

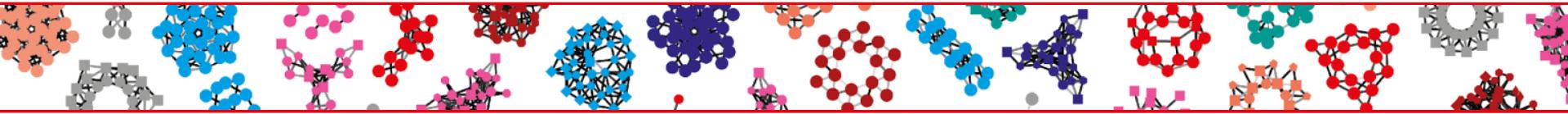
Swiss Institute of  
Bioinformatics

# UniProt

Jerven Bolleman

With the extra help of: Marie Claude-Blatter & Anne Morgat

# Overview



01

• **Biology**

02

• **Data model**

03

• **SPARQL**

04

• **To Rhea**

# Protein sequence data: where does it come from ?

- > 190 billion ‘different’ proteins on earth ( $\Sigma N$  species x M genes)
- ~ 182 million ‘known and public’ protein sequences in now
  - 50 % more by next year !
- About 98% of the protein sequences are derived from the **translation of nucleotide sequences** (mRNA or DNA/genome)
- About 1 % come from direct protein sequencing (Edman, MS/MS...)



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**UniProt consortium :**



**EBI : European Bioinformatics Institute (UK)**

**SIB : Swiss Institute of Bioinformatics (CH)**

**PIR : Protein Information Resource (USA)**



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**www.uniprot.org**

**~180 millions of proteins/entries  
derived from ~550'000 different species**

**Millions unique visitors/year**

**Very regular releases**



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**sparql.uniprot.org**  
**~60 billion triples**

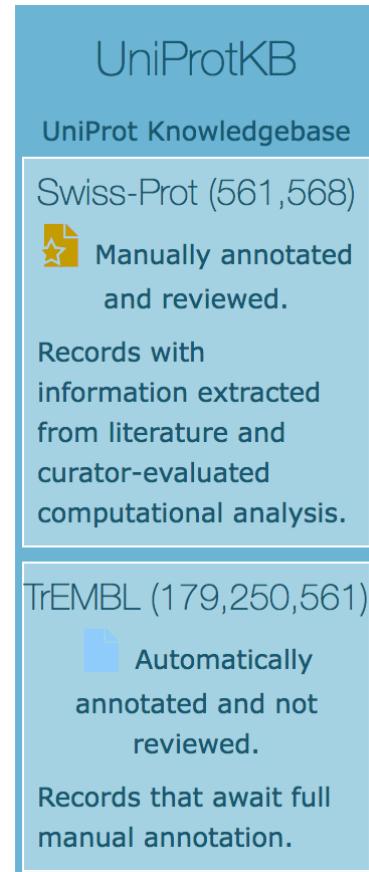
**Thousands unique visitors/year**

**Very regular releases**

# UniProtKB: 2 sections



Major differences in  
the protein sequence  
and  
annotation  
accuracy !



# UniProtKB record

<Q5NUF3> rdf:type up:Protein ;  
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up:modified "2017-07-05"^^xsd:date ;  
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up:mnemonic "HIDB\_SOYNN" ;  
up:citation citation:15734910 ,  
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citation:15734910 rdf:type up:National\_Citation ;  
up:title "Molecular and biochemical characterization of 2-hydroxyisoflavanone dehydratase. Involvement  
up:submittedTo "EMBL/GenBank/DBJB".

*Biological information / annotation*

*protein sequence*

SQ SEQUENCE 319 AA; 35138 MW; E8333CF425FBA4A3 CRC64;  
MAKEIVKELL PLIRVYKDGGS VERLLSSENV AASPEDPQTG VSSKDVIAD NPYVSARIFL  
PKSHHTNNKL PIFLYFHGGC FCVESAFSFF VRHVLNLAS EANIIAISVD FRLPLHHPIP  
AYEDGWTTL KWIASHANNNT NTTNPEPWILL NHADFTKVVY GGETSGANIA HNLLLRAGNE  
SLPGDLKLILG GLLCCCPFWNG SKPIGSSEAVE GHEQLSLAMKV WNFACPDAPG GIDNPWINPC  
VPGAPSILATL ACSKLLVTIT GKDEFRDRDI LYHHTVEQSG WQGELQLFDA GDEEHAFQLF  
KPETHLAKAM IKRLASFLV

**Text search**  
**Training Dataset**  
**Statistics**  
**Genome annotation (Features)**  
**System biology**  
...

**Proteomics**  
**BLAST**  
**Phylogeny**  
**Training datasets**  
**Domains**  
...

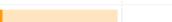
# Source of biological knowledge / annotation

## Subcellular location<sup>i</sup>

- Cell membrane 5 Publications ; Multi-pass membrane protein 1 Publication
- Cytoplasmic vesicle membrane 2 Publications
- Early endosome 1 Publication
- Membrane raft 2 Publications
- Endoplasmic reticulum 2 Publications
- Basolateral cell membrane 1 Publication

**Note:** Colocalized with KCNE3 at the plasma membrane (PubMed:10646604). Upon 17beta-oestradiol treatment, colocalizes with RAB5A at early endosome (PubMed:23529131). Heterotetramer with KCNQ5 is highly retained at the endoplasmic reticulum and is localized outside of lipid raft microdomains (PubMed:24855057). During the early stages of epithelial cell polarization induced by the calcium switch it removed from plasma membrane to the endoplasmic reticulum where it retained and it is redistributed to the basolateral cell surface in a PI3K-dependent manner at a later stage (PubMed:21228319). 4 Publications

## Topology

Feature key	Position(s)	Length	Description	Graphical view	Feature identifier	Actions
Transmembrane <sup>i</sup>	122 – 142	21	Helical; Name=Segment S1 Sequence analysis			 Add  BLAST
Topological domain <sup>i</sup>	143 – 147	5	Extracellular Sequence analysis			
Transmembrane <sup>i</sup>	148 – 168	21	Helical; Name=Segment S2 Sequence analysis		Capture	 Add  BLAST
Topological domain <sup>i</sup>	169 – 196	28	Cytoplasmic Sequence analysis			 Add  BLAST
Transmembrane <sup>i</sup>	197 – 217	21	Helical; Name=Segment S3 Sequence analysis			 Add  BLAST
Topological domain <sup>i</sup>	218 – 225	8	Extracellular Sequence analysis			
Transmembrane <sup>i</sup>	226 – 248	23	Helical; Voltage-sensor; Name=Segment S4 Sequence analysis			 Add  BLAST
Topological domain <sup>i</sup>	249 – 261	13	Cytoplasmic Sequence analysis			 Add

