

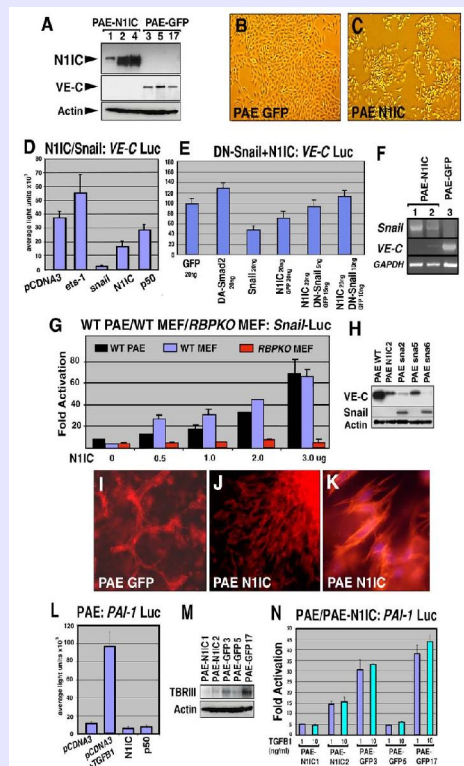
Text mining the Biomedical Literature

“We have here much data, and we must proceed to lay out our campaign”,

Van Helsing in Bram Stockers Dracula

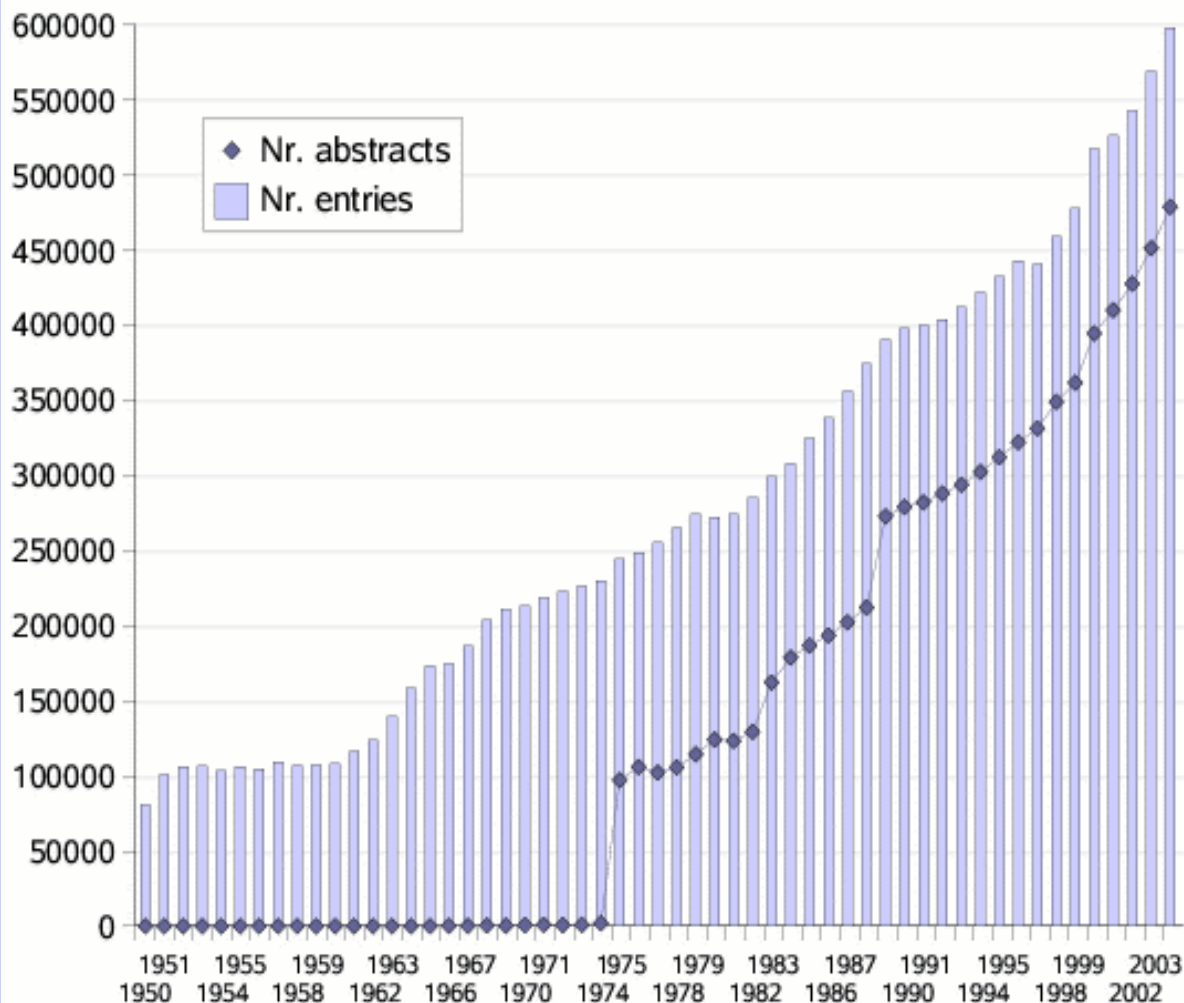
Talk overview:

- The Biomedical literature
- Natural language processing (NLP)
- NLP in the Molecular Biology domain
- Text mining applications
- Evaluation of Text mining tools
- Conclusions and outlook
- Useful links, reviews and articles



PubMed/Medline database at NCBI

PubMed growth



- Developed at the National Center for Biotechnology Information (NCBI).
- The core 'Textome'.
- repository of citation entries of scientific articles.
- PubMed titles and abstracts are primary data source for Bio-NLP.
- ~ 450,000 new abstracts/a
- > 4,800 biomedical journals
- ENTREZ search engine

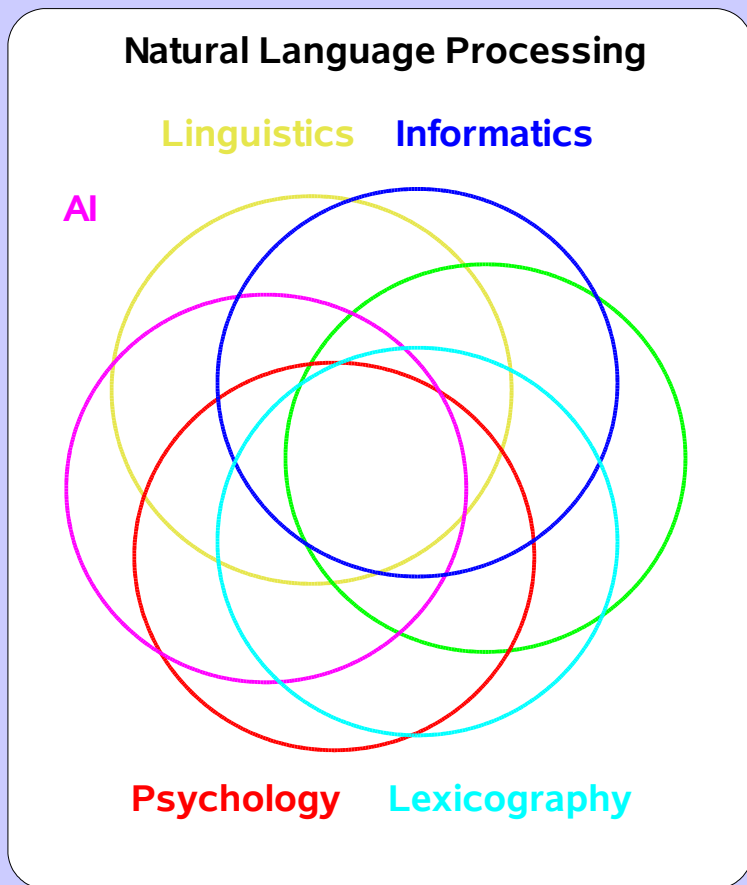
PubMed online

The screenshot shows the PubMed website interface as it appeared in 2005. The browser window is titled "Entrez PubMed - Mozilla <2>". The address bar shows the URL: http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=pubmed&dopt=Abstract&list_uids=15784250&query_hl=7. The search bar contains the text "PubMed".

Annotations on the screenshot include:

- Find similar entries**: Points to the "Find similar entries" link in the top right corner.
- Link to full text**: Points to the "Full Text" link next to the search result.
- Journal and publication date**: Points to the citation information: "J Mol Biol. 2005 Apr 15;347(5):895-902."
- Title**: Points to the article title: "Structure of the connector of bacteriophage T7 at 8A resolution: structural homologies of a basic component of a DNA translocating machinery."
- Authors**: Points to the author list: "Agirrezabala X, Martin-Benito J, Valle M, Gonzalez JM, Valencia A, Valpuesta JM, Carrascosa JL."
- Abstract**: Points to the abstract text: "The three-dimensional structure of the bacteriophage T7 head-to-tail connector has been obtained at 8A resolution using cryo-electron microscopy and single-particle analysis from purified recombinant connectors. The general morphology of the T7 connector is that of a 12-folded toroidal homopolymer with a channel that runs along the longitudinal axis of the particle. The structure of the T7 connector reveals many structural similarities with the connectors from other bacteriophages. Docking of the atomic structure of the varphi29 connector into the three-dimensional reconstruction of T7 connector reveals that the narrow, distal region of the two oligomers are almost identical. This region of the varphi29 connector has been suggested to be involved in DNA translocation, and is composed of an alpha-beta-alpha-beta-beta-alpha motif. A search for alpha-helices in the same region of the T7 three-dimensional map has located three alpha-helices in approximately the same position as those of the varphi29 connector. A comparison of the predicted secondary structure of several bacteriophage connectors, including among others T7, varphi29, P22 and SPP1, reveals that, despite the lack of sequence homology, they seem to contain the same alpha-beta-alpha-beta-beta-alpha motif as that present in the varphi29 connector. These results allow us to suggest a common architecture related to a basic component of the DNA translocating machinery for several viruses."
- Pubmed identifier (unique document ID)**: Points to the PMID: 15784250 [PubMed - indexed for MEDLINE]

Natural Language Processing (NLP) basics



Domain, e.g. Biomedicine/
Molecular Biology

- Techniques that analyse, understand and generate language (free text, speech).
- Linguistic tools, e.g. syntactic analyser and semantic classification.
- Multidisciplinary field.
- Strongly language dependent.
- Create computational models of language.
- Learn statistical properties of language.
- Methods: statistical analysis, machine learning, rule-based, pattern-matching, AI, etc...
- Domain dependent (biomedical) vs generic NLP.

Major NLP tasks

- Information Retrieval (IR).
 - Information extraction/Text mining (IE).
 - Question Answering (QA).
 - Natural Language Generation (NLG).
-
- Automatic summarisation.
 - Machine translation.
 - Text proofing.
 - Speech recognition.
 - Optical character recognition (OCR).

Information Retrieval (IR)

- IR: process of **recovery of those documents** from a collection of documents which satisfy a given information demand.
- Information demand often posed in form of a **search query**.
- Example: retrieval of web-pages using search engines, e.g. Google.
- First step: indexing document collection:
 - Tokenization
 - Case folding
 - Stemming
 - Stop word removal
- Efficient indexing to reduce vocabulary of terms and query formulations.
- Example: 'Glycogenin *AND* binding' and 'glycogenin *AND* bind'.
- Query types: Boolean query and Vector Space Model based query.

Boolean query

- Based on **combination of terms** using Boolean operators.
- Basic **Boolean operators**: AND, OR and NOT.
- Queries matched against the terms in the inverted index file.
- Entrez – Boolean search in PubMed.
- Fast, easy to implement.
- **Search engines**: often stop word removal and case folding.
- Stop word removal : space saving speed improvement.
- Return a **unranked list**.
- Return large list of documents, many not relevant.
- Terms have different information content ->
better weighted query.

Zipf's law



- A small number of words occur very often
- Those high frequency words are often function words (e.g. prepositions)
- Most words with low frequency .

From: Rebholz-Schuhmann D, Kirsch H, Couto F (2005) Facts from Text—Is Text Mining Ready to Deliver? PLoS Biol 3(2): e65

Commonly excluded stop words

after	also	an	and
as	at	be	because
before	between	but	before
for	however	from	if
in	into	of	or
other	out	since	such
than	that	the	these
there	this	those	to
under	upon	when	where
whether	which	with	within
without	.	.	.

Vector space model

- Measure **similarity** between query and documents.
- Query can be a list of terms or whole documents.
- Documents and queries as **vectors of terms**.
- **Term weighting** according to their frequency:
 - within the document
 - within the document collection
- Widespread term weighting: tf x idf.
- Calculate similarity between those vectors.
- Cosine similarity.
- Return a ranked list.
- Example: related article search in PubMed

$$w_{i,j} = tf_{i,j} \times idf_j$$

$$idf_{i,j} = \log \left(\frac{N}{df_j} \right)$$

$$sim(Q, D) = \frac{\sum_{j=1}^V w_{Q,j} \times w_{i,j}}{\sqrt{\sum_{j=1}^V w_{Q,j}^2 \times \sum_{j=1}^V w_{i,j}^2}}$$

PubMed online

The screenshot shows the PubMed website interface in a Mozilla browser window. The address bar displays the URL: http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=pubmed&dopt=Abstract&list_uids=15784250&query_hl=7. The page features the NCBI logo, the PubMed logo, and the National Library of Medicine logo. A search bar is located at the top, and a navigation menu is visible below it. The main content area displays a search result for the article "Structure of the connector of bacteriophage T7 at 8A resolution: structural homologies of a basic component of a DNA translocating machinery." by Agirrezabala X, Martin-Benito J, Valle M, Gonzalez JM, Valencia A, Valpuesta JM, Carrascosa JL. The article is published in J Mol Biol. 2005 Apr 15;347(5):895-902. The abstract text is visible below the title. Annotations with arrows point to various elements: "Find similar entries" points to a button in the top right; "Journal and publication date" points to the citation information; "Title" points to the article title; "Authors" points to the author list; "Abstract" points to the abstract text; "PubMed identifier" points to the PMID: 15784250; and "Link to full text" points to the "Full-text article" link.

