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Zefeng Wang Associate Professor Department of Pharmacology University of North Carolina at Chapel Hill USA

### Biography

• **Prof. Zefeng Wang** received his Ph.D. degree from Johns Hopkins Medical School, worked as a Damon Runyon fellow at Massachusetts Institute of Technology before becoming an assistant professor at University of North Carolina at Chapel Hill. Dr. Wang's research focuses on the regulation of gene expression in RNA level. He has made significant contribution to the filed of RNA biology by developing a series of new methods to study RNA splicing and degradation, and has pioneered the filed of engineering RNA binding proteins. His work was recognized by several research awards.

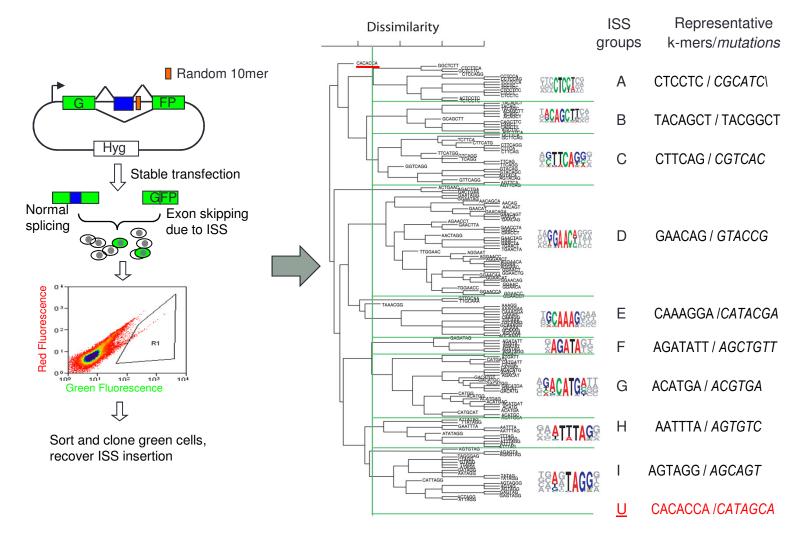
### **Research Interests**

- Systematic study of the regulation of alternative splicing
- Mechanisms and functional roles of splicing mis-regulation in cancers
- Manipulation of RNA processing with artificial protein factors

