

Varistran: Anscombe’s variance stabilizing transformation for RNA-seq gene expression data

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Software Repository: <https://github.com/MonashBioinformaticsPlatform/varistran>

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Summary

RNA-seq measures RNA expression levels in a biological sample using high-throughput cDNA sequencing, producing counts of the number of reads aligning to each gene. Noise in RNA-seq read count data is commonly modelled as following a negative binomial distribution, where the variance is a quadratic function of the expression level. However many statistical, machine learning, and visualization methods work best when the noise in a data set has equal variance. Varistran is an R package that uses Anscombe’s (1948) variance stabilizing transformation for the negative binomial distribution to transform RNA-seq count data, so that the noise has equal variance across all measured gene expression levels. The transformed data may be treated as \log_2 transformed gene expression levels, but with variability reduced at low read counts. Varistran also includes a function to open a Shiny report with simple diagnostic visualizations, including a plot to assess how effective the variance stabilization was, a biplot of samples and genes, and a heatmap. This allows defective samples, sample mislabelling, and batch effects to be easily identified.

References

Anscombe, Francis J. 1948. “The Transformation of Poisson, Binomial and Negative-Binomial Data.” *Biometrika* 35 (3/4): 246–54.