# Wen-Wei Liang, Ph.D.

101 6th Ave, New York, NY wliang@nygenome.org (314) 540-1364

Molecular geneticist and genomics researcher specializing in CRISPR screening and multi-omic profiling, focusing on epigenetic regulation in tumorigenesis through integrated experimental and computational methods.

# **EDUCATION**

Washington University in St. Louis	Ph.D. in Molecular Genetics and Genomics	2014-2020
National Yang-Ming University, Taiwan	M.S. in Microbiology and Immunology	2009-2011
National Tsing-Hua University, Taiwan	B.S. in Life Science	2005-2009

# **FELLOWSHIPS**

Taiwan Ministry of Education - Washington University in St. Louis Fellow	2014-2018
Washington University in St. Louis Precision Medicine Pathway Fellow	2015-2016
Taiwan Ministry of Education Fellowship to Study Abroad	2007-2008

#### RESEARCH POSITIONS

#### **New York University & New York Genome Center**

New York, NY

Postdoctoral research associate, Laboratory of Neville Sanjana

2021-Present

- Transcriptome-scale RNA-targeting CRISPR screens reveal essential lncRNAs in human cells: Conducted Cas13 pooled screens and perturbation assays, coupled with single-cell RNA sequencing, to identify and characterize hundreds of lncRNAs crucial for cell proliferation and their mechanistic roles.
- High-content CRISPR screens identify lncRNAs modulate CAR-macrophages in tumor microenvironment: Implemented *in vitro* and *in vivo* screens to discover lncRNAs that promote a pro-inflammatory microenvironment in BRCA xenograft and BRCA organoid models.

#### Washington University in St. Louis

St. Louis, MO

Ph.D. student and postdoctoral research associate, Laboratory of Li Ding

2015-2021

- *Epigenetic regulation during cancer transitions across 11 tumour types:* Utilized snATAC sequencing, single-cell RNA sequencing, and bulk whole-exome sequencing data from the Human Tumor Atlas Network (HTAN) to profile over a million cells or nuclei. Identified cancer-specific accessible region dynamics correlating with gene expression, uncovering novel regulatory regions and their target genes.
- Integrative multi-omic cancer profiling reveals DNA methylation patterns associated with therapeutic vulnerability and cell-of-origin: Leveraged DNA methylation, RNA sequencing, and proteomic data from the Clinical Proteogenomic Tumor Analysis Consortium (CPTAC) to elucidate how epigenetic changes in cancer cells influence gene expression, protein levels, tumor characteristics, and therapeutic responses.
- *Driver fusions and their implications in the development and treatment of human cancers:* Conducted comprehensive analysis of gene fusions in over 9,000 tumors across 33 cancer types from The Cancer Genome Atlas (TCGA), uncovering how these fusions drive the expression of oncogenes, tumor suppressor genes, and kinases. Identified druggable fusions in 6.0% of cases, highlighting their potential as targets for precision therapies.

Academia Sinica Taipei, Taiwan

Master's student and research assistant, Laboratory of Soo-Chen Cheng

2009-2014

• A novel mechanism for Prp5 function in prespliceosome formation and proofreading the branch site sequence: Identified the role of Prp5 in spliceosome assembly and branch site proofreading, showing how Prp5's interactions with U2 snRNA and subsequent release facilitate accurate spliceosome assembly.

#### **National Tsing-Hua University**

Hsinchu, Taiwan

Undergraduate student, Laboratory of Chung-Yu Lan

2008-2009

• Investigated the reactions of the iron-responsive element under virulence gene expression in Candida albicans.

# **Linköping University**

Linköping, Sweden

Visiting student, Laboratory of Jordi Altimiras

2007-2008

· Analyzed G-protein coupled receptor-dependent contractility in chicken heart tissue.

