# (RNA-) Structuredness in Viruses

#### Roman Ochsenreiter

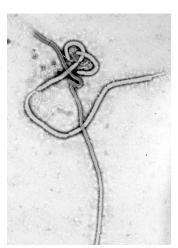
TBI Wien University of Vienna

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### Viruses I

#### Minimal Life Forms

- Obligate intracellular parasites
- No organelles
- (Very) small size
- DNA/RNA genomes



Electron micrograph of an Ebola Virion, size: 1000 nm (image source: wikipedia)



# Viruses II Minimal Life Forms

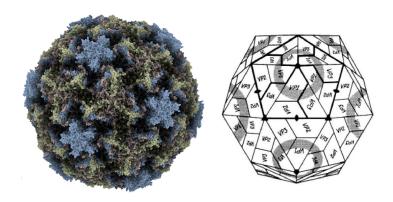


Figure: Crystal Structure and Schematic of Rhinovirus A 1

Genome size (nt): Human:  $3.2 \times 10^9$  Rhinovirus:  $7 \times 10^3$ 

<sup>&</sup>lt;sup>1</sup>Zhao et al. 1996

### Viruses and RNA Structure

- Viruses have tightly packed genomes
- Specific Functions are often provided by RNA Structures

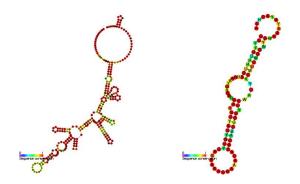
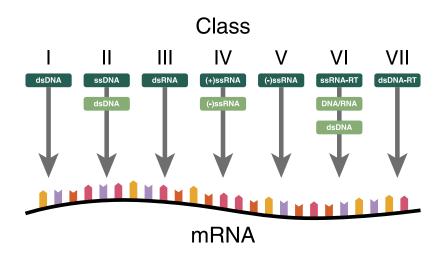


Figure: Pestivirus IRES<sup>1</sup>, Rhinovirus CRE-Element

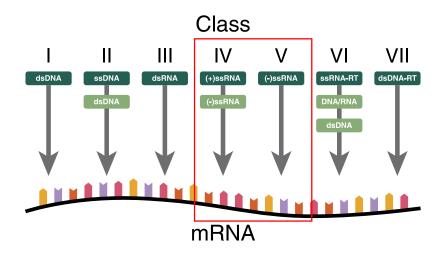


<sup>&</sup>lt;sup>1</sup>Rfam IDs: RF00209, RF00220

**Baltimore Classification** 



**Baltimore Classification** 



### Viral Phylogeny

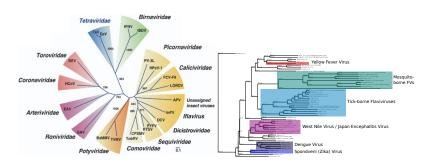
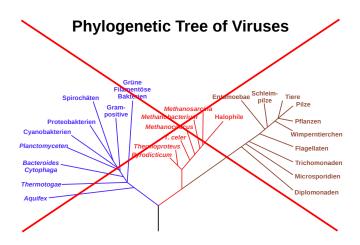


Figure: Picorna-like Viruses (left)1, Flavivirus (right



<sup>&</sup>lt;sup>1</sup>Fauquet, Schrock; 2006

#### Viral Phylogeny



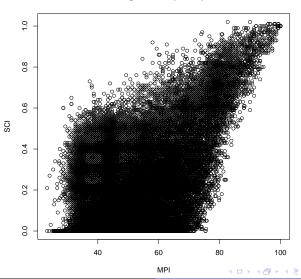
# Virus Ortholog Groups

### Viral Clusters of Ortholog Groups

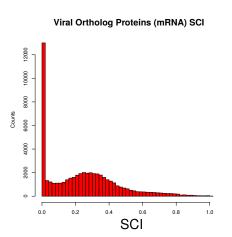
- Groups of Ortholog proteins
- Calculated on Protein alignments
- Based on newest RefSeq Virus Genomes
- Includes all annotated viral species

# Virus Ortholog Groups





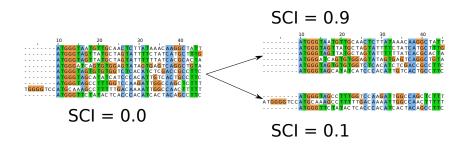
# Virus Ortholog Groups



# Viral Ortholog Proteins (mRNA) MPI 4000 3000 Counts 60 20 40 80 100 MPI

# Viral Ortholog Proteins

### Conserved sub-alignments



- No overall consensus structure but...
- Strong structure conservation in subparts of the alignment!



### Viruses and structured RNAs

More than UTRs...

#### The Problem

- Viruses are a highly diverse group of organisms
- Just a fraction is investigated for ncRNAs
- UTRs get the most attention

#### **Aims**

- Screen all viruses for structured RNAs
- Take a close look at CDS regions

#### Methods

- RNAz
- R-scape



### **Dataset**

#### Isolation Pipeline

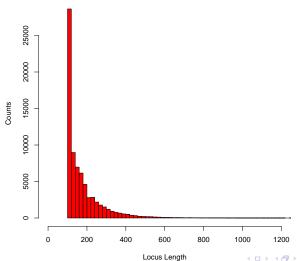
- Fetch Species from NCBI taxonomy database
- Petch Refseq Sequences
- Fetch genomes from NCBI Nucleotide Database
- Align with clustal Omega
- Multiple consistency checks in-between

| Name               | #Species |
|--------------------|----------|
| NCBI Tax DB        | 18939    |
| Refseq DB          | 8299     |
| Consistent Species | 1580     |

| Type       | #Species |  |
|------------|----------|--|
| dsDNA      | 422      |  |
| ssDNA      | 276      |  |
| dsRNA      | 142      |  |
| ssRNA      | 600      |  |
| RT         | 81       |  |
| Satellites | 32       |  |

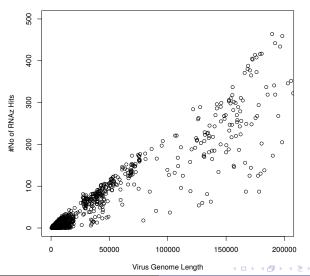
### Results





### Results





### Results

|            | #Hits | w/ Cov (G) | w/ Cov (RAF) |
|------------|-------|------------|--------------|
| Total Hits | 73564 | 4216       | 43554        |
| CDS        | 69941 | 4806       | 41579        |
| UTR        | 3664  | 410        | 1975         |

# Summary & Outlook

### Summary

- Structured Elements are widespread in Viruses
- Lots of Covariation in most elements!
- CDS and UTRs are equally affected

#### Outlook

- Check again with structure-based alignments
- Build covariance models for all significant loci
- Search for 'structural orthologs'

# Acknowledgements

#### Thanks to...

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...and you!