Entrez PubMed - Mozilla <2>

File Edit View Go Bookmarks Tools Window Help

http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?cmd=Retrieve&db=pubmed&dopt=Abstract&list_uids=15784250&query_hl=7 Search

Home Bookmarks Yahoo Google MK Homepage ORF Zope on http://... PubMed Python Zope PyTut OEAW GeneDic biocreative GenomeNet

NCBI PubMed National Library of Medicine NLM

My NCBI Welcome martink [Sign Out]

All Databases PubMed Nucleotide Protein Genome Structure OMIM PMC Journals Books

Search PubMed for Go Clear

Limits Preview/Index History Clipboard Details

Display Abstract Show 20 Sort by Send to

All: 1 Review: 0

Link to full text

1: J Mol Biol. 2005 Apr 15;347(5):895-902. Journal and publication date

ELSEVIER FULL-TEXT ARTICLE

Structure of the connector of bacteriophage T7 at 8A resolution: structural homologies of a basic component of a DNA translocating machinery. Title

Agirrezabala X, Martin-Benito J, Valle M, Gonzalez JM, Valencia A, Valpuesta JM, Carrascosa JL. Authors

Centro Nacional de Biotecnología, CSIC, Campus de la Universidad Autonoma de Madrid, Cantoblanco 28049 Madrid, Spain.

The three-dimensional structure of the bacteriophage T7 head-to-tail connector has been obtained at 8A resolution using cryo-electron microscopy and single-particle analysis from purified recombinant connectors. The general morphology of the T7 connector is that of a 12-folded toroidal homopolymer with a channel that runs along the longitudinal axis of the particle. The structure of the T7 connector reveals many structural similarities with the connectors from other bacteriophages. Docking of the atomic structure of the varphi29 connector into the three-dimensional reconstruction of T7 connector reveals that the narrow, distal region of the two oligomers are almost identical. This region of the varphi29 connector has been suggested to be involved in DNA translocation, and is composed of an alpha-beta-alpha-beta-alpha motif. A search for alpha-helices in the same region of the T7 three-dimensional map has located three alpha-helices in approximately the same position as those of the varphi29 connector. A comparison of the predicted secondary structure of several bacteriophage connectors, including among others T7, varphi29, P22 and SPP1, reveals that, despite the lack of sequence homology, they seem to contain the same alpha-beta-alpha-beta-alpha motif as that present in the varphi29 connector. These results allow us to suggest a common architecture related to a basic component of the DNA translocating machinery for several viruses.

PMID: 15784250 [PubMed - indexed for MEDLINE]

Abstract

Display Abstract Show 20 Sort by Send to

PubMed identifier

Write to the Help Desk NCBI | NLM | NIH Department of Health & Human Services Privacy Statement | Freedom of Information Act | Disclaimer

eTBlast (1)

eTBlast > Search - Mozilla Firefox

Archivo Editar Ver Ir Marcadores Herramientas Ayuda

http://invention.swmed.edu/etblast/etblast.shtml

eT BLAST Biomedical Database Search System

Other Search Tools: [ARGH](#) [RIC](#) [FRISC](#) [TRITE](#)

Currently Searching: MEDLINE Advanced Search offers other databases.

Input a paragraph:

[Advanced Search](#)

OR

upload a "text only" file:

If you would like your results emailed to you, please enter an email address.
Your address will be kept strictly confidential, and will not be used for any other purpose.

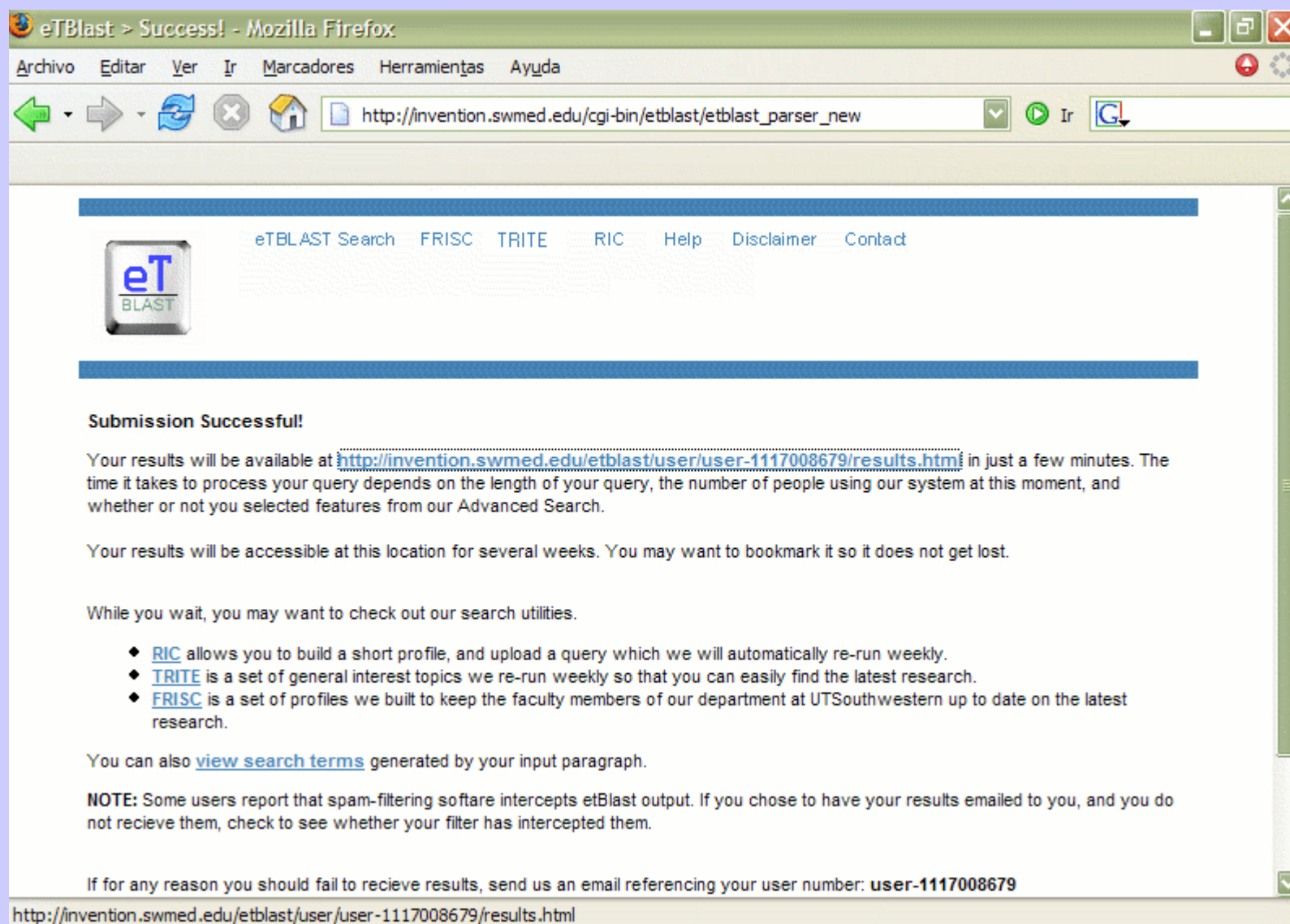
Optional Email:

Terminado

Science, May 14, 2004 issue. Under NetWatch, see the topic, "TOOLS: Just the Right Words".

Text mining biomedical literature (2005)

eTBlast (2)



eTBlast > Success! - Mozilla Firefox

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http://invention.swmed.edu/cgi-bin/etblast/etblast_parser_new

eTBlast Search FRISC TRITE RIC Help Disclaimer Contact

Submission Successful!

Your results will be available at <http://invention.swmed.edu/etblast/user/user-1117008679/results.html> in just a few minutes. The time it takes to process your query depends on the length of your query, the number of people using our system at this moment, and whether or not you selected features from our Advanced Search.

Your results will be accessible at this location for several weeks. You may want to bookmark it so it does not get lost.

While you wait, you may want to check out our search utilities.

- ◆ [RIC](#) allows you to build a short profile, and upload a query which we will automatically re-run weekly.
- ◆ [TRITE](#) is a set of general interest topics we re-run weekly so that you can easily find the latest research.
- ◆ [FRISC](#) is a set of profiles we built to keep the faculty members of our department at UTSouthwestern up to date on the latest research.

You can also [view search terms](#) generated by your input paragraph.

NOTE: Some users report that spam-filtering software intercepts etBlast output. If you chose to have your results emailed to you, and you do not receive them, check to see whether your filter has intercepted them.

If for any reason you should fail to receive results, send us an email referencing your user number: **user-1117008679**


<http://invention.swmed.edu/etblast/user/user-1117008679/results.html>

eTBlast (3)

Results of the search - Mozilla Firefox

Archivo Editor Ver Ir Marcadores Herramientas Ayuda

http://invention.swmed.edu/etblast/user/user-1117008679/results.html



[Submitted paragraph](#) | [Statistics Summary](#) | [Disclaimer](#)

Closest Matches:

- ☐ [Glycogen metabolism in quail embryo muscle. The role of the glycogenin primer and the intermediate proglycogen.](#)
J Lomako ... W J Whelan
Eur J Biochem 1995 Nov;234(1);343-9. Score: 61.496
- ☐ [A new look at the biogenesis of glycogen.](#)
M D Alonso ... W J Whelan
FASEB J 1995 Sep;9(12);1126-37. Score: 54.209
- ☐ [Glycogen synthesis in the astrocyte: from glycogenin to proglycogen to glycogen.](#)
J Lomako ... M D Norenberg
FASEB J 1993 Nov;7(14);1386-93. Score: 50.494
- ☐ [Further studies on the role of glycogenin in glycogen biosynthesis.](#)
C Smythe ... P Cohen
Eur J Biochem 1990 Apr;189(1);199-204. Score: 48.524
- ☐ [Glycogenin-dependent organization of Ascaris suum muscle glycogen.](#)

Terminado

eTBlast (4)

PMID: 8529663 - Mozilla Firefox

Archivo Editar Ver Ir Marcadores Herramientas Ayuda

http://invention.swmed.edu/cgi-bin/etblast/abstract_local?pmid=8529663&use

eTBLAST

Eur J Biochem 1995 Nov;234(1);343-9.

Glycogen metabolism in quail embryo muscle. The role of the glycogenin primer and the intermediate proglycogen.

J Lomako
W M Lomako
W J Whelan

Department of Biochemistry and Molecular Biology, University of Miami School of Medicine, FL 33101, USA.

Cultured quail embryo muscle has proven to be an excellent model system for studying the synthesis of macromolecular glycogen from, and its degradation to, glycogenin, the autocatalytic, self-glucosylating primer for glycogen synthesis. We recently demonstrated that proglycogen, a low-M(r) form of glycogen, is an intermediate in the synthesis. Here we show that proglycogen also functions as an intermediate in macroglycogen degradation and, in one set of circumstances, represents an arrest point in glycogen breakdown, which does not continue to glycogenin. We suggest that in the nutritionally dependent turnover of glycogen in tissues, the molecules cycle between proglycogen and macromolecular glycogen and are not normally degraded to glycogenin. Nevertheless, when this does happen, the released glycogenin is active, capable of re-initiating glycogen synthesis. Under culture conditions where the conversion of proglycogen into glycogenin does take place, the intermediates lying between form a discrete rather than a continuous series, suggestive of a cluster structure for proglycogen and indicating that breakdown is stepwise. Evidence of post-translational modification of glycogenin was obtained by the finding that, in glycogen from cultured muscle, glycogenin is phosphorylated.

MedlineID: 0
PMID: 8529663

IR performance

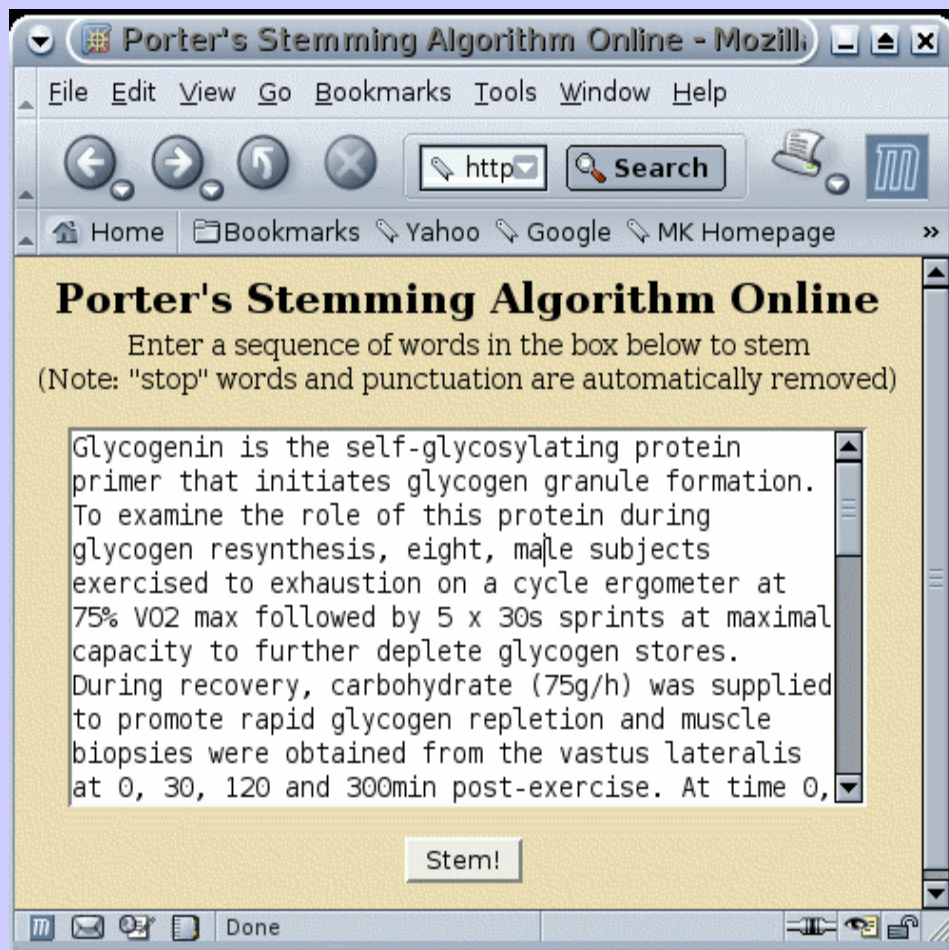
- **Precision:** fraction of relevant documents retrieved divided by the total returned documents
- **Recall:** proportion of relevant documents returned divided by the total number of relevant documents
- **F-score:** the harmonic mean of precision and recall
- Precision-recall curves

Information Extraction and Text mining

- Identification of **semantic structures** within free text.
- Use of syntactic and Part of Speech (POS) information.
- Integration of domain specific knowledge (e.g. ontologies).
- Identification of textual patterns.
- Extraction of predefined **entities** (NER), relations, **facts**.
- Entities like: companies, places or proteins, drugs.
- Relations like: protein interactions
- Methods: heuristics, rule-based systems, machine learning and statistical techniques, regular expressions,...

