2006), by Wendt K.D. and Shilatifard A.

10. Chen, X. (2008) Stem cells- what can we learn from flies? Invited review for Fly. FLY 2-1: 19-28.

- 11. Gan, Q*, Chepelev, I*, Wei, G, Tarayrah, L, Cui, K, Zhao, K and **Chen, X.** (2010) Dynamic regulation of alternative splicing and chromatin structure in *Drosophila* gonads revealed by RNA-seq. <u>Cell Research</u> 20(7): 763-783 (2010). PMCID: PMC2919574. (* equal contribution)
- 12. Gan, Q, Schones, DE, Eun, S, Wei, G, Cui, K, Zhao, K and **Chen, X.** (2010) Monovalent and unpoised status of most genes in undifferentiated cell-enriched *Drosophila* testis. <u>Genome Biology</u> 11(4):R42 (2010). PMCID: PMC2884545.
- 13. Kracklauer, M.P., Wiora, H.M., Deery, W.J., **Chen, X.,** Bolival, B., Romanowicz, D., Simonette, R.A., Fuller, M.T., Fischer, J.A. and Beckingham, K.M. (2010) The *Drosophila* SUN protein Spag4 cooperates with the coiled-coil protein Turi Gagarin to maintain association of the basal body and spermatid nucleus. <u>Journal of Cell Science</u> 123 (16): 2763- 2772. PMCID: PMC2915878.
- 14. Eun, S*, Gan, Q* and **Chen, X.** (2010) Epigenetic regulation of germ cell differentiation. Current Opinion in Cell Biology 22, 737-743. PMCID: PMC2993805. (* equal contribution)
- 15. **Chen, X.***, Lu, C., Morillo, J., Eun, S. and Fuller, M.T.* (2011) Sequential changes at differentiation gene promoters as they become active in a stem cell lineage. <u>Development</u> 138: 2441-2450 (*co-corresponding authors.) PMCID: PMC3100706.
- 16. Tran, V., Gan, Q. and **Chen, X.** (2012) Chromatin immunoprecipitation (ChIP) using *Drosophila* tissue. <u>Journal of Visualized Experiments (JoVE)</u>, NIHMS429932, PMID: 22473446.
- 17. Cuddapah, S.*, Roh, T-Y., Cui, K., Fuller, M.T., Zhao, K. and **Chen, X.*** (2012) A novel human polycomb binding site acts as a functional polycomb response element in *Drosophila*, <u>PLoS One</u> 7(5):e36365, PMCID: PMC3343078. (* co-corresponding authors)
- 18. Lim, C.*, Tarayrah, L.* and **Chen, X.** (2012) Transcriptional regulation during *Drosophila* spermatogenesis. Invited review to <u>Spermatogenesis</u> 2(3):158-166, PMCID: PMC3469439 (* equal contribution).
- 19. Tran, V.*, Lim, C.*, Xie, J. and **Chen, X.** (2012) Asymmetric division of *Drosophila* male germline stem cell shows asymmetric histone distribution. <u>Science</u>, 338(6107): 679-682, PMCID: PMC3532436 (* equal contribution). Featured in Faculty of 1000 Biology.
- 20. Morillo Prado, J. R., **Chen, X.** and Fuller, M.T. (2012) Polycomb group genes *Psc* and *Su(z)*2 maintain somatic stem cell identity and activity in *Drosophila*. <u>PLoS One</u>, 7(12):e52892, PMCID: PMC3528704.
- 21. Eun, S., Stoiber, P.M., Wright, H. J., McMurdie, K.E., Choi, C.H., Gan, Q., Lim, C., **Chen, X.** (2013) MicroRNAs downregulate Bag of marbles to ensure proper terminal differentiation in *Drosophila* male germline lineage. <u>Development</u>, 140, 23-30, PMCID: PMC3513990.
- 22. Tarayrah, L., Herz, H-M., Shilatifard, A. and **Chen, X.** (2013) Histone demethylase dUTX directly antagonizes JAK-STAT signaling to maintain the *Drosophila* testis niche architecture. <u>Development</u>, 140, 1014-1023, PMCID: PMC3583039.
- 23. Chepelev, I. and **Chen, X.** (2013) Alternative splicing switching in stem cell lineages. Invited review to <u>Frontiers in Biology</u>, 8(1):50-59. PMCID: PMC3566875.
- 24. Barckmann, B., **Chen, X.**, Jayaramaiah-Raja, S., Kaiser, S., Rathke, C., Fuller, M.T., Renkawitz-Pohl, R. (2013) Three levels of regulation lead to protamine and Mst77F expression in *Drosophila*. <u>Developmental Biology</u>, 377:33-45. PMCID: PMC4154633.

