**B. Franklin Pugh, Ph.D.**

# Contact

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# Education

Cornell University, Ithaca NY B.S. 1983 Biology

University of Wisconsin-Madison Ph.D. 1987 Molecular Biology

# Professional Employment

* 1. Postdoctoral Research Associate.

*Michael Cox, Dept. of Biochemistry, University of Wisconsin-Madison.*

* 1. Postdoctoral Research Associate.

 *Robert Tjian, Dept. of Molecular & Cellular Biology, University of California-Berkeley.*

* 1. Assistant Professor of Biochemistry & Molecular Biology.

 *The Pennsylvania State University.*

1998-2005 Associate Professor of Biochemistry & Molecular Biology.

 *The Pennsylvania State University.*

2005- present Professor of Biochemistry & Molecular Biology.

 *The Pennsylvania State University.*

# Secondary Titles

2004-present Director – Center for Eukaryotic Gene Regulation

2007-present Willaman Professor of Molecular Biology.

2008-2017 Editor, *Molecular and Cellular Biology*

2014-present Evan Pugh Professor

2014-present Fellow AAAS

# Awards and Service

1988-91 Leukemia Society of America Postdoctoral Fellow

1992-95 Searle Scholar

1996-01 Leukemia & Lymphoma Society Scholar

1996 Daniel Tershak Faculty Teaching Award

1999-03 Member, American Cancer Society Peer Review Committee on Genetic Mechanisms

2001-03 Member, Peer Review Panel – Florida Department of Health

2002-present Member (ad hoc), NIH CDF-2, CDF-1, SEP, GCAT, MGB Study Sections

2001-05 Senator, Penn State Faculty Senate

2004-06 Chair - Penn State University Genomics Futures Committee

2006 Penn State University Faculty Scholars Medal

2006-2007 Chair – Eberly College of Science Committee – Dept Head search

2010 Designed the Center for Eukaryotic Gene Regulation, Penn State University

# Completed Thesis Supervised

17 BS - Honors 2 MS

18 PhD 10 Postdoc

# Classroom Teaching (Enrollment - Limit/Actual/Dept. averages for student ratings of effectiveness)

1992-2000 BMMB 514 – Molecular Biology and Cellular Regulation (>40 - 7.0/**6.2**/5.3)

2003-2007 BMB 251 – Molecular and Cellular Biology I (>200 – 7.0/**5.2**/5.3)

2003-2008 BMB 252 – Molecular and Cellular Biology II (>100 – 7.0/**5.8**/4.7)

2011-2014 BMB 252H – Molecular and Cellular Biology II honors (11 – 7.0/**5.0**/4.7)

2015-2017 BMB 543 – Gene Regulation (14-7/**10**/5.86)

2015-2017 BMB 488 – Communities of Practice in BMB Model System Cell Biology (4-2/**2**/5.86)

# Historical Funding

|  |  |  |  |
| --- | --- | --- | --- |
| ***Years*** | ***Direct ($)*** | ***Agency*** | ***Project*** |
| 1989-1992 | 71,000  | Leuk. Lymph Soc.  | *Eukaryotic Transcription Initiation* |
| 1991-20071992-1997 | 2,000,000158,000  | Penn State Univ.Searle Foundation | *Various Projects**Human Transcription Factor TFIID Complex* |
| 1992-1997 | 697,000 | Natl. Inst. Health  | *Biochemistry of Eukaryotic Gene Regulation* |
| 1998-2000 | 383,000  | Am. Cancer Soc.  | *TBP Associated Factors* |
| 1996-2001 | 200,000  | Leuk. Lymph Soc.  | *Dimerization of the TATA Binding Protein* |
| 1999-2001 | 151,000  | Natl. Sci. Found.  | *Regulation of TATA Binding Protein Dimers* |
| 2004-2007 | 366,000  | Natl. Sci. Found.  | *Computational Modeling of Transcriptional Regulation* |
| 2007-2018 | 1,973,000 | Natl. Inst. Health  | *Mapping of Functional Elements in the Yeast Genome* |
| 2007-2019 | 2,100,000 | Natl. Inst. Health  | *Genome Regulation in Response to Environmental Stress* |
| 2007-20232000-2021 | 1,437,0003,536,000 | Natl. Inst. HealthNatl. Inst. Health | *Mapping of Functional Elements in the Yeast Genome**Genome-wide regulation of the TATA Binding Protein*  |
| 2018-2021 | 840,000 | Natl. Inst. Health | *Genome-wide structural organization within human gene regulatory complexes* |

# Research Highlights (Past)

1987 Deciphered biochemical mechanisms of homologous recombination *(graduate student)*

1991 Discovered transcriptional co-activators *(postdoc)*

1992 Identified subunits of TFIIIB as TBP and TAFs *(assistant professor)*

1996 Discovered autoregulation of TFIID through dimerization *(assistant professor)*

2004 Identified genome regulation as TATA/SAGA/stress and TATA-less/TFIID/housekeeping *(assoc. professor)*

2007 Developed first ChIP-seq assay *(associate professor)*

2008 First high-resolution genome-wide map of Drosophila nucleosome organization *(professor)*

2011 Mapped binding locations of >200 transcription factors across the yeast genome *(professor)*

2011 First genome-wide biochemical reconstitution of chromosome nucleosome organization *(professor)*

2011 Invented the ChIP-exo assay *(professor)*

2012 Defined genomic structure of transcription machinery and chromatin remodelers at single bp resolution.

2014 Identified alternative nucleosome structures and asymmetry in composition on a genomic scale.

2015 Define nucleosome-specific chromatin remodeler organization on a genomic scale in mouse ES cells

2016 Reconstituted nucleosome organization on a genomic scale with purified histones and remodelers.

# Research Goals

Our goal is to understand how all nuclear proteins work together to regulate genomes, and apply this knowledge towards better diagnosis and management of human diseases. We use the well-known budding yeast as both a technological and conceptual model. To date, we have defined the positional organization along the genome of nearly all mappable nuclear proteins (>300) at near single-bp resolution using the ChIP-exo assay that we developed. This provides an understanding of the structural organization of protein complexes along the genome in vivo. Assembly dynamics are next monitored through rapid reprogramming of the genome (e.g., 5 min. of acute heat shock). Function is then assessed through CRISPR/Cas9-engineered depletion of factors, and through biochemical reconstitution of protein/DNA complexes on a genomic scale. Parallel strategies are being conducted in model human cell lines and clinical samples. We intend to identify the mappable protein/DNA interactions that correlate best with disease states and their treatment outcomes so as to develop improved medical diagnostics.

