

Functional Annotation of proteins

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École "Assemblage & Annotation" AVIESAN 2022 - Roscoff



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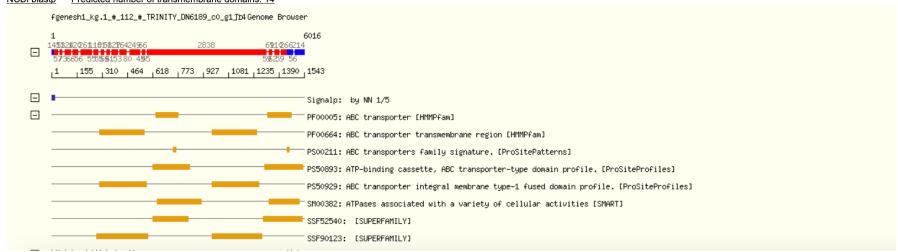
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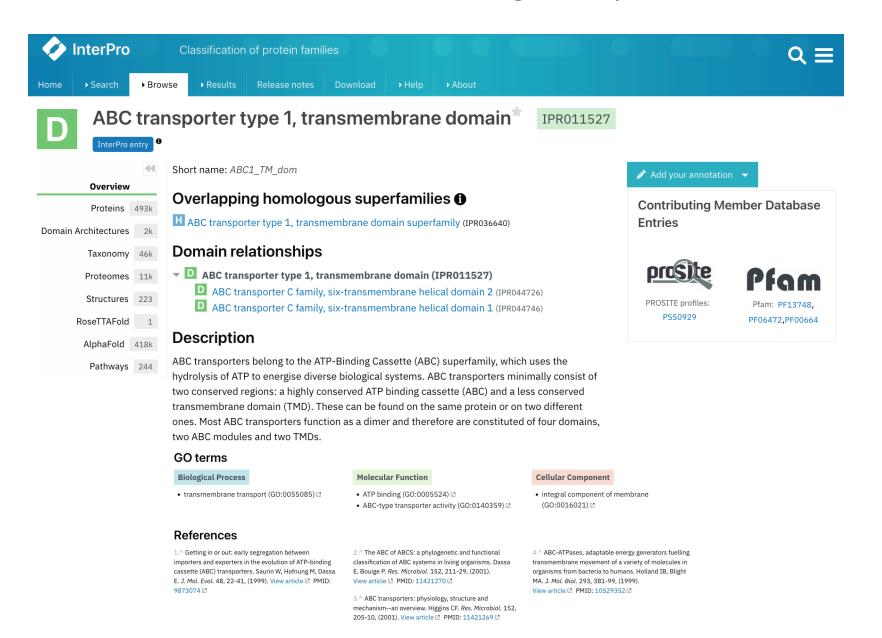
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View/modify manual annotation

View nucleotide and 3-frame translation To Genome Browser

NCBI blastp Predicted number of transmembrane domains: 14







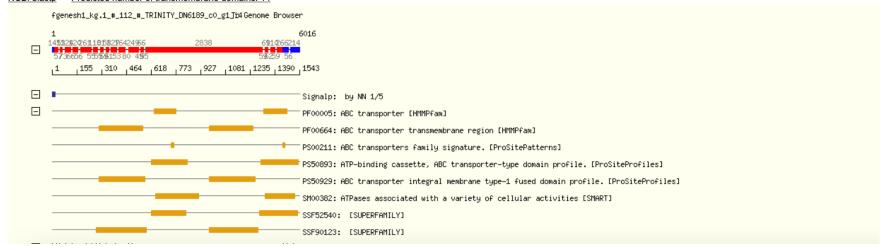
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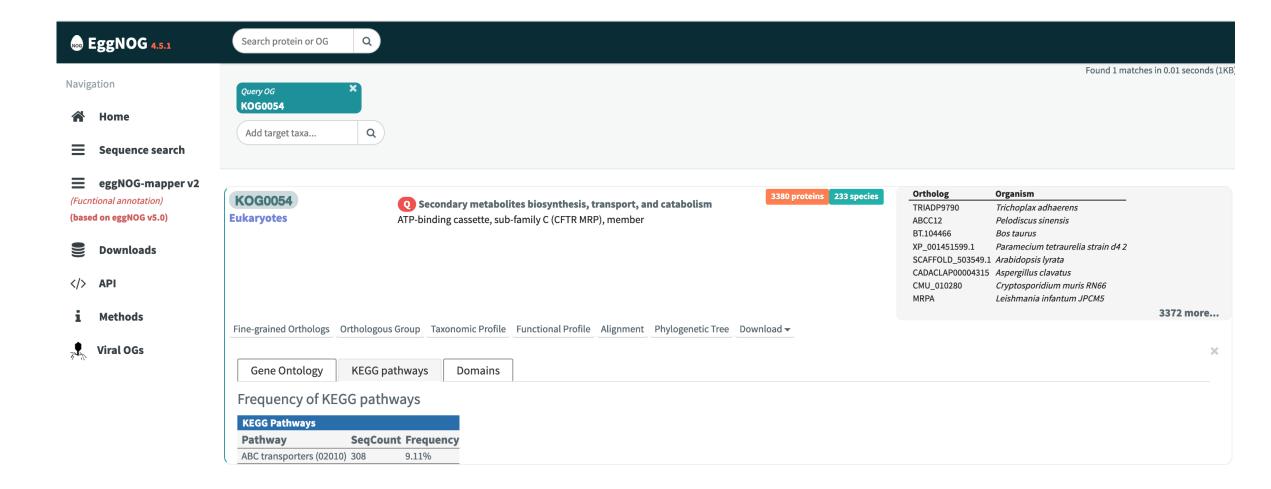
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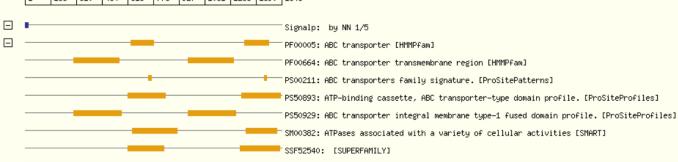
View/modify manual annotation

<u>View nucleotide and 3-frame translation</u> <u>To Genome Browser</u>

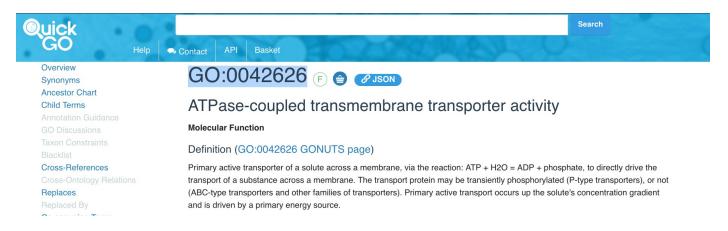
NCBI blastp Predicted number of transmembrane domains: 14

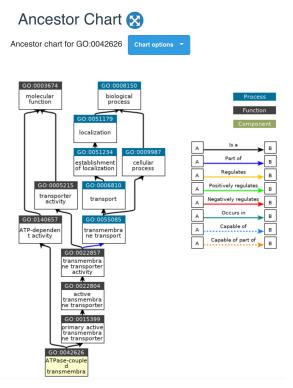
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SSF90123: [SUPERFAMILY]





Child Terms

This table lists all terms that are direct descendants (child terms) of GO:0042626

Child Term	Relationship to GO:0042626
GO:1901514 F 📵 🕁 ATPase-coupled lipo-chitin oligosaccharide transmembrane transporter activity	is_a
GO:0034040 F 🛍 🕁 ATPase-coupled lipid transmembrane transporter activity	is_a
GO:0019829 F 🖨 🕁 ATPase-coupled cation transmembrane transporter activity	is_a
GO:0042625 F 🖨 🕁 ATPase-coupled ion transmembrane transporter activity	is_a
GO:0033225 F 📵 🚯 ATPase-coupled 2-aminoethylphosphonate transporter activity	is_a
GO:0015450 F 🏚 💠 protein-transporting ATPase activity	is_a
GO:0033221 F 📵 🚯 ATPase-coupled urea transmembrane transporter activity	is_a
GO:0098533 C 🏚 🕁 ATPase dependent transmembrane transport complex	capable_of
GO:0043225 F 🖨 🕁 ATPase-coupled inorganic anion transmembrane transporter activity	is_a
GO:0140359 F 🖨 🕁 ABC-type transporter activity	is_a

Functional annotation pipelines



Victoria Dominguez Del Angel 10, Erik Hjerde 10, Lieven Sterck 10,4, Salvadors Capella-Gutierrez 10,5,6, Cederic Notredame^{7,8}, Olga Vinnere Pettersson⁹, Joelle Amselem 10, Laurent Bouri 10, Stephanie Bocs 10,11,13, Christophe Klopp 10,4, Jean-Francois Gibrat 10,15, Anna Vlasova 10,8, Brane L. Leskosek 16, Lucile Soler 17, Mahesh Binzer-Panchal 10,17, Henrik Lantz 10,17

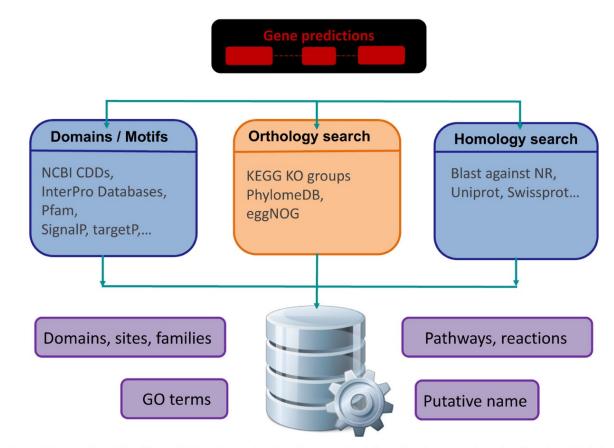
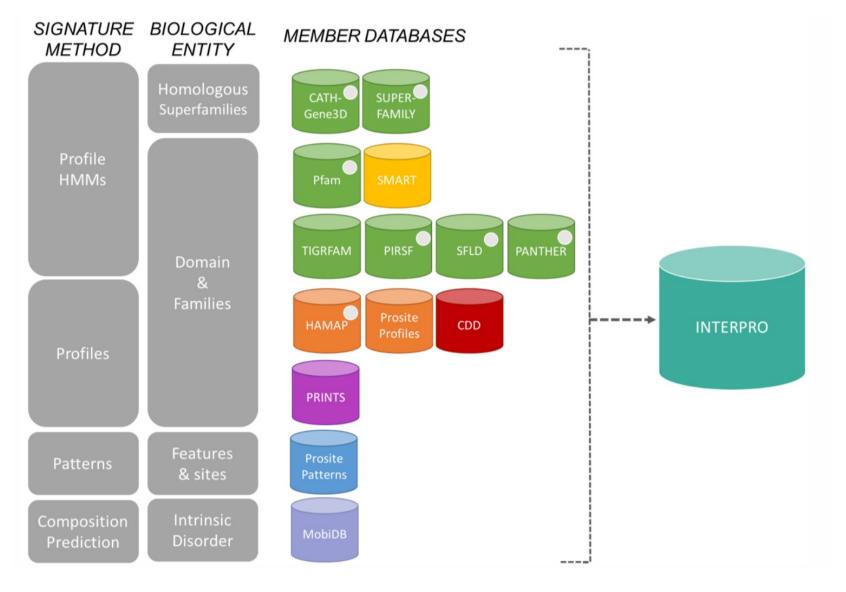


Figure 4. Functional Annotation Pipelines. This schema is showing a typical functional annotation pipeline, in which functional roles are assigned to coding sequences (CDSs) inferred in the gene prediction process. The process implements three parallel routes for the definition of functions. The first refers to proteins domains and motifs, the second for orthology search and finally the third is applied to homology search. At the end, the output from the three different sources is put together for more valuable predictions.

Domains / Motifs NCBI CDDs, InterPro Databases, Pfam, SignalP, targetP,...

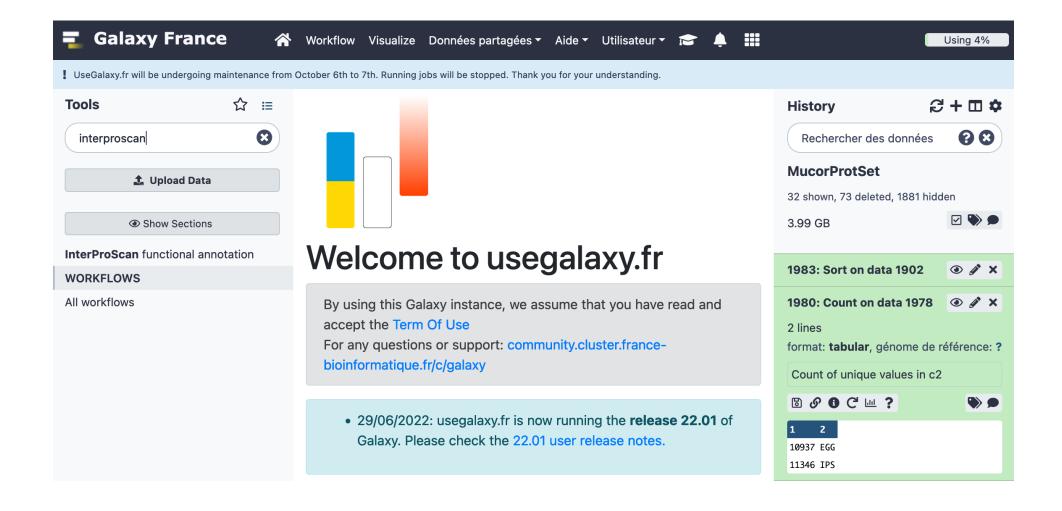
Running InterProScan



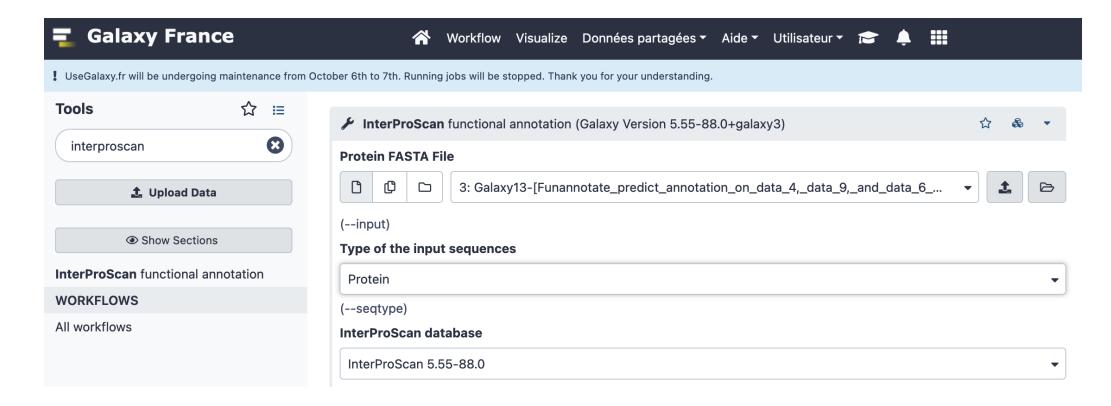
InterProScan is the software package that allows sequences to be scanned against InterPro's member database signatures.

Domains / Motifs

NCBI CDDs, InterPro Databases, Pfam, SignalP, targetP,...

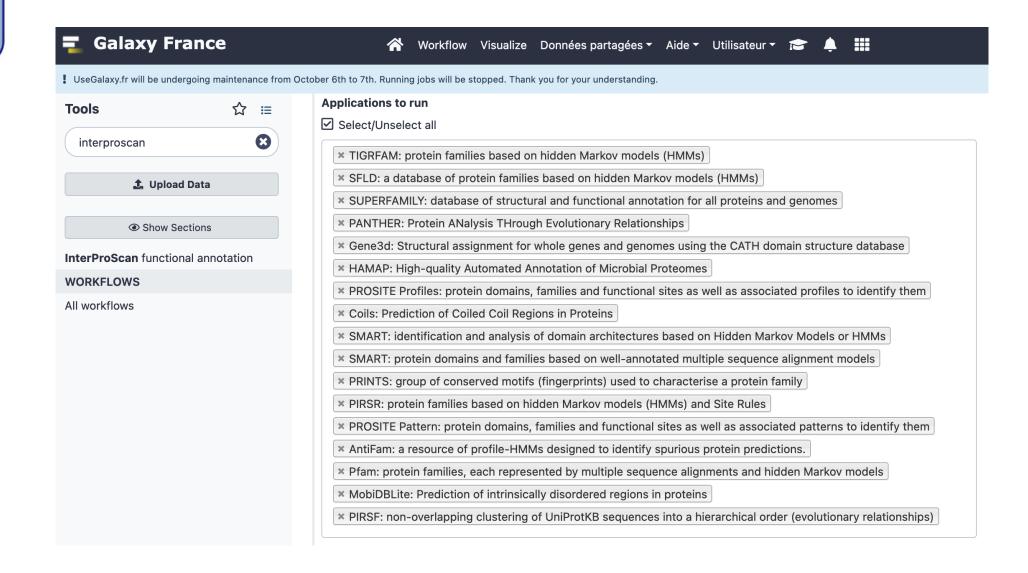


NCBI CDDs, InterPro Databases, Pfam, SignalP, targetP,...



