

High-throughput Sequencing course 2019

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OMICS ERA

What is 'OMICS': derived from Latin suffix “ome” meaning mass or many.

Thus OMICS involve a mass (large number) of measurements per endpoint. (Jackson et al. 2006)

- Genomics
- Transcriptomics
- Proteomics
- Reactomics
- Metabonomics
- Phenomics

APPLICATION OF DIFFERENT OMICS

Genomics	Transcriptomics	Proteomics	Metabolomics	Protein-DNA interactions	Protein-protein interactions	Fluxomics	Phenomics
Genomics (sequence annotation)	• ORF validation • Regulatory element identification⁷⁶	• SNP effect on protein activity or abundance	• Enzyme annotation	• Binding-site identification⁷⁵	• Functional annotation⁷⁹	• Functional annotation	• Functional annotation^{71, 101} • Biomarkers¹²⁵
	Transcriptomics (microarray, SAGE)	• Protein: transcript correlation⁷⁸	• Enzyme annotation¹⁸⁴	• Gene-regulatory networks⁷⁶	• Functional annotation⁸⁹ • Protein complex identification⁵⁷		• Functional annotation¹⁸²
		Proteomics (abundance, post-translational modification)	• Enzyme annotation⁹⁹	• Regulatory complex identification	• Differential complex formation	• Enzyme capacity	• Functional annotation
			Metabolomics (metabolite abundance)	• Metabolic-transcriptional response		• Metabolic pathway bottlenecks	• Metabolic flexibility • Metabolic engineering¹⁰⁸
				Protein-DNA interactions (ChIP-chip)	• Signalling cascades^{89, 182}		• Dynamic network responses⁸⁴
					Protein-protein interactions (yeast 2H, coAP-MS)		• Pathway identification activity⁸⁹
						Fluxomics (isotopic tracing)	• Metabolic engineering
							Phenomics (phenotype arrays, RNAi screens, synthetic lethals)

Genomics

Refer to the use of computational and statistical analysis to decipher biology from genome sequences including both DNA and RNA sequences as other post-genomic data (i.e. Genomic microarrays)

Functional Genomics

Is a field of molecular biology that attempts to make use of the vast wealth of data produced by genomics (i.e. genomic sequences) to describe function and interactions. Unlike Genomics, functional genomics focuses on the dynamic aspects such as transcription, translation, protein-protein interactions. Is a more high-throughput methods rather than a gene by gene approach.

Confusion with the word ontology ?



Controlled Vocabulary:

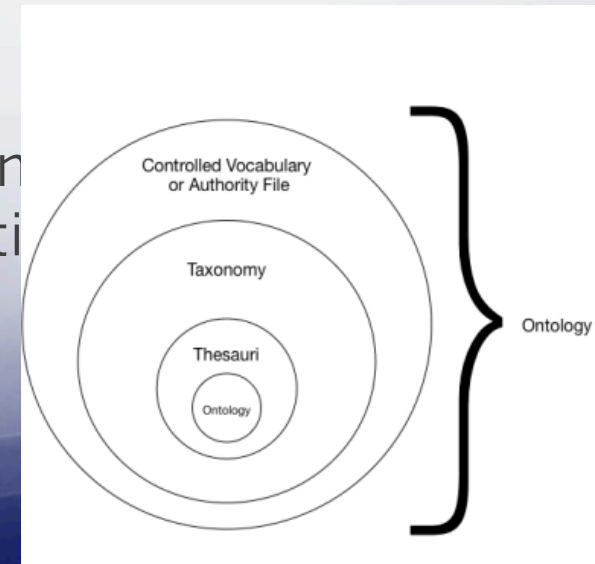
Is a restricted list of words used for labeling, indexing or categorizing

Taxonomy:

Is a classification system. The taxonomy in a controlled vocabulary indicates a hierarchical structure based on an IsA relationship

Ontology:

Is a controlled vocabulary with a taxonomy
complex than complex than just IsA relationship



Using Ontology for data analysis

- Standard identifiers for classes and relations that represent the phenomena within a domain;
- Vocabulary for a domain;
- Metadata that describes the intended meaning of the classes and relations in ontologies;
- Machine-readable axioms and definitions that enable computational access to some aspects of the meaning of classes and relations

The use of standard identifiers for classes and relations in ontologies is a key component in enabling data integration across multiple databases, because the same identifiers can be used across multiple, disconnected databases, files or web sites

These identifiers are widely used in structured file formats, in knowledge bases and data repositories.

