
SEMINAR

***Computational analysis of mRNA isoform expression
using RNA-Seq data***

Yi Xing, Ph.D.

Associate Professor

**Department of Microbiology, Immunology and
Molecular Genetics**

University of California, Los Angeles

ABSTRACT

Alternative splicing is the process by which exons from precursor mRNA transcripts are differentially included during splicing, resulting in different mature mRNA isoforms from a single gene locus. Common genetic variation that affects splicing regulation can lead to differences in alternative splicing between human individuals and consequently impact expression level or protein function. Recently, ultra-deep RNA sequencing (RNA-Seq) has become a powerful technology for genome-wide analysis of alternative splicing. We have developed GLiMMPS, a robust method for detecting genetic variation of alternative splicing from RNA-Seq data. GLiMMPS takes into account the individual variation in sequencing depth and the noise prevalent in RNA-Seq data. Analyses of simulated and real RNA-Seq data demonstrate that GLiMMPS outperforms competing statistical models. Quantitative RT-PCR tests of randomly selected GLiMMPS predictions yielded a validation rate of 100%. As population-scale RNA-Seq studies become increasingly affordable and popular, GLiMMPS provides a useful tool for elucidating the genetic architecture of alternative splicing variation in humans and model organisms.

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