# Commercial activity

**2012** Founded Peconic, LLC – a service-based epigenome mapping company.

**2013** Rhee, H. S. and Pugh, B. F., Methods, Systems and Kits for Detecting Protein-Nucleic Acid Interactions. U.S. Patent No. 8,367,334 to Penn State. Licensed to Active Motif in 2011.

# Publications

1. Rojanaridpiched, C., Gracen, V. E., Everett, H. L., Coors, J. G., Pugh, B.F., and Boutyette, P. **(1984**). Multiple factor resistance in maize to European corn borer*.* ***Maydica*** *29*, 305-315.

2. Cox, M. M., Pugh, B.F., Schutte, B. C., Lindsley, J. E., Lee, J. W., and Morrical, S. W. (**1987**). On the mechanism of recA protein-promoted DNA branch migration. *In* ***DNA Replication and Recombination***, T. Kelly and R. McMacken, eds. (New York: Alan R. Liss, Inc.), pp. 597-607.

3. Pugh, B.F., and Cox, M. M. **(1987**). Stable binding of recA protein to duplex DNA. Unraveling a paradox*.* ***J. Biol. Chem.*** *262*, 1326-36.

4. Pugh, B.F., and Cox, M. M. **(1987**). recA protein binding to the heteroduplex product of DNA strand exchange*.* ***J. Biol. Chem.*** *262*, 1337-43.

5. Pugh, B.F., and Cox, M. M. (**1987**). Salt can functionally mimic DNA in activating the RecA protein ATPase. *In* ***Protein structure, folding, and design***: Alan R. Liss, Inc.), pp. 275-82.

6. Pugh, B.F., and Cox, M. M. **(1988**). General mechanism for RecA protein binding to duplex DNA*.* ***J. Mol. Biol.*** *203*, 479-93.

7. Pugh, B.F., and Cox, M. M. **(1988**). High salt activation of recA protein ATPase in the absence of DNA*.* ***J. Biol. Chem.*** *263*, 76-83.

8. Pugh, B.F., Schutte, B. C., and Cox, M. M. **(1989**). Extent of duplex DNA underwinding induced by RecA protein binding in the presence of ATP*.* ***J. Mol. Biol.*** *205*, 487-92.

9. Hoey, T., Dynlacht, B. D., Peterson, M. G., Pugh, B.F., and Tjian, R. **(1990**). Isolation and characterization of the Drosophila gene encoding the TATA box binding protein, TFIID*.* ***Cell*** *61*, 1179-86.

10. Peterson, M. G., Tanese, N., Pugh, B.F., and Tjian, R. **(1990**). Functional domains and upstream activation properties of cloned human TATA binding protein*.* ***Science*** *248*, 1625-30.

11. Pugh, B.F., and Tjian, R. **(1990**). Mechanism of transcriptional activation by Sp1: evidence for coactivators*.* ***Cell*** *61*, 1187-97.

12. Pugh, B.F., and Tjian, R. **(1991**). Transcription from a TATA-less promoter requires a multisubunit TFIID complex*.* ***Genes Dev.*** *5*, 1935-45.

13. Pugh, B.F., and Tjian, R. (**1991**). Regulation of transcription in animal cells:

factors and mechanisms. *In* ***Nuclear Processes and Oncogenes***, P. Sharp, ed. (New York: Academic Press, Inc.), pp. 201-210.

14. Tanese, N., Pugh, B.F., and Tjian, R. **(1991**). Coactivators for a proline-rich activator purified from the multisubunit human TFIID complex*.* ***Genes Dev.*** *5*, 2212-24.

15. Pugh, B.F., and Tjian, R. **(1992**). Diverse transcriptional functions of the multisubunit eukaryotic TFIID complex. ***J. Biol. Chem.*** *267*, 679-82.

16. Taggart, A. K., Fisher, T. S., and Pugh, B.F. **(1992**). The TATA-binding protein and associated factors are components of pol III transcription factor TFIIIB*.* ***Cell*** *71*, 1015-28.

17. Coleman, R. A., and Pugh, B.F. **(1995**). Evidence for functional binding and stable sliding of the TATA binding protein on nonspecific DNA*.* ***J. Biol. Chem.*** *270*, 13850-9.

18. Coleman, R. A., Taggart, A. K., Benjamin, L. R., and Pugh, B.F. **(1995**). Dimerization of the TATA binding protein*.* ***J. Biol. Chem.*** *270*, 13842-9.

19. Jackson, A. J., Ittmann, M., and Pugh, B.F. **(1995**). The BN51 protein is a polymerase (Pol)-specific subunit of RNA Pol III which reveals a link between Pol III transcription and pre-rRNA processing*.* ***Mol. Cell. Biol.*** *15*, 94-101.

20. Pugh, B.F. (**1995**). Purification of the human TATA-binding protein, TBP. *In* ***In Vitro Transcription and Translation Protocols***, M. J. Tymms, ed. (Totowa, N.J.: Humana Press, Inc.), pp. 359-67.