Stemming

Process of removing affixes of words transforming them to their corresponding morphological base form or root.



<http://maya.cs.depaul.edu/~classes/ds575/porter.html>

Original Word	Stemmed Word
glycogenin	glycogenin
selfglycosylating	selfglycosyl
protein	protein
primer	primer
initiates	initi
glycogen	glycogen
granule	granul
formation	format
examine	examin
role	role
protein	protein
during	dure
glycogen	glycogen
resynthesis	resynthesi
eight	eight

Text mining biomedical literature (2005)

POS-tagging

The screenshot shows a Mozilla browser window titled "POS Tagging - Mozilla". The address bar contains the URL "http://cpk.auc.dk/~tb/tagger/tagit.php". The browser's menu bar includes File, Edit, View, Go, Bookmarks, Tools, Window, and Help. The toolbar shows navigation buttons and a search icon. The bookmarks bar lists Home, Bookmarks, Yahoo, Google, MK Homepage, ORF, Zope on http://..., and PubMed.

The main content area has a blue header with the title "POS Tagging" and links for [UP], [ABOUT THE TAGGER], and [MORE ONLINE]. Below the header, it states "Copyright Tom Brøndsted March 2004, Test version 1.0". A note reads: "NB! Netscape and Mozilla users can toggle between this view and a view without tags by selecting 'view - use style - variant 3'".

The "Tagged text:" section displays the following sentence with POS tags in red: "Glycogenin<Noun, sing. or mass> is<Verb, 3rd person sing. present> the<Determiner> self-glycosylating<Adjective> protein<Noun, sing. or mass> primer<Noun, sing. or mass> that<Prep. or subord. Conjunction> initiates<Verb, 3rd person sing. present> glycogen<Verb sing. present, non-3rd.> granule<Noun, sing. or mass> formation.<Noun, sing. or mass>".

Below this, the "Input (English) text for POS-tagging" section contains a text box with the sentence: "Glycogenin is the self-glycosylating protein primer that initiates glycogen granule formation." and a "submit text" button.

A left sidebar contains a blue background with a logo and the URL "kom.aau.dk/~tb". It lists links for Top, Publications, Teaching, Projects, NLP Suite, Resources, Demos, Online, and Statistics. At the bottom of the sidebar is the email address "tom.brondsted.dk" and an "@ email" icon.

At the bottom center of the main content area is a "CHECK" link. The browser's status bar at the very bottom shows various icons.

Providing each word given a sentence with its corresponding part of speech label, e.g. whether it is a noun, verb, preposition, article, etc.

Question Answering (QA)

- Humans formulate questions using natural language.
- Example: *What are the molecular functions of Glycogenin?*.
- QA: **automatic generation of answers** to queries in form NL expressions from document collections.
- Most systems limited to generic literature or newswire.
- QA difficult: heterogeneous, poorly formalised domain, new scientific terms
- Ad hoc retrieval task of the TREC Genomics Track 2005.
- Galitsky system (semantic skeletons (SSK), logical programming).

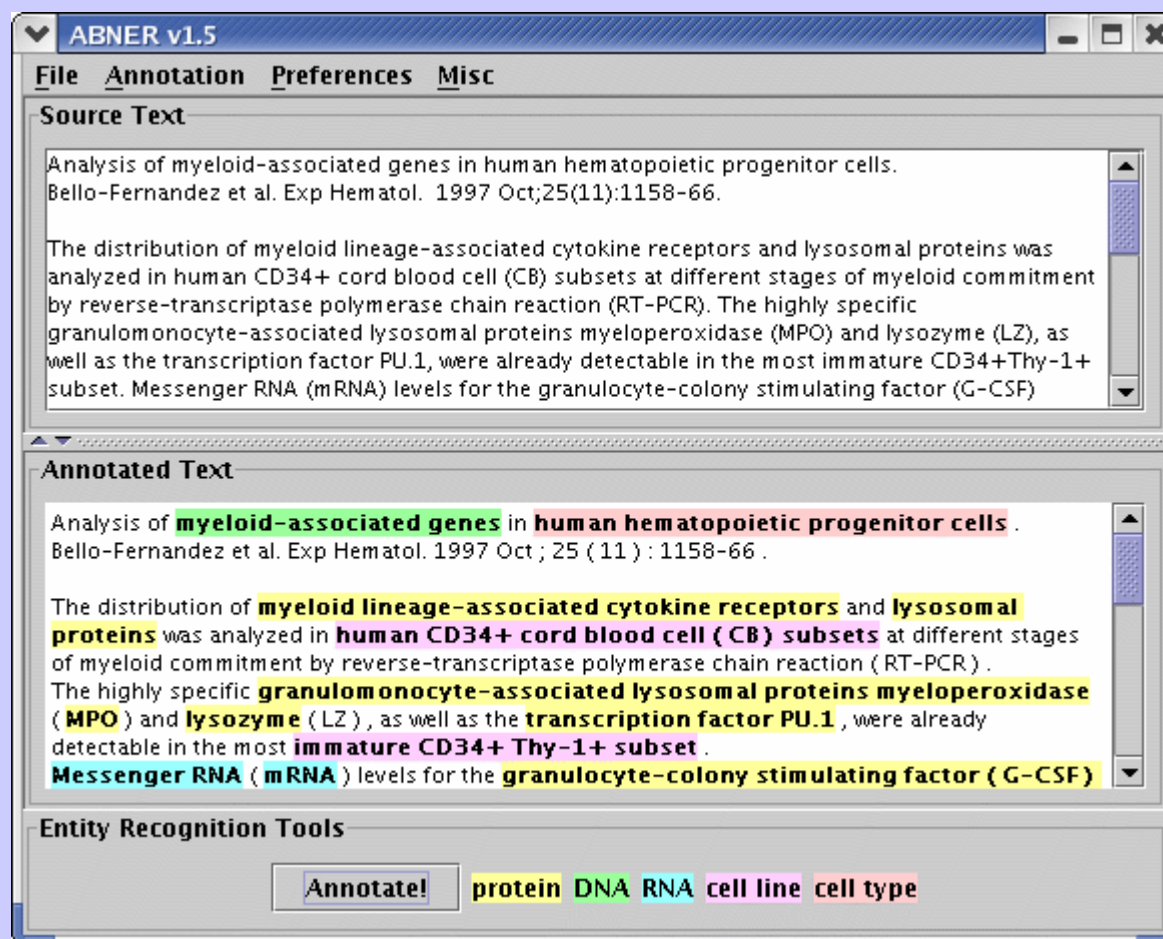
Natural Language Generation (NLG)

- NLG: constructing automatically natural language texts.
- Display the content of databases: reports, error messages.
- Based on semantic input, providing computer-internal representation of the information.
- Different degrees of complexity.
- Biology: modelling the domain language difficult.
- Simpathica/XSSYS trace analysis tool.

Named entity recognition (NER)

- **Identification of entity types** in textual data.
- Semantic tagging.
- Example identification of company names and places
- Mainly identification of proper nouns.
- NER in Molecular Biology: identification of genes, proteins, chemical compounds, diseases,...
- Methods: ad-hoc rule based systems,
- ML techniques (HMM,SVM,...), statistical tools.
- Tools: GAPSCORE, ABNER, AbGene, NLProt

ABNER



Burr Settles. "ABNER: A Biomedical Named Entity Recognizer."
<http://www.cs.wisc.edu/~bsettles/abner/>. 2004.

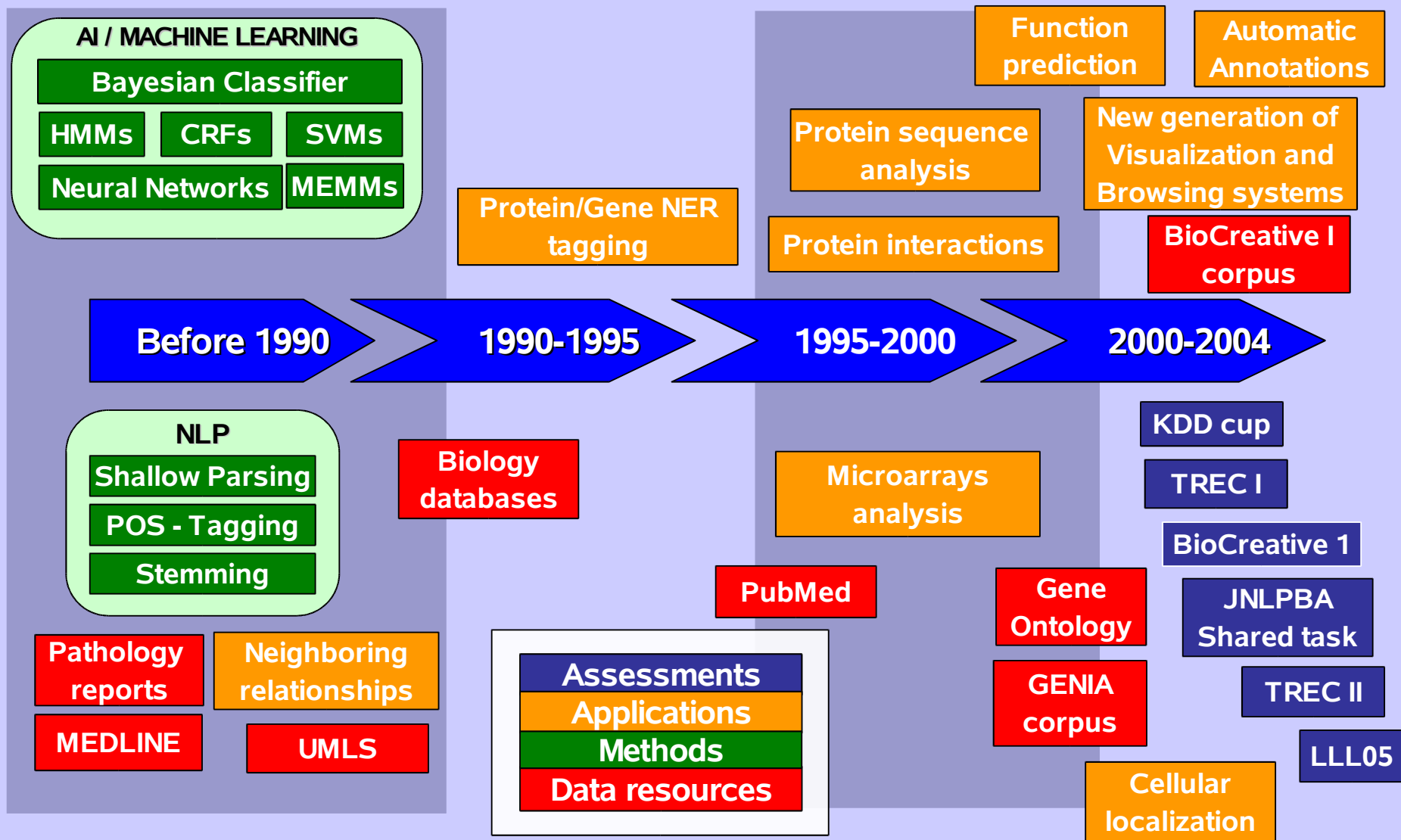
Basic NLP terms

Corpus: collection of documents.

POS tagging: labeling each word in a sentence with its part of speech (verb,noun,..) depending on its context.

Word sense disambiguation: assigning the semantic class (meaning) to a given word depending on its context.

NLP in Biomedicine – Timeline

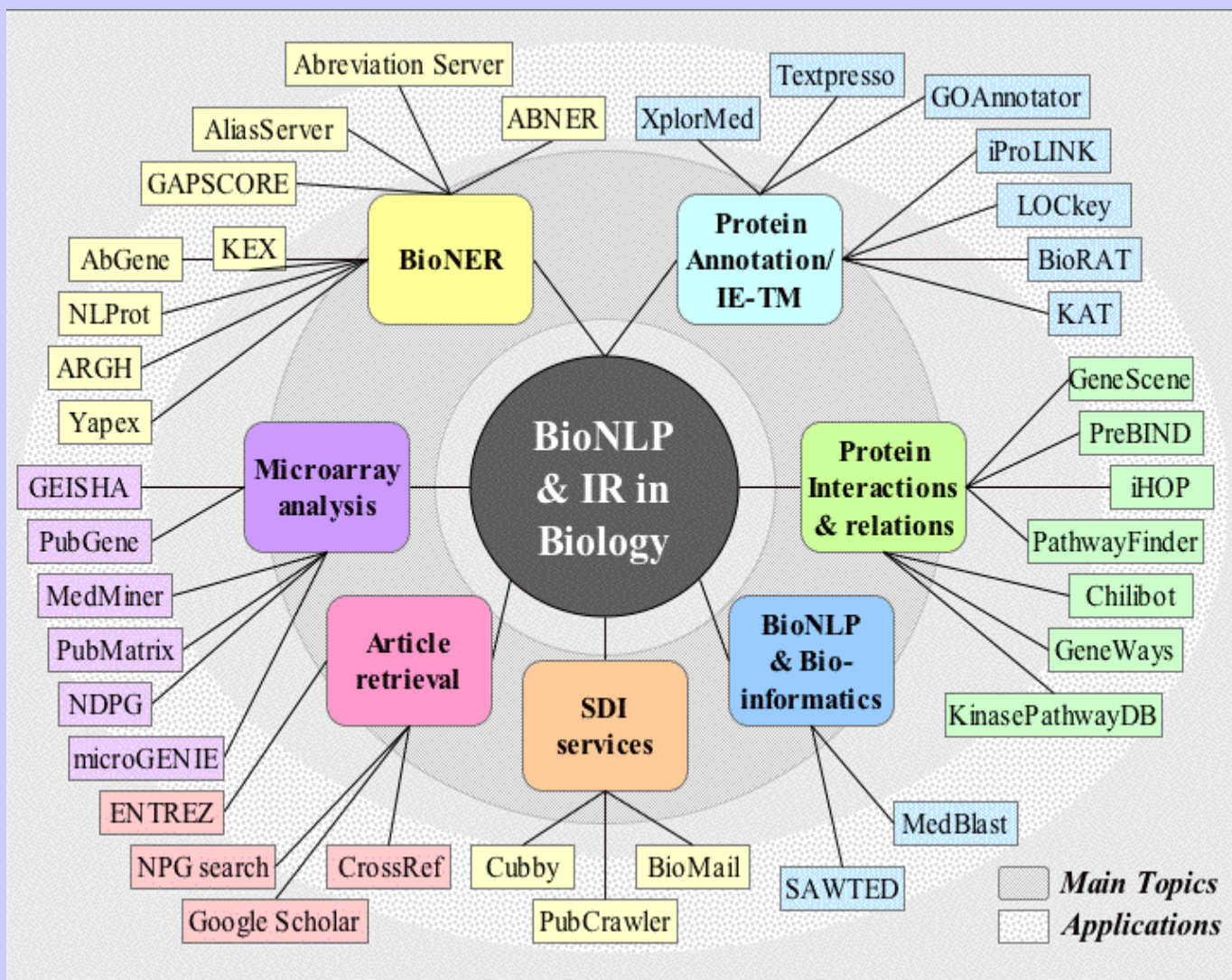


Text mining biomedical literature (2005)

Text mining applications in biology

- NER: tagging biological entities.
- Automatic annotation: associating proteins to functional descriptions.
- Protein interactions: extracting interactions of proteins, genes and drugs.
- Microarray analysis: providing biological context through literature mining
- Protein localisation
- Improving sequence-based homology detection.

