

ComparativeMarkerSelection Documentation

Module name: ComparativeMarkerSelection

Description: Computes significance values for features using several metrics,

including FDR(BH), Q Value, FWER, Feature-Specific P-Value,

and Bonferroni.

Author: Joshua Gould, Gad Getz, Stefano Monti (Broad Institute)

gp-help@broad.mit.edu

The ComparativeMarkerSelection module includes several approaches to determine the features that are most closely correlated with a class template and the significance of that correlation. If the input class template has more than two classes, than a one-versus-all comparison is performed for each class. Note that the p-values obtained from the one-versus-all comparison are not fully corrected for multiple hypothesis testing. The module outputs a file containing the following columns:

- 1. **Rank** The rank of the feature within the dataset based on the value of the test statistic. If a two-sided p-value is computed, the rank is with respect to the absolute value of the statistic.
- 2. Feature The feature name.
- 3. **Description** The description of the feature.
- 4. **Score** The value of the test statistic.
- 5. *Feature P* The feature-specific p-value based on permutation testing.
- 6. Feature P Low The estimated lower bound for the feature p-value.
- 7. **Feature P High** -The estimated upper bound for the feature p-value.
- 8. **FDR (BH)** An estimate of the false discovery rate by the Benjamini and Hochberg procedure (3). The FDR is the expected proportion of erroneous rejections among all rejections.
- 9. **Q Value** An estimate of the FDR using the procedure developed by Storey and Tibshirani (4).
- 10. **Bonferroni** The value of the Bonferroni correction applied to the feature specific p-value.
- 11. **maxT** The adjusted *p*-values for the maxT multiple testing procedure described in (5), which provides strong control of the FWER.
- 12. **FWER** (Family Wise Error Rate) the probability of at least one null hypothesis/feature having a score better than or equal to the observed one. This measure is not feature-specific.
- 13. **Fold Change** The class zero mean divided by the class one mean.
- 14. Class Zero Mean The class zero mean.
- 15. Class Zero Standard Deviation The class zero standard deviation.
- 16. Class One Mean The class one mean.
- 17. Class One Standard Deviation The class one standard deviation.

The results from the ComparativeMarkerSelection algorithm can be viewed with the ComparativeMarkerSelectionViewer.

Parameters:

Name Description



input filename The input file - .res, .gct, .odf type=Dataset

cls filename The class file - .cls

confound variable cls

filename

The class file containing the confounding variable - .cls

test direction The test to perform (up-regulated for class 0, up-regulated for

class 1, two-sided)

test statistic The statistic to use

min std The minimum standard deviation if test statistic includes min std

option

number of permutations The number of permutations to perform (use 0 to calculate

asymptotic p-values)

complete Whether to perform all possible permutations balanced Whether to perform balanced permutations random seed The seed of the random number generator

smooth p values Whether to smooth p-values

phenotype test Tests to perform when cls file has more than two classes (Note:

not fully corrected for multiple hypothesis testing

Return Value:

An odf file of type ComparativeMarkerSelection

References:

- 1. Golub, T.R., et al., *Molecular Classification of Cancer: Class Discovery and Class Prediction by Gene Expression.* Science, 1999. **286**(5439): p. 531-537.
- 2. Slonim, D.K., et al., Class Prediction and Discovery Using Gene Expression Data, in RECOMB 2000: The Fourth Annual International Conference on Research in Computational Molecular Biology. 2000: Tokyo, Japan. p. 263-272.
- 3. Benjamini, Y. and Y. Hochberg, *Controlling the False Discovery Rate: A Practical and Powerful Approach to Multiple Testing.* Journal of the Royal Statistical Society. Series B (Methodological), 1995. **57**(1): p. 289-300.
- 4. Storey, J.D. and R. Tibshirani, *Statistical significance for genomewide studies*. PNAS, 2003. **100**(16): p. 9440-9445.
- 5. Westfall, P.H. and S.S. Young, *Resampling-Based Multiple Testing: Examples and Methods for P-Value Adjustment*. Wiley Series in Probability and Statistics. 1993, New York: Wiley.

Platform dependencies:

Task type: Gene List Selection

CPU type: any
OS: any
Java JVM level: 1.4
Language: Java, R