21. Pugh, B.F. (**1995**). Preparation of HeLa nuclear extracts. *In* ***In Vitro Transcription and Translation Protocols***, M. J. Tymms, ed. (Totowa, N.J.: Humana Press, Inc.), pp. 349-358.

22. Pugh, B.F. **(1996**). Mechanisms of transcription complex assembly*.* ***Curr. Opin. Cell Biol.*** *8*, 303-311.

23. Taggart, A. K., and Pugh, B.F. **(1996**). Dimerization of TFIID when not bound to DNA*.* ***Science*** *272*, 1331-3.

24. Booth Jr., B. L., and Pugh, B.F. **(1997**). Identification and characterization of a nuclease specific for the 3’ end of the U6 snRNA*.* ***J. Biol. Chem.*** ***272***, 984-991.

25. Coleman, R. A., and Pugh, B.F. **(1997**). Slow dimer dissociation of the TATA binding protein dictates the kinetics of DNA binding*.* ***Proc. Natl. Acad. Sci. USA*** *94*, 7221-6.

26. Pugh, B.F. (**1997**). Dynamics of transcription complex assembly. *In* ***Transcription Factors in Eukaryotes***, A. G. Papavassiliou, ed. (Austin, TX: R. G. Landes Co.).

27. Weideman, C. A., Netter, R. C., Benjamin, L. R., McAllister, J. J., Schmiedekamp, L. A., Coleman, R. A., and Pugh, B.F. **(1997**). Dynamic interplay of TFIIA, TBP and TATA DNA*.* ***J Mol Biol*** *271*, 61-75.

28. Chicca, J. J., 2nd, Auble, D. T., and Pugh, B.F. **(1998**). Cloning and biochemical characterization of TAF-172, a human homolog of yeast Mot1*.* ***Mol. Cell. Biol.*** *18*, 1701-10.

29. Jackson-Fisher, A. J., Chitikila, C., Mitra, M., and Pugh, B.F. **(1999**). A role for TBP dimerization in preventing unregulated gene expression*.* ***Mol. Cell*** *3*, 717-727.

30. Jackson-Fisher, A. J., Burma, S., Portnoy, M., Schneeweis, L., Coleman, R. A., Mitra, M., Chitikila, C., and Pugh, B.F. **(1999**). Dimer dissociation and thermosensitivity kinetics of the *Saccharomyces cerevisiae* and human TATA binding proteins*.* ***Biochemistry*** *38*, 11340-11348.

31. Coleman, R. A., Taggart, A. K. P., Burma, S., Chicca II, J. J., and Pugh, B.F. **(1999**). TFIIA regulates TBP and TFIID dimers*.* ***Mol. Cell*** *4*, 451-457.

32. Pugh, B.F. **(2000**). Control of gene expression through regulation of the TATA binding protein*.* ***Gene*** 255, *1-14*.

33. Pugh, B.F. (**2001**). The RNA polymerase II transcription machinery. *In* ***Transcription Factors***, J. Locker, ed. (Oxford: BIOS Scientific Publishers), pp. 1-16

34. Pugh, B.F. and D. Gilmour (**2001**) Genome-wide analysis of protein-DNA interactions in living cells. ***Genome Biol.*** 2, 1013.1-1013.3.

35. Chitikila, C., K. L. Huisinga, J. D. Irvin, M. Mitra, and B. F. Pugh (**2002**) Interplay of TBP inhibitors in global transcriptional control. ***Mol. Cell*** 10, 871-882.

36. Kou, H., Irvin, J. D., Huisinga, K. L., and Pugh, B.F. (**2003**). Structural and functional analysis of mutations along the crystallographic dimer interface of the yeast TATA binding protein*.* ***Mol. Cell. Biol.*** *23*, 3186-3201.

37. Huisinga, K. L., and Pugh, B.F. (**2004**). A genome-wide housekeeping role for TFIID and a highly regulated stress-related role for SAGA in Saccharomyces cerevisiae. ***Mol. Cell*** *13*, 573-585.

38. Basehoar, A. D., Zanton, S. J., and Pugh, B.F. (**2004**). Identification and distinct regulation of yeast TATA box-containing genes. ***Cell*** *116*, 699-709.

39. Pugh, B.F. (**2004**). Is acetylation the key to opening locked gates? ***Nature Struct. Mol. Biol.***  *11*, 298-300.

40. Kou, H., and Pugh, B.F. (**2004**). Engineering dimer stabilizing mutations in the TATA binding protein that reverse biological defects associated with dimer destabilizing mutants*.* ***J. Biol. Chem.*** *279*, 20966-73.

41. Zanton, S. J., and Pugh, B.F. (**2004**). Changes in genome-wide occupancy of core transcriptional regulators during heat stress. ***Proc. Natl Acad. Sci.***  *101*, 16843-16848.

42. Alexander, D. E., Kaczorowski, D., Lowry, D. M., Jackson-Fisher, A. J., Zanton, S. J. and Pugh, B.F. (**2004**). The Brf1 subunit of RNA polymerase III transcription factor TFIIIB induces TBP dimers to dissociate. ***J. Biol. Chem.*** *279*, 32401-32406.

43. Irvin, J. D., and Pugh, B.F. (**2006**). Genome-wide transcriptional dependence on TAF1 functional domains. ***J. Biol. Chem.*** 281, 6404-6412.

44. Durant, M., and Pugh, B.F. (**2006**). Genome-wide relationships between TAF1 and histone acetyltransferases in Saccharomyces cerevisiae. ***Mol. Cell. Biol***. 26, 2791-2802.