### **Recent Publications**

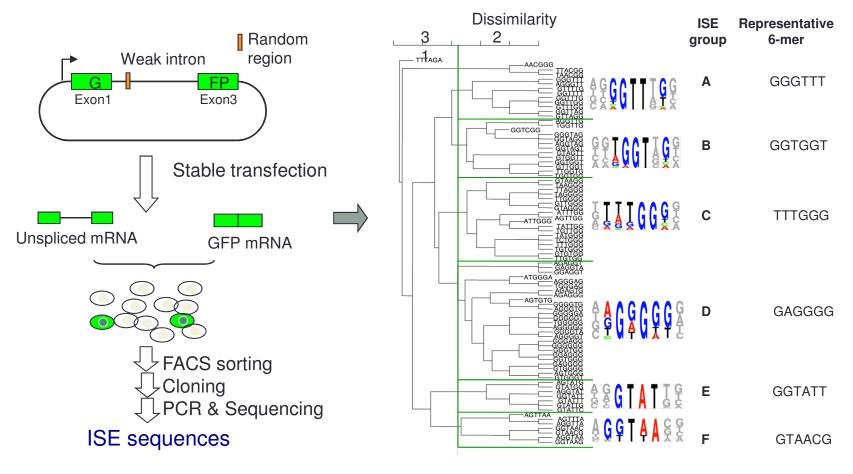
- Wang Y, Cheong CG, Hall TM and Wang Z. Engineering splicing factors with designed specificities (2009). Nature Method. 6(11):825-30. Epub 2009 Oct 4
- Wang Y, Ma M, Xiao XS and **Wang Z**. Intronic splicing enhancers, cognate splicing factors and context dependent regulation rules. (2012) Nature Structure Molecular Biology, doi:10.1038/nsmb.2377. Epub Sep 16.
- Choudhury R, Dominguez D, Wang Y and **Wang Z**. Engineering RNA endonucleases with customized sequence specificities (2012). Nature Communication 23;3:1147. doi: 10.1038/ncomms2154.
- Wang Y, Xiao X, Zhang J, Choudhury R, Robertson A, Li K, Ma M, Burge CB, Wang Z. A complex network of factors with overlapping affinities represses splicing through intronic elements. (2013) Nat Struct Mol Biol. 20(1):36-45. doi: 10.1038/nsmb.2459. PMCID: PMC3537874
- Zhang W, Wang Y, Dong S, Choudhury R, Jin Y and Wang Z Treatment of type 1 Myotonic Dystrophy by engineering site-specific RNA endonucleases that target (CUG)<sub>n</sub> repeats. (2013) Molecular Therapy. Oct 23 doi: 10.1038/mt.2013.251 Epub ahead of print
- Choudhury R, Ghose Roy S, Tsai YS, Tripathy A, Graves LM and Wang Z. The splicing activator DAZAP1 integrates splicing control into MEK/Erk regulated cell proliferation and migration. Nature Communications (2014) Jan 23;5:3078. doi: 10.1038/ncomms4078
- Matera AG\* and Wang Z\* (co-corresponding author). Ribonucleoprotein assembly and dynamics: A day in the life of the spliceosome. Nature Review Molecular Cell Biology (2014) Jan 23;15(2):108-21. doi: 10.1038/nrm3742
- Tsai YS, Gomez SM, and Wang Z. Prevalent RNA recognition motif duplication in the human genome. RNA (2014) May;20(5):702-12. doi: 10.1261/rna.044081.113. Epub 2014 Mar 25
- Wang Y, Chen D, Qian H, Tsai YS, Shao S, Dominguez D and Wang Z. The splicing factor RBM4 controls apoptosis, proliferation, and migration to suppress tumor progression. Cancer Cell (2014) Sep 8; 26(3):374-89. doi: 10.1016/j.ccr.2014.07.010.