#### **PUBLICATIONS**

# First authorship (\* Equal contribution)

- 1. **Liang, W.-W.** \*, Müller, S. \*, Hart, S. K., Wessels, H.-H., Méndez-Mancilla, A., Sookdeo, A., Choi, O., Caragine, C., Corman, A., Lu, L., Kolumba, O., Williams, B. & Sanjana, N. E. Transcriptome-scale RNA-targeting CRISPR screens reveal essential lncRNAs in human cells. *Revision in cell* (2024).
- 2. **Liang, W.-W.**, Lu, R. J.-H., Jayasinghe, R. G., Foltz, S. M., Porta-Pardo, E., Geffen, Y., Wendl, M. C., Lazcano, R., Kolodziejczak, I., Song, Y., *et al.* Integrative multi-omic cancer profiling reveals DNA methylation patterns associated with therapeutic vulnerability and cell-of-origin. *Cancer cell* 41, 1567–1585.e7 (2023). *("Featured in The Scientist magazine")*.
- 3. Terekhanova, N. V. \*, Karpova, A. \*, **Liang, W.-W.** \*, Strzalkowski, A., Chen, S., Li, Y., Southard-Smith, A. N., Iglesia, M. D., Wendl, M. C., Jayasinghe, R. G., *et al.* Epigenetic regulation during cancer transitions across 11 tumour types. *Nature* 623, 432–441 (2023). *(Spotlighted in Nature Genetics Research Highlight)*.
- 4. Gao, Q. \*, **Liang, W.-W.** \*, Foltz, S. M. \*, Mutharasu, G., Jayasinghe, R. G., Cao, S., Liao, W.-W., Reynolds, S. M., Wyczalkowski, M. A., Yao, L., *et al.* Driver fusions and their implications in the development and treatment of human cancers. *Cell rep.* 23, 227–238.e3 (2018). (*Best of Cell Reports 2018*).
- 5. **Liang, W.-W.** & Cheng, S.-C. A novel mechanism for Prp5 function in prespliceosome formation and proofreading the branch site sequence. *Genes dev.* 29, 81–93 (2015).

# Co-authorship

- 6. Li, Y., Dou, Y., Da Veiga Leprevost, F., Geffen, Y., Calinawan, A. P., Aguet, F., Akiyama, Y., Anand, S., Birger, C., Cao, S., et al. Proteogenomic data and resources for pan-cancer analysis. *Cancer cell* 41, 1397–1406 (2023).
- 7. Carrot-Zhang, J., Yao, X., Devarakonda, S., Deshpande, A., Damrauer, J. S., Silva, T. C., Wong, C. K., Choi, H. Y., Felau, I., Robertson, A. G., *et al.* Whole-genome characterization of lung adenocarcinomas lacking the RTK/RAS/RAF pathway. *Cell rep.* 34, 108707 (2021).
- 8. Huang, C., Chen, L., Savage, S. R., Eguez, R. V., Dou, Y., Li, Y., da Veiga Leprevost, F., Jaehnig, E. J., Lei, J. T., Wen, B., *et al.* Proteogenomic insights into the biology and treatment of HPV-negative head and neck squamous cell carcinoma. *Cancer cell* 39, 361–379.e16 (2021).
- 9. Wang, L.-B., Karpova, A., Gritsenko, M. A., Kyle, J. E., Cao, S., Li, Y., Rykunov, D., Colaprico, A., Rothstein, J. H., Hong, R., *et al.* Proteogenomic and metabolomic characterization of human glioblastoma. *Cancer cell* (2021).
- 10. Bailey, M. H., Meyerson, W. U., Dursi, L. J., Wang, L.-B., Dong, G., **Liang, W.-W.**, Weerasinghe, A., Li, S., Kelso, S., MC3 Working Group, *et al.* Retrospective evaluation of whole exome and genome mutation calls in 746 cancer samples. *Nat. commun.* 11, 4748 (2020).
- 11. Gillette, M. A., Satpathy, S., Cao, S., Dhanasekaran, S. M., Vasaikar, S. V., Krug, K., Petralia, F., Li, Y., Liang, W.-W., Reva, B., *et al.* Proteogenomic characterization reveals therapeutic vulnerabilities in lung adenocarcinoma. *Cell* 182, 200–225.e35 (2020).
- 12. Cao, S., Wylie, K. M., Wyczalkowski, M. A., Karpova, A., Ley, J., Sun, S., Mashl, R. J., **Liang, W.-W.**, Wang, X., Johnson, K., *et al.* Dynamic host immune response in virus-associated cancers. *Commun biol* 2, 109 (2019).
- 13. Bailey, M. H., Tokheim, C., Porta-Pardo, E., Sengupta, S., Bertrand, D., Weerasinghe, A., Colaprico, A., Wendl, M. C., Kim, J., Reardon, B., *et al.* Comprehensive characterization of cancer driver genes and mutations. *Cell* 173, 371–385.e18 (2018).
- 14. Jayasinghe, R. G., Cao, S., Gao, Q., Wendl, M. C., Vo, N. S., Reynolds, S. M., Zhao, Y., Climente-González, H., Chai, S., Wang, F., *et al.* Systematic analysis of splice-site-creating mutations in cancer. *Cell rep.* 23, 270–281.e3 (2018).
- 15. Sanchez-Vega, F., Mina, M., Armenia, J., Chatila, W. K., Luna, A., La, K. C., Dimitriadoy, S., Liu, D. L., Kantheti, H. S., Saghafinia, S., *et al.* Oncogenic signaling pathways in the cancer genome atlas. *Cell* 173, 321–337.e10 (2018).
- 16. Foltz, S. M., **Liang, W.-W.**, Xie, M. & Ding, L. MIRMMR: binary classification of microsatellite instability using methylation and mutations. *Bioinformatics* 33, 3799–3801 (2017).
- 17. Niu, B., Scott, A. D., Sengupta, S., Bailey, M. H., Batra, P., Ning, J., Wyczalkowski, M. A., **Liang, W.-W.**, Zhang, Q., McLellan, M. D., *et al.* Protein-structure-guided discovery of functional mutations across 19 cancer types. *Nat. genet.* 48, 827–837 (2016).