45. Zanton, S. J., and Pugh, B.F. (**2006**). Full and partial genome-wide assembly and disassembly of the yeast transcription machinery in response to heat shock. ***Genes Devel***. 20, 2250-2265. PMID: 16912275

46. Ioshikhes, I., Albert, I., Zanton, S. J., and Pugh, B.F. (**2006**). Nucleosome positions predicted through comparative genomics. ***Nature Genetics***. 38, 1210-1215. PMID: 16964265

47. Pugh, B.F. (**2006**). HATs off to PIC assembly. ***Mol Cell*** 23, 776-777.

48. Albert, I., Mavrich, T. N., Tomsho, L. P, Qi, J., Zanton, S. J., Schuster, S. C., and Pugh, B.F. (**2007**) Translational and rotational settings of H2A.Z nucleosomes across the S. cerevisiae genome. ***Nature*** 446, 572-576. PMID: 17392789

49. Huisinga, K.L. and Pugh, B.F. (**2007**) A TATA Binding Protein regulatory network that governs transcription complex assembly. ***Genome Biol*** 8(4): R46. PMID: 17407552

50. Durant, M. and Pugh, B.F. (**2007**) NuA4-directed chromatin transactions throughout the *S. cerevisiae* genome. ***Mol. Cell. Biol***. 15, 5327-35.

51. Venters, B. J., and Pugh, B.F. (**2007**). Chromatin meets RNA polymerase II. ***Genome Biol*** 8, 319.

52. Lee, C., Li, X., Hechmer, A., Eisen, M., Biggin, M.D., Venters, B.J., Jiang, C., Li, J., Pugh, B.F., and Gilmour, D.S. (**2008**). NELF and GAGA factor are linked to promoter proximal pausing at many genes in Drosophila. ***Mol. Cell. Biol.***28, 3290-3300. PMID: 18332113

53. Albert, I., Wachi, S., Jiang, C., and Pugh, B.F. (**2008**). GeneTrack - a genomic data processing and visualization framework. ***Bioinformatics***. *24*, 1305-1306. PMID: 18388141

54. Mavrich, T.N., Jiang, C., Ioshikhes, I.P., Li, X., Venters, B.J., Zanton, S.J., Tomsho, L.P., Qi, J., Glaser, R., Schuster, S.C., Gilmour, D.S., Albert, I., and Pugh, B.F. (**2008**). Nucleosome organization in the *Drosophila* genome. ***Nature.*** 453, 358-362. PMID: 18408708

55. Mavrich, T.N., Ioshikhes, I.P., Venters, B.J., Jiang, C., Albert, I., Tomsho, L.P., Qi, J., Schuster, S.C., and Pugh, B.F. (**2008**). A barrier nucleosome model for statistical positioning of nucleosomes throughout the yeast genome. ***Genome Res.***7, 1073-1083. PMID: 18550805

56. Yao, H., Li, P., Venters, B.J., Zheng, S., Thompson, P.R., Pugh, B.F., and Wang, Y. (**2008**). Histone Arg modifications and p53 regulate the expression of OKL38, a mediator of apoptosis. **J Biol Chem.** *283*, 20060-20068*.* PMID: 18499678

57. Azmy, YY, Gupta, A., and Pugh, B.F. (**2008**). Computational modeling of genome-wide transcription assembly networks using fluidics analogy. ***PLoS ONE*** 3, e3095. PMID: 18769485

58. Venters, B., and Pugh, B.F. (**2009**). A canonical promoter organization of the transcription machinery and its regulators in the Saccharomyces genome. ***Genome Res.*** 19, 360-371. PMID: 19124666

59. Jiang, C., and Pugh, B.F. (**2009**). Nucleosome positioning and gene regulation: advances through genomics. ***Nature Rev. Genet****. 10*, 161-172. PMID: 19204718

60. Venters, B., and Pugh, B.F. (**2009**). How eukaryotic genes are transcribed. ***Crit. Rev. Bioch. Mol. Biol.*** *44*, 117-141. PMID: 19514890

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62. Jiang, C., and Pugh, B.F. (**2009**). A compiled and systematic reference map of nucleosome positions across the Saccharomyces genome. ***Genome Biol.*** *10,* R109*.* PMID: 19814794

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65. Pugh BF (**2010**) A preoccupied position on nucleosomes. ***Nat Struct Mol Biol*.** 17, 923. PMID: 20683475

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86. Pugh, B.F. (**2013**). Molecular biology: The ends justify the means. ***Nature*** *497*, 48-49. PMID: 23615607

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88. Venters, B.J., and Pugh B.F. (**2013**) Genomic organization of human transcription initiation complexes. ***Nature*** *502*, 53-58. PMID: 24048476. *Retraction in: Venters BJ, Pugh BF.* ***Nature****. (****2014****) 513, 444 PMID: 25079311, due to statistical error.* Republished (**2016**) **PloS ONE** 11, e0149339. PMID: 26866362

89. Chang, G.S., Chen, X.A., Park, B., Rhee, H.S., Li, P., Han, K.H., Mishra, T., Chan-Salis, K.Y., Li, Y., Hardison, R.C., Wang Y., Pugh B.F. (**2014**) A comprehensive and high-resolution genome-wide response of p53 to stress. ***Cell Rep.*** *8*, 514-27. PMID: 25043190

90. Rhee, H-S., Bataille, A. R., Zhang, L., and Pugh, B.F. (**2014**) Subnucleosomal Structures and Nucleosome Asymmetry Across a Genome. ***Cell*** *159*, 1377-1388. PMID: 25480300

91. Wang S., Chen X.A., Hu J., Jiang J.K., Li Y., Chen-Salis K.Y., Gu Y., Chen G., Thomas C., Pugh B.F., Wang Y. (**2015**) ATF4 Gene Network Mediates Cellular Response to the Anticancer PAD Inhibitor YW3-56 in Triple Negative Breast Cancer Cells. ***Mol Cancer Ther.*** 14, 877-888. PMID: 25612620