UniProtKB - P51787 (KCNQ1\_HUMAN)

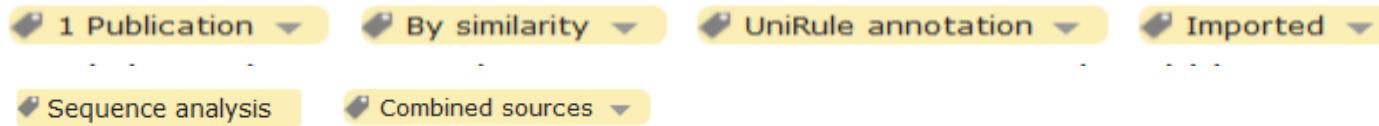
- a summary of knowledge in free text
- structured and machine-readable information

Experimental  
data  
(publication)

Computational  
analysis  
(curator-evaluated or  
not)

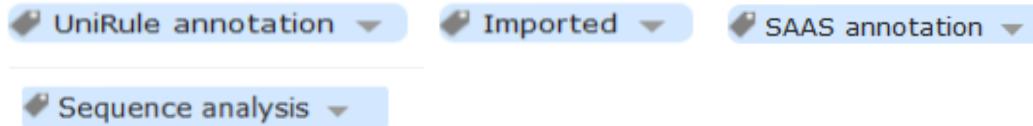
# Source of annotation / Evidence statements

**UniProtKB/Swiss-Prot: Manual insertion, color in yellow**



Computational  
analysis  
(curator-evaluated)

**UniProtKB/TrEMBL: Automated insertion, color in blue**

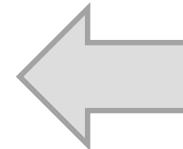


Computational  
analysis  
(NOT curator-  
evaluated)

# Annotation score & evidence for protein existence

## P04150 - GCR\_HUMAN

Protein	Glucocorticoid receptor
Gene	NR3C1
Organism	<i>Homo sapiens (Human)</i>
Sequence features	<a href="#">View only features (sites, domains, PTMs ...)</a>
Status	 Reviewed - Annotation score:  - Experimental evidence at protein level <sup>1</sup>



### Status:

- Reviewed / Unreviewed
- Annotation score: <http://insideuniprot.blogspot.ch/2014/10/introducing-annotation-scores-in-uniprot.html>
- Evidence for protein existence
  - 1. Experimental evidence at protein level
  - 2. Experimental evidence at transcript level
  - 3. Protein inferred from homology
  - 4. Protein predicted
  - 5. Protein uncertain

# UniProtKB/TrEMBL

UniProtKB
UniProt Knowledgebase
Swiss-Prot (561,568)
 Manually annotated and reviewed.
Records with information extracted from literature and curator-evaluated computational analysis.
TrEMBL (179,250,561)
 Automatically annotated and not reviewed.
Records that await full manual annotation.

**One protein sequence per entry**

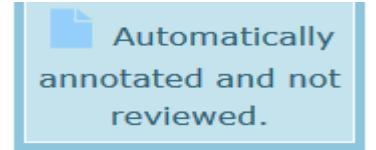
gene-centric /**protein-centric**



**99 % of UniProtKB protein sequences**

# UniProtKB/TrEMBL

## Automatic annotation



### Protein sequence

- The quality of the protein sequences is dependent on the information provided by the submitter of the original nucleotide entry (CDS) or of the gene prediction pipeline (i.e. Ensembl).
- 100% identical sequences (same length, same organism) are merged automatically).

### Biological information

#### Sources of annotation

- Provided by the submitter (EMBL, PDB, TAIR...)
- Automated annotation

# P73722 - P73722\_SYNY3

Protein	Submitted name: <b>SOS function regulatory protein</b>
Gene	<b>lexA</b>
Organism	<i>Synechocystis sp.</i> (strain 6803 / Kazusa)
Status	Unreviewed -  - Protein inferred from homology 

Automatically annotated and not reviewed.

## Function<sup>i</sup>

Represses a number of genes involved in the response to DNA damage (SOS response), including recA and lexA. In the presence of single-stranded DNA, RecA interacts with LexA causing an autocatalytic cleavage which disrupts the DNA-binding part of LexA, leading to derepression of the SOS regulon and eventually DNA repair  [By similarity](#).

 SAAS annotations ▾

## Catalytic activity<sup>i</sup>

Hydrolysis of Ala- $\beta$ -Gly bond in repressor LexA.  SAAS annotations ▾

## Keywords - Molecular function<sup>i</sup>

Hydrolase  SAAS annotations ▾, Repressor  SAAS annotations ▾

## Keywords - Biological process<sup>i</sup>

DNA damage, DNA repair, DNA replication  SAAS annotations ▾, SOS response  SAAS annotations ▾, Transcription, Transcription regulation  SAAS annotations ▾

## Keywords - Ligand<sup>i</sup>

DNA-binding  SAAS annotations ▾

Capture Ctrl+Ins

# UniProtKB/Swiss-Prot

UniProtKB
UniProt Knowledgebase
Swiss-Prot (561,568)
 Manually annotated and reviewed.
Records with information extracted from literature and curator-evaluated computational analysis.
TrEMBL (179,250,561)
 Automatically annotated and not reviewed.
Records that await full manual annotation.

1 % of UniProtKB protein sequences



What is a biocurator?



