# Splicing Factor Mutations and RNA Biology in Cancer Workshop



# HOSTED BY YALE CANCER CENTER AND YALE CENTER FOR RNA SCIENCE AND MEDICINE

May 22-23, 2019 Edward P Evans Hall 165 Whitney Ave, New Haven, CT 06511

MAY 22, 2019 5:00pm-9:00pm Beinecke Terrace Room	
5:00pm	Registration and Poster Set-Up
5:30pm	Welcome Reception and Poster Presentation
6:30pm	Keynote  Joan Steitz, PhD Sterling Professor of Molecular Biophysics and Biochemistry; Investigator, Howard Hughes Medical Institute; Yale University School of Medicine  "The Enigma of Viral Noncoding RNAs"
7:30pm	Dinner
8:15pm	Dessert Reception and Poster Presentation

### **Poster Presentations:**

**Sophia Adamia**, "Mutations in splicing regulatory elements in RHAMM gene may contribute to an increased RHAMM-exon4/RHAMM-FL splice variant ratio in multiple myeloma (MM)".

<u>Yusuf Adeshina</u>, "Rational Design of Small Molecule Inhibitor of Serine Arginine Rich Splicing Factor 2 (SRSF2)".

<u>Olga Anczukow</u>, "Misregulation of splicing factors in breast cancer initiation and metastasis".

<u>Nan Bai</u>, "Designing Inhibitors of the Msi1 Protein-RNA Interaction by RNA Mimicry".

<u>Erson-Bensan</u>, "Unraveling the complexity of breast cancer transcriptome through 3'UTRs".

**Noah Daniels**, "Investigating the Role of DDX41 in pre-messenger RNA Splicing and Leukemogenesis".

<u>Callen Feeney</u>, "A synthetic molecule enhances U2AF2 complexes with splice site RNAs and stalls pre-mRNA splicing".

<u>Courtney Hershberger</u>, "LUC7L2 is a splicing regulatory protein that is frequently mutated in bone marrow neoplasms".

<u>Nathan Leclair</u>, "Dissecting Splicing Factor Poison-Exon Regulation in Breast Cancer".

<u>Wan Yee Leong</u>, "U2AF1 Splicing Factor Mutations Confer Sensitivity to ATR Inhibition".

**Bo Liu**, "Hotspot mutations in SF3B1 promote aberrant splicing and breast tumorigenesis".

<u>Debanjana Maji</u>, "Cancer-associated mutations of the pre-mRNA splicing factor U2AF2 alter splice site signal recognition".

**Nicole Martinez**, "Pseudouridine synthases modify human pre-mRNA cotranscriptionally and affect splicing".

<u>Cameron Soulette</u>, "Nanopore sequencing reveals isoform-specific alterations in human bronchial epithelial cells with U2AF1S34F mutations".

**Alison Tang**, "FLAIR; Full-Length Alternative Isoform analysis of RNA".

**Laura Urbanski**, "MYC-Induced Alternative Splicing in breast cancer".

**Borwyn Wang,** "The Role of Alternatively-Spliced CSF3R in Promoting Myelodysplastic Syndromes".

<u>Chandani Warnasooriya</u>, "U2AF1 and its myelodysplasia-associated S34F mutation alter U2AF2 conformations for splice site RNA recognition".

<u>Emily Wheeler</u>, "Allele-specific enhanced CLIP and RNA-Seq reveal alterations in RNA processing events in hematopoietic progenitor cells derived from isogenic iPSC models of SRSF2 and U2AF1 mutations".

<u>Akhide Yoshimi</u>, "Reciprocal Alterations in RNA Splicing and Epigenetic Regulation Drive Leukemogenesis".

