# XIAOTONG YAO

 $+1(917)833-5729 \diamond xiy2006@med.cornell.edu \diamond xiaotong.yao23@gmail.com$ 

Homepage: xtyao.github.io  $\diamond$  ORCID: 0000-0002-5140-6639

1300 York Ave  $\diamond$  New York, NY 10065

### **EDUCATION**

#### Ph.D., Weill Cornell Medicine, Computational Biology Jul 2015 - expected Apr 2021

- Mentored by Imielinski, M: Illuminating Rearranged Cancer Genome Structures through Genome Graphs
- Courses: Optimization Methods, Data Structure and Algorithms, Population Genetics, Statistical Genetics and Linear Models, Biomedical Machine Learning

### M.S., New York University, Bioinformatics

Overall GPA: 3.9/4.0

Aug 2013-May 2015 New York, NY

- Mentored by Vogel, C: *iSUMO* Integrative Prediction of Functionally Relevant SUMOylated Proteins
- Courses: Linear Regression and Multivariate Analysis, Statistics in Biology, Bioinformatics and Genomes, Biological Databases and Data Mining, Proteomics Informatics

B.S., Zhejiang University, Biotechnology	Aug 2009-May 2013
Overall GPA: 3.4/4.0	New York, NY

• Mentored by Chen, M: Optimization of Streptomycin Production in S. avermitilis by Metabolic Network Analysis

### PUBLICATIONS

Hadi K, Yao X, Behr JM, ..., Imielinski M. Distinct Classes of Complex Structural Variation Uncovered across Thousands of Cancer Genome Graphs. Cell. 2020;183: 197-210.e32.

(Accepted at Nature Communications) Dewhurst SM, Yao X, ..., de Lange T, Imielinski M. Structural Variant Evolution after Telomere Crisis. BioRxiv. 2020. p. 2020.09.29.318436. doi:10.1101/2020.09.29.318436

Carrot-Zhang J., Yao X, Devarakonda S., ..., TCGA Reasearch Network, Meyerson M, Govindan R, Imielinski M (2021). Whole-genome Characterization of Lung Adenocarcinomas Lacking Alterations in the RTK/RAS/RAF Pathway. Cell Rep. 34, 108707.

Wala JA, ..., Zhang C, Imielinski M, Beroukhim R. SvABA: Genome-wide Detection of Structural Variants and Indels by Local Assembly. Genome Res. 2018. doi:10.1101/gr.221028.117

Gerstung M, Jolly C, Leshchiner I, ..., PCAWG Consortium. The Evolutionary History of 2,658 Cancers. Nature. 2020;578: 122-128.

Rheinbay E, Nielsen MM, ..., PCAWG Consortium. Analyses of Non-coding Somatic Drivers in 2,658 Cancer Whole Genomes. Nature. 2020;578: 102–111.

**Yao X**, ..., Vogel C. *iSUMO* - Integrative Prediction of Functionally Relevant SUMOylation Events. bioRxiv. 2017. p. 056564. doi:10.1101/056564

(Submitted) Jungmin Choi, Aranzazu Manzano, Weilai Dong, ..., Richard P. Lifton, Joseph Schlessinger, and Alessandro D. Santin. Integrated mutational landscape analysis of uterine leiomyosarcomas.

### **RESEARCH PROJECTS**

#### Distinct Classes of Complex Structural Variation Uncovered across Thousands of Cancer May 2016 - Oct 2020 Genome Graphs Senior leader: Imielinski M

- Topology of junction copy number reveals novel classes of complex structural variants
- Rigma are deletion "chasms" at fragile sites arising early in GI tumor evolution
- Pyrgo are superenhancer-associated duplication "towers" in breast and ovarian cancer
- Tyfonas are "typhoons" of amplified fold-back inversions in acral melanoma

# Illuminating Rearranged Genome Structure through gGnome

Senior leader: Imielinski M

- Developed R API to genome graph data structures with a series of algorithms
- Designed an interactive web app to visualize genome graphs on arbitrary genomic regions
- Quick exploration of genome graphs built from 1000 Genome Project

Senior leader: de Lange T & Imielinski M

- Screened more than a hundred shallow WGS for prevalent SV regions in clones of telomere crisissurviving cells
- Reconstruct the exact linear allele after rearrangement with deep WGS
- Built consistent phylogeny using both SVs and SNVs
- Proved a single parental allele of chr12 to be the origin of SVs during crisis using allelic imbalances

#### Whole-genome Characterization of Lung Adenocarcinomas Lacking Alterations in RTK / **RAS / RAF Pathways** Dec 2017 - Nov 2020