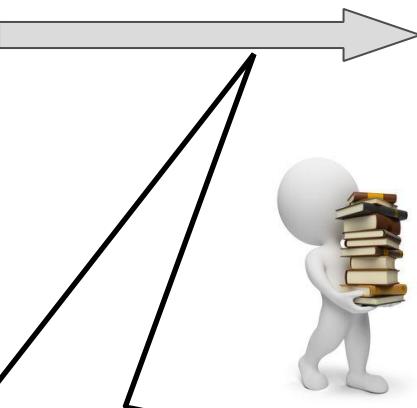
They spend their time behind the scenes, yet biocurators are essential to knowledge maintenance. The Swiss-Prot biocurators

# UniProtKB/Swiss-Prot

## UniProtKB/TrEMBL

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PF	01-JUN-2002	Sequence version 1.0	
PC	00-JAN-2002		
NN	MARCH1	Protein Name=MARCH1; Synonyms=DKF2p54M1682;	
OC	Bivalvia; Mollusca; Chordata; Craniata; Vertebrates; Euteleostomi; Mammalia; Eutheria; Euschontoglires; Primates; Haplorhini; Ceterrhini; Hominoidea; Homo.		
RN	2002-05-10-01-03-0004		
RN	[1]		
RR	REVIEWED; SOURCE;		
RT	T158UP-Brain1		
RG	The German cDNA Consortium;		
RS	Postberg F., Stummeyer P., Schupp T., Wellerreuther R., Mewes H.W., Weil B., Auld C., Osoegawa A., Fobo G., Han M., Wiemann S.; Submitted (SEP-2004) to the EMBL/GenBank/DBJ databases.		
CC	- I - SIMILARITY: Contains 1 RING-type zinc finger.		
DR	GO:0045504 Zinc finger, RING-type		
DR	Copyrighted by the UniProt Consortium, see <a href="http://www.uniprot.org/terms">http://www.uniprot.org/terms</a>		
DR	Distributed under the Creative Commons Attribution-NoDerivs License		
DR	DR: <a href="http://www.ncbi.nlm.nih.gov/entrez/Entrez.cgi?db=pubmed&amp;term=(MARCH1+AND+Homo+sapiens)&amp;cmd=Search">http://www.ncbi.nlm.nih.gov/entrez/Entrez.cgi?db=pubmed&amp;term=(MARCH1+AND+Homo+sapiens)&amp;cmd=Search</a> ; EMBL: <a href="http://www.ebi.ac.uk/embl/submit/march1/MARCH1_EMBL.html">http://www.ebi.ac.uk/embl/submit/march1/MARCH1_EMBL.html</a> ; CAD26529.1; - ; mRNA;		
DR	SMTB: Q7TCQ1; 72-136.		
DR	InterPro: IPR0015416 Homo sapiens.		
DR	HMNC: HGNC:26077; MARCH1.		
DR	ArrayExpress: Q7TCQ1; - .		
DR	KEGG: KEGG0045504; RING1; - .		
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GO	GO:0004315 Zinc ion binding; IEA:UniProtKB-PW.		
GO	GO:0008270 Zinc ion binding; IEA:UniProtKB-PW.		
GO	GO:0009455 Zinc ion homeostasis; Zinc ion cycle; IEA:UniProtKB-PW.		
InterPro	IPR011016; RING1.		
InterPro	IPR001841; Znf_RING1.		
InterPro	IPR004635; RING1.		
SMART	SM00744; RING1; 1.		
SWISSPROT	P39267; RING1; 1.		
KW	Zinc-finger protein; Metal-binding; Ubiquitin conjugation pathway; Zinc-finger		
KW	Zinc-finger		
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Manual annotation of  
the sequence and  
associated biological  
information



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Q7TCQ1_HUMAN UniProtKB/Swiss-Prot 316 AA.
ID Q7TCQ1_HUMAN UniProtKB/Swiss-Prot.
DT 2003-06-20T09:00:00Z.
PF 01-JUN-2002.
PC 00-JAN-2002.
NN MARCH1; Synonyms=DKF2p54M1682;
OC Bivalvia; Mollusca; Chordata; Craniata; Vertebrates; Euteleostomi; Mammalia; Eutheria; Euschontoglires; Primates; Haplorhini; Ceterrhini; Hominoidea; Homo.
RN 2002-05-10-01-03-0004;
RN [1];
RR REVIEWED; SOURCE;
RT T158UP-Brain1;
RS The German cDNA Consortium;
RS Postberg F., Stummeyer P., Schupp T., Wellerreuther R., Mewes H.W., Weil B., Auld C., Osoegawa A., Fobo G., Han M., Wiemann S.; Submitted (SEP-2004) to the EMBL/GenBank/DBJ databases.
CC - I - SIMILARITY: Contains 1 RING-type zinc finger.
DR GO:0004635 Protein ion binding; IEA:UniProtKB-PW.
DR Copyrighted by the UniProt Consortium, see http://www.uniprot.org/terms
CC Distributed under the Creative Commons Attribution-NoDerivs License
DR EMBL: A5713759; CAD26529.1; - ; mRNA;
DR SMTB: Q7TCQ1; 72-136.
DR InterPro: IPR011016; RING1.
DR HMNC: HGNC:26077; MARCH1.
DR ArrayExpress: Q7TCQ1; - .
DR KEGG: KEGG0045504; RING1; - .
DR R2PD-ProteExp; W1571; - .
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GO GO:0008270 Zinc ion binding; IEA:UniProtKB-PW.
GO GO:0009455 Zinc ion homeostasis; Zinc ion cycle; IEA:UniProtKB-PW.
InterPro IPR011016; RING1.
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SMART SM00744; RING1; 1.
SWISSPROT P39267; RING1; 1.
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KW Zinc-finger
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```

- At least 20% of UniProtKB/Swiss-Prot entries required curation effort to “correct” the sequences.
- Typical problems
  - unsolved conflicts;
  - uncorrected initiation sites;
  - frameshifts;
  - other ‘problems’



# UniProtKB/Swiss-Prot

Biological knowledge / annotation / GO annotation

Knowledge:

- comprehensive summary (free text) that provides a complete overview of the information available
- standardized vocabularies to facilitate subsequent retrieval whenever possible

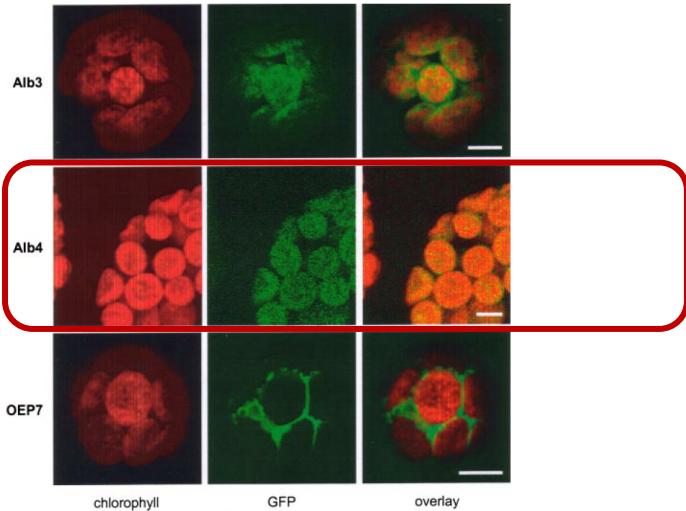
# Source of annotation / Evidence statements