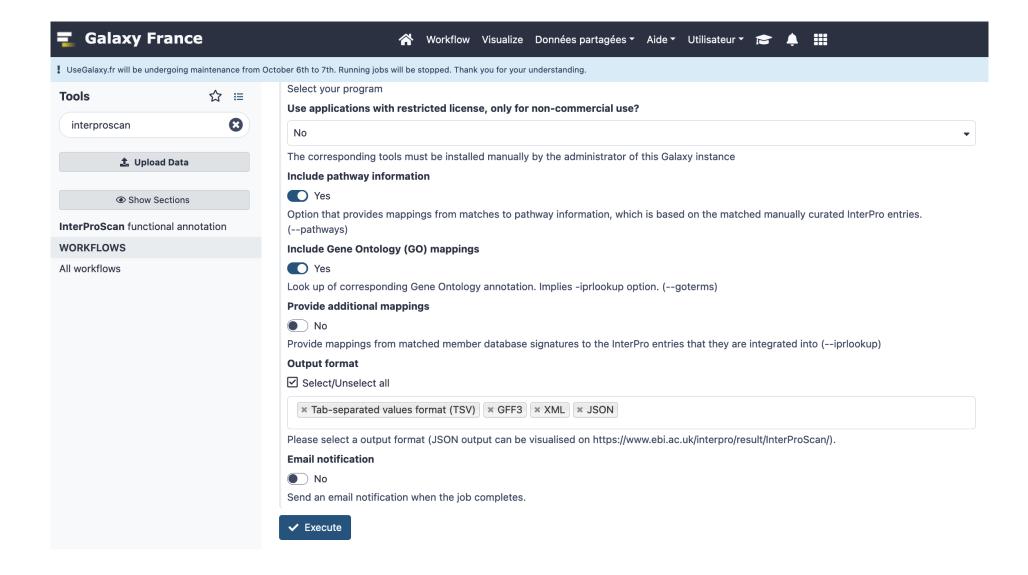
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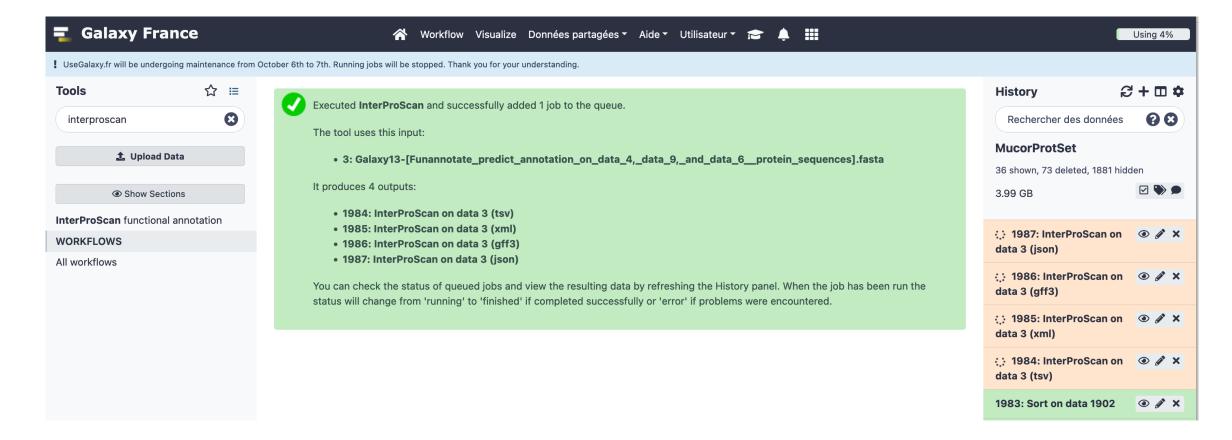


Domains / Motifs

NCBI CDDs, InterPro Databases, Pfam, SignalP, targetP,...



NCBI CDDs, InterPro Databases, Pfam, SignalP, targetP,...



Functional annotation pipelines



Ten steps to get started in Genome Assembly and Annotation

[version 1; peer review: 2 approved]

Victoria Dominguez Del Angel (10, Erik Hjerde (10, Lieven Sterck (10, 3, 4), Salvadors Capella-Gutierrez (10, 5, 6), Cederic Notredame^{7,8}, Olga Vinnere Pettersson⁹, Joelle Amselem (10, Laurent Bouri (10, Stephanie Bocs (10, 11, 11, 11, 11)), Christophe Klopp (10, 14, Jean-Francois Gibrat (10, 11, 15), Anna Vlasova (10, 8), Brane L. Leskosek (16, Lucile Soler (17, Mahesh Binzer-Panchal (10, 17), Henrik Lantz (10, 17)

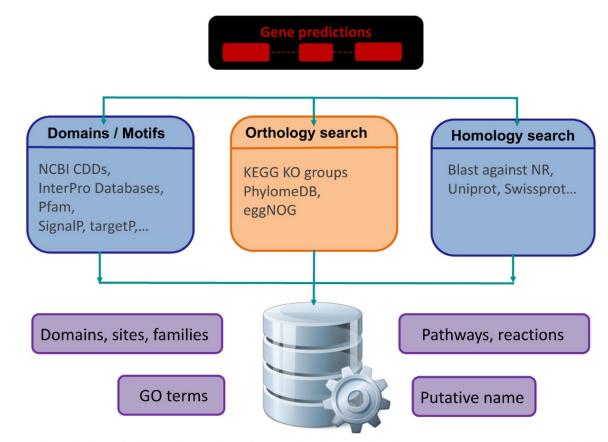


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Orthology search

KEGG KO groups
PhylomeDB,
eggNOG

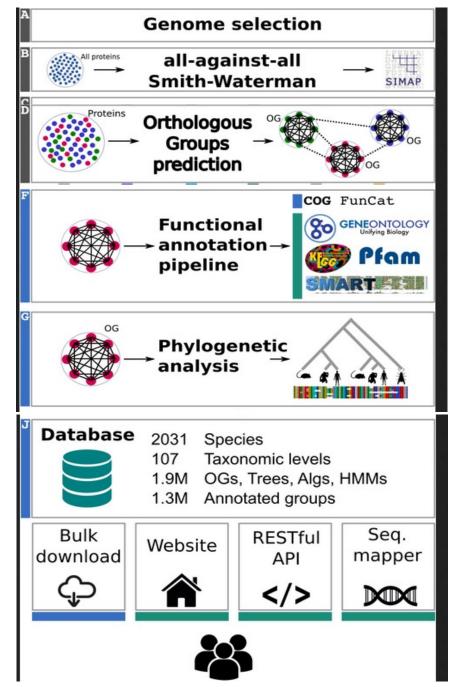
Running EggNOG-mapper

EggNOG v5.0

A database of orthology relationships, functional annotation, and gene evolutionary histories.

Organisms | Viruses | Orthologous Groups | Tree & Algs 5,090 | 2,502 | 4.4M | 4.4M

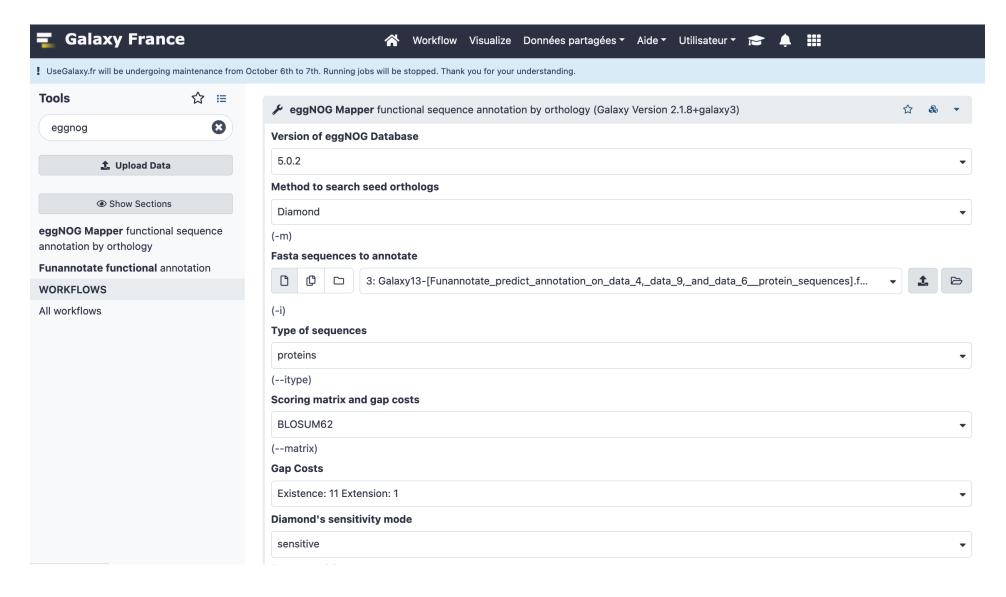
EggNOG-mapper is a tool for fast functional annotation of novel sequences. It uses precomputed Orthologous Groups (OGs) and phylogenies from the EggNOG database (http://eggnog5.embl.de) to transfer functional information from fine-grained orthologs only.



Orthology search KEGG KO groups PhylomeDB,

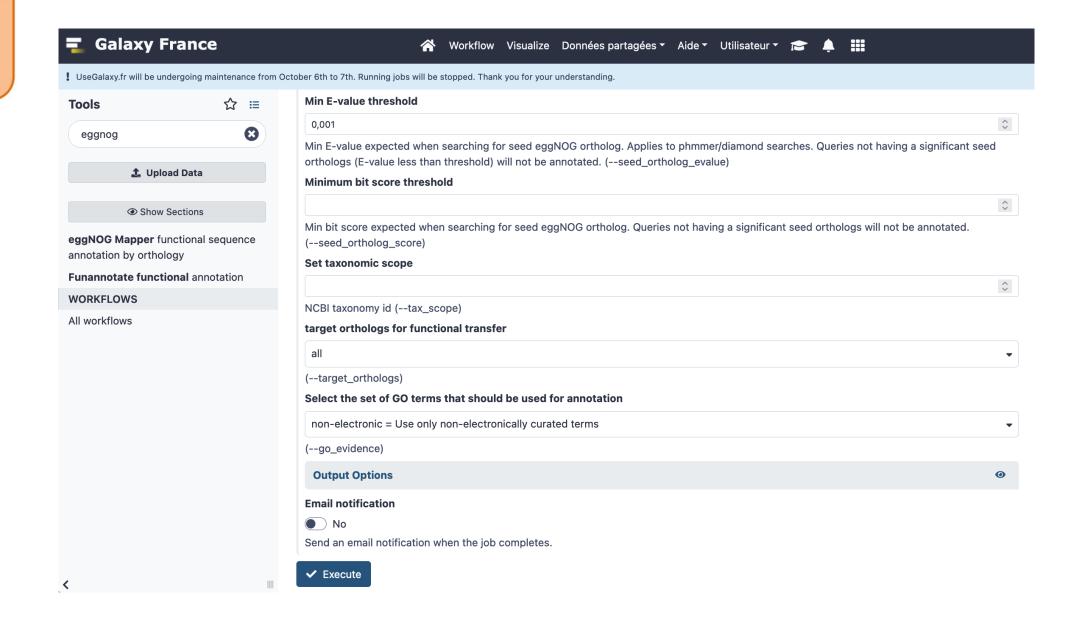
eggNOG

Running EggNOG-mapper



Orthology search

KEGG KO groups PhylomeDB, eggNOG



To be continued ...

How to assign a biological function to a gene/protein?

- Experimental characterization
- Automated prediction:
 - 1/ Search for similar genes/proteins in other organisms
 - 2/ Transfer the annotation to the query sequence

The assumption of function transfer is that the function is retained in proteins that have similar sequences and have evolved from a single ancestor.



Group of genes descended from single gene in LCA of group of species

A. Orthogroup



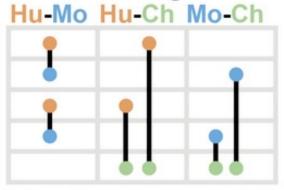
Group of genes descended from single gene in LCA of group of species

A. Orthogroup



Group of genes descended from single gene in LCA of group of species

B. Orthologs



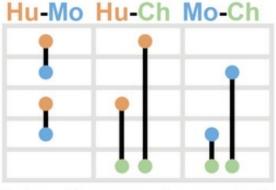
Pairs of genes descended from single gene in LCA of pair of species

A. Orthogroup



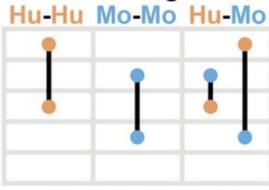
Group of genes descended from single gene in LCA of group of species

B. Orthologs



Pairs of genes descended from single gene in LCA of pair of species

C. Paralogs



Pairs of genes descended from gene duplication event

LCA: Last Common Ancestor

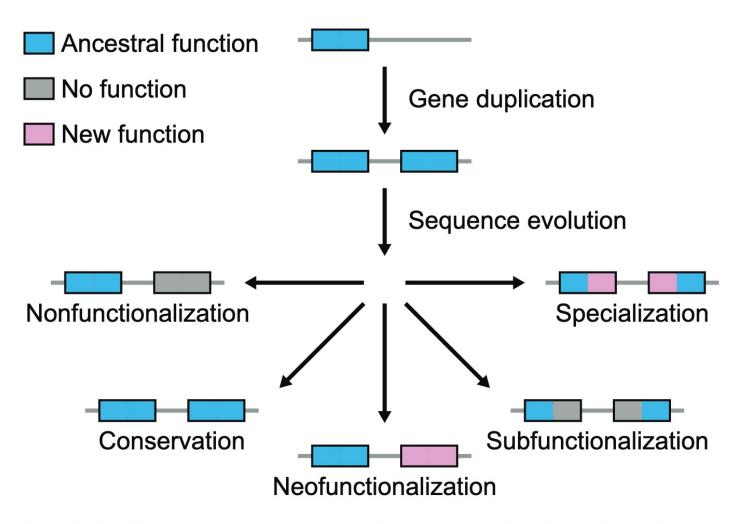
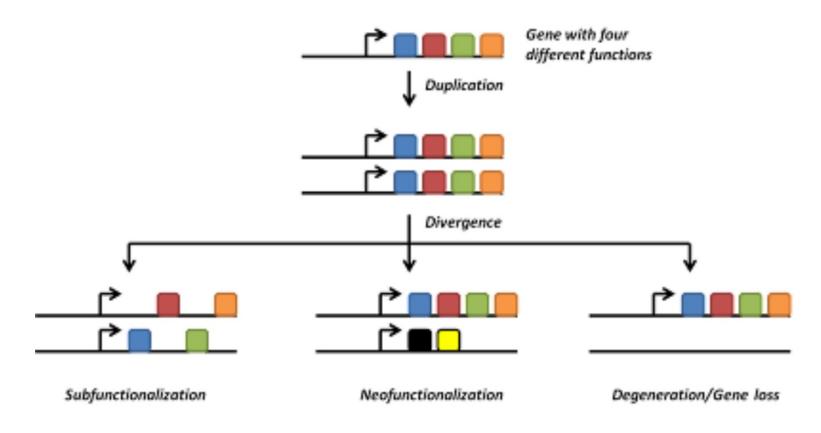
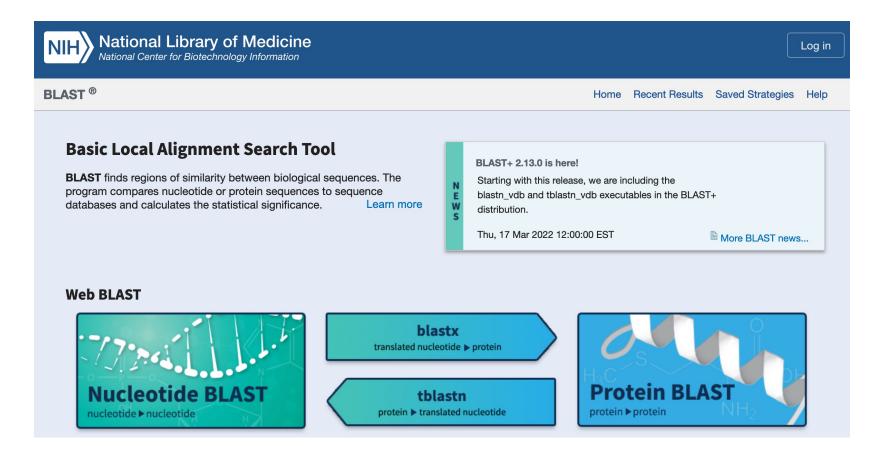


Figure 1: Hypothesized evolutionary trajectories of duplicate genes. Gene duplication results in two copies of an ancestral gene. Evolution may result in the loss of one functional copy by nonfunctionalization, or in the retention of two functional copies by either conservation, neofunctionalization, subfunctionalization, or specialization.

