

# Übungen zum Sequenzanalyse-Praktikum

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<http://wiki.techfak.uni-bielefeld.de/gi/Teaching/2013summer/SequaPrak>

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Multiple sequences usually carry more information than each sequence alone. For RNA, compensating base mutations are a strong evidence for putative base pairs: If a putative G-C base pair mutates to an A-U base pair these positions are likely to pair.

## Exercise 14.1:

**Plan A** Use CLUSTALW <http://bibiserv.techfak.uni-bielefeld.de/clustalw/> to align the following two tRNA sequences. Then fold the resulting alignment with RNAALIFOLD <http://rna.tbi.univie.ac.at/cgi-bin/alifold.cgi>. Report consensus Vienna-Dot-Bracket structure and consensus sequence.

```
>Human_tRNA_38.chr6 (Ile, AAT)
GGCTGGTTAGTTCAGTTGGTtAGAGCGTGGTGCTAATAACGCCAAGGtCG
TGGGTTTCGATCCCCATATCGGCCA
>Human_tRNA_2.chr7 (Pro, AGG)
GGCTCGTTGGTCTAGGGGTATGATTCTCGCTTAGGGTGCGAGAGGTCCCG
GGTTCAAATCCCGGACGAGCCC
```

**Plan B** Simultaneously align and fold the two tRNA sequences with FOLDALIGN <http://foldalign.ku.dk/server/index.html>. Report consensus Vienna-Dot-Bracket structure.

**Plan C** Structurally align the two MFE-structures computed by RNAFOLD <http://rna.tbi.univie.ac.at/cgi-bin/RNAfold.cgi> with RNAFORESTER <http://bibiserv.techfak.uni-bielefeld.de/rnaforester> in multiple mode (you would not get a consensus structure in pairwise mode). Report consensus Vienna-Dot-Bracket structure and consensus sequence. Are there any differences to the alignments from Plan A. and Plan B.?

**Plan D** Plan D is solely based on stochastic properties like covariance or mutual information. It needs a lot of input data and does not answer the question for *one* optimal secondary structure, but gives a stochastically sound overview about compensatory mutation during evolution. Take the 20 tRNA sequences from <http://tinyurl.com/nhq46em>, extracted from RFAM <http://rfam.sanger.ac.uk/>, in multiple Fasta format. Use MATRIXPLOT <http://www.cbs.dtu.dk/services/MatrixPlot/mutualNucl/index.html> to compute the mutual information. Report the dot-plot. Can you detect the clover leaf in the plot?

## Exercise 14.2:

Which plan works best? Which is the fastest one? State your judgements in just a few sentences.