Every piece of knowledge is associated with the source of the information and the type of supporting evidence, using the evidence ontology (ECO)

- Selected Publication (experimental)  1 Publication ▾
- Another UniProtKB entry (orthologs): *by similarity*  By similarity ▾
- An entry from another database: *imported*  Imported ▾
- Curator-evaluated computational analysis  UniRule annotation ▾
- Combined sources  Combined sources ▾

# Selected Publication

PubMed=16595657



**FIGURE 3. Subcellular localization of Alb3-GFP and Alb4-GFP fusion proteins.** *Arabidopsis* mesophyll protoplasts were transiently transformed with constructs for Alb3- or Alb4-GFP. Maximum intensity signals from confocal images are shown for chlorophyll autofluorescence, GFP fluorescence, and an overlay of both. OEP7-GFP is included as a marker for the chloroplast envelope. Bar, 5  $\mu$ m.

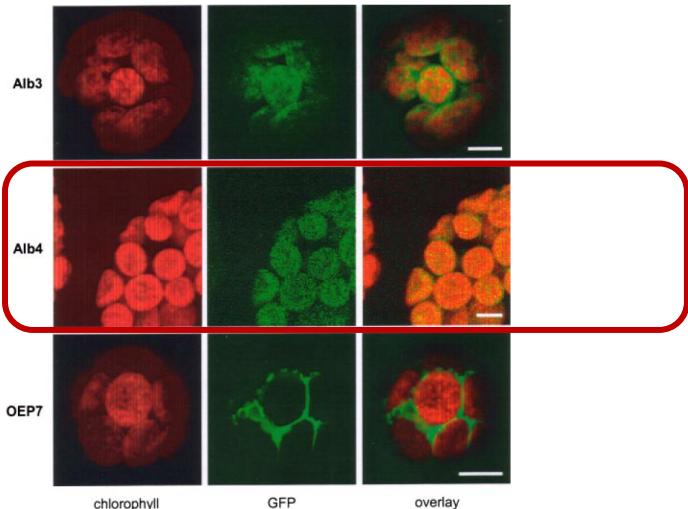
corresponds to the predicted length of the Alb4 mRNA. Even after prolonged exposure of the blots treated with the Alb4 probe, no signal could be found at  $\sim$ 3.5 kb, the predicted size of the Artemis transcript.

**Alb4 Is a Thylakoid Membrane Protein**—Alb4 is predicted to have a chloroplast transit peptide with a processing site after amino acid resi-

comprehensive and computer friendly representation of biological knowledge

due 45 based on the ChloroP prediction program (33). To test this prediction, *Arabidopsis* mesophyll protoplasts were transiently transformed with constructs for Alb3-GFP or Alb4-GFP. Merging of the GFP and autofluorescence images indicated a thylakoid localization of Alb4. The GFP distribution for Alb4 is similar to that of Alb3 and not to that of outer envelope protein AtOEP7 (Fig. 3). To test this assumption, *in vitro* translated radiolabeled Alb4 was imported into isolated pea

PubMed=16595657



**FIGURE 3. Subcellular localization of Alb3-GFP and Alb4-GFP fusion proteins.** *Arabidopsis* mesophyll protoplasts were transiently transformed with constructs for Alb3- or Alb4-GFP. Maximum intensity signals from confocal images are shown for chlorophyll autofluorescence, GFP fluorescence, and an overlay of both. OEP7-GFP is included as a marker for the chloroplast envelope. Bar, 5  $\mu$ m.

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**Alb4 Is a Thylakoid Membrane Protein**—Alb4 is predicted to have a chloroplast transit peptide with a processing site after amino acid resi-

**Subcellular location<sup>i</sup>**

- Plastid > chloroplast thylakoid membrane 1 Publication ; Multi-pass membrane protein 1 Publication

**Topology**

Feature key	Position(s)	Length	Description	Graphical view
Transmembrane <sup>i</sup>	115 – 135	21	Helical Sequence analysis	
Transmembrane <sup>i</sup>	184 – 204	21	Helical Sequence analysis	
Transmembrane <sup>i</sup>	263 – 283	21	Helical Sequence analysis	
Transmembrane <sup>i</sup>	302 – 322	21	Helical Sequence analysis	

**GO - Cellular component<sup>i</sup>**

- chloroplast Source: TAIR
- chloroplast thylakoid membrane Source: TAIR
- integral component of membrane Source: UniProtKB-KW
- thylakoid Source: TAIR

**Controlled vocabulary GO annotation**

**2. "A second thylakoid membrane-localized Alb3/OxaI/YidC homologue is involved in proper chloroplast biogenesis in *Arabidopsis thaliana*."**  
 Gerdes L., Bals T., Klostermann E., Karl M., Philippar K., Huenken M., Soll J., Schuenemann D.  
*J. Biol. Chem.* 281:16632–16642(2006) [PubMed] [Europe PMC] [Abstract]  
**Cited for:** SEQUENCE REVISION, TISSUE SPECIFICITY, SUBCELLULAR LOCATION.