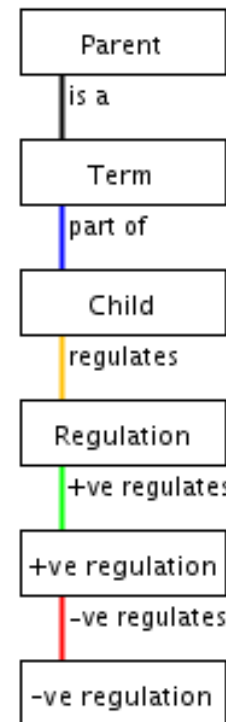
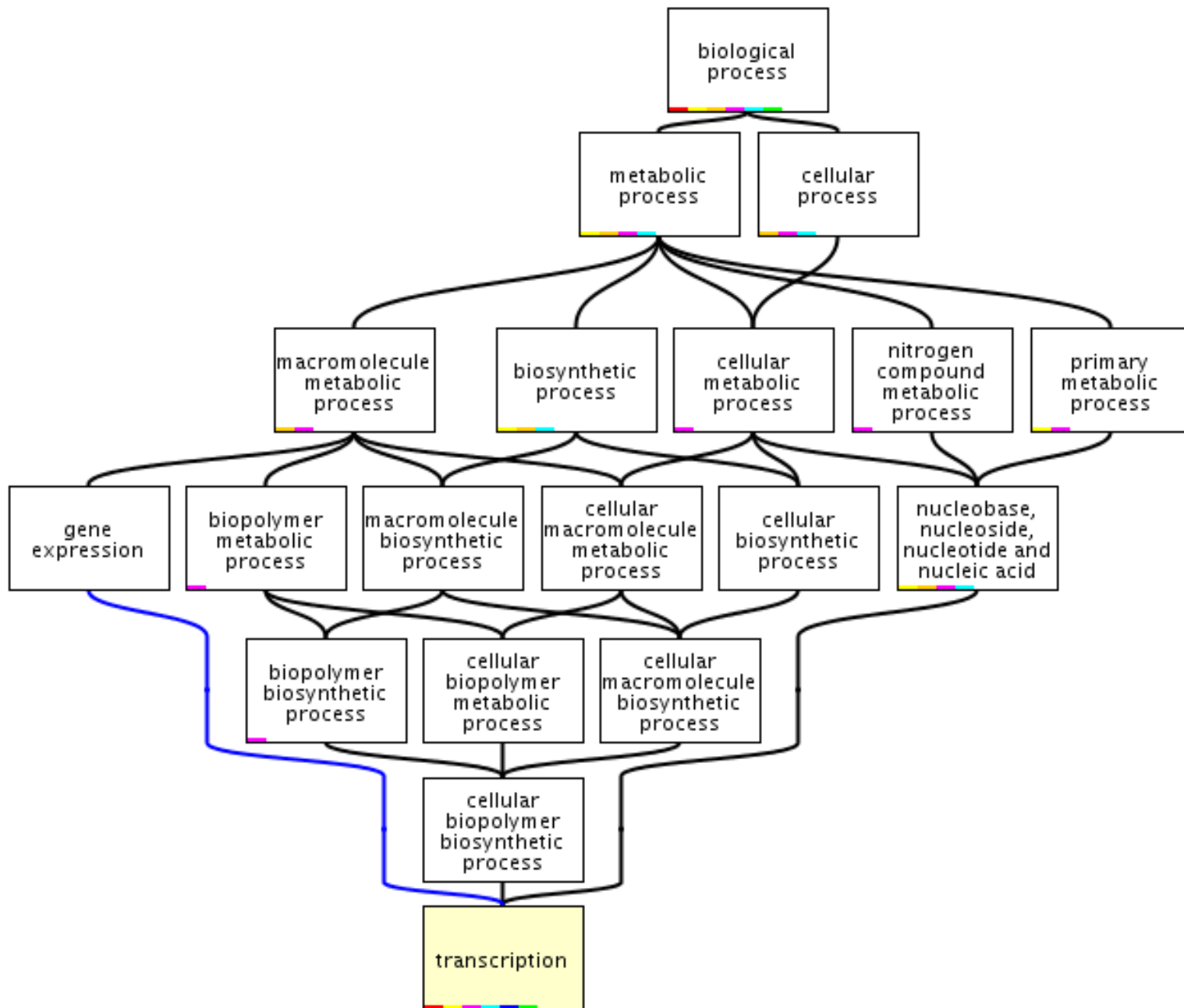
Most successful ontology thus far

