



# **Applied Machine Learning - Intro**

### Annalisa Marsico OWL RNA Bioinformatics, MPI Molgen Berlin Freie Universität Berlin 20.04.16

## Who am I?





### "Laurea" in Physics (5 years)

Thesis: "From conventional tests to adaptive tests: application of artificial neural networks to ability evaluation"

### Master in Bioinformatics (1.5 years)

PPNEMA: database for sorting and classificaiton of rRNA genes in nematodes

### PhD in Bioinformatics (4 years)

Prof. Michael Schroeder (TU Dresden) & Prof. Marino Zerial (MPI MGC)

- **clustering algorithm** for de novo motif discovery
- MeMotif: database of sequence / structure motifs
- pattern classification of unolding pathways from biophysical data

### est-doc at the Max Planck for molecular Genetics Berlin (3 years)

- f. Martin Vingron
  - PROmiRNA: semi-supervised learning of miRNA promoters
  - linear model for miRNA processing from sequence signal
  - RNA-seq and ChIP-seq data for infection processes (SFB TR48)

## The RNA Bioinformatics group

Since July 2014 Junior Professor for High-Throughput Genomics (FU)
& group leader of the RNA Bioinformatics group at MPI Molgen

http://www.molgen.mpg.de/2733742/RNA-Bioinformatics



## **Additional co-lecturers**





Stefan Budach, PhD student

- Bachelor & Master in Bioinformatics, Freie Universität Berlin
- Bachelor thesis in Martin Vingron's lab on statistical modeling of batch effects in RNA-Seq data
- Student assistant in my lab for a year (2014) predictive model of miRNA-eQTLs
- Master thesis in Knut Reinert's lab on parallelization in SeqAn
- Since November 2015, IMPRS PhD student in the RNA Bioinformatics Lab



### Wolfgang Kopp, PhD student

- Bachelor & Master in Biomedical Engineering, Technical University of Graz (Austria)
- student assistant for the ,Computational Intelligence' course
- Since 2012 PhD student of the IMPRS in the lab of Martin Vingron (MPI Molgen) Projects: statistical models for pattern occurrences in DNA sequences, ChIP-seq timeseries analysis

### Work in our Lab



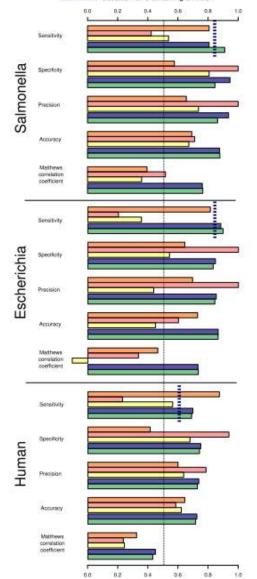
- RNA Binding Proteins (RBPs) prediction
- RBPs target identification
- Target specificity (de novo motif discovery)

## **RBPs prediction**

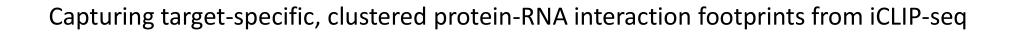


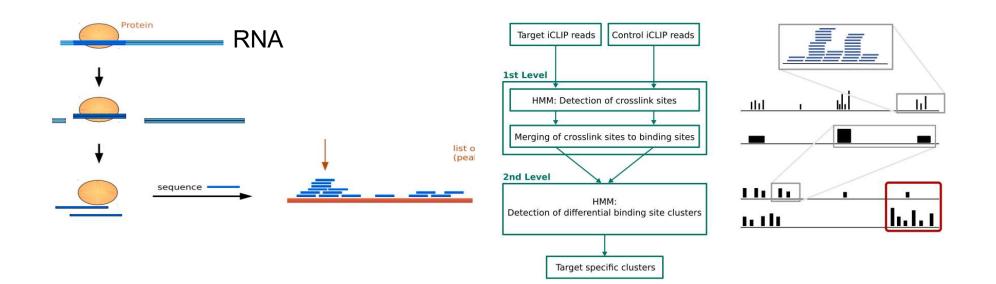
### RNApred = SPOT-Seq-RNA = catRAPID = TriPepSVM = ForestOpt TripSVM - A tool for species-specific RBP prediction ..... classical RBPs containing an RBD 0.2 D.4 Human (9606) Senativity Salmonella (590) mode: normal Uniprot taxon identifier mode: recursive Salmonella Specificity . normal recursive Uniprot (Swiss-Prot) Precision proteom QuickGO QuickGO, (taxon identifier) Uniprot (Taxonomy) Accuracy **RNA-binding annotation RNA-binding annotation** (taxon identifier) proteins with length (all taxa which are hierarchically Mathews consistent > 6000 or < 50 arranged to the specified taxon) coefficient Uniprot (Keywords) Sensitivity. CD-HIT (Li et al., 2001) proteins from RNA associated keyword list (Caland Linet al., 2001) proteins with sequence Escherichia such as RNA-, DNA- or similarity higher than 75 % Specificity Nucleotide-binding Precisión QuickGO positive data set RNA, DNA or nucleotidebinding annotation Accuracy 256 1740 Mathews Pfam correlation proteins containing RNA associated Pfam domains databases/applications Sensitivity collected proteins CD-HIT (Li et al., 2001) proteins with sequence removed proteins Human similarity higher than 75 % negative data set Accuracy 12054 1048