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# Curator-evaluated computational analysis

PubMed=16595657

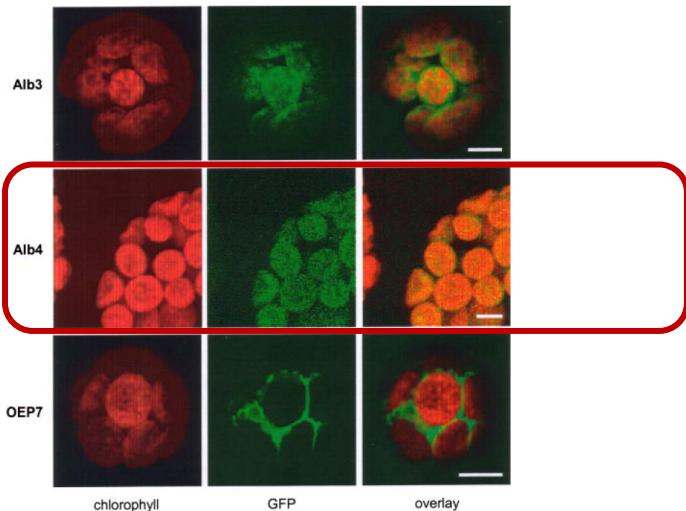


FIGURE 3. Subcellular localization of Alb3-GFP and Alb4-GFP fusion proteins. *Arabidopsis* mesophyll protoplasts were transiently transformed with constructs for Alb3- or Alb4-GFP. Maximum intensity signals from confocal images are shown for chlorophyll autofluorescence, GFP fluorescence, and an overlay of both. OEP7-GFP is included as a

corresponds to the predicted length of the Alb4 mRNA. Even after prolonged exposure of the blots treated with the Alb4 probe, no signal could be found at  $\sim 3.5$  kb, the predicted size of the Artemis transcript.

*Alb4 Is a Thylakoid Membrane Protein*—Alb4 is predicted to have a chloroplast transit peptide with a processing site after amino acid resi-

## UniProtKB Q9FYL3

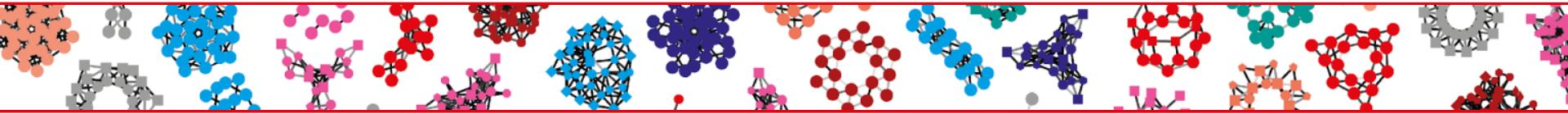
### Molecule processing

Feature key	Position(s)	Length	Description
Transit peptide <sup>i</sup>	1 – 45	45	Chloroplast  Sequence analysis
Chain <sup>i</sup>	46 – 499	454	ALBINO3-like protein 1, chloroplastic



due 45 based on the ChloroP prediction program (33). To test this prediction, *Arabidopsis* mesophyll protoplasts were transiently transformed with constructs for Alb3-GFP or Alb4-GFP. Merging of the GFP and autofluorescence images indicated a thylakoid localization of Alb4. The GFP distribution for Alb4 is similar to that of Alb3 and not to that of outer envelope protein AtOEP7 (Fig. 3). To test this assumption, *in vitro* translated radiolabeled Alb4 was imported into isolated pea

# Overview



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- To Rhea

# The FAIRest format of them all

## UniProt Code

```
<P05067> rdf:type up:Protein ;  
    up:reviewed      true ; #This is a Swiss-Prot entry  
    up:organism      taxon:9606 ; #Talking about a human entry  
    up:classifiedWith
```

```
[ go:GO_0043198 rdf:type      owl:Class ;  
    rdfs:label      "dendritic shaft" ;  
    rdfs:subClassOf go:GO_0044463 ,  
                    go:GO_0097458 ,  
                    go:GO_0030425 ,
```

```
[ owl:restriction
```

```
[ owl:onProperty obo:BFO_0000050 ;  
owl:someValuesFrom go:GO_0030425 .
```

```
]]
```

```
]
```

## GO Code

# UniProt RDF

---

- <https://sparql.uniprot.org>
- 400+ files on FTP
  - Fast loading into your own DB
  - Categorized by taxonomy
  - Or dataset
- Downloadable subsets on our website
  - Slice and dice as you wish

# UniProt RDF

---

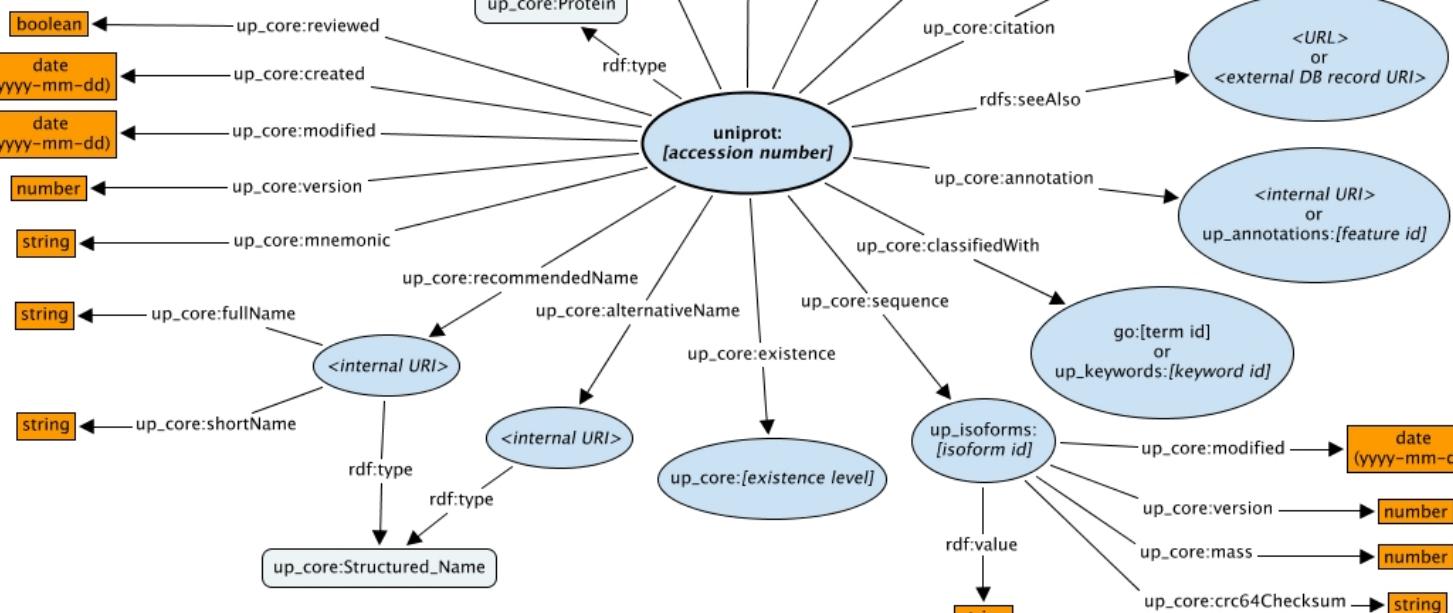
- **It's huge !**
  - **Terabytes !**
- **Query for something**
  - **Get back 10 billion rows**
  - **Protection for this in the HTML view**