Evolutionary fate of duplicate genes



Search for sequence silmilarity BLAST



Which database ? (quality redundancy ...) Which cut-off ? (E-value)

Local alignment searches suffer from some important caveats

Local alignment searches suffer from some important caveats:

1. Excessive transfer of annotations.



- 2. Annotation errors in the source database.
- 3. Threshold relativity.
- 4.Low sensitivity/specificity.
- 5. Paralogs versus orthologs.

AUTOMATED FUNCTION PREDICTION

Functional annotation prediction: All for one and one for all

Local alignment searches suffer from some important caveats

Local alignment searches (BLAST) suffer from some important caveats:

1. Excessive transfer of annotations.



- 2. Annotation errors in the source database.
- 3. Threshold relativity.
- 4.Low sensitivity/specificity.
- 5. Paralogs versus orthologs.
- 6.Retreave essentialy poorly formatted text

AUTOMATED FUNCTION PREDICTION

Functional annotation prediction: All for one and one for all

¹School of Computer Science and Engineering, and ²Department of Biological Chemistry, The Hebrew University of Jerusalem, Jerusalem 91904, Israel

Functional annotation pipelines



F1000Research 2018, 7(ELIXIR):148 Last updated: 31 MAR 2022



OPINION ARTICLE

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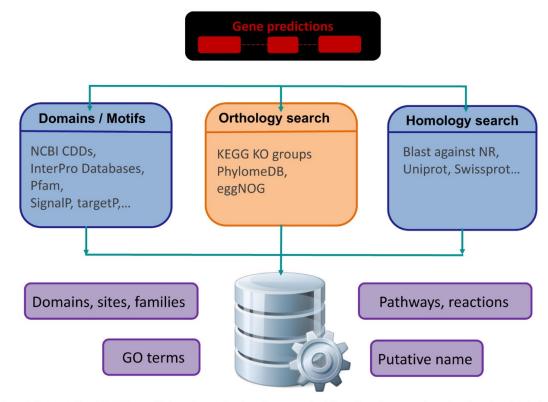


Figure 4. Functional Annotation Pipelines. This schema is showing a typical functional annotation pipeline, in which functional roles are assigned to coding sequences (CDSs) inferred in the gene prediction process. The process implements three parallel routes for the definition of functions. The first refers to proteins domains and motifs, the second for orthology search and finally the third is applied to homology search. At the end, the output from the three different sources is put together for more valuable predictions.

Proteins classification

Proteins can be classified into groups according to sequence or structural similarity.

These groups often contain well characterised proteins whose function is known.

Thus, when a novel protein is identified, its functional properties can be proposed based on the group to which it is predicted to belong.

Proteins can be classified into groups based on

- •the **FAMILIES** to which they belong
- •the **DOMAINS** they contain
- •the **SEQUENCE FEATURES** they possess

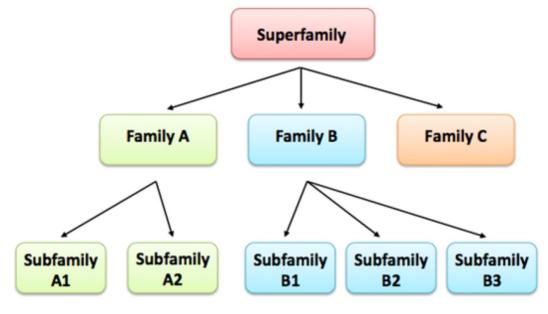


Figure 2 A hypothetical protein family hierarchy showing the relationships between superfamily, family and subfamily members. Directional arrows indicate that one group is a subgroup of another.

Protein families

Families

A protein family is a group of proteins that share a common evolutionary origin, reflected by their related functions and similarities in sequence or structure.

Protein families are often arranged into hierarchies, with proteins that share a common ancestor subdivided into smaller, more closely related groups.

The terms **superfamily** (describing a large group of distantly related proteins) and **subfamily** (describing a small group of closely related proteins) are sometimes used in this context.

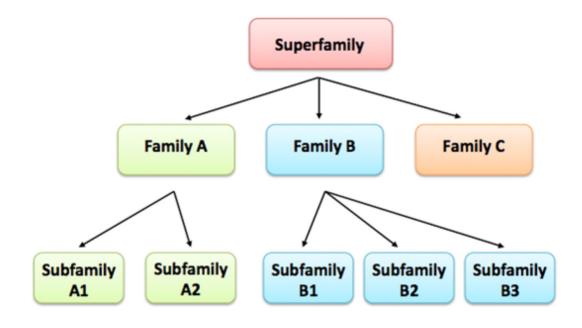


Figure 2 A hypothetical protein family hierarchy showing the relationships between superfamily, family and subfamily members. Directional arrows indicate that one group is a subgroup of another.

Protein domains

Domains

Domains are distinct functional and/or structural units in a protein.

Usually they are responsible for a particular function or interaction, contributing to the overall role of a protein.

Domains may exist in a variety of biological contexts, where similar domains can be found in proteins with different functions

Protein familly/subfamilly can often be defined by an specific domain arrangement

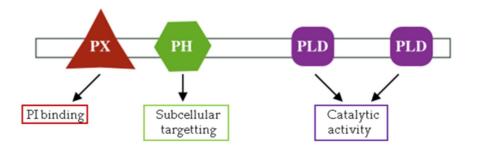
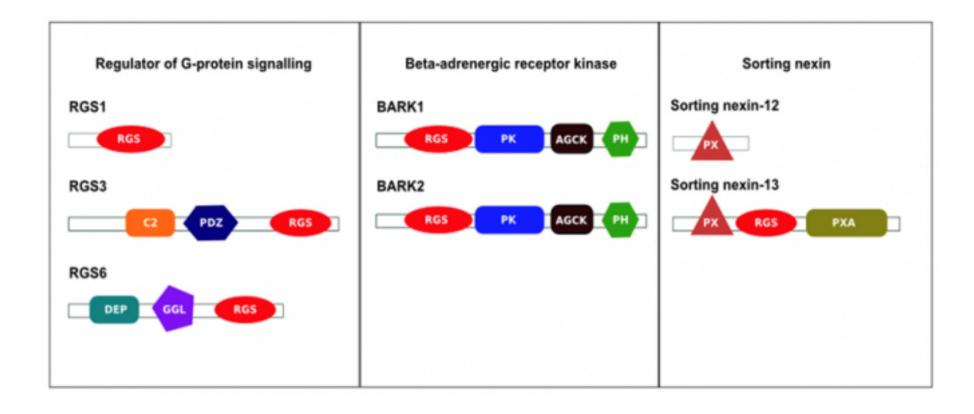


Figure 6 Domain composition of phospholipase D1, which is an enzyme that breaks down phosphatidylcholine.

Protein domains

Family- and domain-based protein classification

Family- and domain-based classifications are not always straightforward and can overlap, since proteins are sometimes assigned to families by virtue of the domain(s) they contain.

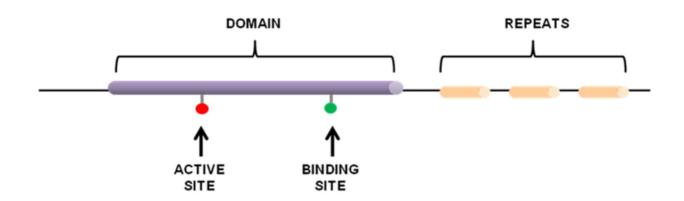


Protein sequence features

Sequence features are groups of amino acids that confer certain characteristics upon a protein

- active sites.
- binding sites
- post-translational modification (PTM) sites.
- repeats.

Sequence features differ from domains in that they are usually quite small (often only a few amino acids long), whereas domains represent entire structural or functional units of the protein



Protein domains and features have signatures

To classify proteins into families and to predict the presence of important domains or sequence features, we need predictive models known as *protein signatures*.

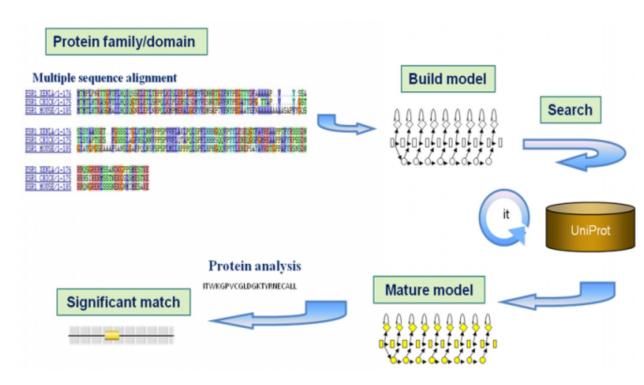
There are different types of signatures, built using different computational approaches.

A common starting point is a multiple sequence alignment of proteins sharing a set of characteristics (e.g. belonging to the same family or sharing a domain).

1/ When building the initial model, the level of amino acid conservation at different positions in the alignment is taken into account.

2/ The model is then used to search a protein database in an iterative manner, refining the model as more distantly related sequences in the database are identified.

3/Once the model is mature, the signature is ready and can be used for protein sequence analysis.

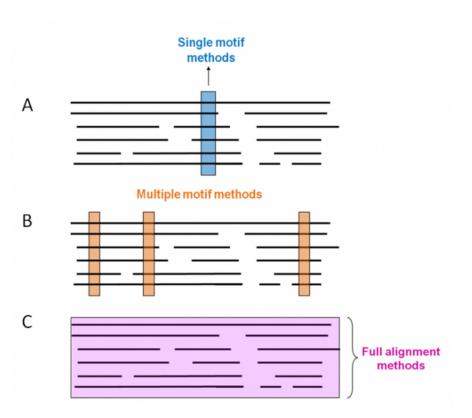


Protein signature types

Different approaches can be used to generate signatures. These include:

- •patterns
- •profiles
- •fingerprints
- hidden Markov models (HMMs)

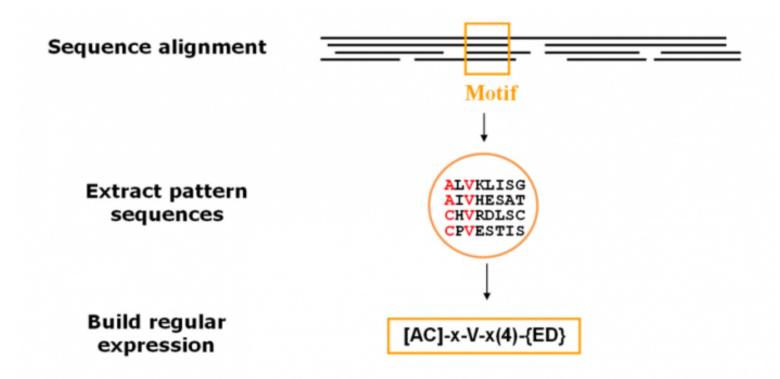
Each approach starts with a protein multiple sequence alignment, and can focus on a **single** conserved sequence region (known as a motif), **multiple** conserved motifs, or the **full** alignment of the entire protein or a particular domain .



Protein patterns

Many important sequence features, such as binding sites or the active sites of enzymes, consist of only a few amino acids that are essential for protein function.

The pattern of conservation within the sequence feature is then modelled as a **regular expression**. An example of a database that uses patterns is **PROSITE**.



Protein profiles

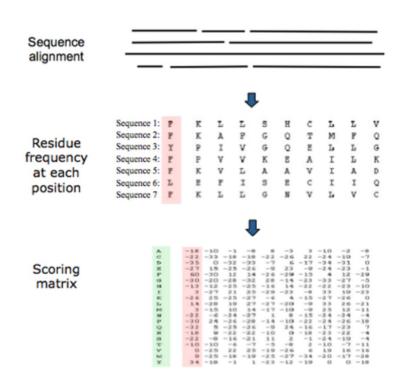
Profiles are used to model protein families and domains.

They are built by converting multiple sequence alignments into **position-specific scoring systems** (PSSMs).

Amino acids at each position in the alignment are scored according to the frequency with which they occur.

Examples of databases

- CDD
- HAMAP
- PROSITE (which produces profiles as well as patterns)
- **The PRODOM** database also uses a related approach, using PSI-BLAST to create its profiles.



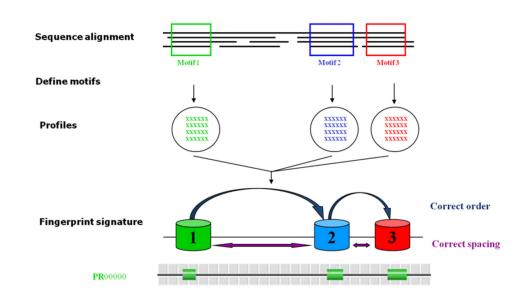
Protein fingerprints

Most protein families are characterised not by one, but by several conserved regions, which occur in a certain order.

Fingerprints are composed of multiple short conserved motifs. Each motif is converted into an individual profile (as described in the previous slide) to create a fingerprint signature.

Fingerprints can distinguish individual subfamillies

Exemple of database: PRINTS



Hidden Markov Models

Hidden Markov models (HMMs) can be used to convert multiple sequence alignments into position-specific scoring systems.

HMMs are adept at representing amino acid insertions and deletions, meaning that they can model entire alignments, including divergent regions.

Databases:

- Pfam
- SMART
- TIGRFAM
- PIRSF
- PANTHER
- SFLD
- Superfamily
- Gene3D

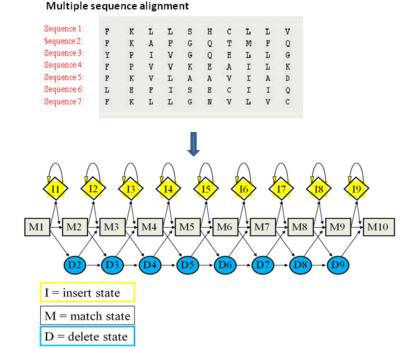


Figure 14 Representation of a Hidden Markov model based on a multiple sequence alignment. Amino acids are given a score at each position in the sequence alignment according to the frequency with which they occur. Transition probabilities (i.e., the likelihood that one particular amino acid follows another particular amino acid) and insertion and deletion states are also modelled.

InterPro is an integrative database

In InterPro, patterns, profiles, fingerprints and HMMs from a number of different databases are brought together into a single searchable resource, offering convenient access to their predictive capabilities without the need to visit the member databases individually.

By combining the different databases and signature types, InterPro capitalises on their individual strengths, producing a powerful tool for the prediction of protein function.

InterPro aims to simplify and rationalise protein sequence analysis for the user by combining and organising information in a consistent manner, removing redundancy, and adding extensive annotation including GO terms and useful links about the signatures and the proteins they match.

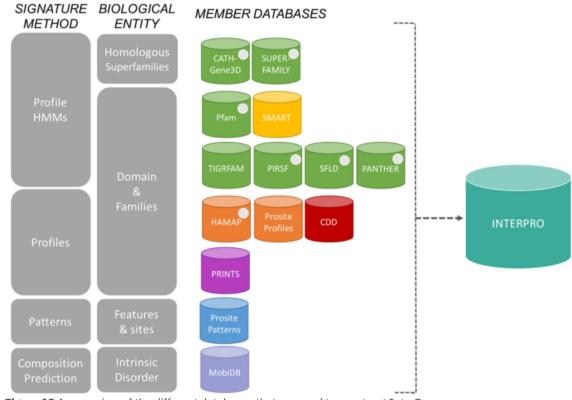


Figure 18 An overview of the different databases that are used to construct InterPro.

InterProSan can scan a query sequence against all the signatures aggregated in InterPro

InterPro Entry types

D Domain

Domains are distinct functional, structural or sequence units that may exist in a variety of biological contexts. A match to an InterPro entry of this type indicates the presence of a domain. Common examples of protein domains are the PH domain, Immunoglobulin domain or the classical C2H2 zinc finger.

Family

A protein family is a group of proteins that share a common evolutionary origin reflected by their related functions, similarities in sequence, or similar primary, secondary or tertiary structure. A match to an InterPro entry of this type indicates membership of a protein family.

Ⅲ Homologous Superfamily

A homologous superfamily is a group of proteins that share a common evolutionary origin, reflected by similarity in their structure. Since superfamily members often display very low similarity at the sequence level, this type of InterPro entry is usually based on a collection of underlying hidden Markov models, rather than a single signature. Homologous superfamilies usually comprise signatures from the SUPERFAMILY and CATH-Gene3D databases.

R Repeat

A short sequence that is typically repeated within a protein. Repeats are often relatively short <50 amino acids in length. Common repeats examples are Leucine Rich Repeats or WD40 repeats.

S Site

InterPro contains data for the following types of sites:

- Active site A short sequence that contains one or more conserved residues, which allow the protein to bind to a ligand and carry out a catalytic activity.
- Binding site A short sequence that contains one or more conserved residues, which form a protein interaction site.
- Conserved site A short sequence that contains one or more conserved residues.
- PTM site A short sequence that contains one or more conserved residues some of which are the site of a Post-translational modification.