# MAY 23, 2019 8:00am-5:00pm

Class of 1980 Classroom / Room 2400

8:00am- Regis 4:00pm

Registration

8:00am Breakfast

8:25am Opening Remarks

# 8:30am-10:00am

# Session 1

### Talk 1/2:

# Robert Bradley, PhD

Associate Member, Public Health Sciences Division; Associate Member, Basic Sciences Division, Fred Hutchinson Cancer Research Center

# Omar Abdel-Wahab, MD

Associate Attending, Leukemia Service, Department of Medicine, Associate Member, Human Oncology and Pathogenesis, Memorial Sloan Kettering Cancer Center

"Spliceosomal Disruption of the non-canonical BAF complex in cancer"

# Talk 3:

Dong-Er Zhang, MD, PhD

Professor, Pathology, UC San Diego

"RUNX1 related post-transcriptional regulation of gene expression"

### Talk 4:

# Nicole Martinez, PhD

Postdoctoral Fellow, Yale University School of Medicine "Pseudouridine synthases modify human pre-mRNA cotranscriptionally and affect splicing"

### Talk 5:

# Dan Larson, PhD

Earl Stadtman Investigator; Head, Systems Biology of Gene Expression Section, Laboratory of Receptor Biology and Gene Expression, National Cancer Institute

"U2AF1 contributes to cancer progression through a noncanonical role in translation"

10:00am-10:30am **Break** 

10:30am-

Session 2

12:05pm

### Talk 1:

# Christine Mayr, MD, PhD

Associate Member, Cancer Biology and Genetics Program, Memorial Sloan Kettering Cancer Center

"Regulation of 3'UTR-mediated protein-protein interactions"

# Talk 2:

# William Fairbrother, PhD

Professor of Biology, Molecular Biology, Cell Biology, & Biochemistry, Brown University

"Pathogenic mutations and the evolution of the splicing code"

### Talk 3:

# Michael Kharas, PhD

Associate Member, Molecular Pharmacology Program, Memorial Sloan Kettering Cancer Center

"RNA Regulators in Normal and Malignant Stem Cells"

### Talk 4:

# Matthew Disney, PhD

Professor, Department of Chemistry, Scripps Research

"Translating RNA Sequence into Lead Small Molecule Medicines"

### Talk 5:

# Erin Ahn, PhD

Associate Professor, University of South Alabama "Oncogenic RNA Splicing in Glioblastoma"

# 12:05pm-1:30pm

**Lunch (+Pharma/Sponsor Exposition)** 

# 1:30pm-3:00pm

### **Session 3**

### Talk 1:

# Matt Simon, PhD

Associate Professor of Molecular Biophysics and Biochemistry "Chemical tools to study the non-coding transcriptome"

# Talk 2:

# Angela Brooks, PhD

Assistant Professor of Biomolecular Engineering, UC Santa Cruz "Full-length alternative isoforms associated with splicing factor mutations identified from long-read nanopore sequencing"

### Talk 3:

### Ernesto Guccione, PhD

Associate Professor, Icahn School of Medicine at Mount Sinai "Therapeutic targeting of RNA splicing catalysis through inhibition of protein arginine methylation"

### Talk 4:

# Luisa Escobar-Hoyos, PhD

Research Scholar, Human Oncology and Pathogenesis, Memorial Sloan Kettering Cancer Center

"Altered mRNA splicing by mutant p53 activates oncogenic KRAS signaling in pancreatic cancer"

# Talk 5:

# Courtney Hershberger, BS

Graduate Student, Cleveland Clinic

"LUC7L2 is a splicing regulatory protein that is frequently mutated in bone marrow neoplasms".

3:00pm-3:20pm **Break** 

# 3:20pm-4:00pm

# Session 3 (cont.)

### Talk 6:

# Clara Kielkopf, PhD

Professor, Department of Biochemistry and Biophysics, University of Rochester Medical Center

"A synthetic molecule stalls pre-mRNA splicing by enhancing cancer-relevant U2AF2 – RNA complexes"

# Talk 7:

# Carl Walkley, PhD

Head, Cancer & RNA Laboratory, St Vincent's Institute/The University of Melbourne

"Understanding how RNA splicing mutations contribute to myelodysplastic syndrome: *Srsf2* P95H/+"

4:00pm-5:00pm **Keynote** 

Samie Jaffrey, PhD

Professor of Pharmacology, Weill Cornell Medical College

"Control of mRNA, snRNA and IncRNA function by regulated nucleotide methylations"

5:00pm

**Closing Remarks** 

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