**Namespaces**

up\_core:<<http://purl.uniprot.org/core/>>  
 uniprot:<<http://purl.uniprot.org/uniprot/>>  
 up\_citations:<<http://purl.uniprot.org/citations/>>  
 up\_taxonomy:<<http://purl.uniprot.org/taxonomy/>>  
 up\_annotations:<<http://purl.uniprot.org/annotation/>>  
 up\_keywords:<<http://purl.uniprot.org/keywords/>>  
 up\_isoforms:<<http://purl.uniprot.org/isoforms/>>

ec:<<http://purl.uniprot.org/enzyme/>>  
 go:<<http://purl.uniprot.org/go/>>

rdf:<<http://www.w3.org/1999/02/22-rdf-syntax-ns#>>  
 rdfs:<<http://www.w3.org/2000/01/rdf-schema#>>  
 owl:<<http://www.w3.org/2002/07/owl#>>  
 skos:<<http://www.w3.org/2004/02/skos/core#>>



Typed resource/node

Class

Typed literal

# Annotation

- 27 subtypes

```
1 PREFIX rdfs:<http://www.w3.org/2000/01/rdf-schema#>
2 PREFIX owl:<http://www.w3.org/2002/07/owl#>
3 PREFIX up:<http://purl.uniprot.org/core/>
4 SELECT (COUNT(?class) AS ?classes)
5 FROM <http://purl.uniprot.org/core/>
6 WHERE
7 {
8     ?class a owl:Class ;
9         rdfs:subClassOf up:Annotation .
10 }
```

# Annotation

<a href="http://biohackathon.org/resource/faldo#location">http://biohackathon.org/resource/faldo#location</a>	faldo:location
<a href="http://purl.uniprot.org/core/catalyticActivity">http://purl.uniprot.org/core/catalyticActivity</a>	up:catalyticActivity
<a href="http://purl.uniprot.org/core/catalyzedPhysiologicalReaction">http://purl.uniprot.org/core/catalyzedPhysiologicalReaction</a>	up:catalyzedPhysiologicalReaction
<a href="http://purl.uniprot.org/core/cofactor">http://purl.uniprot.org/core/cofactor</a>	up:cofactor
<a href="http://purl.uniprot.org/core/conflictingSequence">http://purl.uniprot.org/core/conflictingSequence</a>	up:conflictingSequence
<a href="http://purl.uniprot.org/core/disease">http://purl.uniprot.org/core/disease</a>	up:disease
<a href="http://purl.uniprot.org/core/frameshift">http://purl.uniprot.org/core/frameshift</a>	up:frameshift
<a href="http://purl.uniprot.org/core/locatedIn">http://purl.uniprot.org/core/locatedIn</a>	up:locatedIn
<a href="http://purl.uniprot.org/core/maximum">http://purl.uniprot.org/core/maximum</a>	up:maximum
<a href="http://purl.uniprot.org/core/measuredActivity">http://purl.uniprot.org/core/measuredActivity</a>	up:measuredActivity
<a href="http://purl.uniprot.org/core/measuredAffinity">http://purl.uniprot.org/core/measuredAffinity</a>	up:measuredAffinity
<a href="http://purl.uniprot.org/core/measuredError">http://purl.uniprot.org/core/measuredError</a>	up:measuredError
<a href="http://purl.uniprot.org/core/measuredValue">http://purl.uniprot.org/core/measuredValue</a>	up:measuredValue
<a href="http://purl.uniprot.org/core/method">http://purl.uniprot.org/core/method</a>	up:method
<a href="http://purl.uniprot.org/core/range">http://purl.uniprot.org/core/range</a>	up:range
<a href="http://purl.uniprot.org/core/sequence">http://purl.uniprot.org/core/sequence</a>	up:sequence
<a href="http://purl.uniprot.org/core/substitution">http://purl.uniprot.org/core/substitution</a>	up:substitution
<a href="http://www.w3.org/1999/02/22-rdf-syntax-ns#type">http://www.w3.org/1999/02/22-rdf-syntax-ns#type</a>	a
<a href="http://www.w3.org/2000/01/rdf-schema#comment">http://www.w3.org/2000/01/rdf-schema#comment</a>	rdfs:comment
<a href="http://www.w3.org/2000/01/rdf-schema#seeAlso">http://www.w3.org/2000/01/rdf-schema#seeAlso</a>	rdfs:seeAlso
<a href="http://www.w3.org/2004/02/skos/core#related">http://www.w3.org/2004/02/skos/core#related</a>	skos:related

# Function annotation (web view)

Functions as a cell surface receptor and performs physiological functions on the surface of neurons relevant to neurite growth, neuronal adhesion and axonogenesis. Interaction between APP molecules on neighboring cells promotes synaptogenesis (PubMed:[25122912](#)). Involved in cell mobility and transcription regulation through protein-protein interactions. Can promote transcription activation through binding to APBB1-KAT5 and inhibits Notch signaling through interaction with Numb. Couples to apoptosis-inducing pathways such as those mediated by G(O) and JIP. Inhibits G(o) alpha ATPase activity (By similarity). Acts as a kinesin I membrane receptor, mediating the axonal transport of beta-secretase and presenilin 1 (By similarity). By acting as a kinesin I membrane receptor, plays a role in axonal anterograde transport of cargo towards synapses in axons (PubMed:[17062754](#), PubMed:[23011729](#)). Involved in copper homeostasis/oxidative stress through copper ion reduction. In vitro, copper-metallated APP induces neuronal death directly or is potentiated through Cu<sup>2+</sup>-mediated low-density lipoprotein oxidation. Can regulate neurite outgrowth through binding to components of the extracellular matrix such as heparin and collagen I and IV. The splice isoforms that contain the BPTI domain possess protease inhibitor activity. Induces a AGER-dependent pathway that involves activation of p38 MAPK, resulting in internalization of amyloid-beta peptide and leading to mitochondrial dysfunction in cultured cortical neurons. Provides Cu<sup>2+</sup> ions for GPC1 which are required for release of nitric oxide (NO) and subsequent degradation of the heparan sulfate chains on GPC1.