\$200 mill. invested in literature and database curation using the Gene Ontology (GO) since 1999

over 11 million annotations relating gene products (proteins) described in the UniProt, Ensembl and other databases to terms in the GO

GO provides a controlled system of representations for use in annotating data and literature that is

- multi-species
- multi-disciplinary
- multi-granularity, from molecules to population

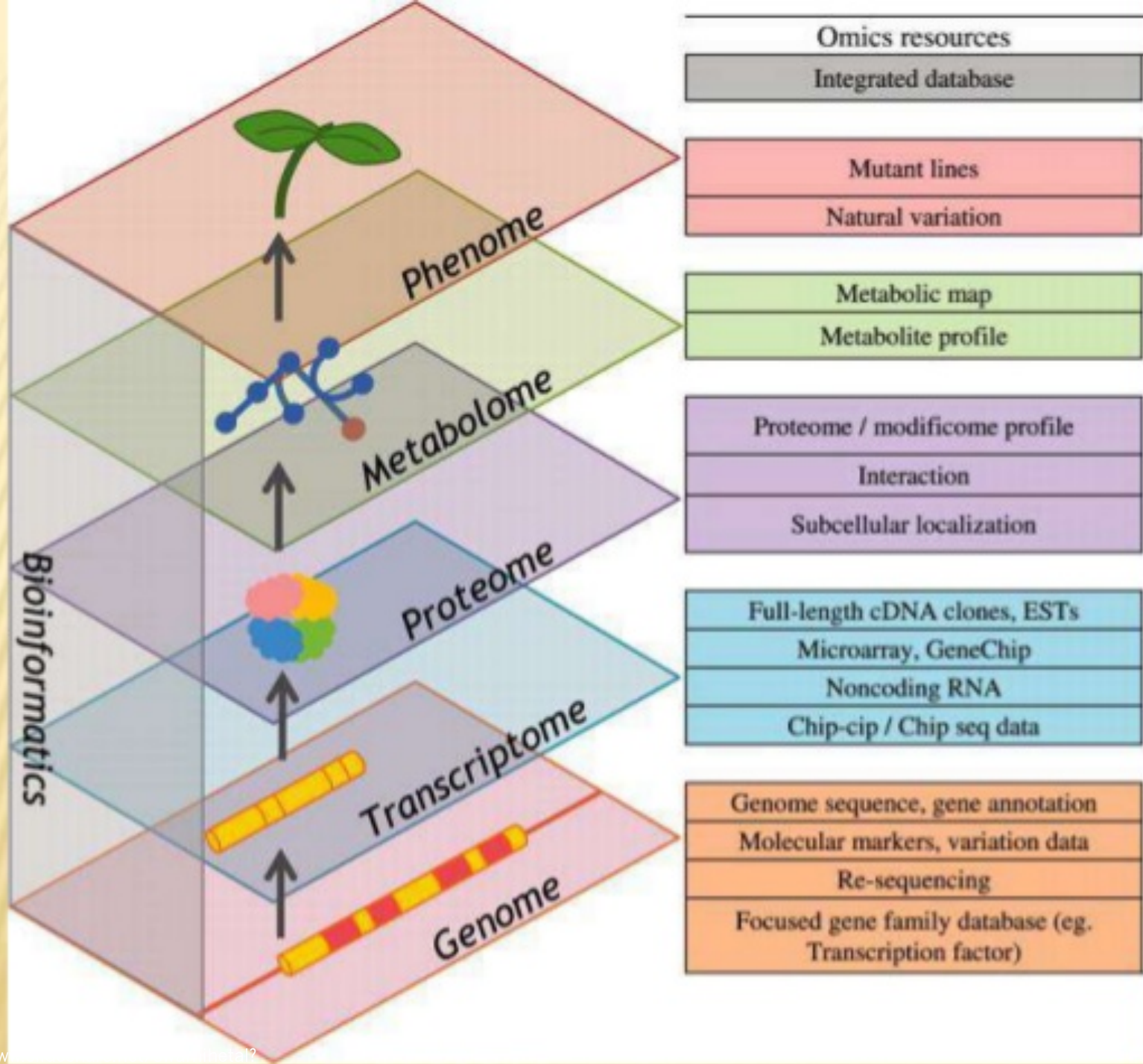


Gene ontology structure

GO MF molecular functions
Molecular activities of a gene product

GO BP biological processes
Specific Objective that the organism is
genetically programmed to achieve

GO CC cellular components
Where gene products are active



High-throughput technologies generated data sets of often thousands of genes, which needed to be interpreted in terms of biological roles.

