Contact Information	Computational OncologyPhone: (607) 351-8429Sage Bionetworkse-mail: bstephenwhite@gmail.comSeattle, WA 98103linkedin: www.linkedin.com/in/brian-s-white	l
Summary	Computational biologist with cross-disciplinary background in machine learning bioinformatics, cancer biology, and statistical modeling. 11+ years of experient developing machine learning methods (e.g., variational Bayesian approximal inference and fuzzy spectral clustering); 7+ years of experience applying tho methods to elucidate mechanisms of cancer (e.g., clonal evolution of colorectal cance and dysregulated splicing in myelodysplastic syndromes) and therapeutic respons (e.g., immune infiltration in tumors and biomarkers of BCL-2 inhibitor response acute myeloid leukemia). Proven ability to work independently and to initiate an lead external multidisciplinary collaborations of experimentalists, clinicians, an computational biologists. Excellent written communication skills evidenced 1 first-author publications in high-impact journals and successful grant applications.	ce te se er se in nd
Education	Cornell University , Ithaca, NY, USA Ph.D., Electrical and Computer Engineering (Physics and Computer Science Minors Mar 2008, GPA: 3.55),
	University of Virginia , Charlottesville, VA, USA Master of Computer Science, May 2002, GPA: 3.81	
	Carnegie Mellon University , Pittsburgh, PA, USA BS, Computer Science (Mathematics Minor), May 1998, GPA: 3.50	
Work Experience	Computational Sciences The Jackson Laboratory , Farmington, CT, USA <i>Senior Computational Scientist</i> , Oct 2020 – Present	
	 Computational Oncology Sage Bionetworks, Seattle, WA, USA Senior Scientist, Oct 2016 – Oct 2020 Co-organizing a DREAM Challenge (a community-wide competition) that comparing deconvolution method predictions of tumor immune infiltration mat from bulk RNA-seq against ground truth derived from single-cell platforms (e.g. CyTOF and single-cell RNA-seq). Predicted expression-based biomarkers of response in acute myeloid leukem (AML) using penalized regression, random forest, and novel Bayesia multi-source approaches, which revealed an insensitivity of monocytic AMLs venetoclax ("Beat AML" collaboration with Oregon Health & Science Universitiand Institute of Molecular Medicine Finland). Mentoring a junior scientist in predicting expression-based biomarkers response to tyrosine kinase inhibitors in chronic myeloid leukemia clinical triated for the second secon	de g., ia an to ty of

(collaboration with Prof. Jerry Radich and Novartis).

- Co-chairing an imaging analysis working group focused on high-dimensional microscopy (e.g., multiplexed/cyclic immunofluorescence, CODEX) within the NCI's Cancer Systems Biology Consortium (CSBC) and the Physical Sciences-Oncology Network (PS-ON).
- Dissected the Consensus Molecular Subtypes (CMS) of colorectal cancer (CRC) according to immune response.
- Defining a pathway- and network-based classification system for CRC that extends CMS to account for immuno-oncological features of disease.
- Integrated multi-modal data (i.e., molecular and clinical features) to predict disease progression in multiple myeloma using Cox proportional hazards model.
- Co-managing a team of six computational biologists and research associates that act as the data coordinating center for the CSBC / PS-ON.

Division of Oncology and McDonnell Genome Institute

Washington University School of Medicine, St Louis, MO, USA

Research Assistant Professor, Sept 2012 - Sept 2016

- Developed SciClone, a widely-used and highly-cited variational Bayesian mixture modeling approach for defining subclonal tumor architecture from whole-exome/whole-genome sequencing (WES/WGS) data, and used it to elucidate tumor heterogeneity in AML and to track progression to metastasis in CRC.
- Designed capture-based sequencing platforms for detecting B-cell receptor rearrangements (including IGH translocations), as well as single nucleotide variants and copy number alterations, in multiple myeloma (MM) and follicular lymphoma (FL). Used these platforms to discover frequent, progression-associated *IGLL5* mutations in MM and recurrent mutations affecting B-cell receptor signaling in FL.
- Inferred alterations in spliceosome/splicing site acceptor affinity induced by leukemia-associated mutations in the splicing factor U2AF1 by analyzing RNA-seq data with generalized linear models (collaboration with Prof. Matt Walter).
- Identified splicing disrupted in *H2AFY* by U2AF1 mutation, which may contribute to abnormal hematopoiesis in mice, and genes simultaneously dysregulated by U2AF1 mutation and spliceosome-targeted drugs, with a cumulative effect that may lead to a cell's sensitivity to pharmacological modulation (collaboration with Prof. Matt Walter).
- Designed a prototype NanoString-based clinical assay to evaluate the efficacy of splice-modulating drugs (collaboration with Prof. Matt Walter).