92. Mahony S, Pugh BF (**2015**) Protein-DNA binding in high-resolution. ***Crit Rev Biochem Mol Biol.*** 3:1-15. PMID: 26038153

93. Han G.C., Vinayachandran V., Bataille A.R., Park B., Chan-Salis K.Y., Keller C.A., Long M., Mahony S., Hardison R.C., Pugh B.F. (**2015**) Genome-Wide Organization of GATA1 and TAL1 Determined at High Resolution. ***Mol Cell Biol*.** 36, 157-172. PMID: 26503782

94. Reja, R., Vinayachandran, V., Ghosh, S. & Pugh, B. F. (**2015**) Molecular mechanisms of ribosomal protein gene coregulation. ***Genes & Devel.*** 29, 1942-1954. PMID: 26385964

95. Schneider M., Hellerschmied D., Schubert T., Amlacher S., Vinayachandran V., Reja R., Pugh B.F., Clausen T., Köhler A. (**2015**) The Nuclear Pore-Associated TREX-2 Complex Employs Mediator to Regulate Gene Expression. ***Cell*** 162, 1016-1028. PMID: 26317468

96. de Dieuleveult M., Yen K.\*, Hmitou I., Depaux A., Boussouar F., Bou Dargham D., Jounier S., Humbertclaude H., Ribierre F., Baulard C., Farrell N.P., Park B., Keime C., Carrière L., Berlivet S., Gut M., Gut I., Werner M., Deleuze J.F., Olaso R., Aude J.C., Chantalat S., Pugh B.F.\*, Gérard M.\* (**2016**) Genome-wide nucleosome specificity and function of chromatin remodellers in ES cells. ***Nature*** 530, 113-116. PMID: 26814966 *\*co-corresponding author.*

97. Baranello L., Wojtowicz D., Cui K., Devaiah BN., Chung HJ., Chan-Salis KY., Guha R., Wilson K., Zhang X., Zhang H., Piotrowski J., Thomas C.J., Singer D.S., Pugh B.F., Pommier Y., Przytycka T.M., Kouzine F., Lewis B.A., Zhao K., Levens D. (**2016**). RNA Polymerase II Regulates Topoisomerase 1 Activity to Favor Efficient Transcription. **Cell** 165, 357–371. PMID: 27058666

98. Iwafuchi-Doi M., Donahue G., Kakumanu A., Watts J.A., Mahony S., Pugh B.F., Lee D., Kaestner K.H., Zaret K.S. (**2016**). The Pioneer Transcription Factor FoxA Maintains an Accessible Nucleosome Configuration at Enhancers for Tissue-Specific Gene Activation. **Mol Cell.** 62, 79-91. PMID: 27058788

99. Aguilar-Gurrieri, C., Larabi, A., Vinayachandran, V., Patel, N. A., Yen, K., Reja, R., Ebong, I. O., Schoehn, G., Robinson, C. V., Pugh, B. F. & Panne, D. (**2016**) Structural evidence for Nap1-dependent H2A-H2B deposition and nucleosome assembly. **EMBO J**. 35, 1465-1482. PMID: 27225933.

100. Krietenstein, N., Wal, M., Watanabe, S., Park, B., Peterson, C.L., Pugh\*, B.F., and Korber\*, P. (**2016**). Genomic nucleosome organization reconstituted with pure proteins. **Cell** 167, 709-721. PMID: 27768892 \**co-corresponding author*.

101. Jeronimo, C., Langelier, M.F., Bataille, A.R., Pascal, J.M., Pugh, B.F., and Robert, F. (**2016**). Tail and Kinase Modules Differently Regulate Core Mediator Recruitment and Function In Vivo. **Mol Cell** 64, 455-466. PMID: 27773677

102. Van Oss, S. B., Shirra, M. K., Bataille, A. R., Wier, A. D., Yen, K., Vinayachandran, V., Byeon, I. L., Cucinotatta, C. E., Heroux, A., Jeon, J., Kim, J., VanDenmark, A. P., Pugh, B. F. & Arndt, K. M. (**2016**) The histone modification domain of paf1 complex subunit rtf1 directly stimulates h2b ubiquitylation through an interaction with rad6. **Mol Cell** 64, 815-825 PMID: 27840029.

103. Lai, W. K. & Pugh, B. F. (**2017**). Genome-wide uniformity of human 'open' pre-initiation complexes. **Genome Res** 27, 15-26PMID: 27927716.

104. Lai WKM, Pugh BF. **(2017)** Understanding nucleosome dynamics and their links to gene expression and DNA replication. **Nat Rev Mol Cell Biol.** PMID: [28537572](https://www.ncbi.nlm.nih.gov/pubmed/?term=28537572)

105. Rossi MJ, Lai WKM, Pugh BF. **(2017)** Correspondence: DNA shape is insufficient to explain binding. **Nat Commun.**PMID: [28580956](https://www.ncbi.nlm.nih.gov/pubmed/?term=28580956)

**Significance of selected publications** -

*11. Pugh, B.F., and Tjian, R.* ***(1990****). Mechanism of transcriptional activation by Sp1: evidence for coactivators.* ***Cell*** *61, 1187-97.* This paper reports the initial discovery of a class of human proteins, termed “coactivators”, that connect transcriptional activators to the transcription machinery, thereby biochemically-defining gene regulatory circuit.

*23. Taggart, A. K., and Pugh, B.F.* ***(1996****). Dimerization of TFIID when not bound to DNA.* ***Science*** *272, 1331-3.* TFIID promotes gene transcription by assembling the transcription machinery via stable interactions with promoters. This paper describes the a mechanism by which human TFIID auto-regulates its own assembly, and suggesgts a novel mechanism by which genes are kept quiescent.