This provided the impetus behind the ongoing functional and structural annotation of gene products, which is now available through the GO database ([AMIGO](#)) and is a mainstay of modern bioinformatics.

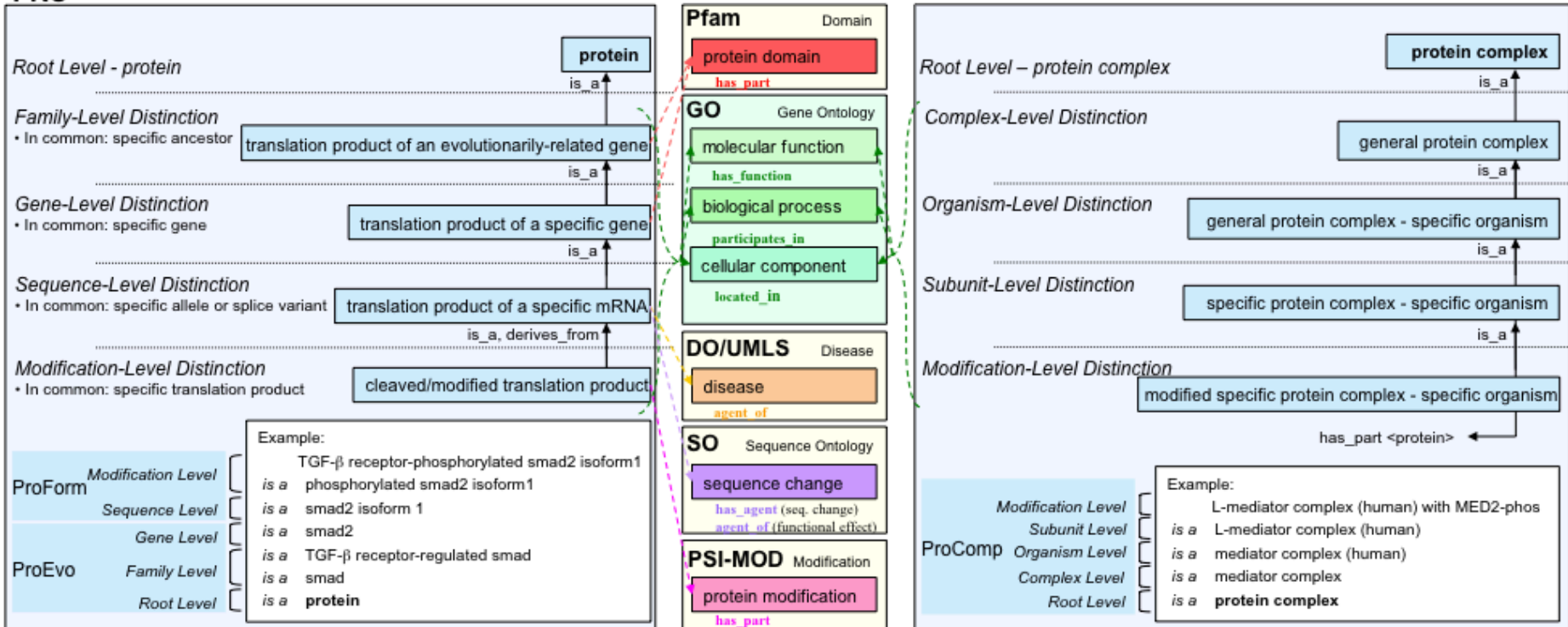
Protein Ontology

<https://proconsortium.org/>

PRO encompasses three sub-ontologies: proteins based on evolutionary relatedness (ProEvo); protein forms produced from a given gene locus (ProForm); and protein-containing complexes (ProComp).