U Unintegrated

InterPro 90.0 • 4th August 2022

Contents and coverage

InterPro protein matches are now calculated for all UniProtKB and UniParc proteins. InterPro release 90.0 contains 40 597 entries (last entry: IPR046937), representing:

Н	Homologous Superfamily	3399
F	Family	23 997
D	Domain	11 954
R	Repeat	327
S	Site	
	Active Site	132
	: Binding Site	75
	Conserved Site	696
	i PTM	17

Interpro cites 53778 publications in PubMed.

InterPro Entries are linked to GO terms

InterPro2GO

A total number of 35 086 GO terms mapped to InterPro entries. These are available through the EBI GO browser QuickGO

The assignment of GO terms to InterPro entries is performed manually, and is an ongoing process

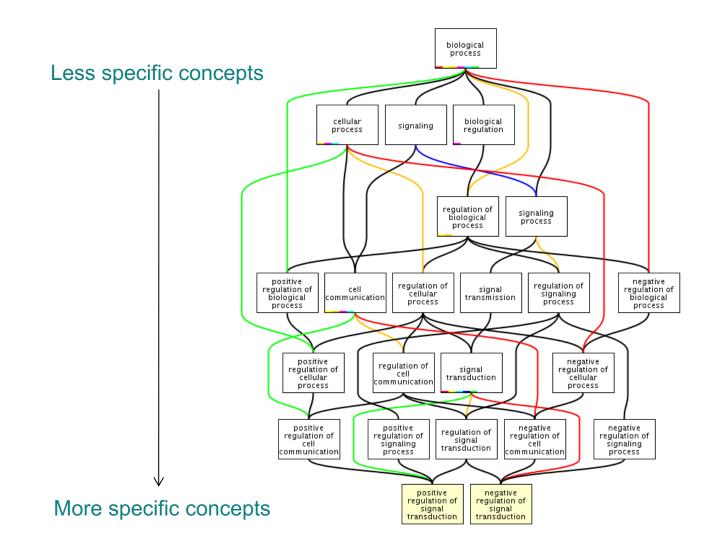
GO annotation in InterPro: why stability does not indicate accuracy in a sea of changing annotations

Amaia Sangrador-Vegas^{1,*}, Alex L. Mitchell¹, Hsin-Yu Chang¹, Siew-Yit Yong¹ and Robert D. Finn¹

The Gene Ontology

 A way to capture biological knowledge for individual gene products in a written and computable form

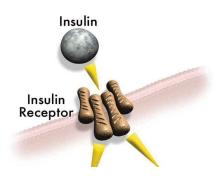
 A set of concepts and their relationships to each other arranged as a hierarchy



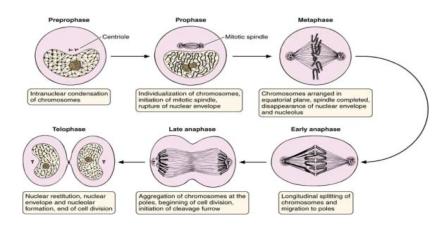
The domains in GO

1. Molecular Function

An elemental activity or task or job



- protein kinase activity
- insulin receptor activity



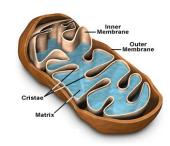
2. Biological Process

A commonly recognised series of events

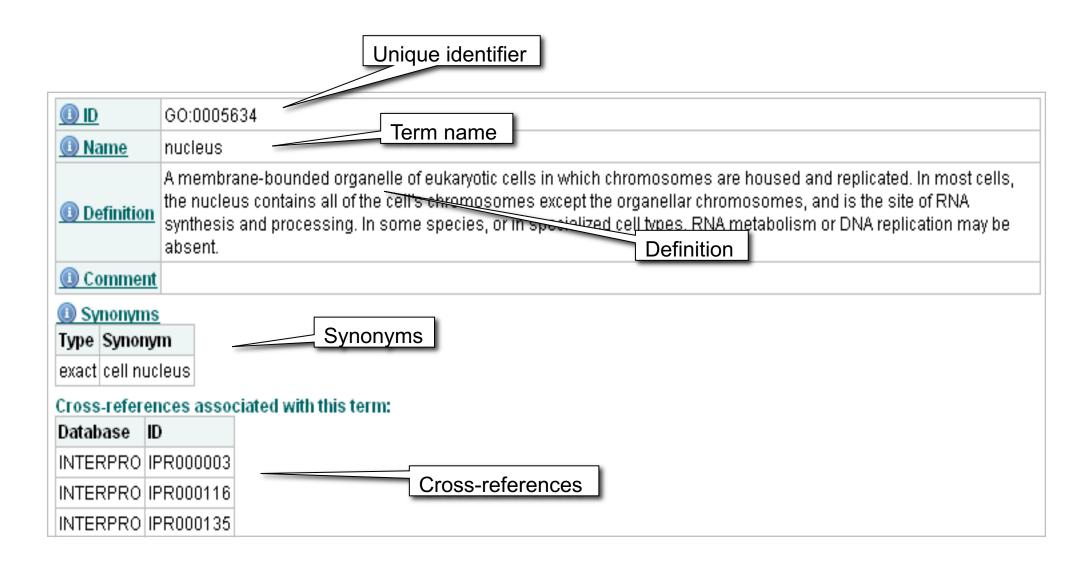
cell division

3. Cellular Component

Where a gene product is located



Anatomy of a GO term



GO evidence codes

Evidence codes fall into six general categories:

- •experimental evidence
- phylogenetic evidence
- •computational evidence
- author statements
- curatorial statements
- automatically generated annotations

- Inferred from Experiment (EXP)
- Inferred from Direct Assay (IDA)
- Inferred from Physical Interaction (IPI)
- Inferred from Mutant Phenotype (IMP)
- Inferred from Genetic Interaction (IGI)
- Inferred from Expression Pattern (IEP)
- Inferred from High Throughput Experiment (HTP)
- Inferred from High Throughput Direct Assay (HDA)
- Inferred from High Throughput Mutant Phenotype (HMP)
- Inferred from High Throughput Genetic Interaction (HGI)
- Inferred from High Throughput Expression Pattern (HEP)

What is Gene Ontology?

Statistics for release 2022-09 V

Ontology	
Property	Value
Valid terms	43335 (Δ = -223)
Obsoleted terms	$4008 \ (\Delta = 238)$
Merged terms	2431 (Δ = 44)
Biological process terms	28056
Molecular function terms	11242

4037

Cellular component terms

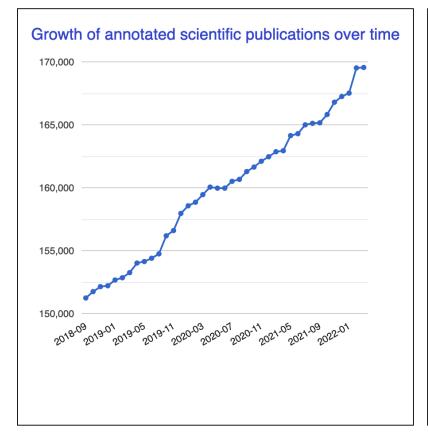
Annotations

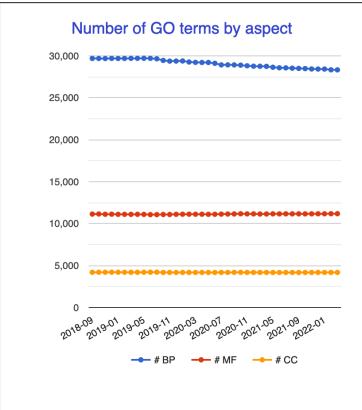
Property	Value
Number of annotations	7,493,159
Annotations for biological process	2,831,298
Annotations for molecular function	2,362,896
Annotations for cellular component	2,298,965
Annotations for evidence PHYLO	3,993,128
Annotations for evidence IEA	1,385,272
Annotations for evidence OTHER	869,051
Annotations for evidence EXP	934,536
Annotations for evidence ND	252,132
Annotations for evidence HTP	59,040
Number of annotated scientific publications	172,379

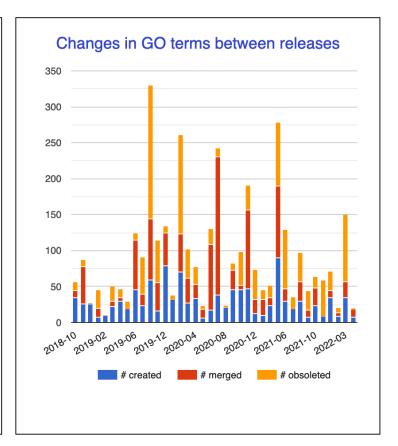
Gene products and species

Property	Value
Annotated gene products	1,483,687
Annotated species	5,257
Annotated species with over 1,000 annotations	185

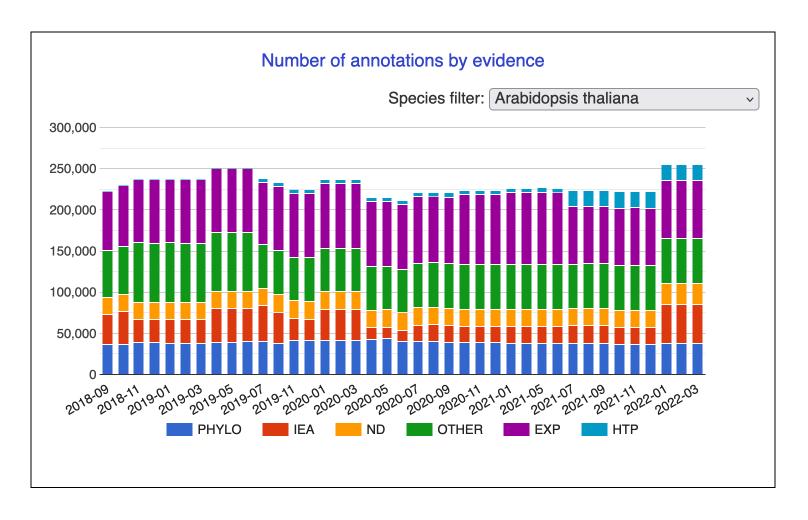
What is Gene Ontology?







What is Gene Ontology?



Inferred from Experiment (EXP)

Inferred from Experiment (EXP)

Electronic (IEA) annotation are not manually reviewed

No biological Data available (ND)

What can we do with Gene Ontology?

Over-representation analysis (ORA) in a nutshell

- Given:
 - Gene list: e.g. RRP6, MRD1, RRP7, RRP43, RRP42 (yeast)
 - Gene annotations: e.g. Gene ontology, transcription factor binding sites in promoter
- ORA Question: Are any of the gene annotations surprisingly enriched in the gene list?
- Details:
 - How to assess "surprisingly" (statistics)
 - How to correct for repeating the tests

What can we do with Gene Ontology?

TopGO:

Some GO-endorsed enrichment tools are:

- BiNGO
- GeneWeaver
- gProfiler
- GOrilla
- Ontologizer

ORIGINAL PAPER Vol. 22 no. 13 2006, pages 1600–1607

Gene expression

Improved scoring of functional groups from gene expression data by decorrelating GO graph structure

Adrian Alexa*, Jörg Rahnenführer and Thomas Lengauer Max-Planck-Institute for Informatics, Stuhlsatzenhausweg 85, D-66123 Saarbrücken. Germany Received on September 28, 2005; revised on March 30, 2006; accepted on April 4, 2006 Advance Access publication April 10, 2006

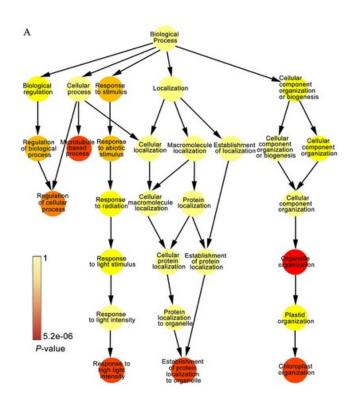
Results:

Associate Editor: Martin Bishop

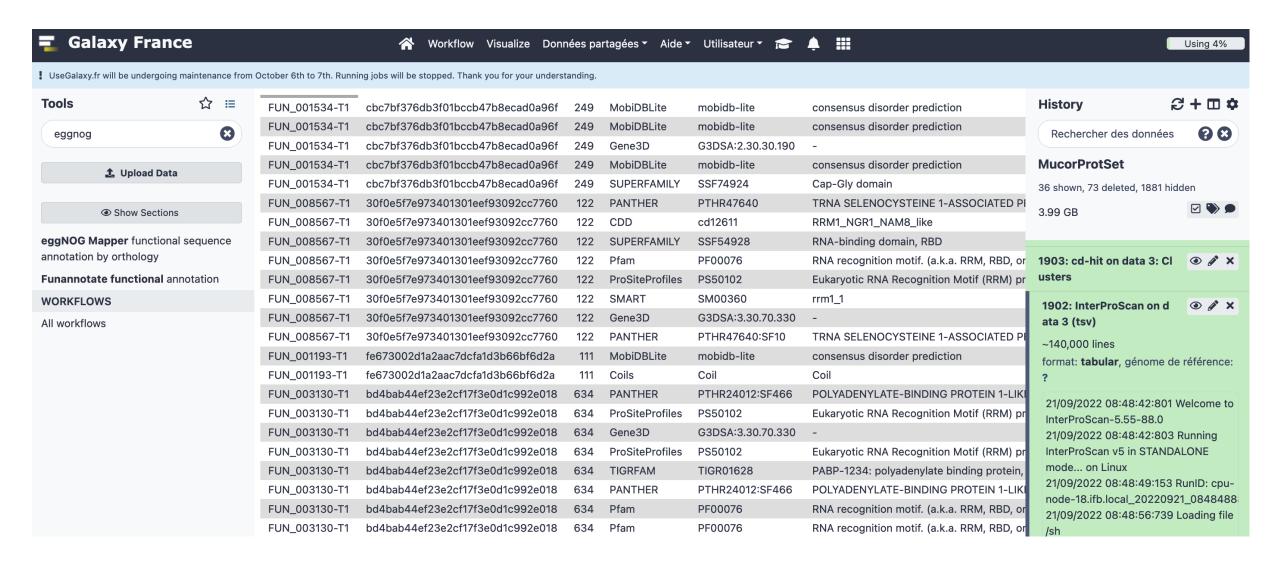
We present two novel algorithms that improve GO group scoring using the underlying GO graph topology.

We show that both methods eliminate local dependencies between GO terms and point to relevant areas in the GO graph that remain undetected with state-of-the-art algorithms for scoring functional terms.

A simulation study demonstrates that the new methods exhibit a higher level of detecting relevant biological terms than competing methods



Let's have a look at InterProScan results

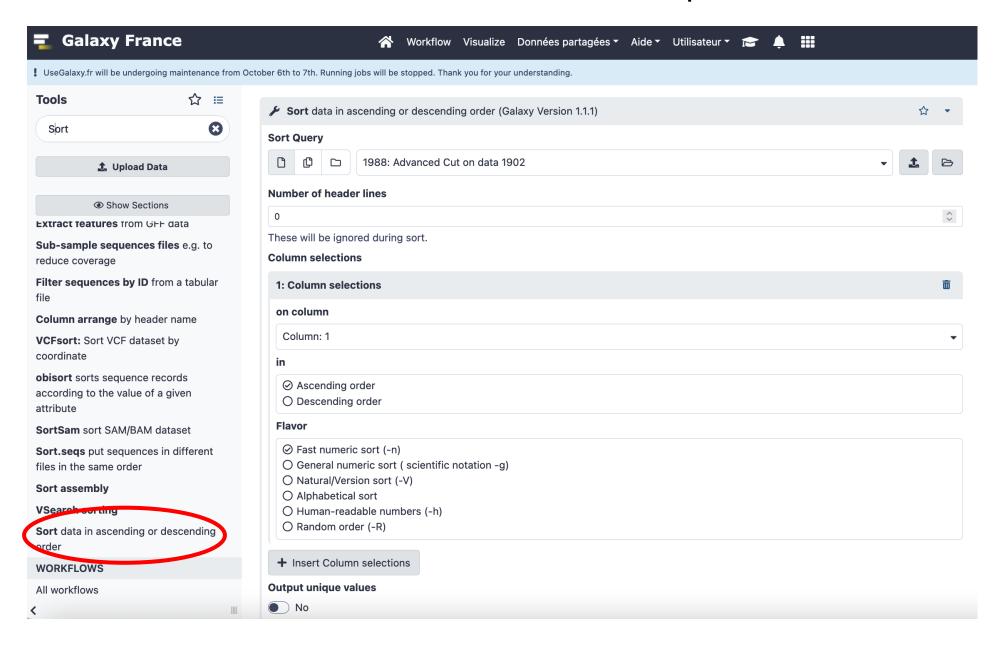


Let's have a look at InterProScan results

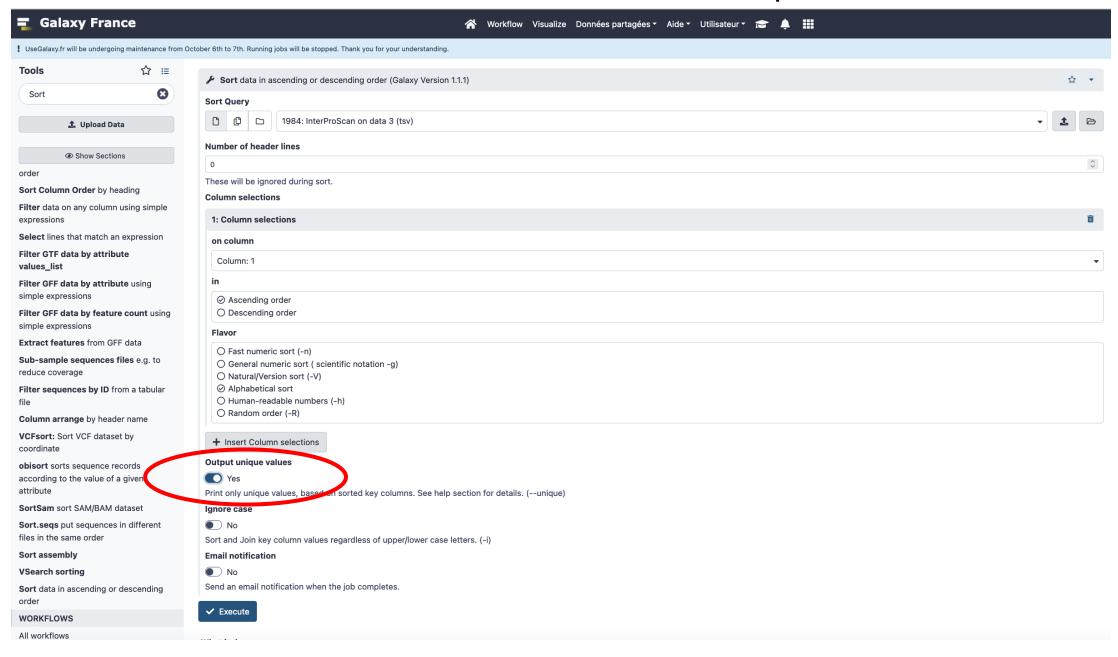
The TSV format presents the match data in columns as follows:

- Protein Accession (e.g. P51587)
- •Sequence MD5 digest (e.g. 14086411a2cdf1c4cba63020e1622579)
- •Sequence Length (e.g. 3418)
- Analysis (e.g. Pfam / PRINTS / Gene3D)
- •Signature Accession (e.g. PF09103 / G3DSA:2.40.50.140)
- Signature Description (e.g. BRCA2 repeat profile)
- Start location
- Stop location
- •Score is the e-value of the match reported by member database method (e.g. 3.1E-52)
- •Status is the status of the match (T: true)
- •Date is the date of the run
- •(InterProScan annotations accession (e.g. IPR002093) optional column; only displayed if -iprscan option is switched on)
- •(InterProScan annotations description (e.g. BRCA2 repeat) optional column; only displayed if -iprscan option is switched on)
- •(GO annotations (e.g. GO:0005515) optional column; only displayed if --goterms option is switched on)
- •(Pathways annotations (e.g. REACT_71) optional column; only displayed if --pathways option is switched on)

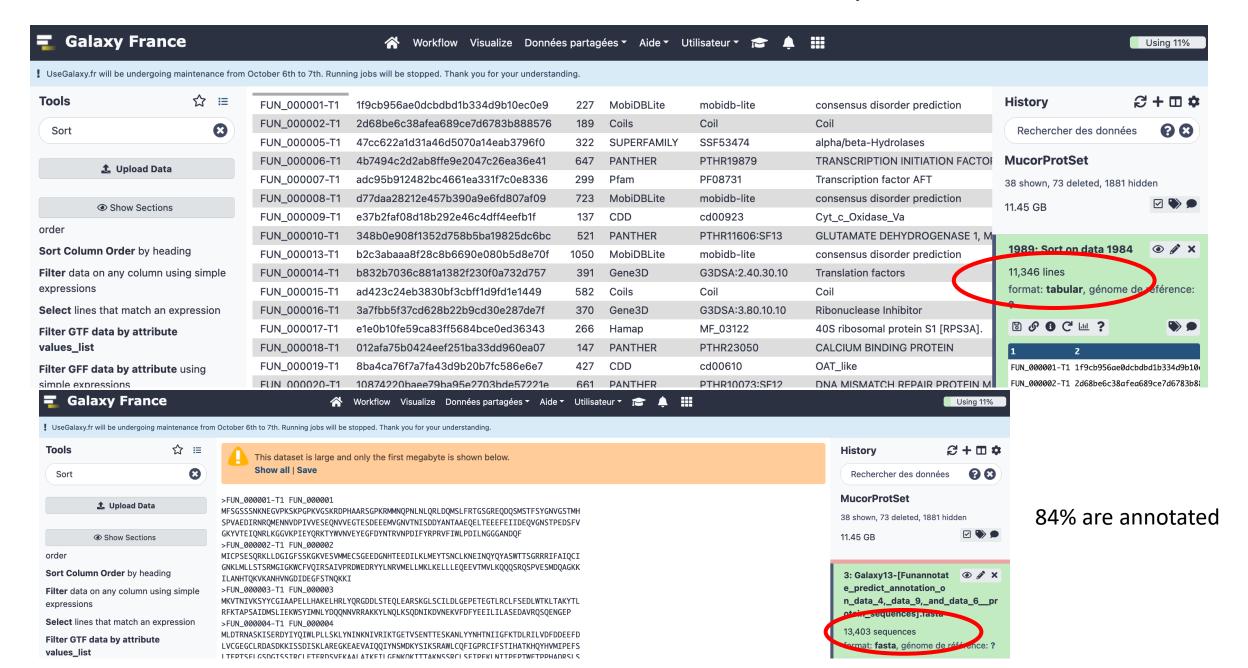
What's the fraction of annotated sequences?



What's the fraction of annotated sequences?



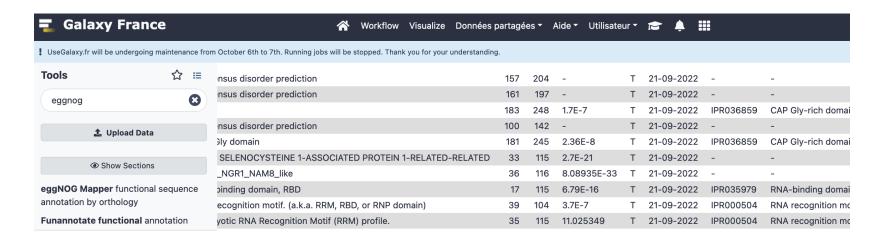
What's the fraction of annotated sequences?



How reliable is an InterProScan Hit?

Why are there no e-values associated with InterPro entries?

The signatures contained within InterPro are produced in different ways by different member databases, so their e-values and/or scoring systems cannot be meaningfully compared or combined. For this reason, we do not show e-values on the InterPro web site. However, e-values can be obtained via the downloadable InterProScan software package, which outputs detailed individual results for each member database sequence analysis algorithm.



The E-value (expectation value) is the number of hits that would be expected to have a score equal to or better than this value, by chance alone. This means that a good E-value which gives a confident prediction is much less than 1. E-values around 1 is what is expected by chance. Thus, the lower the E-value, the more specific the search for domains will be.

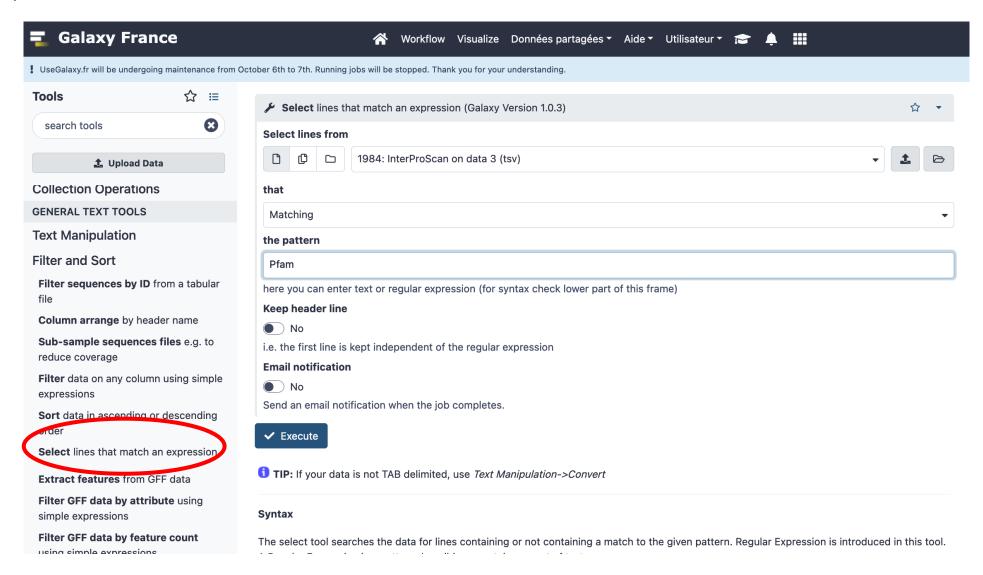
How reliable is an InterProScan Hit?

_				
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FUN_003130-T1	ProSiteProfiles	PS50102	16.807129	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1187 MetaCyc: PWY-2083 MetaCyc: PWY-3542 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4161 MetaCyc: PWY-
FUN_003130-T1	Gene3D	G3DSA:3.30.70.330	4.0E-25	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1187 MetaCyc: PWY-2083 MetaCyc: PWY-3542 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-
FUN_003130-T1	ProSiteProfiles	PS50102	18.137451	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1187 MetaCyc: PWY-2083 MetaCyc: PWY-3542 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-
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FUN_003130-T1	PANTHER	PTHR24012:SF466	2.4E-205	
FUN_003130-T1	Pfam	PF00076	2.0E-22	- MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1187 MetaCyc: PWY-2083 MetaCyc: PWY-3542 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-4061 MetaCyc: PWY-4002 MetaCyc: PW
FUN_003130-T1	Pfam	PF00076	1.1E-20	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1187 MetaCyc: PWY-2083 MetaCyc: PWY-3542 MetaCyc: PWY-402 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5
FUN_003130-T1	Pfam	PF00076	1.8E-18	- MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1187 MetaCyc: PWY-2083 MetaCyc: PWY-3542 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5035 MetaCyc: PWY-5035 MetaCyc: PWY-5035 MetaCyc: PWY-5035 MetaCyc: PWY-5035 MetaCyc: PWY-4021 MetaCyc: PW
FUN_003130-T1	Pfam	PF00076	3.1E-20	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1061 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-406 MetaCyc: PWY-400 MetaCyc: PW
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FUN_003130-T1	CDD	cd12381	1.08824E-45	
FUN_003130-T1	Coils	Coil	-	
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FUN_003130-T1	CDD	cd12380	1.37304E-41	
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FUN_003130-T1	MobiDBLite	mobidb-lite	-	
FUN_003130-T1	MobiDBLite	mobidb-lite	_	
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FUN_003130-T1	ProSiteProfiles	PS50102	20.576372	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1061 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-
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FUN_003130-T1	MobiDBLite	mobidb-lite	-	
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FUN_003130-T1	SMART	SM00361	0.48	MetaCyc: PWY-7511 Reactome: R-CEL-72163 Reactome: R-DDI-156827 Reactome: R-DDI-975956 Reactome: R-DDI-975957 Reactome: R-DME-159236 Reactome: R-DME-6781823 Reactome: R-DME-6782135 Reactome: R-DME-67
FUN_003130-T1	SMART	SM00361	0.33	MetaCyc: PWY-7511 Reactome: R-CEL-72163 Reactome: R-DDI-156827 Reactome: R-DDI-975956 Reactome: R-DDI-975957 Reactome: R-DME-159236 Reactome: R-DME-6781823 Reactome: R-DME-6782135 Reactome: R-DME-67
FUN_003130-T1		SM00361	0.2	MetaCyc: PWY-7511 Reactome: R-CEL-72163 Reactome: R-DDI-156827 Reactome: R-DDI-975956 Reactome: R-DDI-975957 Reactome: R-DME-159236 Reactome: R-DME-6781823 Reactome: R-DME-6782135 Reactome: R-DME-67
FUN_003130-T1	SUPERFAMILY	SSF54928	2.45E-49	MetaCyc: PWY-102 MetaCyc: PWY-4161 MetaCyc: PWY-4161 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-4801 MetaCyc: PWY-402 MetaCyc: PWY-4801 MetaCyc: PWY-402 MetaCyc: PWY-4801 MetaCyc: PWY-402 MetaCyc: PWY-4801 MetaCyc
FUN_003130-T1		SM00517	1.1E-36	MetaCyc: PWY-7511 Reactome: R-DDI-156827 Reactome: R-DDI-975956 Reactome: R-DDI-975957 Reactome: R-DME-156827 Reactome: R-DME-429947 Reactome: R-DME-450408 Reactome: R-DME-75649 Reactome: R-DME-7564
FUN_003130-T1	SMART	SM00360	3.2E-22	MetaCvc: PWY-102lMetaCvc: PWY-106lMetaCvc: PWY-1187lMetaCvc: PWY-2083lMetaCvc: PWY-3542lMetaCvc: PWY-402lMetaCvc: PWY-4161lMetaCvc: PWY-4202lMetaCvc: PWY-5035lMetaCvc: PWY-5036lMetaCvc: PWY-5036lMetaCvc: PWY-402lMetaCvc: PWY-40
FUN_003130-T1	SMART	SM00360	2.1E-25	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-4181 MetaCyc: PWY-4020 MetaCyc: PWY-
FUN_003130-T1	SMART	SM00360	1.3E-27	MetaCyc: PWY-102 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-
FUN_003130-T1		SM00360	3.0E-28	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-1187 MetaCyc: PWY-2083 MetaCyc: PWY-3542 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5035 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-
FUN_003130-T1		PS50102	19.757713	MetaCyc: PWY-102 MetaCyc: PWY-1061 MetaCyc: PWY-4181 MetaCyc: PWY-5036 MetaCyc: PWY-5036 MetaCyc: PWY-4021 MetaCyc: PWY-4161 MetaCyc: PWY-4202 MetaCyc: PWY-5036 MetaCyc: PWY-
FUN_003130-T1		Coil	-	, , , , , , , , , , , , , , , , , , ,
FUN_003130-T1		PTHR24012	2.4E-205	
FUN_003130-T1		SSF81995	2.17E-6	
FUN_003130-T1		cd12378	3.79251E-50	
FUN_003130-T1		PS51309	21.821808	MetaCvc: PWY-7511 Reactome: R-DDI-156827 Reactome: R-DDI-975956 Reactome: R-DDI-975957 Reactome: R-DME-156827 Reactome: R-DME-450408 Reactome: R-DME-72649
FUN_003130-T1		PTHR24012	2.4E-205	measys
1 014_003130-11	PANTHER	FIRKZ4UIZ	Z.4E-ZU0	

How reliable is an InterProScan Hit?