#### **HONORS & AWARDS**

• Best of Cell Reports 2018	2018
<ul> <li>Outstanding Merit Research Award National Yang-Ming University</li> </ul>	2011
<ul> <li>Dean's List, Department of Life Science, National Tsing-Hua University</li> </ul>	2007
<ul> <li>Outstanding student from Low-income, Lu Feng-Zhang Memorial Scholarship</li> </ul>	2006

#### SELECTED TALKS

# CSHL Meeting: The Biology of Genomes Cold Spring Harbor, NY

Transcriptome-scale RNA-targeting CRISPR screens reveal essential lncRNAs in human cells

Virtual

2024

2021

# The CPTAC Annual Scientific Symposium

Multiomic profiling reveals tumorigenic DNA methylation associated with therapeutic vulnerability and cell-of-origin

The CPTAC Site Visit – NYU-WU-BYU PGDAC New York University, NY

Tumorigenic DNA methylation revealed by integrative transcriptomic and proteomic profiling

2019

#### **CSHL Meeting: Eukaryotic mRNA Processing**

Cold Spring Harbor, NY ite sequence 2013

A novel mechanism for Prp5 function in prespliceosome formation and proofreading the branch site sequence 202

Taiwan Yeast Meeting Taiwan

The DEAD-box ATPase Prp5 mediates splicing fidelity control by counteracting tri-snRNP binding

2013

# TEACHING, MENTORING & OUTREACH

**Simons foundation-NYU biology summer undergraduate research program, New York, NY** Summer 2024 Mentored an undergraduate student, Sebastián H. Díaz-Rodríguez, from a diverse background, providing high-quality research training and opportunities that are often inaccessible to underrepresented students.

## Postdoc Seminar Series, New York Genome Center, New York, NY

2021-Present

Organized monthly seminars featuring presentations by postdoctoral researchers.

#### Laboratory of Neville Sanjana, New York University, New York, NY

2021-2024

Guided a master's student, Breanna Williams (recipient of the Wasserman Center Internship Grant), and an undergraduate, Olivia Choi (recipient of the NYU Dean's Undergraduate Research Fund), in completing their thesis.

#### Biology Club, Rutgers University-Camden, Camden, NJ

2022

Participated as a panelist to discuss career development opportunities in science and technology with over 20 undergraduate students.

#### Midwest Taiwanese Biotechnology Association, Chicago, IL

2017-2021

Co-founded the association, served in various leadership roles including Finance (2018), Promotion (2019), and President (2021), and organized events to foster dialogue among young scientists across Midwest cities.

# Laboratory of Li Ding, Washington University in St. Louis, St. Louis, MO

2015-2020

Mentored three undergraduate students (Carolyn Lou, Terrence Tsou, Jessika Baral), three rotation students (Yang Li, Wing Hing Wang, Winston Anthony), and one graduate research assistant (Rita Jui-Hsien Lu) in genome research.

#### Department of Biology, Washington University in St. Louis, St. Louis, MO

Spring 2016

Delivered lectures, provided assistance in molecular biology experiments, and assessed student performance as a teaching assistance in the Microbiology Laboratory course (Biol 3491) for over 30 undergraduate students.

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