Text mining applications in biology



Tagging Biological entities

Aim: **Identify** biological entities in articles and to **link** them to entries in biological databases.

- Generic NER: corporate names and places (0.9 f-score).
- Biology NER: more complex (synonyms, disambiguation, typographical variants, official symbols not used,...).
- Bioinformatics vs NLP approach.
- Performance organism dependent.
- Methods: POS tagging, rule-based, flexible matching, statistics, ML (naïve Bayes, ME, SVM, CRF, HMM).

GAPSCORE (1)

Gene and Protein Search - Mozilla Firefox

Archivo Editar Ver Ir Marcadores Herramientas Ayuda

http://acronym.stanford.

Search for gene and protein names in some text.

Glycogenin is the self-glycosylating protein primer that initiates glycogen granule formation. To examine the role of this protein during glycogen resynthesis, eight, male subjects exercised to exhaustion on a cycle ergometer at 75% VO2 max followed by 5 x 30s sprints at maximal capacity to further deplete glycogen stores. During recovery,

SEARCH

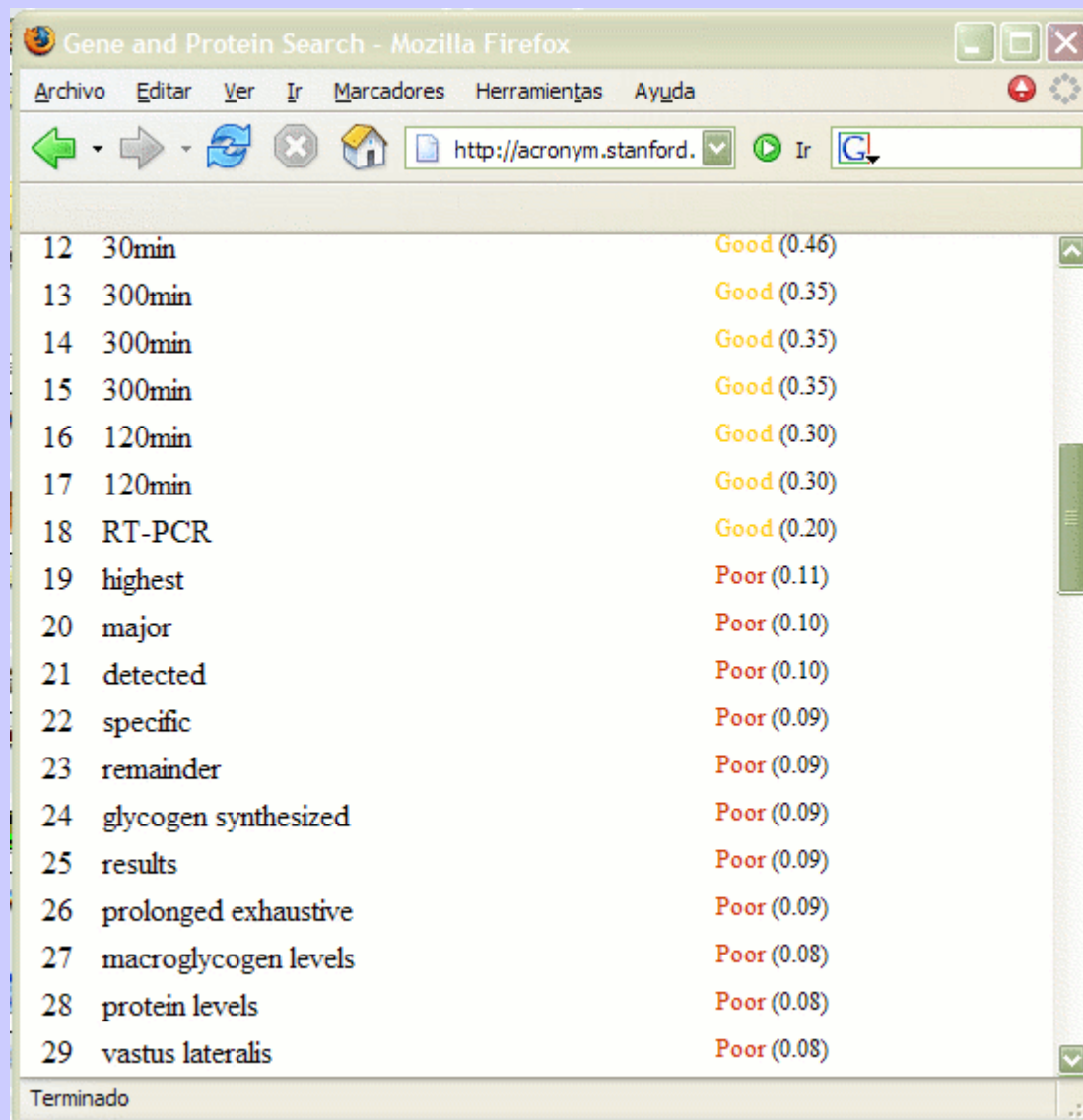
	Gene or Protein Name	Quality (Score)
1	75% VO2	Good (0.70)
2	Glycogenin	Good (0.67)
3	Glycogenin	Good (0.67)
4	Glycogenin	Good (0.67)
5	elevated glycogenin	Good (0.67)
6	free (deglycosylated) glycogenin	Good (0.67)
7	glycogenin	Good (0.67)
8	glycogenin	Good (0.67)
9	glycogenin	Good (0.67)
10	glycogenin	Good (0.67)

Terminado

- Scores words based on a statistical model of gene names
- Quantifies:
 - Appearance
 - Morphology
 - Context.
- Online.

<http://bionlp.stanford.edu/gapscore/>

GAPSCORE (2)



Item	Score
12 30min	Good (0.46)
13 300min	Good (0.35)
14 300min	Good (0.35)
15 300min	Good (0.35)
16 120min	Good (0.30)
17 120min	Good (0.30)
18 RT-PCR	Good (0.20)
19 highest	Poor (0.11)
20 major	Poor (0.10)
21 detected	Poor (0.10)
22 specific	Poor (0.09)
23 remainder	Poor (0.09)
24 glycogen synthesized	Poor (0.09)
25 results	Poor (0.09)
26 prolonged exhaustive	Poor (0.09)
27 macroglycogen levels	Poor (0.08)
28 protein levels	Poor (0.08)
29 vastus lateralis	Poor (0.08)

Terminado

- Choose cut-offs.
- Online.
- Based on Medline analysis
- Score new words using SVM
- Statistical analysis of PubMed words.

Chang JT, Schütze H, and Altman RB.
 GAPSCORE: Finding Gene and Protein
 Names One Word at a Time.
Bioinformatics. 2004 Jan 22;20(2):216-25.

NLProt

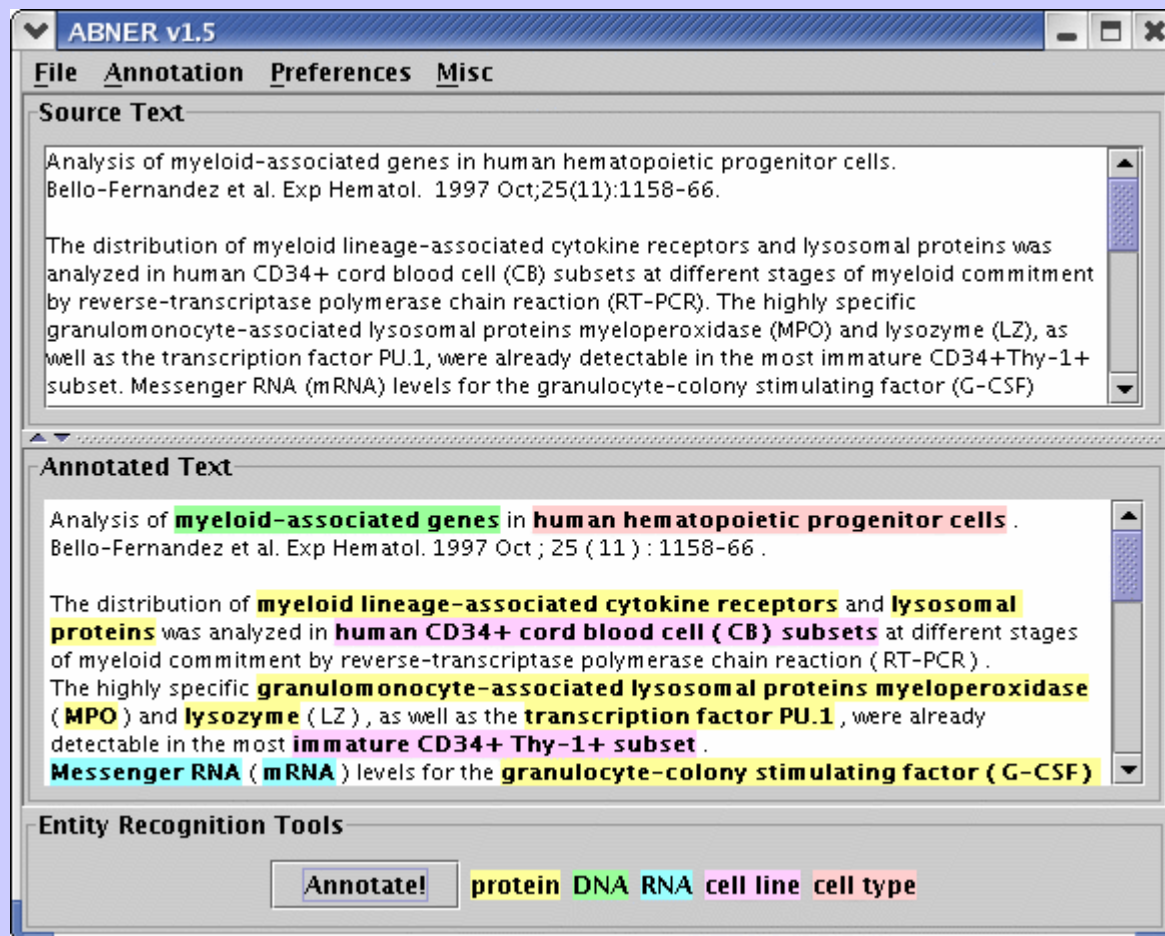
NAME	ORGANISM	TXT-POS	SCORE	METHOD	DB-ID (S)
Glycogenin	homo sapiens	1	1.040	SVM	GYG2 HUMAN (86%)
glycogenin	homo sapiens	96	0.856	SVM	GYG2 HUMAN (91%)
glycogenin	homo sapiens	103	1.040	SVM	GYG2 HUMAN (91%)
Glycogenin	homo sapiens	109	0.871	SVM	GYG2 HUMAN (86%)
glycogenin	homo sapiens	138	0.980	SVM	GYG2 HUMAN (91%)
Glycogenin	homo sapiens	157	0.971	SVM	GYG2 HUMAN (86%)
glycogenin	homo sapiens	161	0.311	SVM	GYG2 HUMAN (91%)
glycogenin	homo sapiens	214	0.819	SVM	GYG2 HUMAN (91%)
glycogenin	homo sapiens	234	0.747	SVM	GYG2 HUMAN (91%)

- Online (e-mail alert).
- Downloadable.
- SVM-based
- Pre-processing dictionary
- Rule-based filtering step
- PubMed words.
- Precision of 75%
- Recall of 76%

<http://cubic.bioc.columbia.edu/services/nlprot/>

Chang JT, Schutze H, Altman RB. GAPSCORE: finding gene and protein names one word at a time. *Bioinformatics*. 2004 Jan 22;20(2):216-25.

ABNER



- A Biomedical Named Entity Recogniser
- Downloadable.
- CRF-based
- Trained on BioCreative and GENIA
- orthographic and contextual features
- Can be trained on new corpora

Burr Settles. "ABNER: A Biomedical Named Entity Recognizer."
<http://www.cs.wisc.edu/~bsettles/abner/>. 2004.

Extracting functional annotations

- **Manual annotation** extraction by database curators.
 - Scientific literature analysis.
 - Time-consuming & labour-intensive.
 - Example: Gene Ontology annotation (GOA).

- **Text mining** to assist annotation extraction:
 - Identification of annotation relevant sentences.
 - Identification of protein-term associations.

Function extraction – applications

- Andrade et. (1997)
- iHOP
- Textpresso system
- Gene Information System (GIS)
- Medical Knowledge Explorer (MeKE)
- GO engine,...

Andrade et al. (1997)

- Extracts sentences from PubMed which contain functional information.
- Statistical analysis of the word frequencies.
- Analysis in protein families.
- Background frequencies of those words.