Dept. of Molecular Biology and Genetics, Cornell University, Ithaca, NY USA

Research Associate, Apr 2011 – Sept 2012

Advisor: David Shalloway

• Mathematically modeled and experimentally validated RNA/protein binding in SELEX (Systematic Evolution of Ligands by Exponential Enrichment) experiments, leading to an optimized experimental protocol that is actively used by our molecular biology collaborators.

Developed a bioinformatic pipeline to integrate next-generation sequencing into • SELEX experiments, resulting in improved sensitivity compared to traditional, Sanger sequencing-based techniques.

Dept. of Molecular Biology and Genetics, Cornell University, Ithaca, NY USA *Postdoc*, Mar 2008 – Apr 2011

Advisor: David Shalloway

- Developed a fuzzy (i.e., probabilistic), spectral (i.e., eigenvector-based) clustering algorithm modeled on statistical coarse-graining of "diffusive" transitions between similar data items, with improved efficiency and applicability over previous methods.
- Mathematically modeled migration of stem cells in collaboration with mouse geneticists, revealing an unexpected organization of the surrounding tissue.
- Quantified stem cell replication dynamics by analyzing fluorescence-activated cell sorting (FACS) data using Bayesian Gaussian mixture modeling, resulting in the discovery that the cell cycle regulator p21 represses stem cell proliferation.

Center for Advanced Scientific Computing, Lawrence Livermore National Laboratory, Livermore, CA, USA

Summer Student Guest, May 2004 – Aug 2005; Jun 2005

School of Computing, University of Utah, Salt Lake City, UT, USA Visiting Graduate Research Assistant, Sept 2001 – Sept 2002

Computing Sciences Research Center, Bell Laboratories, Murray Hill, NJ, USA (Summer) Member of Technical Staff, May 2000 - Aug 2000

Transarc Corporation, Pittsburgh, PA, USA Summer Intern 1997 – 1998

- SKILLS Machine learning: supervised learning LASSO/ridge/elastic net penalized regression, logistic regression), unsupervised learning (e.g., fuzzy spectral clustering, k-means, hierarchical clustering, expectation maximization), ensemble methods (e.g., random forest), dimensionality
 - reduction (e.g., PCA, MDS) Statistical modeling: generalized linear models, variational Bayesian inference, probabilistic graphical models, Monte Carlo techniques, survival analysis (e.g., Cox proportional hazards, Kaplan-Meier)

(e.g.,

SVMs,

neural

networks.

Pathway analysis: module detection (e.g., WGCNA, SPIA), gene-set enrichment (e.g., GSEA, GSVA, GOseq, DAVID, hypergeometric)

Genomic analysis: (pseudo-)alignment (e.g., bwa, tophat, STAR, cufflinks, kallisto), variant calling of whole-genome and exome/capture sequencing data (single nucleotide variants, insertions/deletions, structural variants, and copy number alterations; e.g., VarScan 2, pindel, lumpy), differential gene/splicing expression of RNA-seq and microarray expression data (e.g., cuffdiff, edgeR, DESeq, DEXSeq, rMATS, limma)