### Systematic Identification of Intronic Splicing Silencers with FAS-ISS



Yang Wang et al, NSMB 2013

### Identify intronic splicing enhancers by FAS-ISE

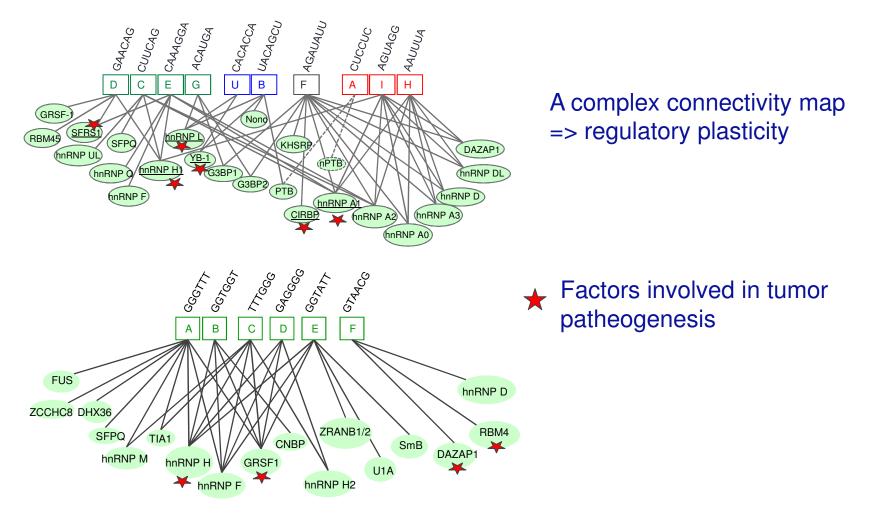


117 ISE decamers, 109 unique

All ISE function as ESS (i.e., inhibit splicing) when inserted in exons

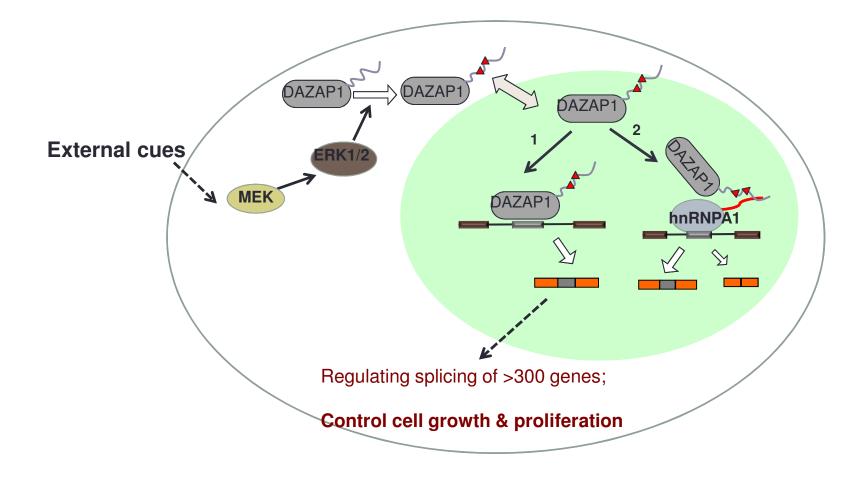
Wang Y et al, 2012 NSMB

### An Overlapping Network of *trans*-factors Recognize ISSs and ISEs



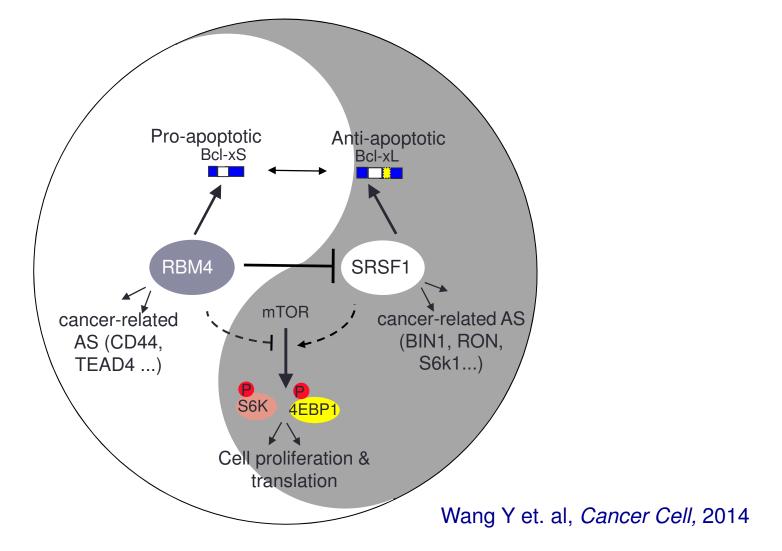
Wang Y et. al, *Nat Struc & Mol Bio* 2012 Wang Y et. al, *Nat Struc & Mol Bio* 2013

## DAZAP1 integrates splicing control into MEK/Erk regulated cell proliferation and migration

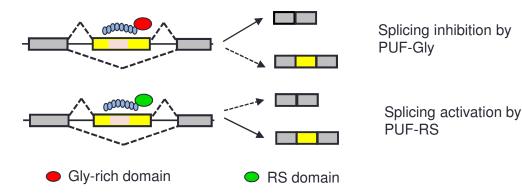


Chundhury et, al. 2014 Nature Communications

# RBM4-mediated splicing regulation balance controls cell growth/proliferation

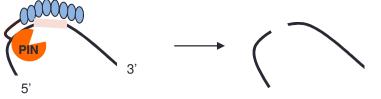


#### Engineered splicing factors: Restore normal splicing in cancer



Artificial site-specific RNA endonuclease: Cleave toxic RNA

Gene silencing in mitochondrion & chloroplast



Wang Y et al, 2009 Nature Methods Chundhury et al, 2012, Nature Communications

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