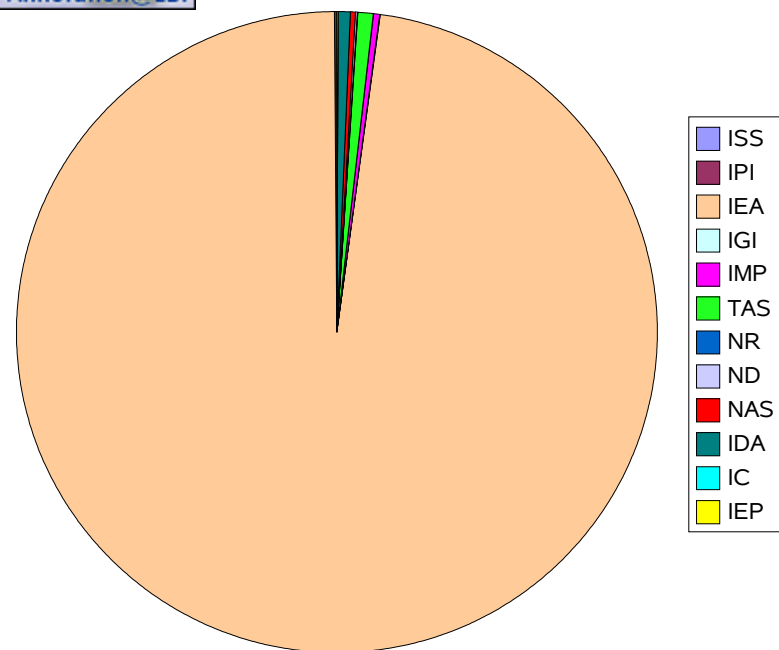
GENE ONTOLOGY (GO)

- Ontology direct acyclic graph structure.
- Controlled vocabulary of concepts.
- Three main categories:
 - Molecular Function
 - Cellular Component
 - Biological Process
- Describe relevant biological aspects of gene products
- Synonyms, links to external keywords.
- Currently most important source annotation terms.

<http://www.geneontology.org/>



Gene Ontology Annotation



Ev.C.	Annot	Perc.	
IEA	6421817	0.97529	Electronic/
ISS	19576	0.00297	sequence-
NR	2191	0.00033	based
ND	4433	0.00067	annotation
IPI	7130	0.00108	
IGI	3014	0.00046	Experimental
IMP	19072	0.00290	evidence
IDA	38862	0.00590	
IEP	1495	0.00023	
IC	831	0.00013	
TAS	49630	0.00754	Curator
NAS	16456	0.00250	knowledge

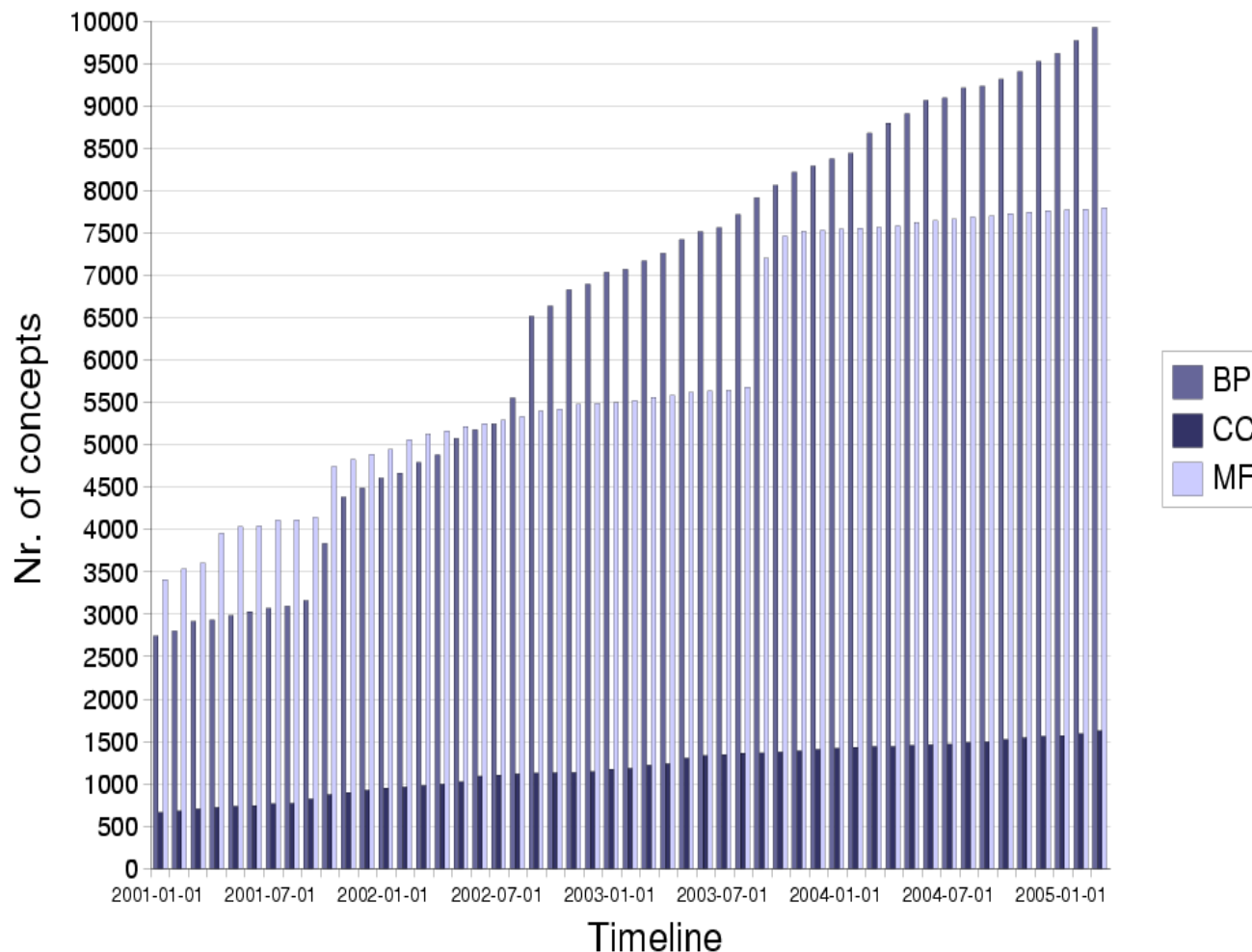
TAS: Traceable Author Statement; IDA: Inferred by direct assay; IC: Inferred by curator ; ND:No data;
 IMP:Inferred from mutant phenotype; IGI: Inferred from genetic interaction; 3.8) IPI :Inferred from physical
 interaction; ISS: Inferred from sequence similarity; IEP: Inferred from expression pattern; NAS: Non traceable
 author statement; IEA: Inferred by electronic annotation; NR: Not recorded;

<http://www.ebi.ac.uk/GOA/> 04/22/05

Text mining biomedical literature (2005)

Gene Ontology Growth

GO growth



- MF: Molecular Function
- CC: Cellular Component
- BP: Biological Process

iHOP

- Protein centric: nucleates the literature around protein name.
- For a range of model organisms (e.g. Human, yeast,..)
- Hyperlinks proteins through co-occurrence
- Highlight direct associations between proteins and functional terms.
- Online, fast, easy to use.


Hoffmann R, Valencia A. A gene network for navigating the literature *Nat Genet.* 2004 Jul;36(7):664.

iHOP

iHOP - Information Hyperlinked over Proteins - Mozilla Firefox

Archivo Editar Ver Ir Marcadores Herramientas Ayuda

http://www.pdg.cnb.uam.es/UniPub/iHOP/



iHOP

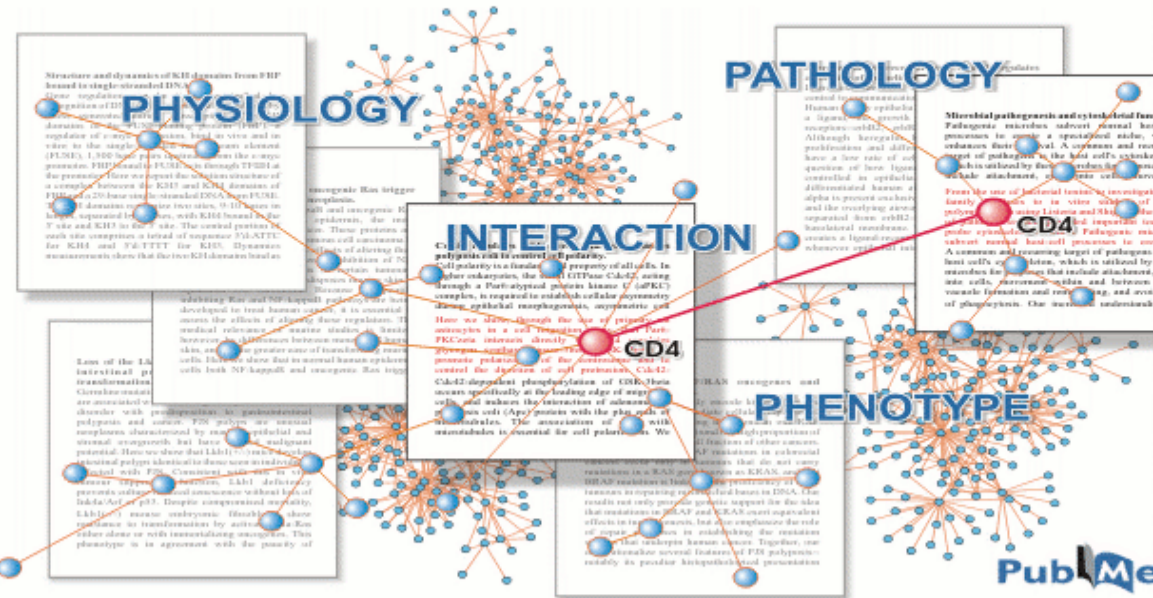
information hyperlinked
over proteins

Search Gene

Gene Model
Developer's Zone **new**
Contact
Help

COCON

Concept & Implementation



PHYSIOLOGY

INTERACTION

PATHOLOGY

PHENOTYPE

PublMe

Hoffmann, R., Valencia, A. A Gene Network for Navigating the Literature. *Nature Genetics* 36, 664 (2004)

Search for a gene synonym or accession number...

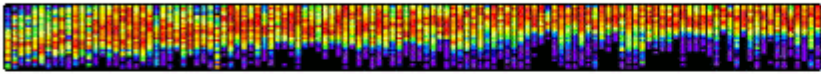
Terminado

iHOP

iHOP - Information Hyperlinked over Proteins - Mozilla Firefox

Archivo Editar Ver Ir Marcadores Herramientas Ayuda

http://www.pdg.cnb.uam.es/UniPub/iHOP/index.html?field=all&search=glycog

Symbol	Name	Synonym/ DB-reference	Organism
 Life cycles of successful genes			
GYG	glycogenin-1		Homo sapiens
GYG2	glycogenin-2		Homo sapiens
GYG2P	glycogenin 2 pseudogene		Homo sapiens
TRIM7	Tripartite motif protein 7	glycogenininteracting protein	Homo sapiens
Gyg1	glycogenin-1		Mus musculus
glycogenin	glycogenin		Drosophila melanogaster
4J165	glycogenin family member (4J165)		Caenorhabditis elegans
5R226	glycogenin family member (33.6 kD) (5R226)		Caenorhabditis elegans
At1g08990	glycogenin glucosyltransferase (glycogenin)-related		Arabidopsis thaliana
At1g54940	glycogenin glucosyltransferase (glycogenin)-related		Arabidopsis thaliana
At1g77120	glycogenin glucosyltransferase		Arabidopsis thaliana

Terminado

iHOP - Information Hyperlinked over Proteins [GYG] - Mozilla

File Edit View Go Bookmarks Tools Window Help

http://www.pdg.cnb.uam.es/UniPub/iHOP/gs/88913.html?IN=1

Home Bookmarks Yahoo Google MK Homepage ORF Zope on http://... PubMed Python Zope PyTut OEAW GeneDic biocreative GenomeNet

iHOP
Information hyperlinked
Over Proteins

Search Gene

Show overview **new**
Find in this Page

Filter and options
Gene Model

Developer's Zone
new
Help

Concept & Implementation
by Robert Hoffmann

Symbol	Name	Synonyms	Organism
GYG	Glycogenin-1	glycogenin, GYG1	Homo sapiens
UniProt	P46976, Q9UNV0		
OMIM	603942		
NCBI Gene	2992		
NCBI RefSeq	NP_004121		
NCBI Accession	AAB00114, AAB09752, AAD31084		

Homologues of GYG ... new

Definitions for GYG ...

Enhanced PubMed/Google query ... new

WARNING: Please keep in mind that gene detection is done automatically and can exhibit a certain error. [Read more.](#)

Find in this Page

Mutation of Tyr-196 in [glycogenin-2](#) to a Phe residue abolished the ability of [glycogenin-2](#) to self-glucosylate but not to **interact** with **glycogenin-1**.

Mutational analysis of the coding regions of the genes encoding protein kinase B-alpha and -beta, phosphoinositide-dependent protein kinase-1, phosphatase targeting to [glycogen](#), [protein phosphatase inhibitor-1](#), and [glycogenin](#): lessons from a search for genetic variability of the insulin-stimulated [glycogen](#) synthesis pathway of [skeletal muscle](#) in [NIDDM](#) patients.

Effects of [exercise](#) on [GLUT-4](#) and [glycogenin](#) gene expression in human [skeletal muscle](#).

The third [cDNA](#) encoded a polypeptide of unknown function and was designated [GNIP](#) ([glycogenin](#) interacting protein).

[GNIP](#), a novel protein that binds and activates [glycogenin](#), the self-glucosylating initiator of [glycogen](#) biosynthesis.

Overall, [GN-2](#) has 40-45% identity to muscle [glycogenin](#) but is 72% identical over a 200-residue segment thought to contain the catalytic domain.

[Glycogenin-1](#) and [glycogenin-2](#) interact with one another, based on in vitro interactions and co-immunoprecipitation from liver and cell extracts.

Mouse [glycogenin-1](#) has a predicted molecular mass of 37kDa, and the deduced amino acid sequence exhibited 87% homology with human [glycogenin-1](#).

For the first time, we report that a single bout of [exercise](#) is sufficient to cause upregulation of [GLUT-4](#) and [glycogenin](#) gene expression in human [skeletal muscle](#).

Fasting plasma [insulin](#) concentrations, muscle [creatine](#), [glycogen](#) and [GLUT-4](#) protein content as well as GLUT-4, [glycogen](#) synthase-1 (GS-1) and [glycogenin-1](#) (Gln-1) [mRNA](#) expression were determined.

In conclusion, the co-expression of [glycogenin](#) with [GLUT3](#) might enable glycogen-storing cells to exchange glucose quite effectively according to prevailing metabolic demands of glycogen synthesis or degradation.

The discovery of a second human gene, [GYG2](#), encoding a liver-specific isoform of [glycogenin](#), the self-glucosylating initiator of [glycogen](#) biosynthesis, raised the possibility for differential controls of this protein in [liver](#) and muscle.

The present study investigated the expression of [glycogenin](#), the protein primer for glycogen synthesis, and the high affinity glucose transporter isoform [GLUT3](#) as a further potential regulator of cellular glycogen metabolism, in first trimester and term human placenta using immunohistochemistry and

Transferring data from www.pdg.cnb.uam.es...

