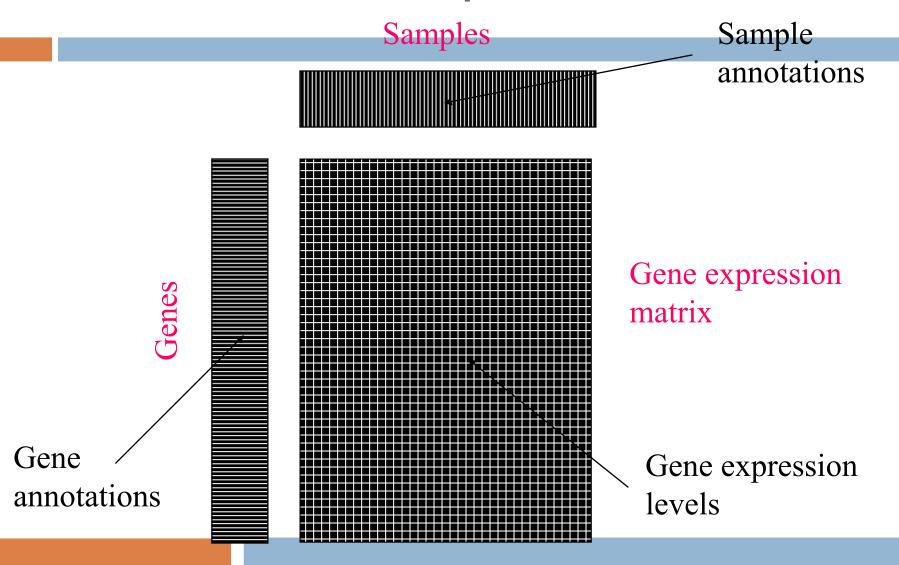
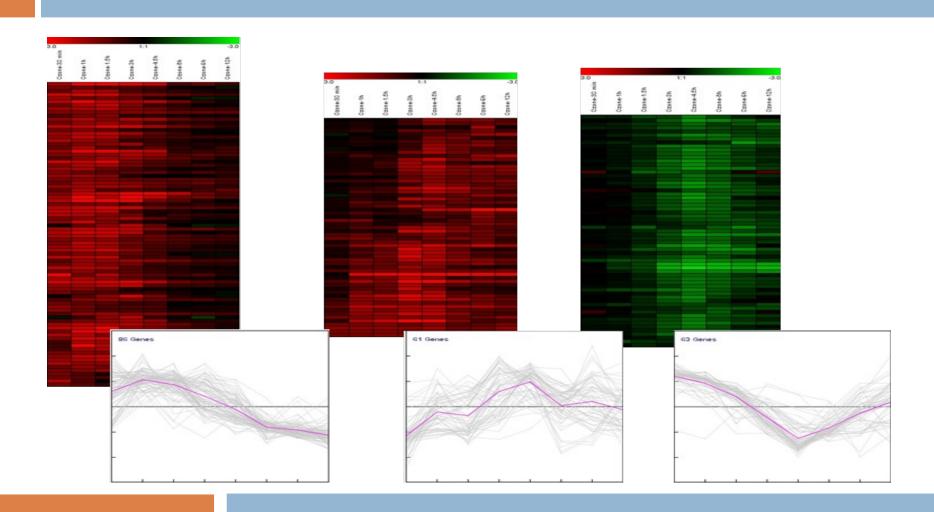
Bioinformatika Gene expression data analysis

Michael Anděl (some slides are courtesy of Mark Craven, U. of Wisconsin)

GE data – conceptual view



GE data – image view



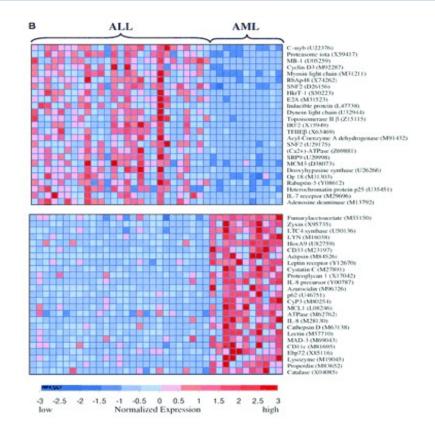
Classification task

Challenge:

- \square samples (10¹) x features (10³)
- False hypotheses, overfitting
- Interpretability: are the expressed genes the causal ones?

What to do?

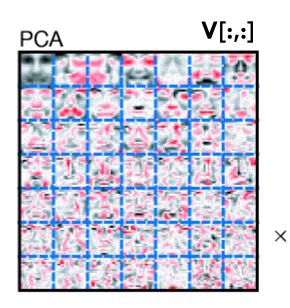
- Decrease number of hypotheses
- Analyze more abstract entities than genes, eg. principal components

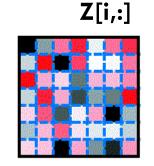


Golub et al.: Molecular classification of cancer: Class discovery and class prediction by gene expression monitoring. Science, 1999

PCA – motivation

- □ M ... # genes
- N ... # samples
- X (N x M) ... GE data in the space of genes





- V (M x K) ... transformation basis, eigengenes
- Z (N x K) ... transformed GE data in the space of eigengenes
- K ... # of eigengenes,
 i.e. the number of underlying

concepts



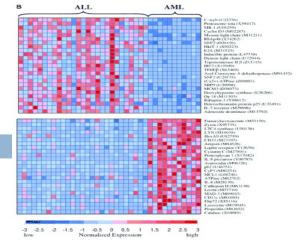
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Lee et al.: Learning the parts of objects by non-negative matrix factorization. Science, 1999

Assignement

Data

7,129 GE profiles of 72 patients



Golub et al.: Molecular classification of cancer: Class discovery and class predi-ction by gene expression monitoring. Science, 1999

- 25 samples ... acute myeloid leukemia (AML)
- 47 samples ... acute lymphoblastic leukemia (ALL)

Task

Construct decision tree model to differentiate these types of tumours.



Workflow

- 1) Learn a decision tree on subjected data. Use Matlab class ClassificationTree, for learning use method fit().
- 2) Visualize resulting model (function view()) and enumerate the training error (function predict())
- 3) Estimate the real error of the model by the means of crossvalidation. Use script **cv11.m**.
- 4) Compare the training error and real error estimation. Why are so different?



Analyzing in terms of PCAs

Transform the data to the reduced space of eigengenes. Use attatched functio **pca.m**:

- Do the transformation for the number of cmponents K=5:5:64, i.e. Z = X V(1:K,:).
- 2) For each of these transformed data Z learn the decision tree.
- 3) Choose an appropriate model according to its training error and model complexity.
- 4) Evaluate the chosen model by the crossvalidation
- 5) Compare the error estimation with the one of simple tree.



Interpreting the results

- Have the genes from your model been already reported in previous studies as associated with leukemia in terms of mutations or polymorphisms? Use web tool Phenopedia HuGE Navigator.
- Try to find some gene from most important PCAs of the tree under the disease terms 'Leukemia, Myelocytic, Acute' and 'Leukemia, Lymphocytic, Acute'.
- 3) To mine the genes from the important components of the tree use attached function **mineGenes.m** and **geneNames.mat**.
- 4) Simmilarly find some genes from the **component tree**.