- 25. Tran, V.*, Feng, L.J.* and **Chen, X.** (2013) Asymmetric distribution of histones during *Drosophila* male germline stem cell asymmetric divisions. Invited review to <u>Chromosome Research</u>, 21:255-269. PMCID: PMC4008969. (* equal contribution)
- 26. Chen, H-Y., **Chen, X.** and Zheng, Y-X. (2013) The nuclear lamina regulates germline stem cell niche organization via modulation of EGFR signaling. <u>Cell Stem Cell</u>, 13 (1):73–86, PMCID: PMC3703100.
- 27. Tarayrah, L.* and **Chen, X.*** (2013) Epigenetic regulation in adult stem cells and cancers. Invited review to <u>Cell and Bioscience</u>, 3:41. PMCID: PMC3852361. (* equal contribution)
- 28. Eun, S., Shi, Z., Cui, K., Zhao, K. and **Chen, X.** (2014) A non-cell autonomous role of E(z) to prevent germ cells from turning on a somatic cell marker. <u>Science</u>, 343(6178):1513-1516, PMCID: PMC4040133. Featured in Faculty of 1000 Biology.
- 29. Feng, L.J. and **Chen, X.** (2015) Epigenetic Regulation of Germ Cells— Remember or Forget? Invited review to Current Opinion in Genes and Development, 31:20-27. PMCID: PMC4470759.
- 30. Lim, C., Gandhi, S., Biniossek, M., Feng, L., Schilling, O., Urban, S. and **Chen, X.** (2015) An aminopepetidase acts in the *Drosophila* testicular niche for germline stem cell maintenance and spermatogonial dedifferentiation, <u>Cell Reports</u>, 13(2):315-325. PMCID: PMC4607668.
- 31. Xie, J., Wooten, M., Tran, V., Chen, B-C., Pozmanter, C., Simbolon, C., Betzig, E. and **Chen, X.** (2015) Histone H3 Threonine phosphorylation regulates asymmetric histone inheritance in the *Drosophila* male germline, <u>Cell</u>, 163(4): 920–933. PMCID: PMC4636931. Featured in Faculty of 1000 Biology. Previewed by Pirrotta, V. (2015) Histone Marks Direct Chromosome Segregation, <u>Cell</u>, 163(4): 792–793. Highlighted by Strzyz, P. (2015) Stem cells: Histone mark of stemness. <u>Nat Rev Mol Cell Biol.</u>, 16(12):703.
- 32. Tarayrah, L.*,*, Li, Y. *, Eun, S., Shi, Z., Gan, Q. and **Chen, X.*** (2015) Histone demethylase Lid maintains germline stem cells through regulating JAK-STAT signaling pathway activity, <u>Biology Open</u>, 4(11):1518-1527 (* equal contribution; * co-corresponding authors). PMCID: PMC4728359.
- 33. Feng, L., Shi, Z. and **Chen, X.** (2017) Enhancer of Polycomb coordinates multiple signaling pathways to promote both cyst and germline stem cell differentiation in *Drosophila* adult testis, <u>PLoS Genetics</u> 13:e1006571. PMCID: PMC5308785.
- 34. Xie, J.*, Wooten, M.*, Tran, V. and **Chen, X.** (2017) Breaking symmetry—asymmetry histone inheritance in stem cells, invited review to <u>Trends in Cell Biology</u>, PMID: 28268050, PMCID: in progress (* equal contribution).
- 35. Eun, S.*, Feng, L.*, Cedeno-Rosario, L., Gan, Q., Wei, G., Cui, K., Zhao, K., and **Chen, X.** (2017) Polycomb group gene E(z) is required for spermatogonial dedifferentiation in *Drosophila* adult testis, <u>Journal of Molecular Biology</u>, PMID: 28434938, PMCID: in progress (* equal contribution).
- 36. Feng, L., Shi, Z. and **Chen, X.** Enhancer of Polycomb maintains germline activity and genome integrity in *Drosophila* testis, resubmitted to <u>Nature Communications</u>.
- 37. Snedeker, J. *, Wooten, M. * and **Chen, X.** Invited review for Volume 33 on Asymmetric DNA and Histone Segregation to be published in October 2017, <u>Annual Review of Cell and Developmental Biology</u>, in revision for resubmission. (* equal contribution)
- 38. Gleason, R.J.*, Anand, A.*, Kai T.§ and **Chen, X.**§ Protecting and diversifying the germline, invited review to <u>Genetics</u> FlyBook, submitted. (* equal contribution; § co-corresponding authors).

D. Research Support Ongoing Research Support

R01GM112008, Chen (PI), Xiao (co-PI) 9/16/14-7/31/18

NIH, NIGMS

Molecular and cellular mechanisms underlying asymmetric histone inheritance

The major goals of this project are to investigate how preexisting and newly synthesized histone H3 are differentially incorporated into two sets of sister chromatids in S-phase germline stem cells; and to determine the mechanisms by which sister chromatids with preexisting *vs.* newly synthesized H3 are asymmetrically segregated in mitotic germline stem cells.

Role: Principal Investigator

R21HD084959, Chen (PI) 08/03/15-07/31/17

NIH, NICHD

Study the generality of asymmetric histone inheritance

The major goals of this project are to investigate the inheritance mode for all canonical histones, in other asymmetrically dividing cells in Drosophila, and in mouse embryonic stem cells that are induced to divide asymmetrically.

Role: Principal Investigator

Faculty Scholar, Chen (PI) 11/01/16- 10/31/21

Howard Hughes Medical Institute, Bill & Melinda Gates Foundation, and the Simons Foundation Investigate the mechanisms of asymmetric histone inheritance in Drosophila male germline stem cell asymmetric division, as well as the generality of this phenomenon in other cell types and in other organisms Role: Principal Investigator

Packard Fellow, Chen (PI) 10/15/09- 10/14/14, No cost extension from 10/15/14- 10/14/18

The David and Lucile Packard Foundation

Single-cell analyses of gene expression and chromatin structure in programming germline stem cells The major goals of this project are to analyze both transcriptome and epigenome of male germ cell cysts at each single stage during spermatogenesis.

Role: Principal Investigator

Completed Research Support (during the last three years)

1R01HD065816-01A1, Chen (PI) 8/24/11 – 4/30/16

NIH, NICHD

Epigenetic Regulation of Germ Cell Differentiation from a Stem Cell Lineage

Role: Principal Investigator