*38. Basehoar, A. D., Zanton, S. J., and Pugh, B.F. (****2004****). Identification and distinct regulation of yeast TATA box-containing genes.* ***Cell*** *116, 699-709.* The TATA box DNA element resides at the core of eukaryotic gene regulation, but it was unknown which genes contained a TATA box. This paper describes the first assignment of TATA boxes to genes on a genomic scale. This identification also precipitated the discovery of a deep dichotomy in the regulatory mechanism of eukaryotic genes. It defined stress-induced genes as having a distinct core transcription initiation complex and distinct mechanism of regulation compared to “housekeeping” genes. This discovery allowed scientists to infer how any gene of interest was likely to be regulated.

*48. Albert, I., Mavrich, T. N., Tomsho, L. P, Qi, J., Zanton, S. J., Schuster, S. C., and Pugh, B.F. (****2007****) Translational and rotational settings of H2A.Z nucleosomes across the S. cerevisiae genome.* ***Nature*** *446, 572-576.* Whether the nucleosomes that package eukaryotic chromosomes are randomly distributed across a genome or organized into specific patterns was unknown. This paper reports the first-ever ChIP-seq experiment, a technology that now dominates the field of functional genomics. Equally important, it demonstrated that every eukaryotic gene is packaged into an array of nucleosomes, all essentially organized in the same way (except for the TATA-class).

*54. Mavrich, T.N., Jiang, C., Ioshikhes, I.P., Li, X., Venters, B.J., Zanton, S.J., Tomsho, L.P., Qi, J., Glaser, R., Schuster, S.C., Gilmour, D.S., Albert, I., and Pugh, B.F. (****2008****). Nucleosome organization in the Drosophila genome.* ***Nature.*** *453, 358-362.* It was not known whether higher eukaryotes packaged their chromosomes as in yeast. This paper describes the first genome-wide map of individual nucleosome positions in flies. It also revealed the organization of the transcription machinery and DNA regulatory elements around nucleosomes on a genomic scale.

*71. Zhang, Z., Wippo, C.J., Wal, M. Ward, E., Korber, P., Pugh, B.F. (****2011****) A Packing Mechanism for Nucleosome Organization Reconstituted Across a Eukaryotic Genome.* ***Science****. 332, 977-980.* How nucleosomes become organized on genes was unclear. This manuscript reports the astonishing feat of biochemical reconstitution of proper nucleosome organization on a genomic scale using pure DNA, pure histones, ATP and a cell-free extract. It revealed the chromatin remodelers are responsible for organizing nucleosomes on a genomic scale. This opens the door to creating synthetic chromosomes that can be studied in vitro as enzymological substrates (see pub. 100).

*75. Rhee, H-S., and Pugh, B.F. (****2011****) Comprehensive genome-wide protein-DNA interactions detected at single nucleotide resolution.* ***Cell****. 147, 1408-19.* Technologies for determining where proteins bind along a genome had been rather low resolution, thereby limiting our understanding of chromosome regulation. This paper reports the development of the ChIP-exo assay. Its single-nucleotide resolution allows the structural organization of protein-DNA interactions to be defined on a genome scale.

*76. Rhee, H-S., and Pugh, B.F. (****2012****) Genome-wide structure and organization of eukaryotic pre-initiation complexes.* ***Nature****. 483, 295-301.* Here, the ChIP-exo assay was applied to define the first structure of transcription pre-initation complexes located at core promoters across a genome.

*81. Yen, K., Vinayachandran, V., Batta, K., Koerber, R.T., and Pugh, B.F. (****2012****). Genome­wide nucleosome specificity and directionality of chromatin remodelers.* ***Cell.*** *149, 1461-1473.* How chromatin remodelers organize nucleosomes on chromosomes was unknown. High-resolution mapping was used to show that remodelers use a “division-of-labor” approach to bind specific nucleosome positions on genes and slide them into an organized state.

*89. Rhee, H-S., Bataille, A. R., Zhang, L., and Pugh, B.F. (****2014****) Subnucleosomal Structures and Nucleosome Asymmetry Across a Genome.* ***Cell*** *159, 1377-1388.* This work reports on the widespread subnucleosomal structures in dynamic chromatin, including novel half-nucleosomes, regulated interactions of H3 tails with linker DNA and asymmetric placement of histone variants and modifications on nucleosomes in relation to the direction of transcription.

*100. Krietenstein, N., Wal, M., Watanabe, S., Park, B., Peterson, C.L., Pugh\*, B.F., and Korber\*, P. (****2016****). Genomic nucleosome organization reconstituted with pure proteins. Cell 167, 709-721.* \**co-corresponding author*. This work was performed equally between the two labs, and reports on the genome-wide reconstitution of promoter nucleosome organization with purified proteins. This represents the first-ever biochemical reconstitution of an entire epigenome (chromatin on a genomic scale) using only pure components: histones, DNA, six remodelers, and two sequence-specific DNA binding proteins. From this, a four-stage genome-wide assembly mechanism was deduced, that now paves the way for detailed mechanistic understanding of chromatin regulation through a blend of redundancy and specialization.