Textpresso

Textpresso - Mozilla Firefox

Archivo Editar Ver Ir Marcadores Herramientas Ayuda

http://www.textpresso.org/

Home

Simple Retrieval

Query should be met in a ☒ sentence ☐ publication.

Type in keywords to be searched for, separated by **white spaces**:

☐ Exact match

Specify categories that should also be met (optional):

,

Search in

☒ Abstract ☐ Author ☒ Full ☐ Title ☐ Year

News and Messages

Welcome to *Textpresso* ! The *Textpresso* search engine for *C. elegans* abstracts and fulltexts was developed at [Wormbase](#) to service the *C. elegans* community, and is being expanded to serve other communities.

● **March 21st, 2005:** A new build that contains over 1,000 new full text papers and an updated Textpresso ontology (version 1.1) is released. The new ontology version has four new categories; Reporter Gene, Restriction Enzyme, Second Messenger and Vector. In addition, new terms have been added to the Drugs and Small Molecules and Organism categories. Details of the updated ontology can be found by following the "Ontology" link on the menu to the left of this page.

● **March 1st, 2005:** We are current testing a new build that contains 4 new categories and over 1,000 new full text papers on the [Development Site](#). This site will be under a heavy barrage of testing and may experience

Terminado

Gene Information System (GIS)

- Focus on 4 types of gene-related info:
 - Biological function
 - Associated disease
 - Related genes
 - Gene-gene relations
- Gene information screening
- Gene-gene relation extraction.
- Downloadable

<http://iir.csie.ncku.edu.tw/~yuhc/gis/>

GIS

Genes' Information Discovery System

File(F) Edit(E) View(V) System Function(S) Dictionary(D) Experiment(P) Help(H)

G **GG** **G-G** **K** **D** **G** **R** **S** **N**

Gene Memo Conclusion

Doc. in PubMed Doc. retrieved Conclusion sent.

Bio-Function & Disease & Gene List

Biological Function list	Disease list	Gene list
phosphorylation 11	breast cancer 2	akt 10
synthesis 3	prostate cancer 1	erk 2
receptor 3		enos 2
protein synthesis 3		src 1
proliferate 3		sh3 1
transduce 2		ras 1
transcript 2		pkc 1
cell-death 2		pka 1
secretion 1		p38 1
oncogene 1		mapk 1

Sentence Information

Num. of sent. PMID Sent. count Pre. Next

Add sample1 Add sample2 Add sample3 View doc.

This survival signal appears to be mediated by activation of the **akt**, as detected by its phosphorylation in ser473 upon **cd53** ligation.

Chiang JH, Yu HC, Hsu HJ. GIS: a biomedical text-mining system for gene information discovery. *Bioinformatics*. 2004 Jan 1;20(1):120-1.

Annotate a protein sequence from a set of papers - Mozilla

File Edit View Go Bookmarks Tools Window Help

http://www.bork.embl-heidelberg.de/kat/ Search

Home Bookmarks Yahoo Google MK Homepage ORF Zope on http://... PubMed »

Annotate a protein sequence from a set of scientific references

[ABOUT](#) [SUPPLEMENT](#) [GLOSSARY](#)

Input several PMID identifiers (a maximum of 10) or a SwissProt identifier
 glycogenin
 (example: 3538478 3774547 3510187) (example: TETX_CLOTE)

<input checked="" type="checkbox"/> Derive SwissProt keywords from the MeSH terms of the abstracts	<input checked="" type="checkbox"/> Derive SwissProt Keywords from the words of the abstracts	<input checked="" type="checkbox"/> Derive Gene Ontology terms from the MeSH terms of the abstracts
Inclusion value >= 0.8 Support >= 5	Inclusion value >= 0.8 Support >= 20	Inclusion value >= 0.8 Support >= 20

Using lower thresholds on the [inclusion](#) and [support](#) values you will obtain more [keywords](#) or [GO-terms](#) but they will not be so reliable. See supplementary material for details.

Annotate Clear

[Bork Group](#) at [EMBL-Heidelberg](#) and [Genetics Dept.](#) of the [Universidad de Málaga](#)

Keyword Annotation Tool (KAT)

- Extraction of mappings between related terms using a model of fuzzy associations
- Mesh terms/SwissProt keywords/GO terms

Perez AJ, Perez-Iratxeta C, Bork P, Thode G, Andrade MA. Gene annotation from scientific literature using mappings between keyword systems. *Bioinformatics*. 2004 Sep 1;20(13):2084-91. Epub 2004 Apr 1.

Medical Knowledge Explorer (MeKE)

- Ontology-based text mining system.
- Methods of sentence alignment.
- Sentence classification methods.
- Flexible matching, stemming and indexing.
- Create new GO-term synonyms from text.
- Edit distance calculation
- Learn sentence motifs via sentence alignment
- Naïve Bayes sentence classifier

<http://gen.csie.ncku.edu.tw/meke3/>

Chiang JH, Yu HC. MeKE: discovering the functions of gene products from biomedical literature via sentence alignment. *Bioinformatics*. 2003 Jul 22;19(11):1417-22

GO engine

- Computational platform for GO annotation.
- Correlation of text info with specific GO nodes.
- Combines: homology info, protein clustering and text analysis.
- Calculate frequency of association of terms to GO nodes.

Xie H, Wasserman A, Levine Z, Novik A, Grebinskiy V, Shoshan A, Mintz L. Large-scale protein annotation through gene ontology. Genome Res. 2002 May;12(5):785-94.

Protein interactions

- Advances in experimental large scale protein interaction analysis
- Exp. Methods for protein interaction characterization:
 - protein arrays
 - mRNA expression microarrays
 - Yeast two-hybrid
 - Affinity purification with MS
 - X-ray, NMRFRET, chemical cross-linking,...
- Bioinformatics methods for protein characterization:
 - Genome-based
 - Sequence-based

Protein interaction databases

Database Name	Reference	URL
BIND	(Bader <i>et al.</i> , 2003)	http://bind.ca
DIP	(Xenarios <i>et al.</i> , 2002)	http://dip.doe-mbi.ucla.edu
GRID	(Breitkreutz <i>et al.</i> , 2003)	http://biodata.mshri.on.ca/grid
HPID	(Han <i>et al.</i> , 2004)	http://www.hpid.org
HPRD	(Peri <i>et al.</i> , 2004)	http://www.hprd.org
IntAct	(Hermjakob <i>et al.</i> , 2004)	http://www.ebi.ac.uk/intact
MINT	(Zanzoni <i>et al.</i> , 2002)	http://cbm.bio.uniroma2.it/mint
STRING	(vonMering <i>et al.</i> , 2003)	http://string.embl.de
ECID	(Juan <i>et al.</i> , 2004)	http://www.pdg.cnb.uam.es/ECID

Text mining and Protein interactions

- Extract automatically those interactions from articles.
- NL used to characterise the nature of the interaction and its directionality.
- Literature-derived interaction networks:
 - power law distribution
 - scale free topology
- Visualised using network graphs.
- Methods range from: simple occurrence, expert derived word patterns (frames) to machine learning.

PubGene

- Use the co-occurrence of protein and gene names.
- Assumption: co-occurrence imply biological relationship
- Indexing PubMed abstracts and titles with human proteins.
- Construction of interaction networks.
- Build upon binary interactions between co-occurring proteins

Jenssen TK, Laegreid A, Komorowski J, Hovig E. A literature network of human genes for high-throughput analysis of gene expression. Nat Genet. 2001 May;28(1):21-8.

<http://www.pubgene.org/>

iHOP - Information Hyperlinked over Proteins / Gene Model - Moz


iHOP
information hyperlinked
over proteins

Search Gene

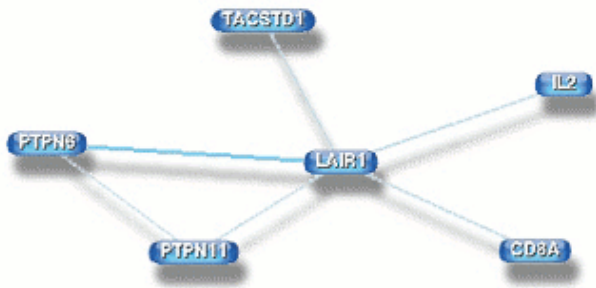
Save/ Load **new**
Help

Concept & Implementation
by Robert Hoffmann

Gene Model - the logbook

In the course of your navigation through iHOP, interesting sentences can be added to the *Gene Model* by clicking on the  icon beside the sentence. The Gene Model stores these sentences and represents their relation in a graph. [More about the Gene Model...](#)

e.g.



```
graph TD; LAIR1 --- TACSTD1; LAIR1 --- IL2; LAIR1 --- CD8A; LAIR1 --- PTPN11; LAIR1 --- PTPN6;
```

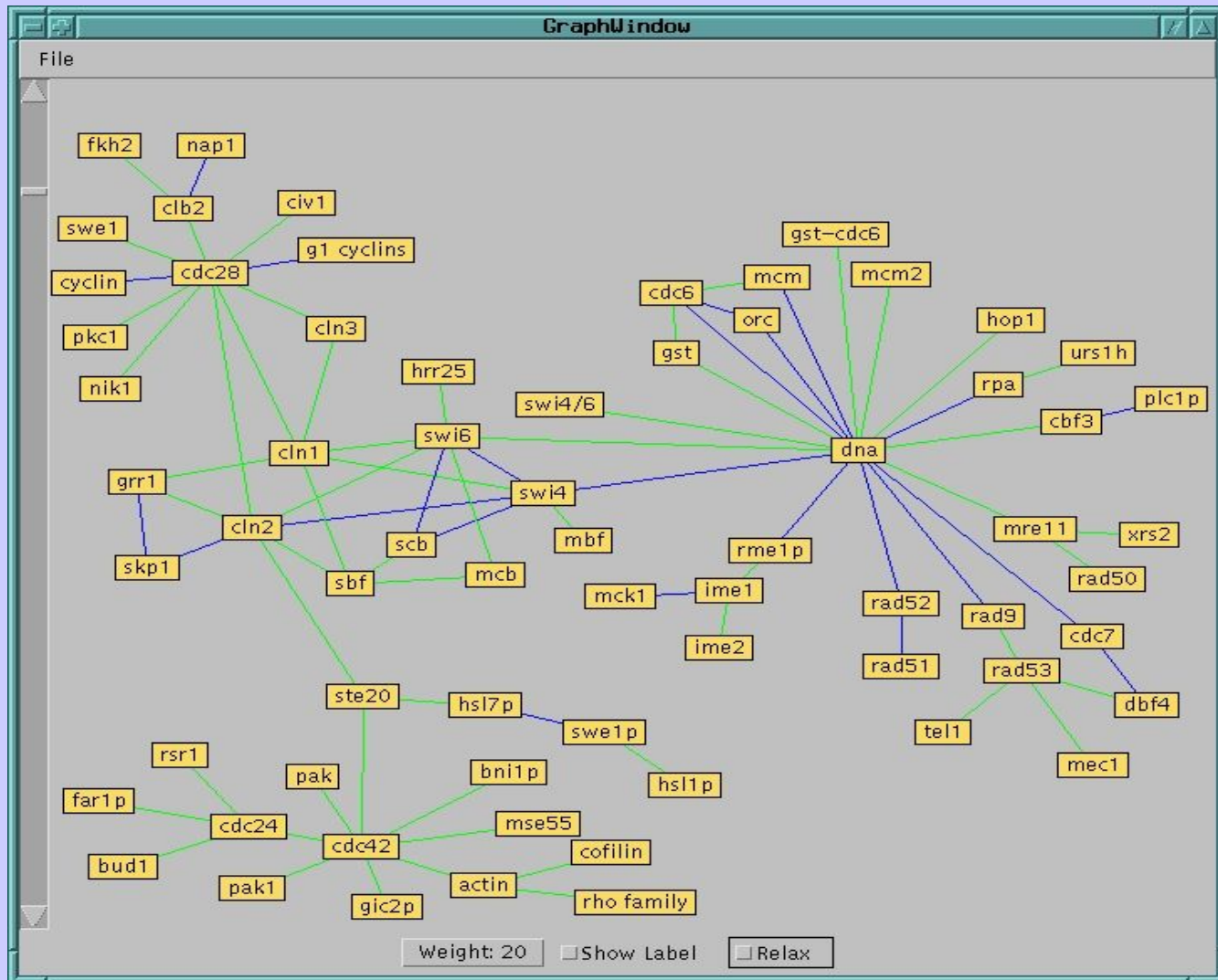
The diagram illustrates a network graph where LAIR1 is the central hub, connected to five other proteins: TACSTD1, IL2, CD8A, PTPN11, and PTPN6. The connections are represented by lines, indicating interactions between the proteins.

iHOP:
Visualization
of protein
interactions
using network
graphs

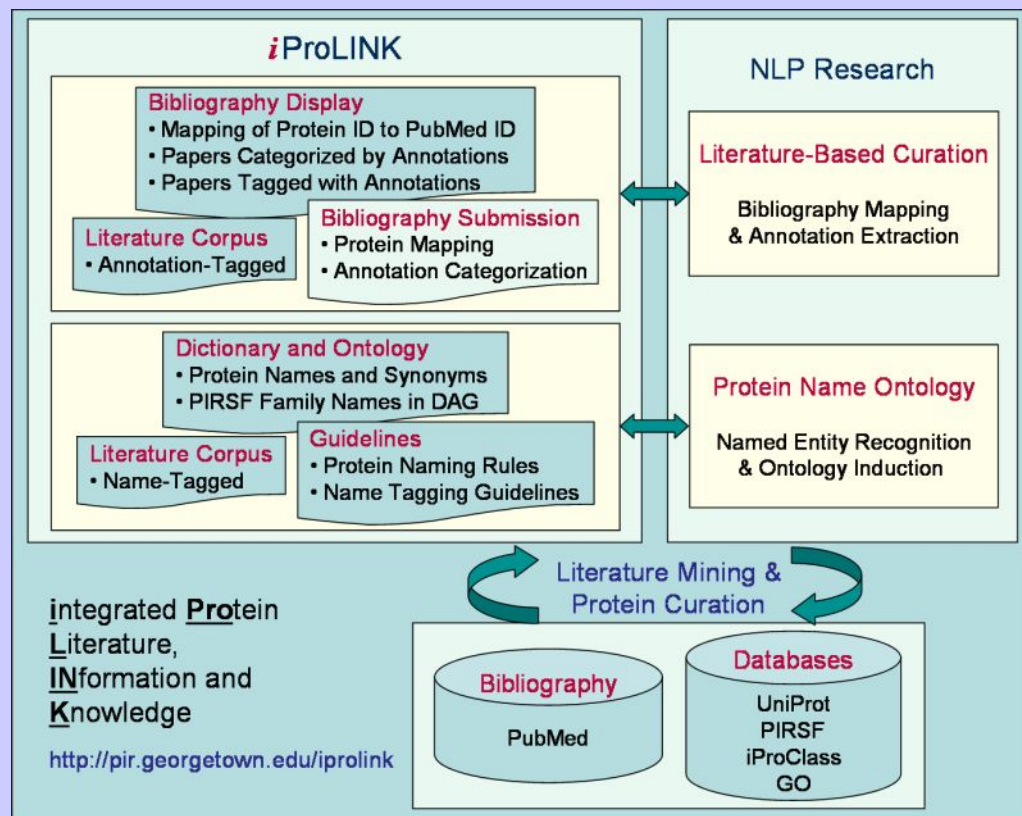
SUISEKI

- Relationship between the co-occurring proteins using **frames**
- Frames: **textual patterns** used to express interactions
- Initial set of 14 interaction words based on domain knowledge.
- Examples: *activate*, *bind*, *suppress*
- Analysed the **order** of protein names within sentences.
- Take into account **distance** (off-set) between protein names.
- System effective for simple interaction types.
- Difficult cases: long sentences with complex grammatical structures

SUISEKI interaction network



iProLINK



- Mapped citations
- Annotation tagged literature corpora
- Data source for protein name ontology development

<http://pir.georgetown.edu/iprolink/>

Hu ZZ, Narayanaswamy M, Ravikumar KE, Vijay-Shanker K, Wu CH. Literature mining and database annotation of protein phosphorylation using a rule-based system.