	 Single-cell RNA-seq analysis: scater (novice), scran (novice) Databases and repositories: KEGG, Reactome, Biocarta, GO, CCLE, TCGA, ImmPort/ImmuneSpace Natural language processing: feature representation (bag of words, tf-idf, word2vec, n-grams), topic modeling (LSA, LDA) Programming languages: R/Bioconductor, python (novice), C/C++, perl, latex Data visualization: R/ggplot2, python/Matplotlib (novice) Scientific programming: NumPy (novice), pandas (novice), scikit-learn (novice) Computing environments/software development: git/github, Docker, AWS Experimental techniques: basic molecular biology (e.g., qPCR, protein quantitation)
Mentoring	Juan Vizcarra (Jun 2020 – Present) Project: Active Learning in Analysis of Highly-Multiplexed Images
	Thomas Corey (Aug 2019 – Feb 2020) Project: CSBC/PSON Knowledge Portal: Semi-automating Curation and Enhancing Semantic Connectivity
	Dr. Matthew Wall (Jun 2019 – Oct 2020) Project: <i>Predicting TKI response and discontinuation in CML</i>
	Andrew Lamb (Jan 2018 – Oct 2020) Project: <i>A tumor deconvolution DREAM Challenge</i>
	Dr. Milen Nikolov (Jul 2018 – Jan 2019) Project: <i>Genotype-phenotype characterization of a cancer cell-line panel</i>
	Dr. Irena Lanc (Oct 2014 – Sept 2016) Project: A Second Generation, Multiple Myeloma-Specific, Targeted Sequencing Platform for Detecting Translocations, Copy Number Alterations, and Single Nucleotide Variants
Awards	NIH K12 Calabresi Career Development Award (Clinical Oncology), 2014 – 2016 International Myeloma Foundation Brian D. Novis Junior Research Grant, 2014 SPORE in Leukemia Career Development Award, 2013 – 2014 Dept. of Energy Krell High-Performance Computer Science Fellowship, 2003 – 2007 University of Virginia Dean's Scholarship, 1998 – 1999 Electronic Data Systems Textbook Scholarship (Carnegie Mellon University), 1994
Open-Source Software	 bmm: variational Bayesian mixture modeling Performs variational Bayesian inference with mixtures of Gaussian, beta, or binomial distributions. Available on <u>http://github.com/genome/bmm</u> (R package)
	sciclone: statistical inference of tumor clones from next-generation sequencing

• Detects evolution of tumor clones using variational Bayesian approaches

- Available on <u>http://github.com/genome/sciclone</u> (R package)
- PATENTS White BS, Lanc I, Fulton R, Auclair D, Tomasson MH, Automated exposition of known and novel multiple myeloma genomic variants using a single sequencing platform. United States Patent Application 20180126354. May, 2018.
- PUBLICATIONS IN SUBMISSION
 White BS*, Khan SA*, Ammad-ud-din M, Potdar S, Mason MJ, Tognon CE, Druker BJ, Heckman CA, Kallioniemi O, Kurtz SE, Porkka K, Tyner JW, Aittokallio T, Wennerberg K, Guinney J, Comparative analysis of independent *ex vivo* functional drug screens identifies predictive biomarkers of BCL-2 inhibitor response in AML. (*: co-first authors)
 - Kim S, Srivatsan N, Chavez M, Duncavage E, Vij K, Shirai CL, **White BS**, Jayasinghe R, Ahmed T, Shao J, Nunley R, Ding L, Pehrson JR, Walter MJ, *H2afy* (macroH2A1) Haploinsufficiency Induces Hematologic Malignancies and its Alternative Splicing Regulates B-Lymphopoiesis.

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- White BS*, Lanc I*, O'Neal J, Gupta H, Fulton RS, Schmidt H, Fronick C, Belter EA Jr, Fiala M, King J, Ahmann GJ, DeRome M, Mardis ER, Vij R, DiPersio JF, Levy J, Auclair D, Tomasson MH, A multiple myeloma-specific capture sequencing platform discovers novel translocations and frequent, risk-associated pointed mutations in IGLL5, Blood Cancer J 2018; 8(3):35. (*: co-first authors)
- Lal N*, **White BS***, Goussous G, Pickles O, Mason MJ, Beggs AD, Taniere P, Willcox BE, Guinney J, Middleton GW, KRAS Mutation and Consensus Molecular

Subtypes 2 and 3 Are Independently Associated with Reduced Immune Infiltration and Reactivity in Colorectal Cancer, Clin Cancer Res 2018; 24(1):224-233. (*: co-first authors)

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Teaching	Division of Oncology, Washington University , St Louis, MO, USA
Experience	Genomics and the Era of Personalized Medicine (2015)
	Guest Lecturer: Analysis of the Transcriptome

Dept. of Molecular Biology and Genetics, Cornell University, Ithaca, NY, USA Mathematical Modeling and Computational Statistics of the Molecular Cell (2011) *Guest Lecturer*

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