# Invited Lectures (accepted)

|  |  |
| --- | --- |
| 1987 | University of California - Berkeley |
| 1988 | Rockefeller University |
| 1990 | European Molecular Biology Laboratory (EMBL), Heidelberg, Germany |
|  | University of Lausanne, Switzerland |
|  | Institut de Chimie Biologique, Strasbourg, France |
|  | University of Chicago |
|  | University of Wisconsin-Madison |
| 1991 | Albert Einstein College of Medicine |
|  | Rutgers University |
|  | Harvard University Medical School |
|  | The Pennsylvania State University |
|  | Johns Hopkins University |
| 1992 | Keystone meeting on Molecular and Cellular Biology, Fundamental Mechanisms of Transcription. |
|  | The Pennsylvania State University, Hershey Medical Center. |
| 1993 | 15th International tRNA Workshop (Cap d'Agde, France). |
|  | Wistar Institute, Philadelphia. |
|  | University of Medicine and Dentistry of New Jersey. |
|  | Biochimie et Génétique Moléculaire, Centre d'Estude de Saclay, France. |
|  | Bowling Green State University, OH. |
| 1994 | Keystone meeting on Molecular and Cellular Biology, Fundamental Mechanisms of Transcription. |
|  | FASEB Meeting, Santa Cruz, CA |
|  | University of Vermont. |
| 1995 | Searle/Chicago Community Trust. |
|  | Penn State Summer Symposium in Molecular Biology on Chromosomal Controls of Gene Expression. |
|  | Cold Spring Harbor Meeting on Cancer Cells Cold Spring Harbor Laboratory. |
|  | Pennsylvania State University, Dept. of Chemistry. |
| 1996 | Symposium on Protein-Protein Interactions, Penn State University |  |
|  | University of Texas - Austin. |
|  | University of North Carolina - Chapel Hill. |
| 1997 | The Pennsylvania State University, Hershey Medical Center. |
|  | Cold Spring Harbor Gene Regulation Course. |
| 1998 | Keystone meeting on Transcriptional Mechanisms, Taos, NM |
| 1999 | Penn State University, Summer Symposium,  |
|  | ASBMB Fall Symposium, Lake Tahoe, CA |
|  | Towsen University, MD |
| 2001 | Penn State University, Hershey, PA |
|  | Eukaryotic Transcription Meeting, Cold Spring Harbor Laboratory, NY |
| 2002 | Integrated Bioinformatics Meeting, Zurich, Switzerland |
|  | Mid-Atlantic Yeast Conference, Cornell University, NY |
|  | New York University, School of Medicine, NY |
| 2003 | Keystone Meeting on Transcription, Santa Fe, NM |
|  | Summer Symposium, Penn State University |
|  | Michigan State University, East Lansing, MI |
|  | Ohio State University - OARDC, Wooster, OH |
| 2004 | University of California, Davis, CA |
|  | Systems Biology Meeting, Cold Spring Harbor Laboratory, NY |
|  | University of South Carolina – Columbia, SC |
|  | Center for Cellular & Molecular Biology, Hyderabad, India |
|  | Department of Biotechnology, New Delhi, India  |
|  | Indian Transcription Assembly Meeting, Pune, India |
| 2005 | Michigan State University, East Lansing, MI |
|  | Systems Biology Meeting, Cold Spring Harbor Laboratory, NY |
|  | Summer Symposium, Penn State University |

2006 MD Anderson Cancer Center, Houston, TX

 University of Rochester, Rochester, NY

 ASBMB Meeting on Chromatin, Kiawah, SC

2007 Eukaryotic Transcription Meeting, Cold Spring Harbor Laboratory, NY

 Center for Cell and Molecular Biology, Hyderabad, India

 Indian Institute of Sciences, Bangalore, India

 Bioconvene Conference on Bioinformatics, Hyderabad, India

 Vimta Labs, Hyderabad, India

 University of Southern Illinois, Carbondale, IL

 University of Missouri, Kansas City, MO

 Stowers Institute, Kansas City, MO

2008 State University of New York, Stony Brook, NY

 EMBO Yeast transcription meeting, St. Feliu de Guixol, Spain

 BioMaPS Summer School in Chromatin, Rutgers University, NJ

 Transcription Regulation by Chromatin meeting, Michigan State University, MI

 EMBL Transcription meeting, Heidelberg, Germany

 Scottish Chromatin Group meeting, Dundee, Scotland

 ASBMB meeting on Transcriptin Regulation, Granlibakken, Lake Tahoe, NV

 Chromatin and Transcription Regulation meeting, Weizmann Institute, Israel

 Pittsburgh Area Chromatin Meeting, University of Pittsburgh, PA

 Dept. of Biology, Duquesne University, Pittsburgh, PA

 Dept. of Biochemistry, State University of New York, Buffalo, NY

2009 CEA Genomics Institute, Paris, France

 ASMBMB meeting on Gene Regulation, New Orleans, La (Venters)

 24th International Conference on Yeast Genetics & Molecular Biology, Manchester, UK (Yen)

 16th Conversation on nucleosome positioning, Albany, NY

 Cold Spring Harbor Course on Eukaryotic Gene Regulation, CSHL, NY

 Mechanisms of Eukaryotic Transcription Meeting, Cold Spring Harbor Laboratory, NY