Annkatrin Bressin, Benedikt Beckmann



## **RBPs and their targets**



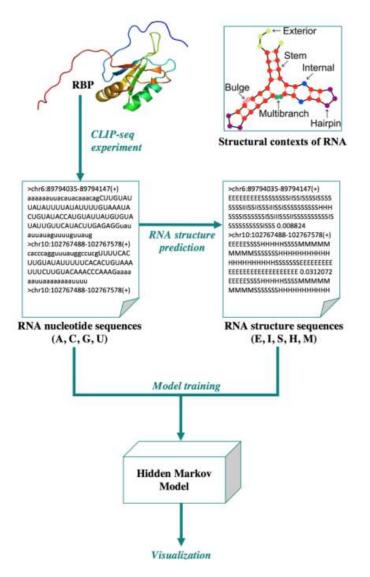


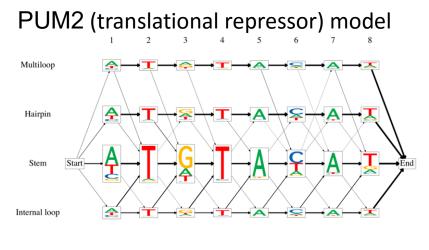
Sabrina Krakau, Hugues Richard (PMC Paris)

## **RBPs and their targets**

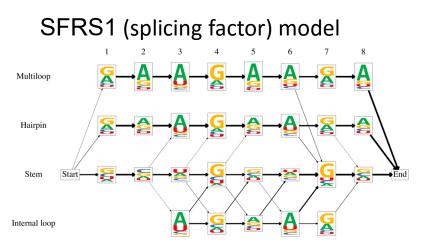


### A tool for de novo sequence-structure motif discovery





Exterior loop



Exterior loop

David Heller, Ralf Krestel (Uni Potsdam), Uwe Ohler (MDC)

## Work in our Lab

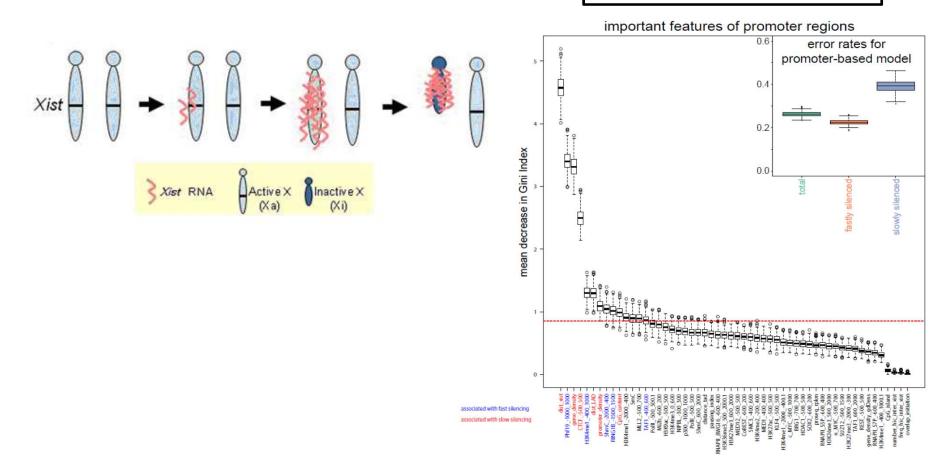


- Regulation and function of IncRNAs
- microRNA-eQTL prediction





### Random Forest prediction



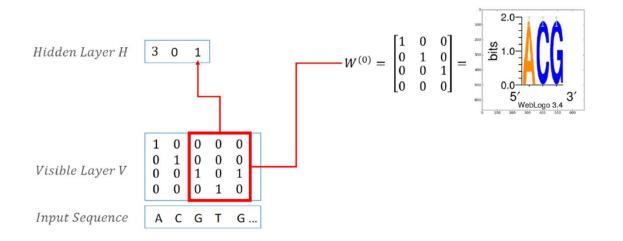
Lisa Barros, Edda Schulz

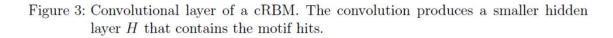
## **Deep Learning**



Predicting DNAse I Hypersensitivity Sites from Dna Sequences Using Deep Learning Techniques

- Convolutional Boltzmann Machines





Roman Schulte-Sasse, Wolfgang Kopp

## **The Plan**



Lecture	Application	Method
8 June	Cancer classification from gene expression	Partial Least Square Regression (PLSR)
15 June	Cancer classification from clinical data / DNA methylation	SVMs
22 June	RBP binding site prediction from RNA sequences	String kernel SVMs
29 June	Promoter prediction from epigenetic features	Semi-supervised learning: co- training algorithm
06 July	Cancer location prediction from microRNAs	Neural Networks (multi-class classification)
13 July	Applications of DP in Bioinformatics	Concepts of deep Learning
20 July	Discuss applications of DP in Bioinformatics	Discuss applications of DP in Bioinformatics

## **Deep Learning Papers**



- 1. Deep Architectures for Protein Contact Map Prediction <u>http://www.ncbi.nlm.nih.gov/pubmed/22847931</u>
- 2. Toxicity Prediction Using Deep Learning http://arxiv.org/pdf/1503.01445.pdf
- 3. Deep Learning of tissue-regulated splicing code <u>http://www.ncbi.nlm.nih.gov/pubmed/24931975</u>
- Predicting effects of non-coding variants with deep-learning-based sequence model <u>http://www.ncbi.nlm.nih.gov/pubmed/26301843</u>
- Predicting the sequence specificities of DNA- and RNA-binding proteins by deep learning <u>http://www.ncbi.nlm.nih.gov/pubmed/26213851</u>

### Link to course material



http://www.molgen.mpg.de/3434810/Applied-Machine-Learning

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