# GRECO TFBS Benchmarking Initiative Towards a representative set of TFs

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## **GitHub repository**

https://github.com/oriolfornes/GRECO

## **Outline**

• Aim

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#### **Aim**

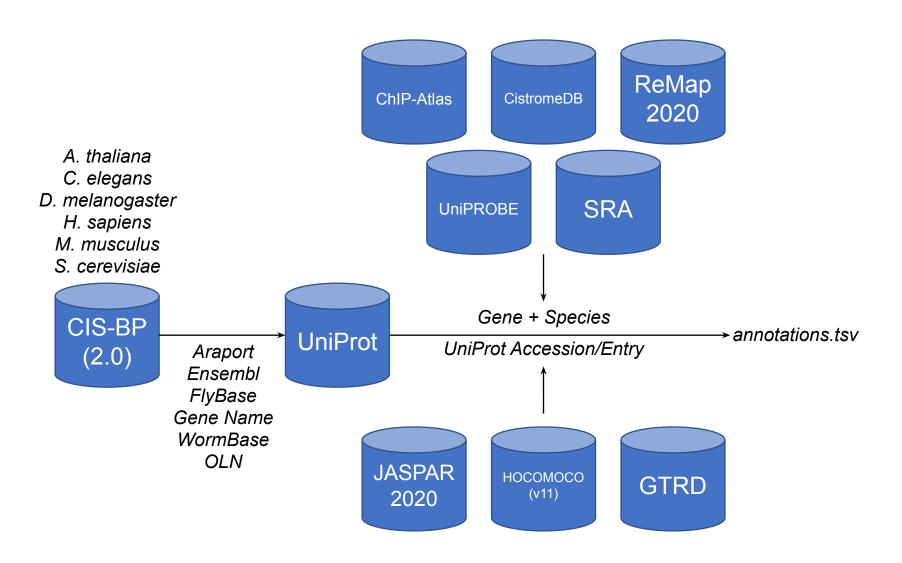
Obtain a representative set of TFs:

1. Supported by high-quality experimental data;

2. From different model organisms; and

3. From different structural families.

#### **Annotation**



## **Annotation**

https://raw.githubusercontent.com/oriolfornes/GRECO/master/annotations.tsv

## Clustering

- For each TF (i.e. query)... (sort by amount of experimental evidence):
  - 1. Identify the query's Pfam DBD(s) using hmmscan with the "--domtblout" option and E-value thresholds for models and domains of 10<sup>-5</sup> and 10<sup>-2</sup>, respectively;
  - 2. Search for TFs homologous to the query using BLAST+;
  - 3. Select homologs:
    - with the same DBD composition than the query; and
    - whose BLAST+ alignment with the query is above the Rost's sequence identity curve;
  - 4. For each selected homolog, if the amino acid sequence identity of the query and homolog DBDs is greater than the DBD-specific motif inference thresholds from CIS-BP, cluster the TFs together.

## Clustering

https://github.com/oriolfornes/GRECO/blob/master/Data/Clusters/TFs.json

#### Results

- Triads: i.e. TFs with support by in vivo and at least two in vitro methods
- Species:
  - 1. Drosophila melanogaster 1
  - 2. Homo sapiens 57
  - 3. Mus musculus 28
- Families:
  - 1. C2H2 ZF **15**
  - 2. C2H2 ZF, MADF 1
  - 3. CUT, Homeodomain 1
  - 4. DM **1**
  - 5. E2F 1
  - 6. Ets **4**
  - 7. Forkhead 6
  - 8. GATA 2
  - 9. Homeodomain 13
  - 10. Homeodomain, POU 1
  - 11. Homeodomain, Paired box 1
  - 12. Nuclear receptor 14
  - 13. RFX **2**
  - 14. Rel **2**
  - 15. SAND 1
  - 16. Sox **4**
  - 17. bHLH **9**
  - 18. bZIP 8

#### Results

- <u>Duos</u>: *i.e.* TFs with support by *in vivo* and *in vitro* methods
- Species:
  - 1. Arabidopsis thaliana 98
  - 2. Caenorhabditis elegans 30
  - 3. Drosophila melanogaster 39
  - 4. Homo sapiens 279
  - 5. Mus musculus 130
  - 6. Saccharomyces cerevisiae 32

Families: 62

#### Results

- Multiple evidence: i.e. TFs with support by at least two methods
- Species:
  - 1. Arabidopsis thaliana 98
  - 2. Caenorhabditis elegans 30
  - 3. Drosophila melanogaster 40
  - 4. Homo sapiens 290
  - 5. Mus musculus 141
  - 6. Saccharomyces cerevisiae 32

Families: 62

### **Next steps**

 Ensure that the representative set of TFs contains only sequence-specific DNA-binding TFs

 For each representative TF, ensure that the mapped experimental data correspond to that TF