**Title:** Novel statistical approaches for data analysis in RNA interference high-throughput Screening technology

## Abstract:

RNA interference (RNAi) high throughput screening (HTS) allows the identification of genes associated with specific biological phenotypes. This technology has been hailed as the second genomics wave following the first genomics wave of gene expression microarrays and single nucleotide polymorphism (SNP) discovery platforms. HTS experiments generate a huge amount of data. In order to use these data to identify the most effective siRNAs tested, it is critical to adopt and develop appropriate statistical methods for quality control (QC) and hit selection. For QC in HTS assays, J-H Zhang et al (1999) proposed the popular Z factor and XD Zhang et al (2006) proposed strictly standardized mean difference (SSMD) and coefficient of variability in difference (CVD). For hit selection, mean  $\pm k$  SD (or z score) was initially used. To be more robust to outliers and skewness, Brideau, Gunter and Pikoulis et al (2003) proposed B-score method, XD Zhang et al (2006) proposed a quartile-based method and suggested using median  $\pm k$  MAD or quartile-based methods. Malo and Hanley *et al* (2006) reviewed the methods developed before early 2006. Recently, XD Zhang has developed an SSMDbased process for hit selection in RNAi HTS experiments. This process maintains a flexible and balanced control of both false negatives and false positives. Here, XD Zhang first present SSMD,  $\beta$ -probability, the SSMD-based process for hit selection, and their applications and then compare the SSMD-based methods with other methods in RNAi HTS experiments.