Galaxy France	:	<page-header> Workflow Visualize Données</page-header>	partagé	es 🕶	Aide 🔻 Utilisateા	ır 🕶	*	!!!	
UseGalaxy.fr will be undergoing maintenance from October 6th to 7th. Running jobs will be stopped. Thank you for your understanding.									
Tools	☆ ≔	nsus disorder prediction	157	204	-	Т	21-09-2022	-	-
eggnog	8	nsus disorder prediction	161	197	-	Т	21-09-2022	-	-
0991109			183	248	1.7E-7	Т	21-09-2022	IPR036859	CAP Gly-rich don
1 Upload Data		nsus disorder prediction	100	142	-	Т	21-09-2022	-	-
es opioda sata		3ly domain	181	245	2.36E-8	Т	21-09-2022	IPR036859	CAP Gly-rich don
Show Sections		SELENOCYSTEINE 1-ASSOCIATED PROTEIN 1-RELATED-RELATED	33	115	2.7E-21	Т	21-09-2022	-	-
Sillow Sections		_NGR1_NAM8_like	36	116	8.08935E-33	Т	21-09-2022	-	-
eggNOG Mapper functional se	equence	oinding domain, RBD	17	115	6.79E-16	Т	21-09-2022	IPR035979	RNA-binding dor
annotation by orthology		ecognition motif. (a.k.a. RRM, RBD, or RNP domain)	39	104	3.7E-7	Т	21-09-2022	IPR000504	RNA recognition
Funannotate functional anno	tation	yotic RNA Recognition Motif (RRM) profile.	35	115	11.025349	Т	21-09-2022	IPR000504	RNA recognition
WORKFLOWS		1	36	109	2.3E-6	Т	21-09-2022	IPR000504	RNA recognition
All workflows			34	114	9.7E-18	Т	21-09-2022	IPR012677	Nucleotide-bindi
		SELENOCYSTEINE 1-ASSOCIATED PROTEIN 1-RELATED	33	115	2.7E-21	Т	21-09-2022	-	-
		nsus disorder prediction	73	111	-	Т	21-09-2022	-	-
			7	27	-	Т	21-09-2022	-	-
		ADENYLATE-BINDING PROTEIN 1-LIKE	49	494	2.4E-205	Т	21-09-2022	-	-
		yotic RNA Recognition Motif (RRM) profile.	52	130	16.807129	Т	21-09-2022	IPR000504	RNA recognition
			129	221	4.0E-25	Т	21-09-2022	IPR012677	Nucleotide-bindi
		yotic RNA Recognition Motif (RRM) profile.	140	217	18.137451	Т	21-09-2022	IPR000504	RNA recognition
		-1234: polyadenylate binding protein, human types 1, 2, 3, 4 family	52	625	6.1E-222	Т	21-09-2022	IPR006515	Polyadenylate bir
		ADENYLATE-BINDING PROTEIN 1-LIKE	546	628	2.4E-205	Т	21-09-2022	-	-
		ecognition motif. (a.k.a. RRM, RBD, or RNP domain)	235	303	2.0E-22	Т	21-09-2022	IPR000504	RNA recognition
		ecognition motif. (a.k.a. RRM, RBD, or RNP domain)	142	210	1.1E-20	Т	21-09-2022	IPR000504	RNA recognition
		ecognition motif. (a.k.a. RRM, RBD, or RNP domain)	54	123	1.8E-18	Т	21-09-2022	IPR000504	RNA recognition
		ecognition motif. (a.k.a. RRM, RBD, or RNP domain)	338	407	3.1E-20	Т	21-09-2022	IPR000504	RNA recognition
			222	320	4.2E-29	Т	21-09-2022	IPR012677	Nucleotide-bindi
			321	431	19F-30	Т	21-09-2022	IPR012677	Nucleotide-hindi

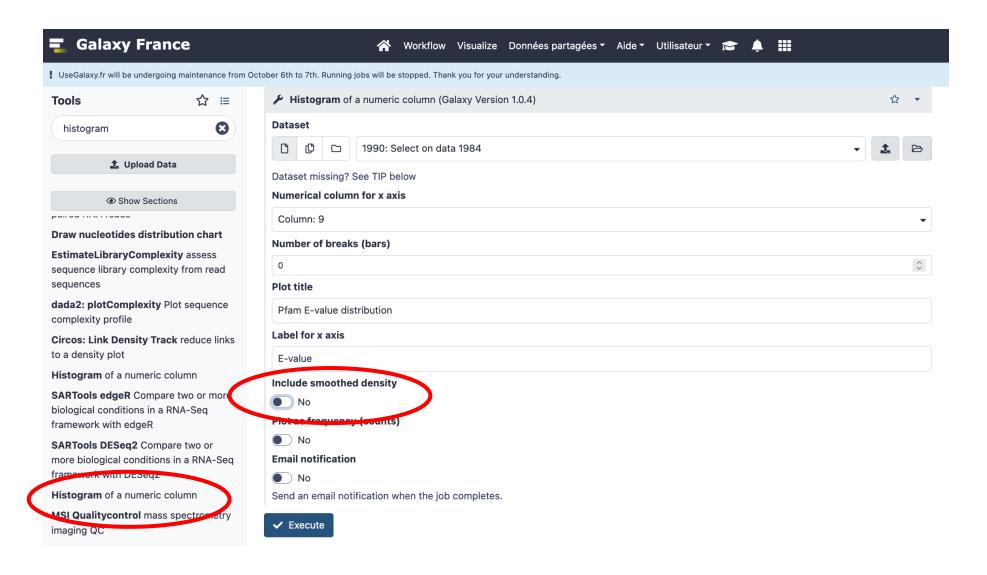
1/ select Pfam hit lines

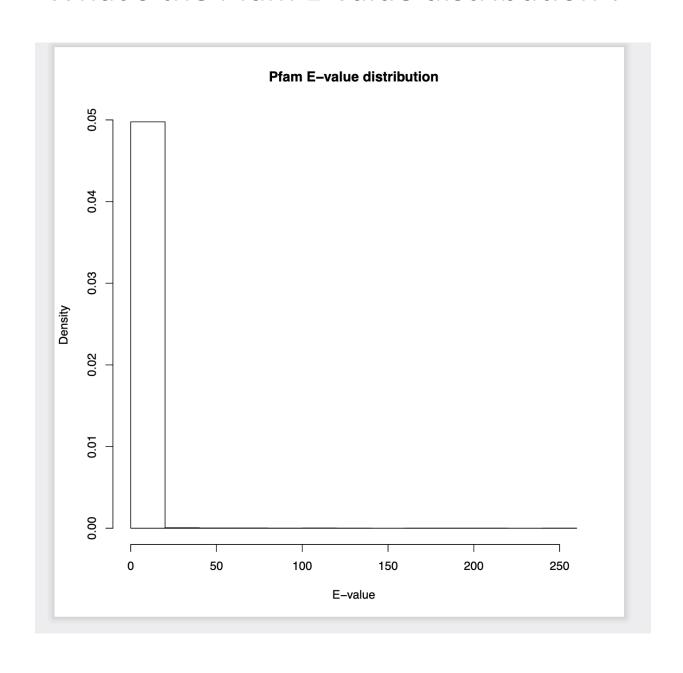


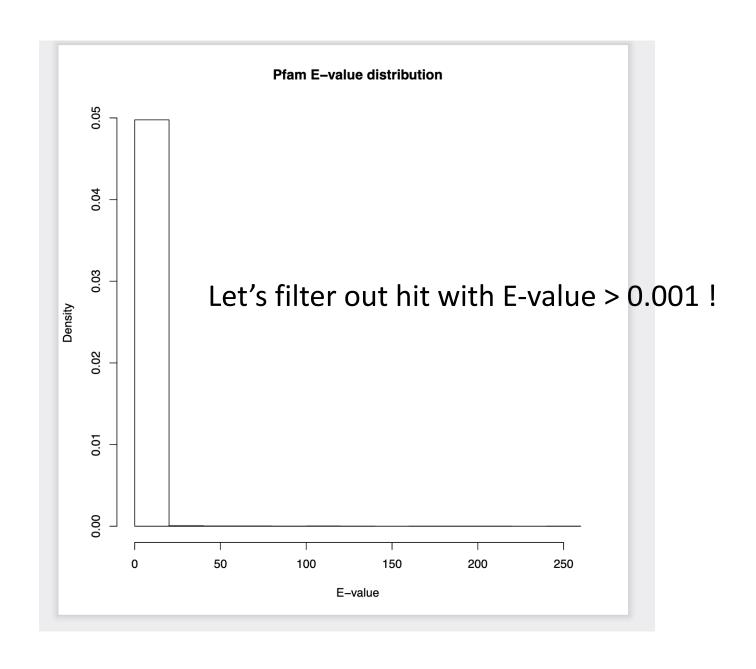
1/ select Pfam hit lines

Galaxy France		☆ Workflow Visualize Dor	nées partagé	ées ▼ Ai	ide ▼ Util	isateur ▼ 🞓 💄 🎹
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Tools ☆	≡ FUN_008567-T1	30f0e5f7e973401301eef93092cc7760	122	Pfam P	PF00076	RNA recognition motif. (a.k.a. RRM, RBD, or RNP domain)
search tools	FUN_003130-T1 FUN_003130-T1	bd4bab44ef23e2cf17f3e0d1c992e018 bd4bab44ef23e2cf17f3e0d1c992e018			PF00076	RNA recognition motif. (a.k.a. RRM, RBD, or RNP domain) RNA recognition motif. (a.k.a. RRM, RBD, or RNP domain)
1 Upload Data	FUN_003130-T1	bd4bab44ef23e2cf17f3e0d1c992e018	634	Pfam P	PF00076	RNA recognition motif. (a.k.a. RRM, RBD, or RNP domain)
Collection Operations	FUN_003130-T1 FUN_003130-T1	bd4bab44ef23e2cf17f3e0d1c992e018 bd4bab44ef23e2cf17f3e0d1c992e018			PF00076 PF00658	RNA recognition motif. (a.k.a. RRM, RBD, or RNP domain) Poly-adenylate binding protein, unique domain
GENERAL TEXT TOOLS	FUN_003678-T1	bd8e2066e65d046fe39d3ea94f2543	ac 603	Pfam P	PF02779	Transketolase, pyrimidine binding domain
Text Manipulation	FUN_003678-T1 FUN_003678-T1	bd8e2066e65d046fe39d3ea94f2543abd8e2066e65d046fe39d3ea94fe2646fe39d4ea94fe2646fe39d4ea94fe2646fe39d4ea94fe2646fe39d4ea94fe2646fe39d4ea946fe39d4ea94fe2646fe39d6ea94fe2646fe39d6ea94fe2646fe39d6ea94fe2666fe466fe466fe466fe466fe466fe466fe4			PF16870 PF00676	2-oxoglutarate dehydrogenase C-terminal Dehydrogenase E1 component
Filter and Sort	FUN_006750-T1	1c2b18a6e6d05afaf94e5d0b32a30128			PF01853	MOZ/SAS family
Filter sequences by ID from a tabul file	FUN_006750-T1	1c2b18a6e6d05afaf94e5d0b32a30128			PF17772 PF00628	MYST family zinc finger domain PHD-finger
Column arrange by header name	FUN_002211-T1	353fc52143176928bca800dc491d4e8			PF10516	SHNi-TPR
Sub-sample sequences files e.g. to		5d490c0e068c14797213b1e58c84703			PF13193	AMP-binding enzyme C-terminal domain
reduce coverage Filter data on any column using sim	FUN_002293-T1 ple FUN_002293-T1	5d490c0e068c14797213b1e58c84703 5d490c0e068c14797213b1e58c84703			PF16177 PF00501	Acetyl-coenzyme A synthetase N-terminus AMP-binding enzyme
expressions	FUN_005342-T1	2931a1f5db4fe7f7cb61223895be49c6	177	Pfam P	PF05255	Uncharacterised protein family (UPF0220)
Sort data in ascending or descending order	FUN_007314-T1 FUN_007400-T1	014faa6d38986eada2186be73a052c1 883d2aec020c728e5f15a6c339b9e5c			PF00704 PF03707	Glycosyl hydrolases family 18 Bacterial signalling protein N terminal repeat
A 1 - 12 - 11 - 1 - 1 - 1	ELINI 007/00_T1	992d2aaa020a729aEf1Ea6a220b0aEa	Q Q21	Dfam D	DEN27N7	Pactorial cignalling protain N terminal report

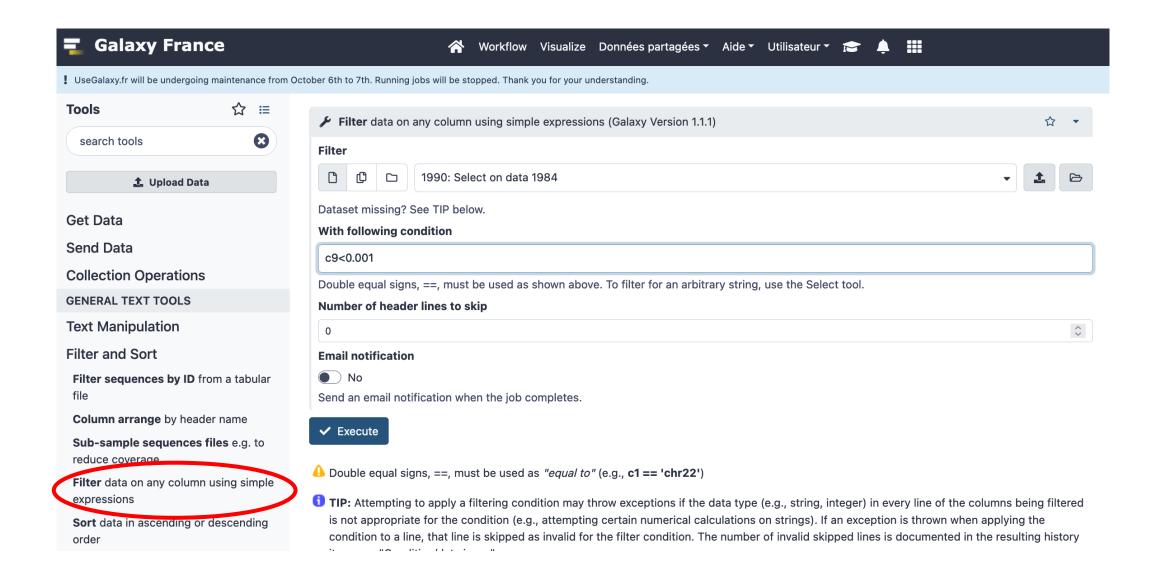
1/ Compute and draw a histogram

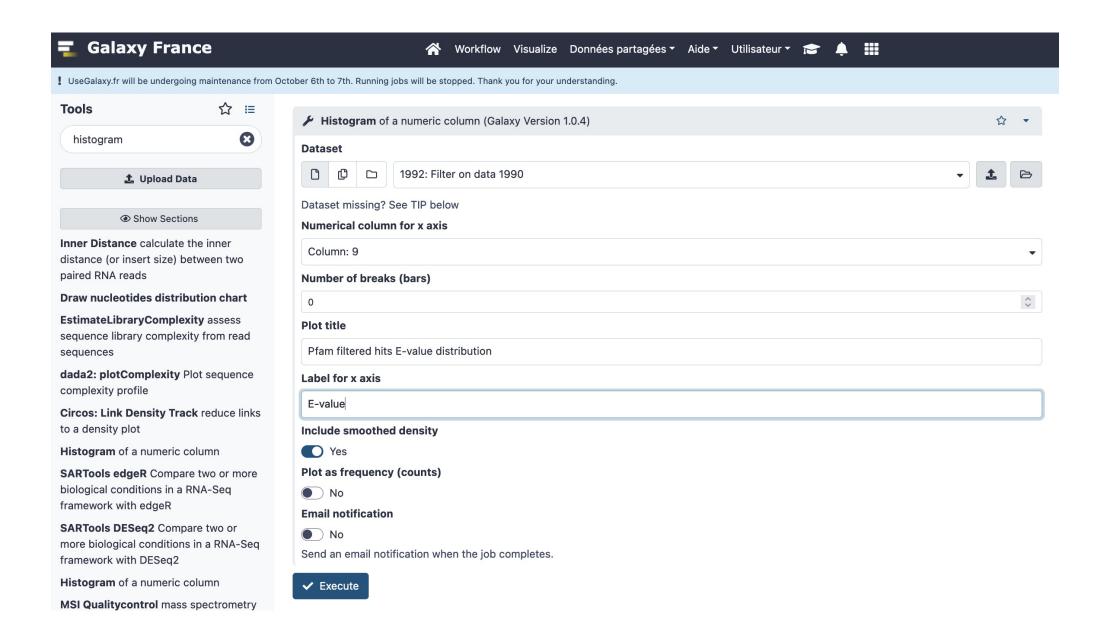


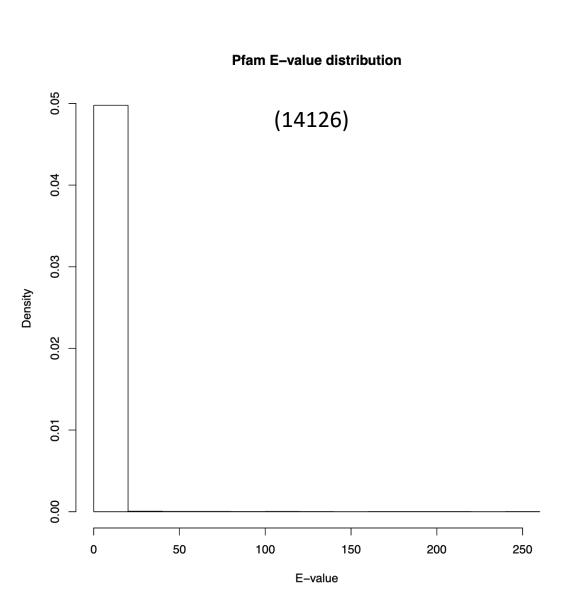


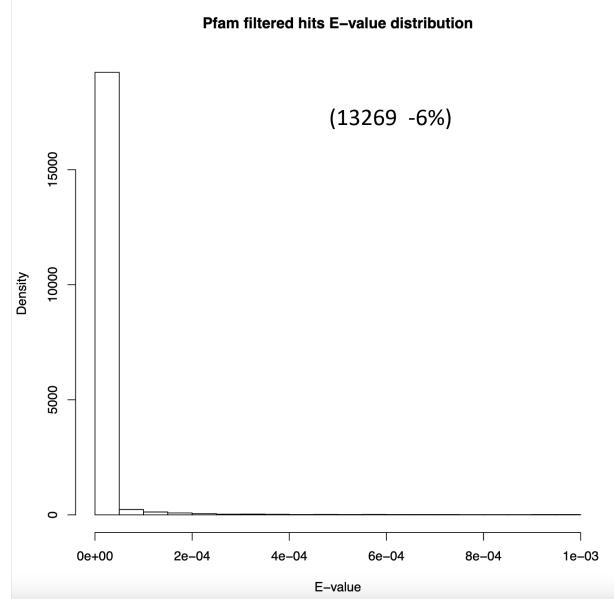


Let's filter out hit with E-value > 0.001!