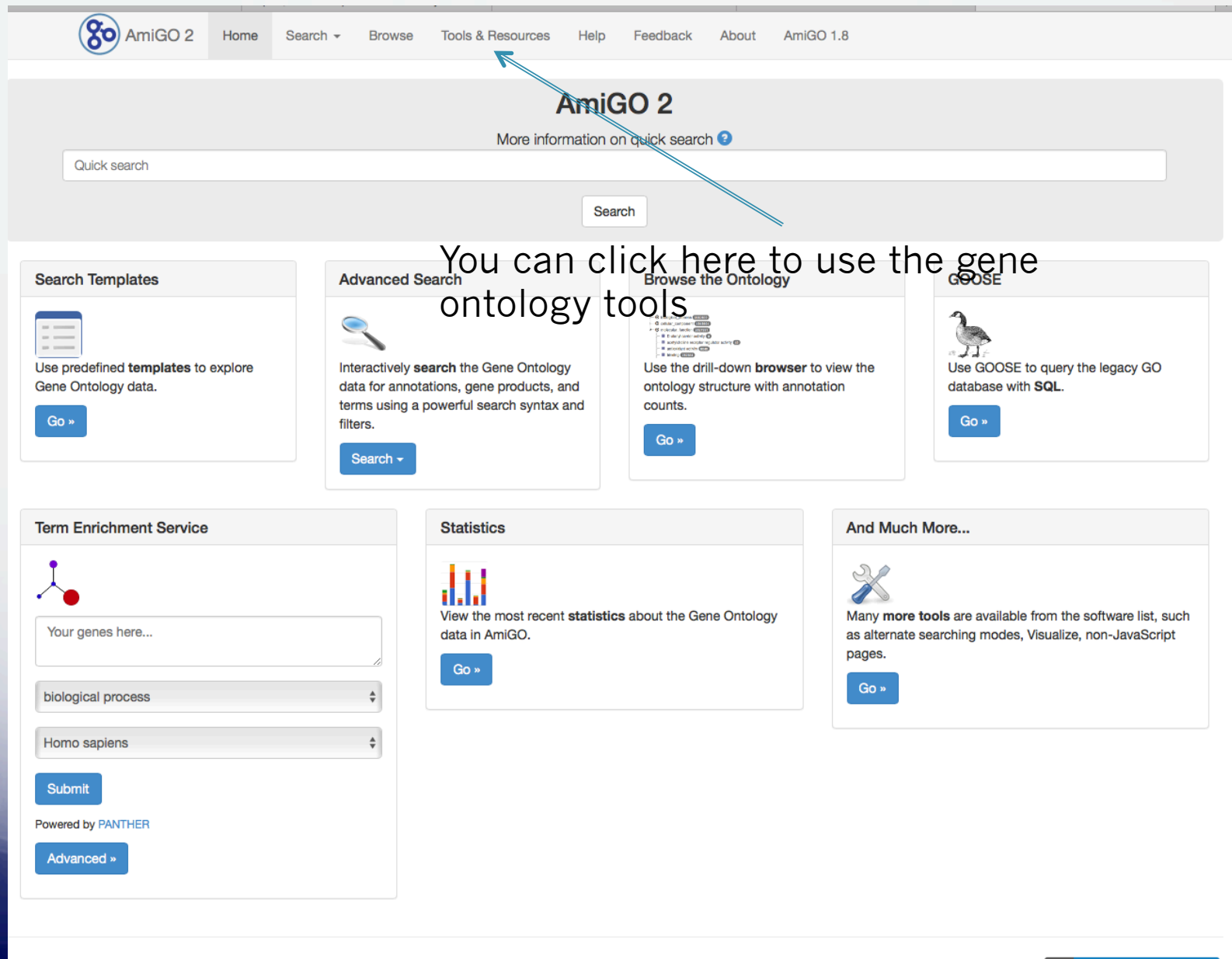
(Explanation of the figure)

PRO



Go to the [AMIGO](#) database and describe it

Amigo2 home page <http://amigo.geneontology.org/amigo>



The screenshot shows the AmiGO 2 home page. At the top, there is a navigation bar with the following links: Home, Search, Browse, Tools & Resources, Help, Feedback, About, and AmiGO 1.8. A blue arrow points to the 'Tools & Resources' link. Below the navigation bar, the main heading 'AmiGO 2' is displayed, followed by the text 'More information on quick search ?'. A search bar with the placeholder text 'Quick search' and a 'Search' button is located below this. The page is divided into several sections:

- Search Templates:** Use predefined templates to explore Gene Ontology data. Includes a 'Go »' button.
- Advanced Search:** Interactively search the Gene Ontology data for annotations, gene products, and terms using a powerful search syntax and filters. Includes a 'Search' button.
- Browse the Ontology:** Use the drill-down browser to view the ontology structure with annotation counts. Includes a 'Go »' button.
- GOOSE:** Use GOOSE to query the legacy GO database with SQL. Includes a 'Go »' button.
- Term Enrichment Service:** A section for term enrichment analysis. It includes a text input field labeled 'Your genes here...', a dropdown menu for 'biological process', another dropdown menu for 'Homo sapiens', a 'Submit' button, and a link to 'Powered by PANTHER'. It also includes an 'Advanced »' button.
- Statistics:** View the most recent statistics about the Gene Ontology data in AmiGO. Includes a 'Go »' button.
- And Much More...:** Many more tools are available from the software list, such as alternate searching modes, Visualize, non-JavaScript pages. Includes a 'Go »' button.



AmiGO 2

[Home](#)[Search ▾](#)[Browse](#)[Tools & Resources](#)[Help](#)[Feedback](#)[About](#)[AmiGO 1.8](#)

Explore GO data

[Home](#) allows simple quick searches using autocomplete and the new GOlr backend.

Interactively search and filter the GO data as you type. These pages allow you to use a powerful search syntax (including boolean operators).

[Annotations](#): Associations between GO terms and genes or gene products.

[Ontology](#): Gene Ontology Term, Synonym, or Definition.

[Genes and gene products](#): Genes and gene products associated with GO terms.

[Search templates](#) (a search "wizard") can be used to quickly answer common questions using a fill-in-the-blank approach.

[Visualization](#) creator for GO graph. Freely make your own ontology graph, including label and color changes. Please see the documentation for more [details](#).

[Drill-down browser](#) can be used to explore the ontologies and annotation from the top levels by opening direct descendant terms individually to create a custom view.

[Base statistics](#) can give a high-level view of what kind of data is available in AmiGO.

PubMed ID search takes you to a page with summary information and all GO annotations using the inputted ID.

Medial search returns match counts for the different search personalities and allows users to select which one they wish to continue with.

Analysis of GO data

[Term enrichment services](#) offers access to term enrichment analysis, finding significant shared GO terms or parents of those GO terms to help discover what input genes may have in common.

[Term enrichment](#) [AmiGO 1.8](#) finds significant shared GO terms or parents of those GO terms, used to describe the genes in the query/input set to help discover what those genes may have in common.

[GO slimmer](#) [AmiGO 1.8](#) allows you to map the granular annotations of the query set of genes to one or more high-level, broader parent terms referred to as GO Slim terms.

[BLAST](#) [AmiGO 1.8](#) search for gene products. The sequence search is performed using either BLASTP or BLASTX (from the WU-BLAST package), depending on the type of the input sequence.

[Term matrix](#) for exploring gene product (bioentity) co-annotation information.

You can click here for the
gene ontology enrichment
analysis

Directly query GO data

[GOOSE](#) is the GO Online SQL Environment. Using this, users may query the a GO SQL database directly using SQL, or edit one of the already made templates.

[Gannet](#) is the Solr analog to GOOSE.

Instance information

[Schema details](#) what fields are recognized by AmiGO 2 on the Solr server.

[Load details](#) about which files are being used in this instance of AmiGO 2 and when they were loaded.

[OWLTools information](#) for the bundled version with this instance of AmiGO.

[Xrefs](#) lists the cross-references currently understood by AmiGO.

UPL14.1 New! PANTHER14.1 is generated from the 2018_04 release of [ReferenceProteome dataset](#)

Search

All

Go

Quick links

[Whole genome function views](#)

[Genome statistics](#)

[Data Version](#)

[How to cite PANTHER](#)

NEW! [Recent publication describing PANTHER](#)

News

PANTHER 14.1 Released

[Click for additional info.](#)

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Subscribe



Gene List Analysis

Browse

Sequence Search

cSNP Scoring

Keyword Search

Please refer to our article in [Nature Protocols](#) for detailed instructions on how to use this page.

Error parsing request, no input specified

Help Tips

Steps:

- 1. Select list and list type to analyze
- 2. Select Organism
- 3. Select operation

1. Enter ids and or select file for batch upload. Else enter ids or select file or list from workspace for comparing to a reference list.