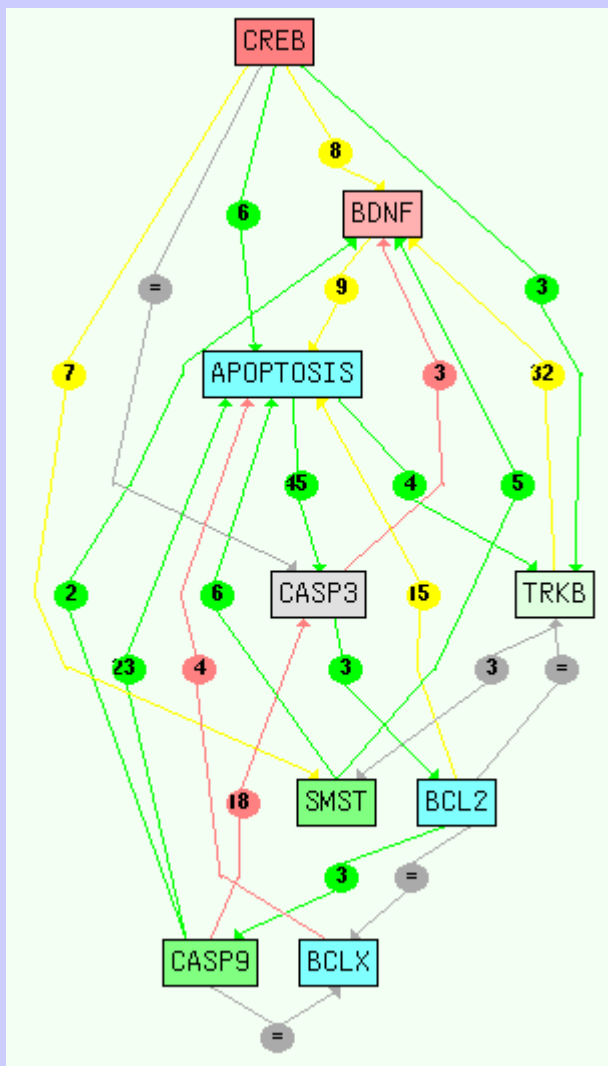
Bioinformatics. 2005 Jun 1;21(11):2759-65. Epub 2005 Apr 6

Chilibot

- NLP-based text mining approach.
- Content-rich relationship networks among biological
- Concepts, genes, proteins or drugs.
- Nature of the relationship: inhibitory, stimulative, neutral and simple co-occurrence.
- Internet-based application with graphical visualisation
- Sentence as unit, POS tagging, shallow parsing and rules.

Chen H, Sharp BM. Content-rich biological network constructed by mining PubMed abstracts. BMC Bioinformatics. 2004 Oct 8;5(1):147.

<http://www.chilibot.net/>



Chilibot (2)

- Need registration.
- Hypothesis generation.

Chen H, Sharp BM.

Content-rich biological network constructed by mining PubMed abstracts.

BMC Bioinformatics. 2004 Oct 8;5(1):147.

<http://www.chilibot.net/>

Welcome to Blueprint - Mozilla

File Edit View Go Bookmarks Tools Window Help

http://www.blueprint.org/prc Search

Home Bookmarks Yahoo Google MK Homepage ORF Zope on http://... »

Blueprint MOUNT SINAI HOSPITAL

News	Search	Contact Us	Products	Exhibits	Help
About	BIND	SeqHound	Services	Research	Jobs

PreBIND Search

Search for a protein by name and organism:
Enter the name of a protein (one word) and the organism it comes from.

Any organism

For example, type "Ras1p" and select *Saccharomyces cerevisiae*.

OR

Search for a protein by accession:
Enter the REFSEQ accession number of a protein.

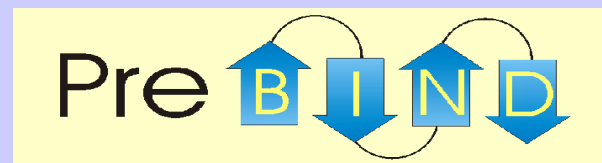
For example, type "NP_014744" to find interaction papers about Ras1p of yeast. Look for REFSEQ accession numbers [here](#).

OR

Search for a PubMed paper by PMID:
Enter the PubMed Identifier (PMID) of the paper.

For example, type "10747882" to view a list of potential interactions in this paper.

[Policies](#)



- Based on SVM.
- Query protein or accession number.
- Assist the Biomolecular Interaction Network Database (BIND)

Donaldson I, Martin J, de Bruijn B, Wolting C, Lay V, Tuekam B, Zhang S, Baskin B, Bader GD, Michalickova K, Pawson T, Hogue CW. PreBIND and Textomy—mining the biomedical literature for protein-protein interactions using a support vector machine. *BMC Bioinformatics*. 2003 Mar 27;4(1):11.

<http://www.blueprint.org/products/prebind>

Microarray data analysis

- Co-ordinated expression of genes.
- Functional co-regulation within biological processes.
- Mine micro array data using the associated biomedical literature.
- Characterise groups of genes extracting functional keywords.
- Score the coherence of gene clusters.
- Group genes based on their associated literature and functional descriptions.

GEISHA

- Text mining tool for microarray analysis.
- Analyse the correlation between:
 - the increase of the level of expression patterns and
 - the significance of functional information derived from the literature.
- Extract functional information from the literature linked to the microarray genes.
- Calculates statistical significance of terms from documents associated to genes of each cluster.

Protein localization

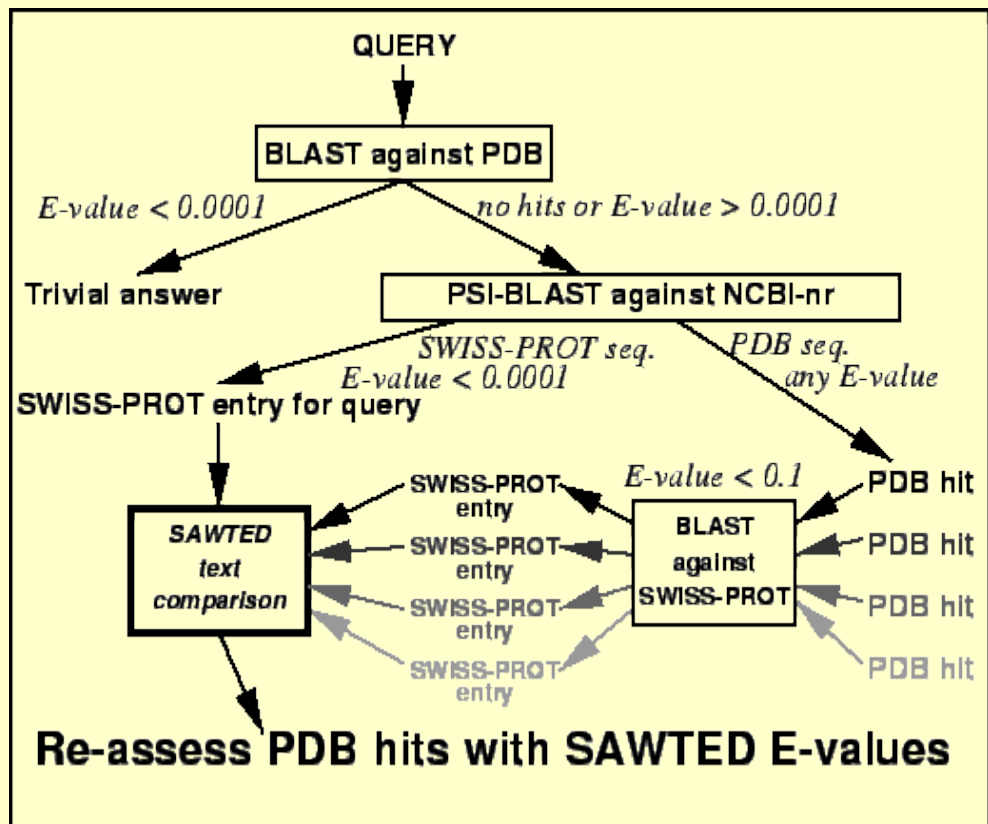
- Protein activity -> specific cellular environments.
- Localisation determination:
 - Experimental techniques.
 - Bioinformatics techniques (PSORT).
 - Text mining.
- Nair and Rost: lexical information in annotation database records.
- Stapley et al: Use SVM to classify proteins according to their subcellular localisation, extracted from PubMed abstracts.

NLP and sequence analysis: MedBlast

- Use NLP techniques to retrieve the related articles for a given sequence (online).
- Related articles:
 - those describing the query sequence (protein) or
 - Its redundant sequences and close homologues
- Direct search with the sequence.
- Indirect search with gene symbols.
- Use Blast against GenBank.
- Use Eutilities toolset to retrieve documents

<http://medblast.sibsnet.org/>

NLP and sequence analysis: SAWTED



Sequence similarity
the base for identifying
structure templates
for query sequence

Structure Assignment
With Text Description

Document comparison
algorithms

<http://www.bmm.icnet.uk/~sawted/>

SAWTED Web Server - Mozilla

File Edit View Go Bookmarks Tools Window Help

http://www.bmm.icnet.uk/servers/sawtec Search

Home Bookmarks Yahoo Google MK Homepage ORF Zope on http://... PubMed

SAWTED

[home](#) [submit sequence](#) [bmm](#) [help](#)

SAWTED PSI-BLAST web server submission form

your **email address** (format: xyz@abc.def...) your **reference** for this submission

describe the **function** of your protein in English * give **keywords** for your protein *

paste the amino acid **sequence** here (in one-letter code - *no headers, no numbers* - just the sequence)

maximum number of **iterations** **-h option**

please note that sequences longer than 800 residues will get no more than 2 iterations this is the E-value threshold for inclusion into the PSI-BLAST iterated profile

please complete all shaded areas * needed if no SWISS-PROT homologue can be found for your query
results will be returned by email

please note that this server only reports hits against the PDB

copyright [ICRF](#) 1999 [disclaimer](#) server admin: maccallr @ cancer.org.uk

Use information
contained in text
descriptions of
SwissProt
annotations

identification of
remote homologues

<http://www.bmm.icnet.uk/~sawted/>

Community wide evaluations

BIOINFORMATICS

CASP

CAMDA

CAPRI

GASP

GAW

PTC

BIO-NLP

KDD cup

BioCreative

JNLPBA shared task

TREC Genomics track

LLL05 challenge

NLP/IR/IE

MUC

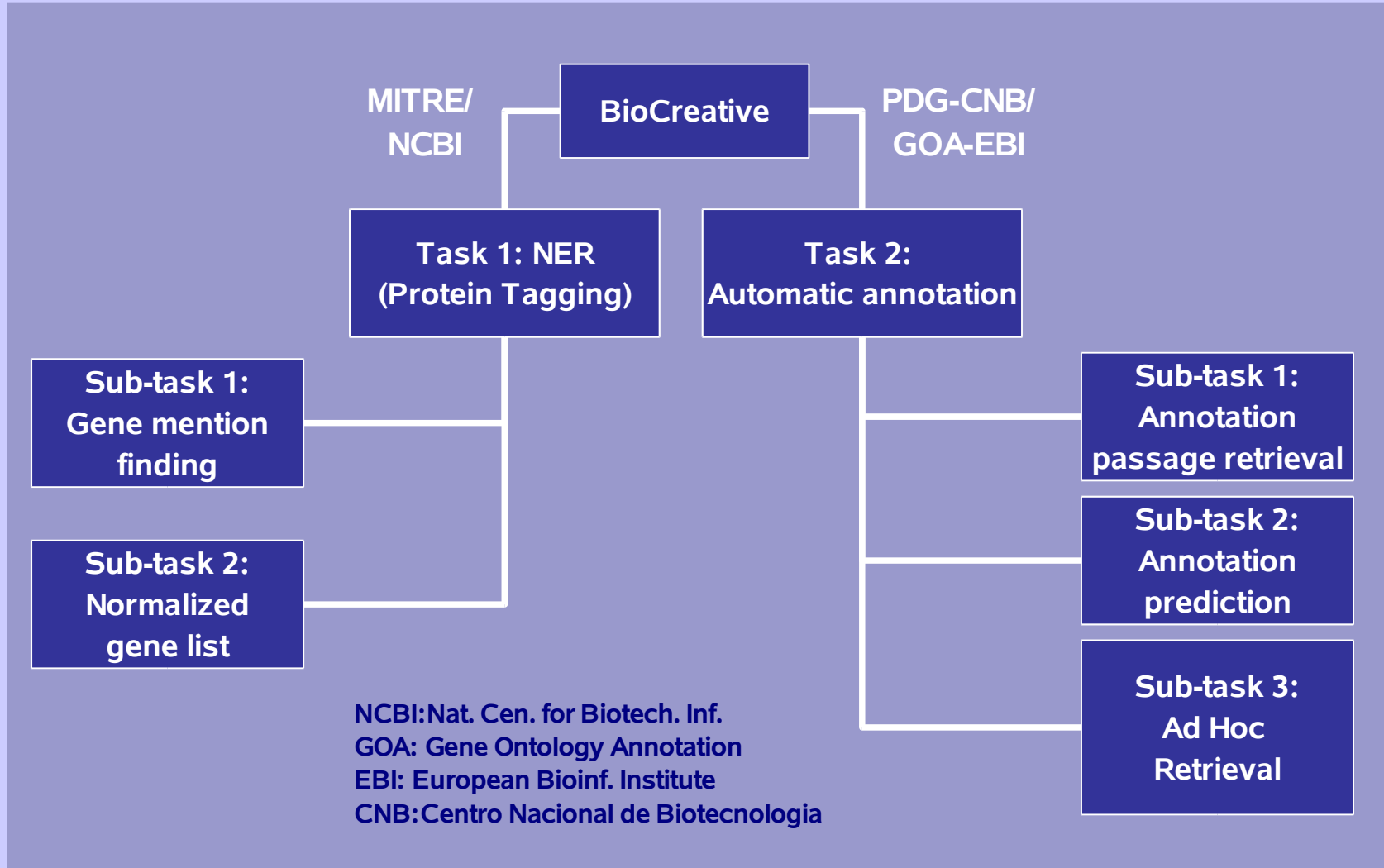
TREC

CASP: Critical assessment of Protein Structure Prediction
 CAMDA: Critical Assessment of Microarray Data Analysis
 CAPRI: Critical Assessment of Prediction of Interactions
 GASP: Genome Annotation Assessment Project
 GAW: Genome Access Workshop

PTC: Predictive Toxicology Challenge
 KDD: Knowledge Discovery and Data mining
 JNLPBA: Joint workshop on Natural Language Processing in Biomedicine
 TREC: Text Retrieval conference
 MUC: Message Understanding conference
 LLL05: Genic interaction extraction challenge

Text mining biomedical literature (2005)

Overview: BioCreative tasks



BioCreative – Why?