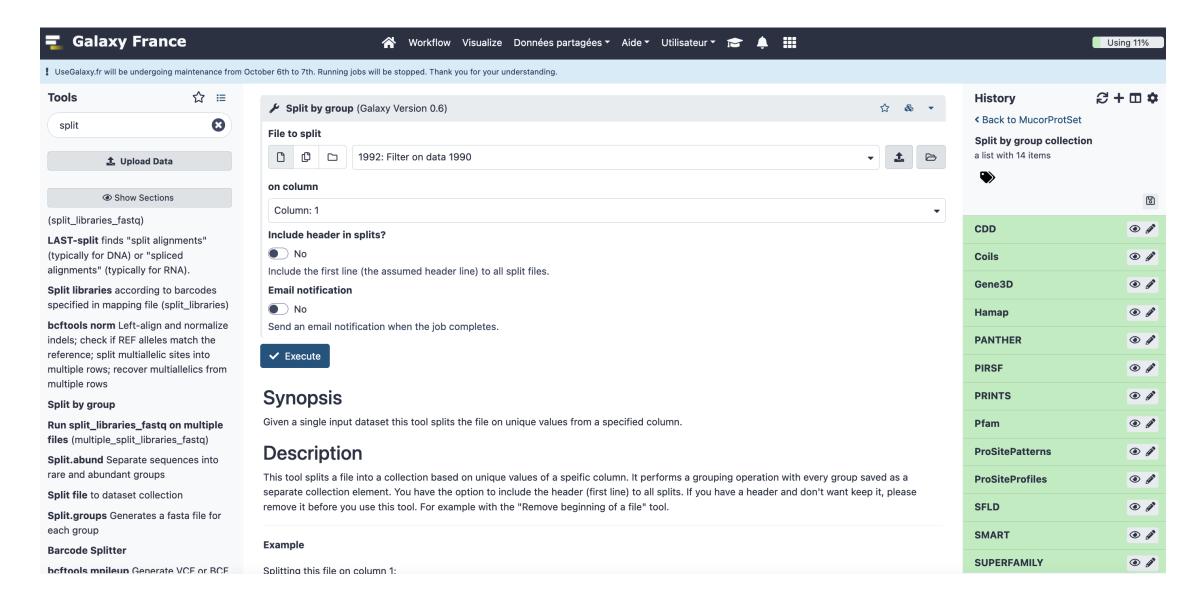


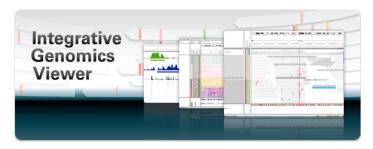




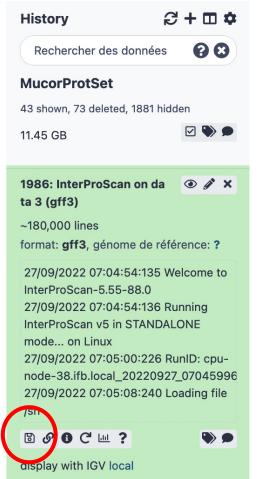


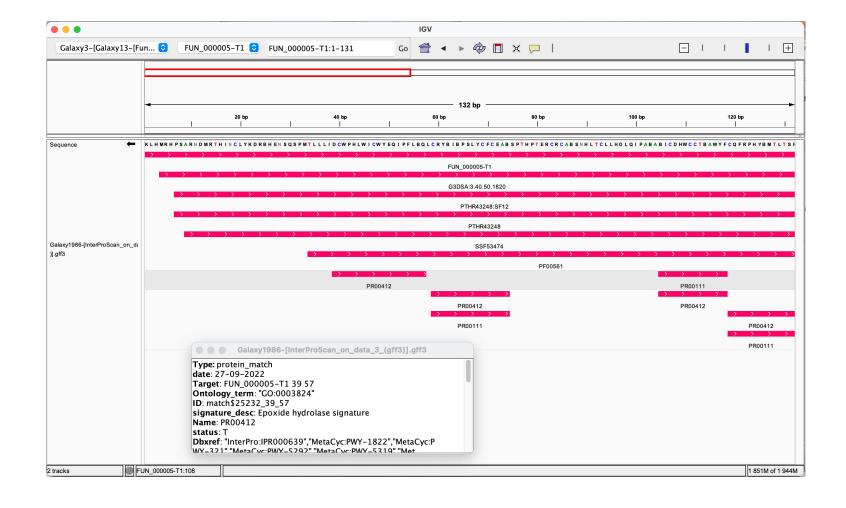
TIPS: the Split by group tool: Split hits into one file per database

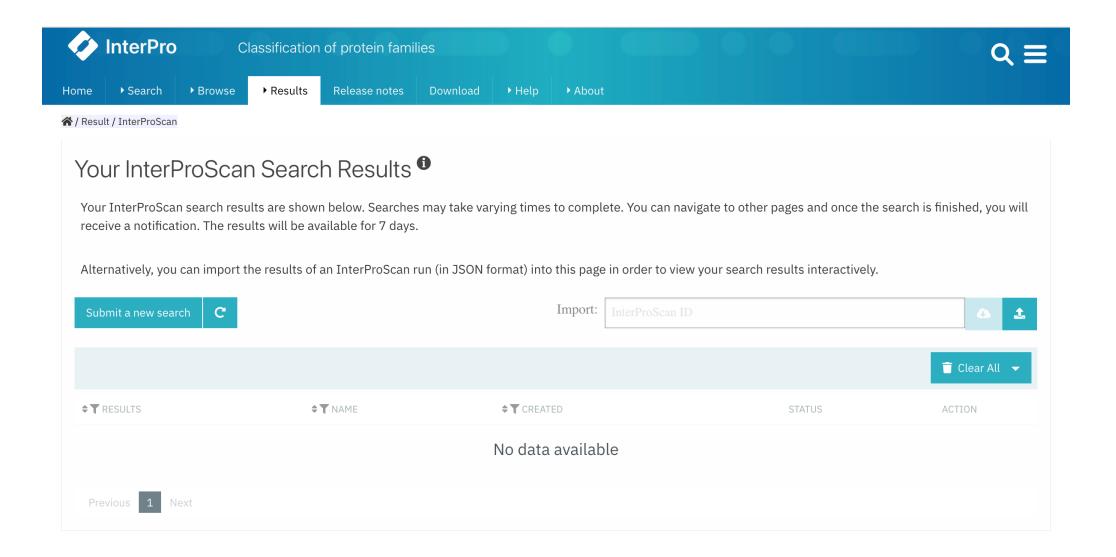


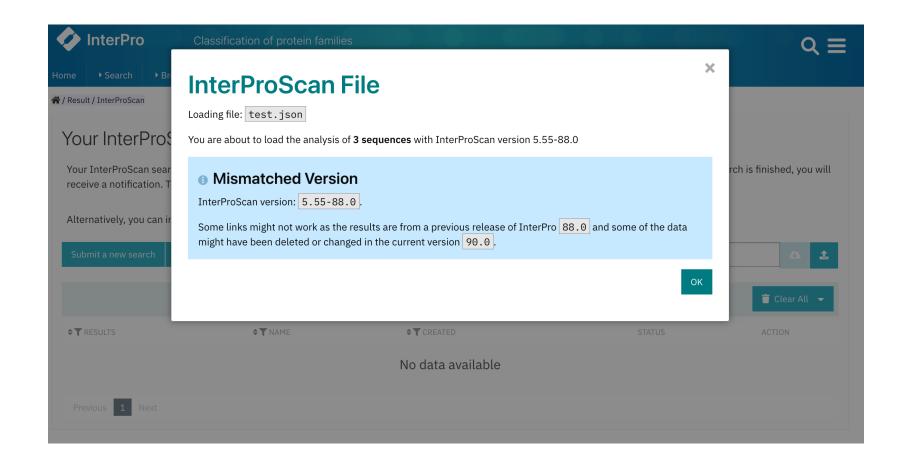


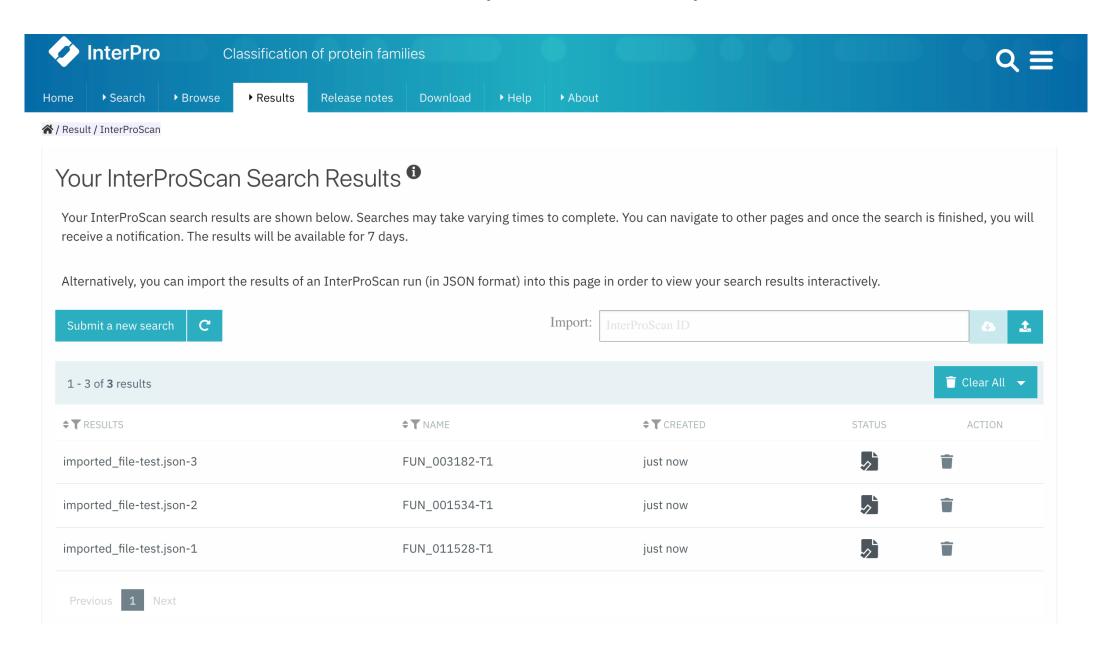
https://software.broadinstitute.org/software/igv/

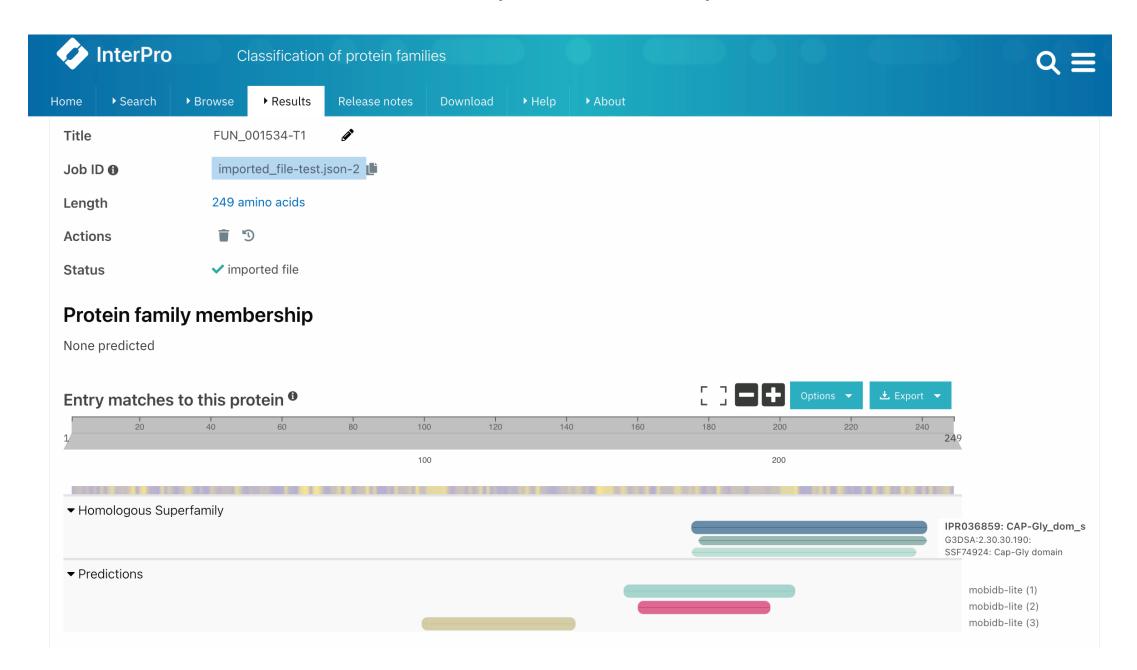


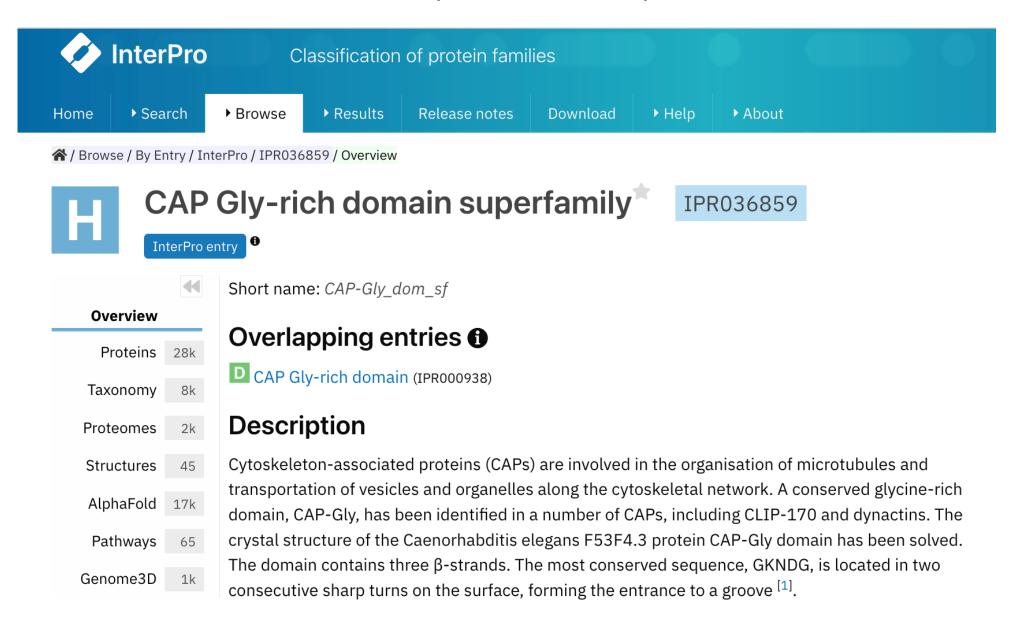












Functional annotation pipelines



F1000Research 2018, 7(ELIXIR):148 Last updated: 31 MAR 2022



OPINION ARTICLE

Ten steps to get started in Genome Assembly and Annotation

[version 1; peer review: 2 approved]

Victoria Dominguez Del Angel 10, Erik Hjerde 10, Lieven Sterck 10,4, Salvadors Capella-Gutierrez 10,5,6, Cederic Notredame^{7,8}, Olga Vinnere Pettersson⁹, Joelle Amselem 10, Laurent Bouri 10, Stephanie Bocs 10,11,13, Christophe Klopp 10,14, Jean-Francois Gibrat 10,15, Anna Vlasova 10,8, Brane L. Leskosek 16, Lucile Soler 17, Mahesh Binzer-Panchal 10,17, Henrik Lantz 10,17

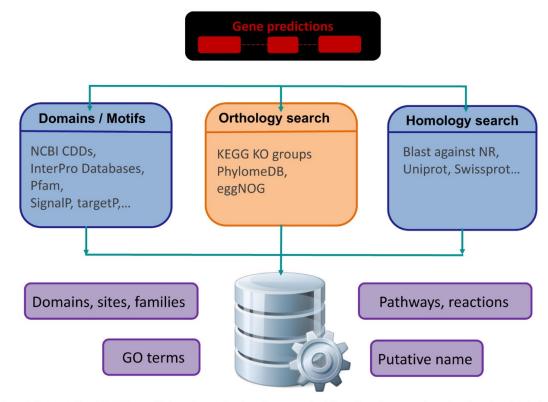


Figure 4. Functional Annotation Pipelines. This schema is showing a typical functional annotation pipeline, in which functional roles are assigned to coding sequences (CDSs) inferred in the gene prediction process. The process implements three parallel routes for the definition of functions. The first refers to proteins domains and motifs, the second for orthology search and finally the third is applied to homology search. At the end, the output from the three different sources is put together for more valuable predictions.