Senior leader: Meyerson M, Govindan R, Imielinski M, TCGA genomic data analysis network

- Discovered KRAS or RTK/RAS/RAF alterations (RPA) from WGS previously missed by whole exome sequencing
- Found higher TP53 loss of function frequency in RPA- cancers
- Delinated diverse complex structural variation mechanisms creating amplification of oncogenes

# Integrated mutational landscape analysis of uterine leiomyosarcomas Dec 2018 - Dec 2020 Senior leader: Richard Lifton, Joseph Schlessinger, Alessandro Santin

• Identified the spectrum of complex SV events in uterine leiomyosarcomas whole genomes and found prevalent chromothripsis and chromoplexy, some disrupting tumor suppressors

#### Somatic Whole Genome Dynamics of Precancer in Barrett's Esophagus Mar 2018 - now Senior leader: Patricia Galipeau, Brian Reid, Nicolas Robine, Marcin Imielinski

- Breakage-fusion-bridge cycles enrich in patient's Barrett's esophagus (BE) who eventually developed esophageal adenocarcinoma
- Rigma is an early, prevalent SV event type in BE regardless of cancer outcome

#### Mar 2020 - now Structural Variation Signatures of Testicular Germ Cell Tumors Senior leader: Victoria Cortessis, Katherine Hoadley, TCGA genomic data analysis network

- Classified mechanisms amplifying *KRAS* in seminoma
- Discovered elevated burden of deletions and rigma in various non-seminoma

# **OPEN SOURCE SOFTWARE**

New York, NY

p 2020 New York, NY

Dec 2017 - Now New York, NY

# Main author

- <u>JaBbA</u> junction balance analysis
- gGnome an R API to genome graphs

# Contributor

- gGnome.js an interactive web-based genome browser for genome graphs
- gUtils elegant and fast genomic interval operations
- gTrack static genome browser style plots
- <u>fishHook</u> Gamma-Poisson regression for count data on genomic intervals
- $\underline{GxG}$  interaction matrices between genomic bins

# SKILLS

Biology	Cancer Genomics, Computational Biology, Systems Biology
Sequencing Informatics	Illumina, Oxford Nanopore WGS
Statistics & Machine Learning	Generalized Linear Models, Random Forests
Computer Languages	R, Shell, Python, Java
Optimization	CPLEX, Gurobi
Data Visualization	ggplot, shiny, plotly
Scientific Communication	LaTeX, Adobe Illustrator, (R)markdown
Databases	SQLite, MySQL
Other Tools	Git, Emacs, Docker, Nextflow, Hugo

### WORK EXPERIENCE

Graduate Research Assistant Weill Cornell Medicine Jul 2015 - Present New York, NY

- Discovered three new patterns of complex structural variations in cancer genomes
- Designed, implemented, benchmarked Junction Balance Analysis to reconstruct genome graphs
- Developed *gGnome* for genome graph data structure
- Captured ongoing SV evolution in post-telomere crisis cell lines
- Characterized whole genome landscape in lung adenocarcinomas without RTK/RAS/RAF pathways alterations
- Contributed to large cancer sequencing consortiums including The Cancer Genome Atlas (TCGA) and the Pan-Cancer Analysis of Whole Genomes (PCAWG)

## Graduate Research Assistant

New York University

• Trained predictive models for post-translational modification from protein function databases

# **Teaching Adjunct**

New York University

- Taught R statistical programming for Introduction to Biostatistics
- Tutored techniques for Biological Databases and Data Mining

# **Bioinformatics Intern**

 $3E \ Bioenergy$ 

• Compared crop genomes for candidate genes linked to drought resistance in sweet sorghum

Jun 2014 - May 2015 New Brunswick, NJ

Feb 2014 - May 2015

December 2013 - May 2015

New York, NY

New York, NY

# AWARDS

Asia Regional Winner, World 2nd Runner Up & Best	New Application Nov 2011	
2011 International Genetically Engineered Machines	Hong Kong, China; Cambridge, MA	
• Team member of ZJU-China		
• Designed multicolor fluorescent expression system in biofilm responsive to gradients of oxygen level		
Master's Student Research Grant	2014 & 2015	
NYU Biology Master's Program	New York, NY	
• Funding for protien sumoylation prediction by mining public protein databases		
Broad Institute Workshop Travel Grant	2015	
Broad Institute	Cambridge, MA	
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• Travel grant for single cell genomics workshop

New Algorithm Reveals Patterns In How Tumors Extensively Rewrite Their DNA. Available: Weill Cornell Medicine News

Anderson A. Algorithm Uncovers New Complex Rearrangement Classes in Cancer Genomes. 1 Oct 2020. Available: Genomeweb.com