 By similarity ▾

 3 Publications ▾

# Function annotation (RDF)

---

<P05067#SIP8BC30A3A03BD108B> rdf:type up:Function\_Annotation ;  
rdfs:comment "Functions as a cell surface receptor and performs physiological functions on the surface of neurons relevant to neurite growth, neuronal adhesion and axonogenesis. Interaction between APP molecules on neighboring cells promotes synaptogenesis (PubMed:25122912). Involved in cell mobility and transcription regulation through protein-protein interactions. Can promote transcription activation through binding to APBB1-KAT5 and inhibits Notch signaling through interaction with Numb. Couples to apoptosis-inducing pathways such as those mediated by G(O) and JIP. Inhibits G(o) alpha ATPase activity (By similarity). Acts as a kinesin I membrane receptor, mediating the axonal transport of beta-secretase and presenilin 1 (By similarity). By acting as a kinesin I membrane receptor, plays a role in axonal anterograde transport of cargo towards synapses in axons (PubMed:17062754, PubMed:23011729). Involved in copper homeostasis/oxidative stress through copper ion reduction. In vitro, copper-metallated APP induces neuronal death".

# Sequence

---

- Isoforms and Canonical materialized
- IUPAC code
  - No spaces

# Cross-reference

---

- 173 databases `rdfs:seeAlso` → `up:database db:{DB}`
- Ensembl (like)
  - `up:translatedTo`
  - `up:transcribedFrom`
- PDB
  - `http://rdf.wwpdb.org/pdb/1AAP`
- All others
  - `rdfs:comment`
- Try to use their (your? IRI) else `purl.uniprot.org/{DB}/{ID}`
- HAMAP, InterPro, PFam etc.
  - `up:signatureSequenceMatch`

# Cross-reference: Ensembl (and ensembl like)

---

```
<http://rdf.ebi.ac.uk/resource/ensembl.transcript/ENST00000233072> ↴
  rdf:type ↴
    up:Transcript_Resource ;
  up:database <http://purl.uniprot.org/database/Ensembl> ;
  up:translatedTo <http://rdf.ebi.ac.uk/resource/ensembl.protein/ENSP00000233072> ;
  up:transcribedFrom <http://rdf.ebi.ac.uk/resource/ensembl/ENSG0000021826> ;
  rdfs:seeAlso isoform:P31327-1 .
```

# Cross-reference: PDB

---

```
<http://rdf.wwpdb.org/pdb/2YVQ> rdf:type up:Structure_Resource ;
  up:database <http://purl.uniprot.org/database/PDB> ;
  up:method up:X-Ray_Crystallography ;
  up:resolution "1.98"^^xsd:float ;
  up:chainSequenceMapping ↴
    <http://purl.uniprot.org/isoforms/P31327-1#PDB_2YVQ_tt1343tt1478> .
<http://purl.uniprot.org/isoforms/P31327-1#PDB_2YVQ_tt1343tt1478> up:chain ↴
  "A=1343-1478" .
```

# Evidence

---

Reifies the statement ↗

Triple ?s ?p ?o ↘

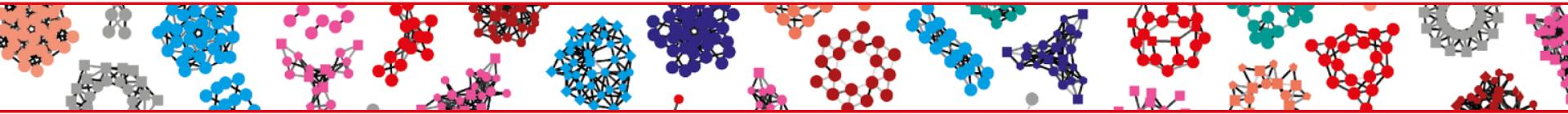
```
[] a rdf:Statement ;  
    rdf:subject ?s ;  
    rdf:predicate ?p ;  
    rdf:object ?o ;  
    up:attribution ?whoWhatWhy .
```

# Evidence (example)

---

```
<P05067#attribution-7B6976B7E1FAA17744B06EA4C9F47A94>^
dcterms:creator^
<http://purl.uniprot.org/goa-projects/ARUK-UCL> ;
up:manual true ;
up:evidence ECO:0000314 ;
up:source citation:18723082 .
```

# Overview



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- To Rhea

# Annotation

- 27 subtypes

```
1 PREFIX rdfs:<http://www.w3.org/2000/01/rdf-schema#>
2 PREFIX owl:<http://www.w3.org/2002/07/owl#>
3 PREFIX up:<http://purl.uniprot.org/core/>
4 SELECT (COUNT(?class) AS ?classes)
5 FROM <http://purl.uniprot.org/core/>
6 WHERE
7 {
8     ?class a owl:Class ;
9         rdfs:subClassOf up:Annotation .
10 }
```

# Sequence

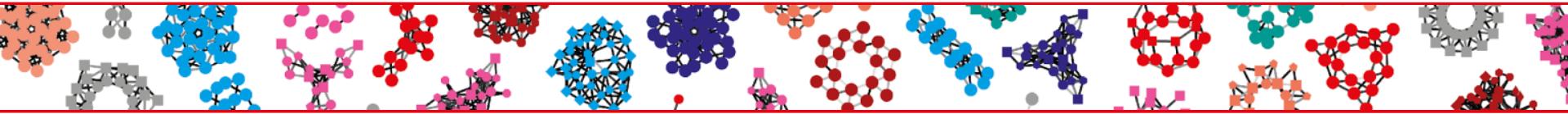
```
1 BASE <http://purl.uniprot.org/uniprot/>
2 PREFIX rdf:<http://www.w3.org/1999/02/22-rdf-syntax-ns#>
3 PREFIX rdfs:<http://www.w3.org/2000/01/rdf-schema#>
4 PREFIX owl:<http://www.w3.org/2002/07/owl#>
5 PREFIX up:<http://purl.uniprot.org/core/>
6 SELECT ?protein ?aaSequence
7 FROM <http://sparql.uniprot.org/uniprot>
8 WHERE
9 {
10   BIND (<P05067> AS ?protein)
11   ?protein up:sequence ?sequence .
12   ?sequence rdf:value ?aaSequence .
13 }
```