Enter IDs:
[Supported IDs](#)

separate IDs by a space or comma

Upload IDs:
[File format](#)

Choose File no file selected

Please [login](#) to be able to select lists from your workspace.

Select List Type:

- ☒ ID List
- ☐ Previously exported text search results
- ☐ Workspace list
- ☐ PANTHER Generic Mapping
- ☐ ID's from Reference Proteome Genome

Organism for id list:

☐ VCF File Flanking region:

Paste your gene list or upload your file

2. Select organism.

Homo sapiens
Mus musculus
Rattus norvegicus
Gallus gallus
Danio rerio

Choose the specie

3. Select Analysis.

- ☒ Functional classification viewed in gene list
- ☐ Functional classification viewed in graphic charts ☐ Bar chart ☐ Pie chart
- ☐ Statistical overrepresentation test ☐ Use default settings
- ☐ Statistical enrichment test ☐ Use default settings

submit

Choose the analysis type

UPL14.1 New! PANTHER14.1 is generated from the 2018_04 release of [ReferenceProteome dataset](#)

PANTHER BAR CHART

[Pie Chart](#) [Export](#)

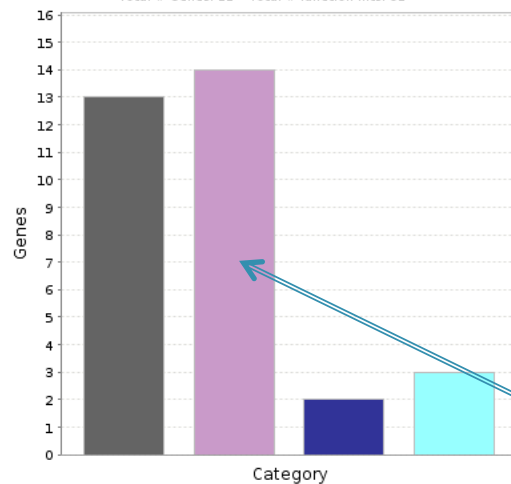
Features:

- Mouse-over bar chart category to see category name and statistics
- Click on a bar chart category to drill down to child categories
- Click on chart legend link to retrieve gene list for each category
- Click on a color key in chart legend to choose your favorite color for the category **NEW!**

Select Ontology: Molecular Function View: 100%

PANTHER GO-Slim Molecular Function

Total # Genes: 22 Total # function hits: 32



Click to get gene list for a category:

- [binding \(GO:0005488\)](#)
- [catalytic activity \(GO:0003824\)](#)
- [molecular function regulator \(GO:0098772\)](#)
- [translation regulator activity \(GO:0045182\)](#)

Color picker powered by 

**Chart tooltips are read as: Category Name, Accession, # genes, Percent of gene hit against total # genes; Percent of gene hit against total # function hits

Click on the bar to see more specific terms

Fungi Databases

[Eupath DB](#)

[FungiDB](#)

[YeastMine](#)

Select gene ids and paste your gene list than
Get Answer

You will get a list of the Gene Id; organism;
transcript IDs; location etc.

You can

Add Step

Transform to pathway

Find Orthologous

Go to

[https://yeastmine.yeastgenome.org/
yeastmine/begin.do](https://yeastmine.yeastgenome.org/yeastmine/begin.do)

.

Homolog

A gene related to a second gene by descent from a common ancestral DNA sequence. The term, homolog, may apply to the relationship between genes separated by the event of speciation (see Ortholog) or to the relationship between genes separated by the event of genetic duplication.

Ortholog

Orthologs are genes in different species that evolved from a common ancestral gene by speciation. Normally, orthologs retain the same function in the course of evolution. Identification of orthologs is critical for reliable prediction of gene function in newly sequenced genomes.

Paralog

Paralogs are genes related by duplication within a genome. Orthologs retain the same function in the course of evolution, whereas paralogs evolve new functions, even if these are related to the original one.

STEPS 1

1. Search for genes and organism
Cryptococcus neoformans var. grubii H99
2. Paste the list of the all Cryptococcus neoformans var. grubii H99 and **RUN**
3. You will find genes that are present in the database
4. Than do save list a new page will open

STEPS 2

5. Than under Orthologous click *S.cerevisiae* and you will get a list of orthologous
6. Than under Orthologous you can click on human or mouse orthologous of *S.cerevisiae*
7. Than you can use that list for search Gene Ontology or other human or mouse databases.

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Thank you

Any questions contact me:
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