- Open evaluation to determine the state of the art.
- Compare the performance of different methods.
- Produce a gold standard training set.
- Monitor improvements in the field.
- Produce useful evaluation tools/metrics.

BioCreative Task 1.1 summary

- Finding gene mentions in abstracts (NER).
- 15 teams, 3-4 submissions per team.
- Data and evaluation software provided by the NCBI.
- Performance: over 80% F-score (balanced precision and recall).
- Top scoring participants used some type of markov modelling (ME,HMM,CRF), SVM or manual rules.

BioCreative Task 1.2 summary

- Gene identifier list task.
- 8 teams, 3 submissions per team.
- Given an abstract from a specific model organism (Fly, Mouse, yeast) create the list of unique gene identifiers.
- F-score: yeast 0.92, fly 0.82 and mouse 0.79.
- Difficulties: ambiguity, complex names, distinguish between multiple identifiers.
- Methods: matching against lexical resources (e.g. exhaustive matching) or task 1.1 type systems.

BioCreative Task 2 description

- Automatic extraction and assignment of GO annotations for human proteins using full text articles.
- Based on triplets: protein – GO term – article passage.
- Task 2.1: Passage retrieval task, find the text passage which support a protein – GO term annotation.
- Task 2.2: text categorization task, predict protein – GO term associations and the corresponding text passage.
- Task 2.3*: ad hoc information retrieval, retrieve annotation relevant articles

Data sets and evaluation strategy

- GO: Gene Ontology: controlled vocabulary (concepts) within an ontology (DAG), 3 categories, MF: Molecular function, BP: Biological Process and CC: Cellular Component.
- GO concepts used for annotation purposes: GOA.
- Training set: 803 GOA derived full text articles from JBC journal.
- Test set: 113 articles for task 2.1 and 99 for 2.2 and triplets.
- Triplets: GO-term - protein – article -> return passage (task 2.1).
- Evaluation by GOA annotators from the EBI.
- Manually evaluation of the predicted passages within its context in the paper using a highlighting tool.
- Evaluation types: High:correct, Generally: OK but to general for practical use and Low:wrong.

BioCreative Task 2 data set

Description	Training set	Test set 2.1	Test set 2.2
Full text articles	803	113	99
Total of GO annotation	2317	1076	1227
Nr of proteins in the GO annot	939	138	138
Nr GO terms used for annot	776	580	544
Average nr of annot/protein	2.467	7.797	8.891
Annotations of MF GO terms	709	330	356
Annotations of BP GO terms	1061	544	701
Annotations of CC GO terms	547	182	170
MF terms in the annotations	343	173	179
BP terms in the annotations	339	334	314
CC terms in the annotations	94	57	51

Data sets and evaluation strategy

- GO: Gene Ontology: controlled vocabulary (concepts) within an ontology (DAG), 3 categories, MF: Molecular function, BP: Biological Process and CC: Cellular Component.
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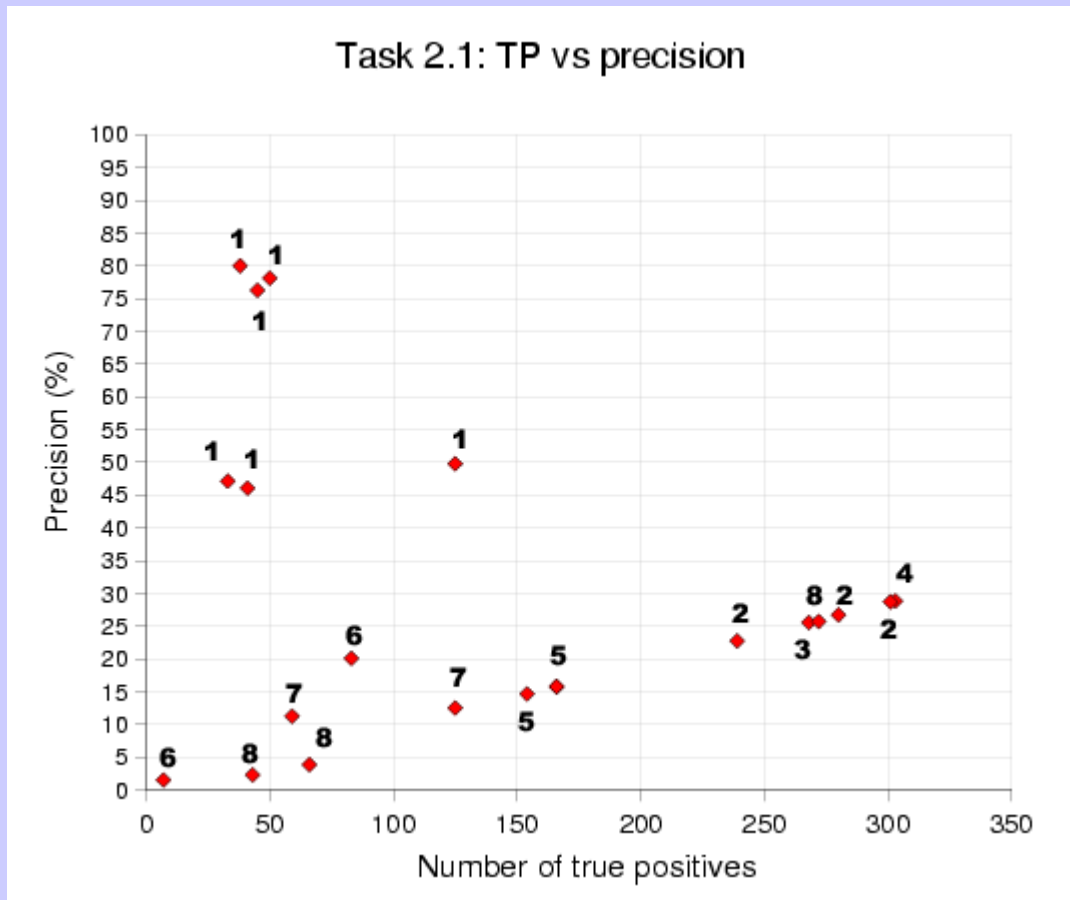
BioCreative Task 2.1 sample submission

```
<protein>
  <namefile>JBC_2001-2/bc4501042445.gml</namefile>
  <idTask>2.1</idTask>
  <participant>user14</participant>
  <nameProtein></nameProtein>
  <dbId>015023</dbId>
  <sourceDb>Swiss-Prot</sourceDb>
  <goCode>
    <name>phosphatidylinositol binding</name>
    <code>0005545</code>
    <evidenceText>In addition, a single point mutation in the
FYVE finger motif at cysteine residue 753 (C753S) is sufficient to
abolish its endosomal association. Its endosomal localization is also
sensitive to the phosphatidylinositol 3-kinase inhibitor, wortmannin.
Using in vitro liposome binding assays, we demonstrate that Myc-tagged
endofin associates preferentially with phosphatidylinositol 3-
phosphate, whereas the C753S point mutant was unable to do so. We also
show that endofin co-localizes with SARA but that they are not
associated in a common complex because they failed to co-
immunoprecipitate in co-expressing cells.</evidenceText>
  </goCode>
</protein>
```

BioCreative Task 2 participating systems

- 8 groups, max. 3 runs.
- Three strategies:
 - (1) GO term centred, pattern matching, GO words
(IC, word weight), recall centred
 - (2) Machine learning techniques.
 - (3) High precision, pattern matching and template
extraction
- Tendency: sentence level, pattern matching, regular expressions and
use of external resources (e.g. HuGO, UMLS), but:
- In general little overlap between the methods and the used resources.

BioCreative Task 2.1 results



TP: prediction evaluated as protein and GO terms correct

Precision: TP / Total nr. of evaluated submissions

Teams:

1: Chiang et al.

2: Couto et al.

3: Ehrler et al.

4: Krallinger et al.

5: Krymolowski et al.

6: Ray et al.

7: Rice et al.

8: Verspoor et al.

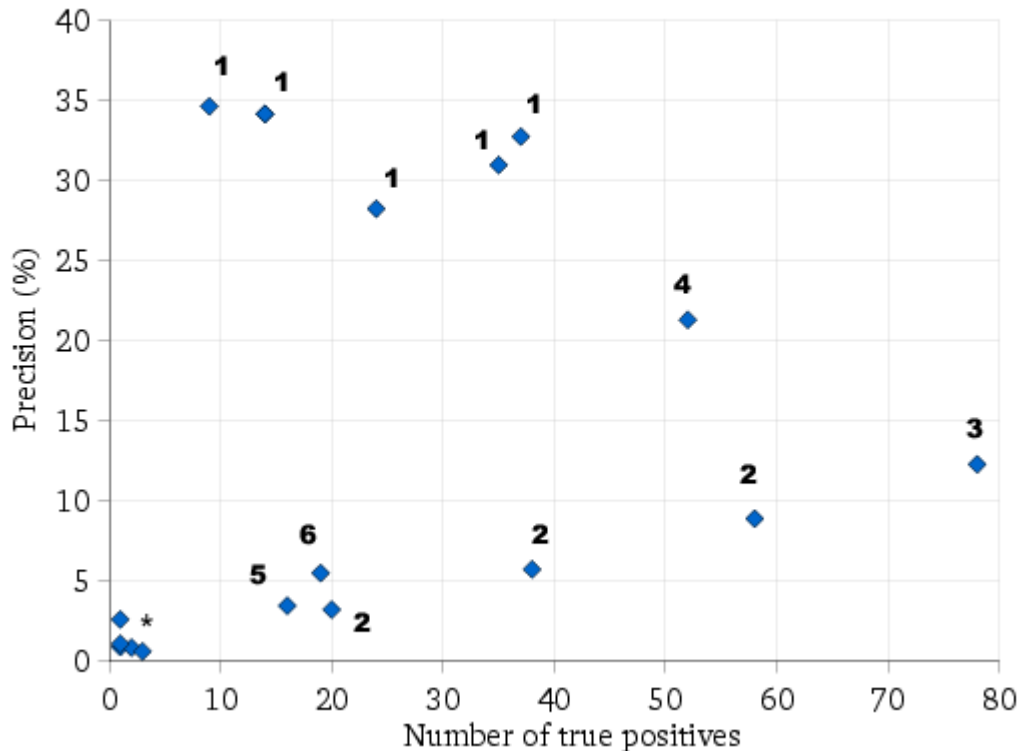
BioCreative Task 2.1 examples

Query_id	Q96PH1_0000910_11483596
PMID:	11483596
UserId:	user20_1
UserName:	Couto et al.
ProteinName:	NADPH oxidase 5 gamma
AccessionNr:	Q96PH1
GO_term:	cytokinesis
GO_id:	0000910
EvalProtein:	high
EvalGO:	high
EvalAnnot:	highhigh
Len_GO:	1
GO_cat:	P

EvidenceText: Thus, <GLOSREF RID="G8">NOX5</GLOSREF> might have a function in the early stages of spermatogenesis such as cell division, induction of apoptosis, or DNA compaction.

BioCreative Task 2.2 results

Task 2.2: TP vs. precision



TP: prediction evaluated as protein and GO terms correct

Precision: TP / Total nr. of evaluated submissions

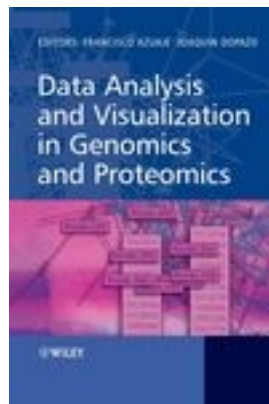
- 1: Chiang et al.
- 2: Couto et al.
- 3: Ehrler et al.
- 4: Ray et al.
- 5: Rice et al.
- 6: Verspoor et al.

Selected review references

R. Hoffmann, M. Krallinger, E. Andres, J. Tamames, C. Blaschke and A. Valencia. Text Mining for Metabolic Pathways, Signaling Cascades, and Protein Networks. Science STKE 283, pe21 (2005).

M. Krallinger, R. Alonso-Allende Erhardt and A. Valencia. Text-mining approaches in molecular biology and biomedicine. Drug Discovery Today 10, 439-445 (2005).

M. Krallinger and A. Valencia. Applications of Text Mining in Molecular Biology, from name recognition to Protein interaction maps. In Data Analysis and Visualization in Genomics and Proteomics, chapter 4, Wiley.



Selected links

http://www.pdg.cnb.uam.es/martink/LINKS/bionlp_tools_links.htm

<http://www.pdg.cnb.uam.es/martink/links.htm>

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