# Evidence

---

- 4,258,699,129
  - Often needs at least 3 joins
  - Probably slowest

# Overview



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- To Rhea

## Query: Retrieve the evidences of Rhea reactions annotated in UniProtKB/Swiss-Prot

```
1 PREFIX rdf:<http://www.w3.org/1999/02/22-rdf-syntax-ns#>
2 PREFIX up:<http://purl.uniprot.org/core/>
3 PREFIX skos:<http://www.w3.org/2004/02/skos/core#>
4
5 SELECT DISTINCT
6     ?upProtein
7     ?rheaReaction
8     ?rheaReaction_evt
9     ?source
10 WHERE {
11     ?upProtein up:reviewed true .
12     #
13     ?upProtein up:annotation ?annotation .
14     ?annotation a up:Catalytic_Activity_Annotation ;
15         up:catalyticActivity ?catalyticActivity .
16     ?catalyticActivity up:catalyzedReaction ?rheaReaction .
17     #
18     ?upProtein up:attribution ?attribution .
19     ?attribution up:evidence ?rheaReaction_evt .
20     [] rdf:subject ?annotation ;
21         rdf:predicate up:catalyticActivity ;
22         rdf:object ?catalyticActivity ;
23         up:attribution ?attribution .
24     OPTIONAL {?attribution up:source ?source .}
25 }
```

# Result: Retrieve the evidences of Rhea reactions annotated in UniProtKB/Swiss-Prot



The image shows the UniProt homepage. At the top left is the UniProt logo. Below it is a navigation bar with links for "SPARQL", "Downloads", "Documentation/Help", and "Contact". The background features a blue banner with a molecular structure graphic.

## Results

Sparql XML Sparql JSON CSV Show query Share

upProtein	rheaReaction	rheaReaction_evt	source
<a href="http://purl.uniprot.org/uniprot/Q9UXZ0">http://purl.uniprot.org/uniprot/Q9UXZ0</a>	<a href="http://rdf.rhea-db.org/10164">http://rdf.rhea-db.org/10164</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000255">http://purl.obolibrary.org/obo/ECO_0000255</a>	<a href="http://purl.uniprot.org/hemap-rule/MF_00318">http://purl.uniprot.org/hemap-rule/MF_00318</a>
<a href="http://purl.uniprot.org/uniprot/A0A087WNH6">http://purl.uniprot.org/uniprot/A0A087WNH6</a>	<a href="http://rdf.rhea-db.org/21708">http://rdf.rhea-db.org/21708</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000255">http://purl.obolibrary.org/obo/ECO_0000255</a>	<a href="http://purl.uniprot.org/hemap-rule/MF_00563">http://purl.uniprot.org/hemap-rule/MF_00563</a>
<a href="http://purl.uniprot.org/uniprot/A0A078BQP2">http://purl.uniprot.org/uniprot/A0A078BQP2</a>	<a href="http://rdf.rhea-db.org/13665">http://rdf.rhea-db.org/13665</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000250">http://purl.obolibrary.org/obo/ECO_0000250</a>	<a href="http://purl.uniprot.org/uniprot/Q19187">http://purl.uniprot.org/uniprot/Q19187</a>
<a href="http://purl.uniprot.org/uniprot/B0YJ81">http://purl.uniprot.org/uniprot/B0YJ81</a>	<a href="http://rdf.rhea-db.org/45812">http://rdf.rhea-db.org/45812</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000269">http://purl.obolibrary.org/obo/ECO_0000269</a>	<a href="http://purl.uniprot.org/citations/18554506">http://purl.uniprot.org/citations/18554506</a>
<a href="http://purl.uniprot.org/uniprot/B0YJ81">http://purl.uniprot.org/uniprot/B0YJ81</a>	<a href="http://rdf.rhea-db.org/45812">http://rdf.rhea-db.org/45812</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000269">http://purl.obolibrary.org/obo/ECO_0000269</a>	<a href="http://purl.uniprot.org/citations/23933735">http://purl.uniprot.org/citations/23933735</a>
<a href="http://purl.uniprot.org/uniprot/A3BF39">http://purl.uniprot.org/uniprot/A3BF39</a>	<a href="http://rdf.rhea-db.org/12132">http://rdf.rhea-db.org/12132</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000269">http://purl.obolibrary.org/obo/ECO_0000269</a>	<a href="http://purl.uniprot.org/citations/22123790">http://purl.uniprot.org/citations/22123790</a>
<a href="http://purl.uniprot.org/uniprot/A3BF39">http://purl.uniprot.org/uniprot/A3BF39</a>	<a href="http://rdf.rhea-db.org/20621">http://rdf.rhea-db.org/20621</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000269">http://purl.obolibrary.org/obo/ECO_0000269</a>	<a href="http://purl.uniprot.org/citations/22123790">http://purl.uniprot.org/citations/22123790</a>
<a href="http://purl.uniprot.org/uniprot/A0A024B7W1">http://purl.uniprot.org/uniprot/A0A024B7W1</a>	<a href="http://rdf.rhea-db.org/60860">http://rdf.rhea-db.org/60860</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000255">http://purl.obolibrary.org/obo/ECO_0000255</a>	<a href="http://purl.uniprot.org/prosite-prorule/PRU00924">http://purl.uniprot.org/prosite-prorule/PRU00924</a>
<a href="http://purl.uniprot.org/uniprot/A0A024B7W1">http://purl.uniprot.org/uniprot/A0A024B7W1</a>	<a href="http://rdf.rhea-db.org/23680">http://rdf.rhea-db.org/23680</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000250">http://purl.obolibrary.org/obo/ECO_0000250</a>	<a href="http://purl.uniprot.org/uniprot/Q32ZE1">http://purl.uniprot.org/uniprot/Q32ZE1</a>
<a href="http://purl.uniprot.org/uniprot/A0A024B7W1">http://purl.uniprot.org/uniprot/A0A024B7W1</a>	<a href="http://rdf.rhea-db.org/60856">http://rdf.rhea-db.org/60856</a>	<a href="http://purl.obolibrary.org/obo/ECO_0000255">http://purl.obolibrary.org/obo/ECO_0000255</a>	<a href="http://purl.uniprot.org/prosite-prorule/PRU00924">http://purl.uniprot.org/prosite-prorule/PRU00924</a>

[...]



# UniProt Team

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