Orthology-based functional annotation with eggNOG

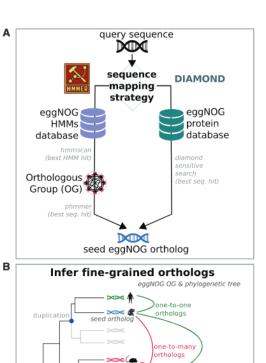
Fast Genome-Wide Functional Annotation through Orthology Assignment by eggNOG-Mapper

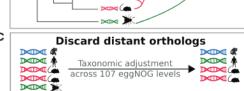
Jaime Huerta-Cepas,^{†,1} Kristoffer Forslund,^{†,1} Luis Pedro Coelho,¹ Damian Szklarczyk,^{2,3} Lars Juhl Jensen,⁴ Christian von Mering,^{2,3} and Peer Bork*,^{1,5,6,7}

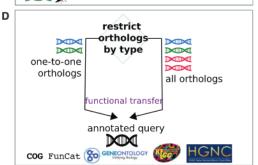
Associate editor: Hongzhi Kong

Abstract

Orthology assignment is ideally suited for functional inference. However, because predicting orthology is computationally intensive at large scale, and most pipelines are relatively inaccessible (e.g., new assignments only available through database updates), less precise homology-based functional transfer is still the default for (meta-)genome annotation. We, therefore, developed eggNOG-mapper, a tool for functional annotation of large sets of sequences based on fast orthology assignments using precomputed clusters and phylogenies from the eggNOG database. To validate our method, we benchmarked Gene Ontology (GO) predictions against two widely used homology-based approaches: BLAST and InterProScan. Orthology filters applied to BLAST results reduced the rate of false positive assignments by 11%, and increased the ratio of experimentally validated terms recovered over all terms assigned per protein by 15%. Compared with InterProScan, eggNOG-mapper achieved similar proteome coverage and precision while predicting, on average, 41 more terms per protein and increasing the rate of experimentally validated terms recovered over total term assignments per protein by 35%. EggNOG-mapper predictions scored within the top-5 methods in the three GO categories using the CAFA2 NK-partial benchmark. Finally, we evaluated eggNOG-mapper for functional annotation of metagenomics data, yielding better performance than interProScan. eggNOG-mapper runs ~15× faster than BLAST and at least 2.5× faster than InterProScan. The tool is available standalone and as an online service at http://eggnog-mapper.embl.de.







¹Structural and Computational Biology Unit, European Molecular Biology Laboratory, Heidelberg, Germany

²Institute of Molecular Life Sciences, University of Zurich, Zurich, Switzerland

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⁴The Novo Nordisk Foundation Center for Protein Research, Faculty of Health and Medical Sciences, University of Copenhagen, Copenhagen, Denmark

⁵Germany Molecular Medicine Partnership Unit (MMPU), University Hospital Heidelberg and European Molecular Biology Laboratory, Heidelberg, Germany

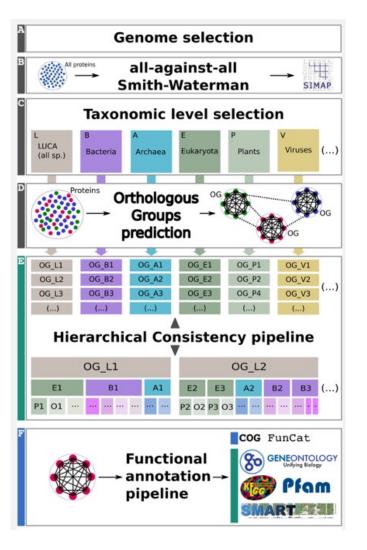
⁶Max Delbrück Centre for Molecular Medicine, Berlin, Germany

⁷Department of Bioinformatics, Biocenter University of Würzburg, Würzburg, Germany

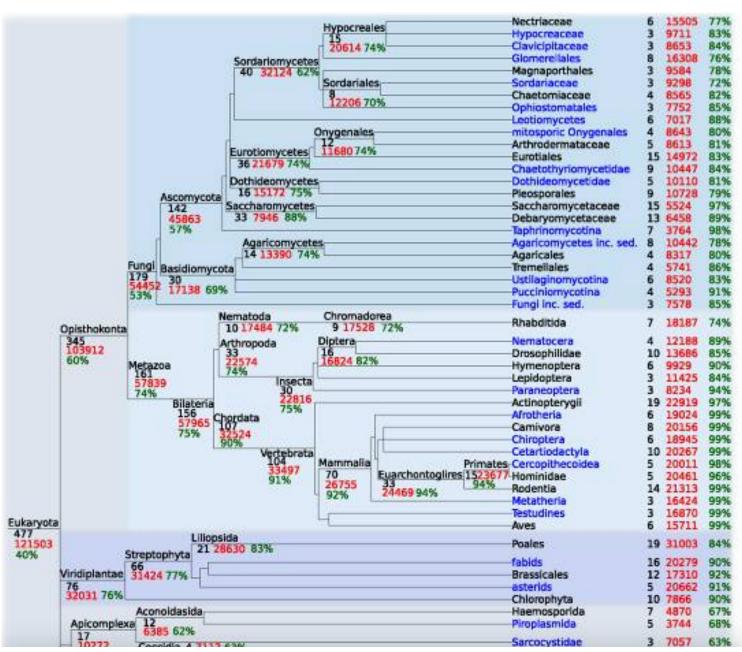
[†]These authors contributed equally to this work.

^{*}Corresponding author: E-mail: bork@embl.de.

eggNOG taxonomy level

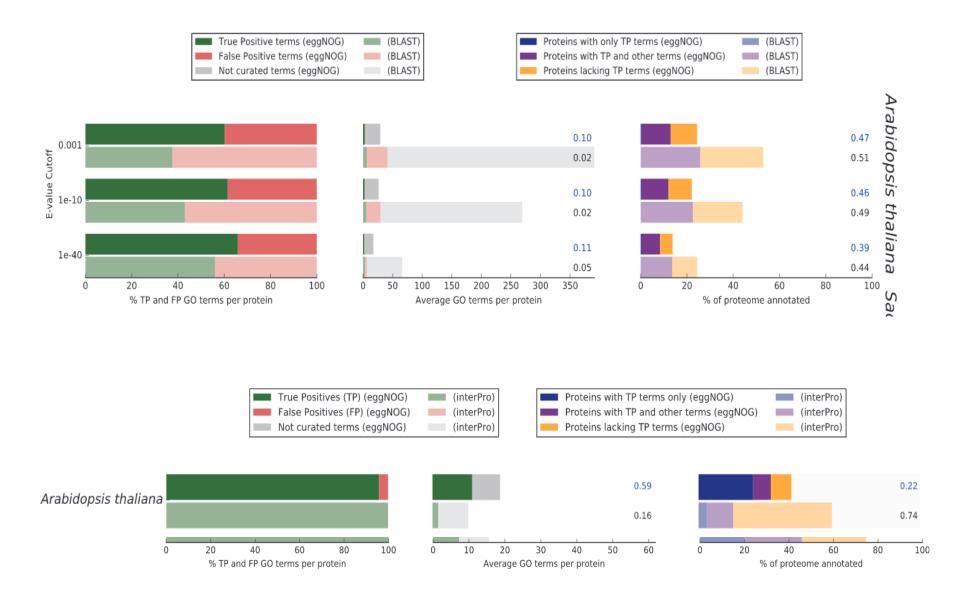


Searches through eggNOG-mapper can be restricted to a taxonomy level => Trade off between specificity and sensitivity



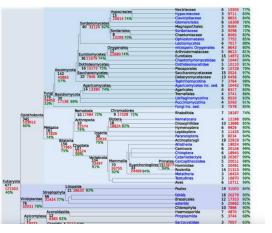
Orthology-based functional annotation with eggNOG

eggNOG is more specific but less sensitive than similarity (blast) or domains-based (InterProScan) methods



Let's have a look at eggNOG-mapper results

				🧩 Workflow Visualize Données partagées ▼ Aide ▼ Utilisateur ▼ 😂 💄 🎹			
October 6th to 7th. Running jobs will be stopped. Thank you for your understanding.							
FUN_000115-T1	36080.S2J093	1.52e-51	187.0	KOG1039@1 root,KOG1039@2759 Eukaryota,3958Y@33154 Opisthokonta,3Q2CW@4751 Fungi,1GXEC@112252 Fungi incertae sedis	4751 Fungi	S	protein ubiquitination
FUN_000116-T1	36080.S2JKD5	4.51e-41	163.0	KOG2992@1 root,KOG2992@2759 Eukaryota,38DH8@33154 Opisthokonta,3NUEP@4751 Fungi,1GTTM@112252 Fungi incertae sedis	4751 Fungi	Υ	SRP40, C-terminal domain
FUN_000117-T1	36080.S2J4G6	0.0	1245.0	KOG3836@1 root,KOG3836@2759 Eukaryota,39JZT@33154 Opisthokonta,3P07N@4751 Fungi,1GVA9@112252 Fungi incertae sedis	4751 Fungi	K	ig-like, plexins, transcription factors
FUN_000118-T1	36080.S2JWH6	5.42e-25	115.0	2CWJA@1 root,2RUPX@2759 Eukaryota,39PS2@33154 Opisthokonta,3Q60Z@4751 Fungi,1GV0X@112252 Fungi incertae sedis	36080.S2JWH6 -	-	-
FUN_000119-T1	36080.S2K4Z5	7.46e-185	556.0	293MF@1 root,2RAI8@2759 Eukaryota,3AWXX@33154 Opisthokonta,3Q1QD@4751 Fungi,1GVJ5@112252 Fungi incertae sedis	4751 Fungi	-	-
FUN_000120-T1	556268.OFAG_00919	1.17e-07	62.4	COG0790@1 root,COG0790@2 Bacteria,1MWPA@1224 Proteobacteria,2VM3A@28216 Betaproteobacteria	1224 Proteobacteria	KLT	PFAM Sel1 domain protein repeat-containing protein
FUN_000121-T1	36080.S2JT78	6.53e-171	487.0	COG0564@1 root,KOG1919@2759 Eukaryota,38G5C@33154 Opisthokonta,3NVMQ@4751 Fungi,1GUC6@112252 Fungi incertae sedis	4751 Fungi	Α	RNA pseudouridylate synthase
FUN_000122-T1	36080.S2IYX5	5.74e-196	556.0	KOG4733@1 root,KOG4733@2759 Eukaryota,38GX0@33154 Opisthokonta,3NWWZ@4751 Fungi,1GUJ5@112252 Fungi incertae sedis	4751 Fungi	Α	RNA recognition motif
FUN_000123-T1	936053.I1CPJ9	3.37e-39	151.0	2DKYR@1 root,2S63V@2759 Eukaryota,3A7H5@33154 Opisthokonta,3P555@4751 Fungi	4751 Fungi	S	Spizellomyces punctatus DAOM BR117
FUN_000127-T1	36080.S2JU99	7.61e-139	413.0	KOG1819@1 root,KOG1819@2759 Eukaryota,39204@33154 Opisthokonta,3PHJW@4751 Fungi,1GVQF@112252 Fungi incertae sedis	4751 Fungi	S	negative regulation of epidermal growth factor-activated
FUN_000128-T1	36080.S2JUA4	1.79e-103	305.0	2CXIS@1 root,2RXWC@2759 Eukaryota,3A19Z@33154 Opisthokonta	33154 Opisthokonta	S	DUF2407 C-terminal domain
FUN_000129-T1	36080.S2IZZ8	9.73e-294	813.0	KOG1087@1 root,KOG1087@2759 Eukaryota,38DR4@33154 Opisthokonta,3NV9 @4751 Fungi,1GSCY@112252 Fungi incertae sedis	4751 Fungi	U	Adaptin C-terminal domain
FUN_000130-T1	36080.S2J5C2	4.29e-222	617.0	KOG1164@1 root,KOG1164@2759 Eukaryota,38BQ1@33154 Opisthokonta,3NUPD@4751 Fungi,1GV4X@112252 Fungi incertae sedis	4751 Fungi	T	Protein tyrosine kinase
FUN_000136-T1	36080.S2K6E0	5.88e-163	491.0	KOG4628@1 root,KOG4628@2759 Eukaryota,3AX63@33154 Opisthokonta,3Q20B@4751 Fungi,1GWRB@112252 Fungi incertae sedis	4751 Fungi	0	zinc-RING finger domain
FUN_000139-T1	36080.S2J1M7	1.29e-231	649.0	KOG3780@1 root,KOG3780@2759 Eukaryota,3AWX9@33154 Opisthokonta,3Q1PR@4751 Fungi,1GVFT@112252 Fungi incertae sedis	4751 Fungi	S	Arrestin (or S-antigen), C-terminal domain
FUN_000140-T1	36080.S2JXA7	8.04e-66	234.0	KOG4339@1 root,KOG4339@2759 Eukaryota,39S5X@33154 Opisthokonta,3NVE2@4751 Fungi,1GUWE@112252 Fungi incertae sedis	4751 Fungi	S	to Saccharomyces cerevisiae BNI4 (YNL233W)
FUN_000141-T1	36080.S2JLS1	1.94e-167	476.0	KOG0588@1 root,KOG0588@2759 Eukaryota,38BGN@33154 Opisthokonta,3NW2V@4751 Fungi,1GWAH@112252 Fungi incertae sedis	4751 Fungi	D	Serine/Threonine protein kinases, catalytic domain
FUN_000142-T1	36080.S2JKN5	7.38e-157	446.0	COG0445@1 root,KOG2667@2759 Eukaryota,39RP0@33154 Opisthokonta,3NUJ6@4751 Fungi,1GUF6@112252 Fungi incertae sedis	4751 Fungi	U	Endoplasmic reticulum vesicle transporter
FUN_000143-T1	936053.I1BJ13	4.18e-60	194.0	2D8F6@1 root,2T9ZJ@2759 Eukaryota,394TC@33154 Opisthokonta,3Q1W1@4751 Fungi,1GW90@112252 Fungi incertae sedis	4751 Fungi	-	-
FUN_000144-T1	36080.S2J4M1	1.23e-180	503.0	COG0090@1 root,KOG2309@2759 Eukaryota,38BCB@33154 Opisthokonta,3NU8K@4751 Fungi,1GSN6@112252 Fungi incertae sedis	4751 Fungi	J	Mortierella verticillata NRRL 6337 60S ribosomal protein
FUN_000145-T1	36080.S2JQW6	4.03e-281	775.0	COG2940@1 root,KOG4056@1 root,KOG2084@2759 Eukaryota,KOG4056@2759 Eukaryota,3A1HC@33154 Opisthokonta,3P2QG@4751 Fungi,1GRT3@112252 Fungi incertae sedis	4751 Fungi	BU	MAS20 protein import receptor
FUN 000146-T1	36080.S2.JU43	4.48e-224	622.0	COG3000@1lroot.KOG0539@2759lEukarvota.38DTY@33154lOpisthokonta.3NV5W@4751lFungi.1GS8X@112252lFungi incertae sedis	4751IFunai	I	Ceramide hydroxylase involved in the alpha-hydroxylation



annotations

This file provides final annotations of each query. Tab-delimited columns in the file are:

- •query_name: query sequence name
- seed_eggNOG_ortholog: best protein match in eggNOG
- •seed_ortholog_evalue: best protein match (e-value)
- •seed_ortholog_score: best protein match (bit-score)
- •predicted_taxonomic_group
- •predicted_protein_name: Predicted protein name for query sequences
- •GO_terms: Comma delimited list of predicted Gene Ontology terms
- •EC number
- •KEGG KO
- •KEGG Pathway: Comma delimited list of predicted KEGG pathways
- •KEGG Module
- •KEGG Reaction
- •KEGG rclass
- •BRITE
- •KEGG TC
- CAZy
- •BiGG Reactions
- •Annotation_tax_scope: The taxonomic scope used to annotate this query sequence
- •Matching_OGs: Comma delimited list of matching eggNOG Orthologous Groups
- •best_OG|evalue|score: Best matching Orthologous Groups (deprecated, use smallest from eggnog OGs)
- •COG_functional_categories: COG functional category inferred from best matching OG
- •eggNOG_free_text_description

INFORMATION STORAGE AND PROCESSING

- [J] Translation, ribosomal structure and biogenesis
- [A] RNA processing and modification
- [K] Transcription
- [L] Replication, recombination and repair
- [B] Chromatin structure and dynamics

CELLULAR PROCESSES AND SIGNALING

- [D] Cell cycle control, cell division, chromosome partitioning
- [Y] Nuclear structure
- [V] Defense mechanisms
- [T] Signal transduction mechanisms
- [M] Cell wall/membrane/envelope biogenesis
- [N] Cell motility
- [Z] Cytoskeleton
- [W] Extracellular structures
- [U] Intracellular trafficking, secretion, and vesicular transport
- [0] Posttranslational modification, protein turnover, chaperones

METABOLISM

- [C] Energy production and conversion
- [G] Carbohydrate transport and metabolism
- [E] Amino acid transport and metabolism
- [F] Nucleotide transport and metabolism
- [H] Coenzyme transport and metabolism
- [I] Lipid transport and metabolism
- [P] Inorganic ion transport and metabolism
- [Q] Secondary metabolites biosynthesis, transport and catabolism

POORLY CHARACTERIZED

- [R] General function prediction only
- [S] Function unknown