 SKMB Gene Regulation workshop, Lausanne, Switzerland

 Dept of Biochemistry, UCLA, CA

 Dept of Biological Science, Columbia U., New York

 Kansas University Medical Center, Kansas City, KS

 Ohio Plant Biotechnology Consortium

2010 Dept. Biochemisty and Molecular Biology, University of Florida, Gainesville, FL

 University of Sherbrooke, Sherbrooke, Canada

 Albert Einstein University, New York

 EMBO Yeast transcription meeting, St. Feliu de Guixol, Spain

 IRCB, Montreal, Canada

 Emory University, Atlanta, GA

 NIEHS, NIH, Chapel Hill, NC

 U. Pennsylvania, Philadelphia, PA

 Chromatin and epigenetics meeting, Munich, Germany

 ASBMB meeting on transcription, Granlibakken, NV

 Gordon Conference on chromatin, NH

 Keystone meeting on transcription, Big Sky, MT

 Intelligent Systems and Molecular Biology, Boston, MA

2011 EMBL meeting on chromatin, Heidelberg, Germany

 Fred Hutchinson Cancer Research Center, Seattle, WA

 Keystone Symposium Histone Code: Fact or Fiction?, Midway, UT

 NIH Workshop: NCI Center of Excellence in Chromatin Biology, Bethesda, MD

 University of Texas Southwestern, Dallas, TX

 NIH Symposium: Chromosome Structure and Function Symposium, Bethesda, MD

 Northwestern University, Evanston, IL

 Cold Spring Harbor Course on Eukaryotic Gene Regulation, CSHL, NY

 Cold Spring Harbor meeting on Eukaryotic Gene Regulation, CSHL, NY

 Chromatin and Systems Biology Meeting, Spetses, Greece

 High Throughput Approaches to Epigenomics Meeting, Bordeaux, France

2012 Memorial Sloan Kettering Cancer Research, NY, NY

 U. Virginia, Charlottesville, VA

 Cold Spring Harbor meeting on Systems Biology, CSHL, NY

 Cold Spring Harbor course on Eukaryotic Gene Regulation, CSHL, NY

 Gordon Research Conference on Chromatin Structure and Function, Il Ciocco, Italy

 Rafael University, Milan, Italy

 Genomic Symposium, New York University, NY

 U. Utrecht, Netherlands

 EMBO Yeast transcription meeting, St. Feliu de Guixol, Spain

 Beyond the Genome, Harvard Medical School, Boston

 Cornell University, Ithaca, NY

 10th EMBL meeting “Transcription and Chromatin”, Heidelberg, Germany

 Yeast Genetics Meeting, Princeton, NJ

 MD Anderson Cancer Center, Houston, TX

 Conference on Gene Regulation, Athens, Greece *(Keynote speaker)*

 EMBL meeting “From Functional Genomics to Systems Biology”, Heidelberg, Germany

 Meeting on Genetics Symposium, U. Pennsylvania, Philadelphia, PA *(Keynote speaker)*

 Temple University, Philadelphia, PA

 University of Michigan, Ann Arbor, MI

 Harvard Medical School, Boston, MA *(invited by students)*

2013 Pacific Symposium on Biocomputing, Kona, HI

 Epigenitic and Chromatin meeting, Harvard Medical School, MA

 Keystone meeting on Epigenetic Marks and Cancer Drugs, Santa Fe, NM

 University of Colorado, Boulder, CO

 Meeting on high-throughput DNA sequencing and the central dogma, Insitut Curie, Paris, France

 Meeting on Chromatin and Epigenetics, Heidelberg, Germany

 FASEB meeting on transcription, chromatin and epigenetics, Nassau, Bahamas

 Cold Spring Harbor course on Eukaryotic Gene Regulation, CSHL, NY

 Stonybrook University, Stonybrook, NY *(invited by students)*

 Cold Spring Harbor meeting on Eukaryotic Gene Regulation, CSHL, NY

 Ludwig-Maximillian University, Munich, Germany *(Octoberfest lecture)*

Meeting on chromatin and the DNA damage response, Boston, MA *(Keynote speaker)*

 EMBO meeting on nuclear structure and dynamics, Avignon, France

 Vienna Biocenter, Vienna, Austria

 Meeting on chromatin structure and function, Grand Cayman Islands

 Stanford University, Palo Alto, Ca

 University of California, San Francisco, CA

2014 Keystone meeting on nuclear receptors and biological networks, Taos, NM *(Keynote speaker)*

 Keystone meeting on transcription regulation, Santa Fe, NM

 ASBMB Annual Meeting, San Diego, CA

 Meeting on Nucleosomes and Chromatin, Hinxton, UK

 Gordon Conference on Chromatin Structure and Function, Waltham, MA

EMBO Conference on Gene Transcription in Yeast, San Felieu, Spain

FASEB Meeting on Yeast Chromatin Structure, Steamboat Springs, CO

EMBL Conference on Transcription and Chromatin, Heidelberg, Germany

 Meeting on Total Transcription, Hinxton, UK

 Cold Spring Harbor Meeting on Chromatin and Epigenetics, CSHL, NY

 Stowers Institute, Kansas City, MO

 Symposium on Epigenetics, University of Chicago, IL

 ASBMB meeting on Transcription, Snowbird, UT

 Nobel Conference on Systems Biology, Karolinska, Sweden

 MD Anderson Cancer Center, Houston, TX

2015 Tongji University, Shanghai, China

 Southern Medical University, Guangzhou, China

 Genomics Meeting, Lorne, Australia

 Pennsylvania State University (*Huck Distinguished Lecture series*)

Vanderbilt University, Nashville, TN

ENCODE Workshop, Bethesda, MD

Indiana University, Indianapolis, IN

Keystone Meeting on DNA Methylation/Epigenomics, Keystone, CO

Genomic Symposium, Northwestern University, Chicago, IL (*Keynote Lecture*)

Cold Spring Harbor Meeting on Eukaryotic Transcription, CSHL, NY

Chromatin Structure Course, Wellcome Trust Genome Campus, Hinxton (*Keynote Lecture*)

Transcription Symposium, Strasbourg, France

Symposium on Chromatin in Development and Disease, Marburg, Germany

AACR Conference on Chromatin and Cancer, Atlanta, GA

Conference on noncoding RNA, Gainesville, FL

Virginia Tech, Blacksburg, VA

2016 Gordon Research Conference, Les Diablerets, Switzerland

EMBO Conference on Gene Transcription in Yeast, San Felieu, Spain

Netherlands Cancer Institute, Amsterdam, Netherlands

Jackson labs, Bar Harbor, ME

 ASBMB meeting on Transcription, Snowbird, UT

 Duquesne University, Pittsburgh, PA

2017 University of Rochester, Rochester, NY

 National Institutes of Health, Bethesda, MD

 University of Texas, San Antonio, TX

Cold Spring Harbor Meeting on Eukaryotic Transcription, CSHL, NY

 Ludwig-Maximillian University, Munich, Germany