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Amos Marc Bairoch, Brigitte Boeckmann

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The SWISS-PROT protein sequence data bank

Amos Bairoch and Brigitte Boeckmann¹

Department of Medical Biochemistry, University of Geneva, 1 rue Michel Servet, 1211 Geneva 4, Switzerland and ¹European Molecular Biology Laboratory, Heidelberg, Germany

INTRODUCTION

SWISS-PROT is an annotated protein sequence database established in 1986 and maintained collaboratively, since 1988, by the Department of Medical Biochemistry of the University of Geneva and the EMBL Data Library [1].

SOURCES OF THE SEQUENCE DATA

Sequence data in SWISS-PROT originates from three different sources:

- From the Protein Sequence Database of the Protein Identification Resource (PIR) [2]
- From translation of entries from the EMBL Nucleotide Sequence Database [1]

From the literature

FORMAT

The SWISS-PROT protein sequence data bank is composed of sequence entries. Each sequence entry is composed of lines. Different types of lines, each with their own format, are used to record the various data which make up the entry. For standardization purposes the format of SWISS-PROT follows as closely as possible that of the EMBL Nucleotide Sequence Database [3]. A sample SWISS-PROT entry is shown in Figure 1.

WHAT DISTINGUISHES SWISS-PROT FROM OTHER PROTEIN SEQUENCE DATABASES?

Annotation

To be useful to the majority of users a protein sequence database should contain as much data as possible on each of the proteins that it describes. In SWISS-PROT, as in most other sequence databases, two classes of data can be distinguished: the core data and the annotation. For each sequence entry the core data consist of the following items:

Sequence data

Citation information (bibliographical references)

Taxonomic data (description of biological source of the protein)

The annotation consists of the description of the following items: Function(s) of the protein

Post-translational modification(s). For example carbohydrates, phosphorylation, acetylation, GPI-anchor, etc.

- Domains and sites. For example calcium binding regions, ATPbinding sites, zinc fingers, homeobox, kringle, etc.
- Secondary structure (using information from the DSSP database) [4]

Quaternary structure

Similarities to other proteins

Disease(s) associated with deficiencie(s) in the protein Sequence conflicts, variants, etc.

We try to include as much annotation information as possible in SWISS-PROT. To obtain this information we use, in addition to the publications that reports new sequence data, review articles to periodically update the annotations of families or groups of proteins. We also make use of external experts who send us their comments and updates concerning specific groups of proteins about which they are knowledgeable. We believe that our having systematic recourse both to publications other than those reporting the core data and to subject referees represents a unique and beneficial feature of SWISS-PROT.

In SWISS-PROT, annotation is mainly found in the comment lines (CC), in the feature table (FT) and in the keyword lines (KW). Most comments are classified by 'topics'; this approach permits the easy retrieval of specific categories of data from the database.

Minimal redundancy

We try as much as possible to minimize the redundancy of the database. Many sequence databases contain, for a given protein sequence, separate entries which correspond to different literature reports. Typically one finds a report that corresponds to a fragment of the protein sequenced at the level of the polypeptide, one or more reports reflecting the results of laboratories that have sequenced that protein at the cDNA level, and finally reports from data provided by genomic sequencing. This state of affairs has many drawbacks; for example it is not easy to obtain an overall view of the current state of the knowledge about a given protein many times during searches. In SWISS-PROT we try as much as possible to merge all these data and, if conflicts exist between various sequencing reports, we indicate them in the feature table.

Integration with other databases

It is important to provide the users of biomolecular databases with a degree of integration between the three types of sequencerelated databases (nucleic acid sequences, protein sequences and protein tertiary structures) as well as with specialized data collections. So as to provide tools that will allow software developers to implement such an integrated approach we have cross-referenced SWISS-PROT with many other databases. Currently cross-references are provided for the following databases:

EMBL Nucleotide Sequence Database [1]

PDB, the Brookhaven Protein Data Bank [5] which stores crystallographic coordinates of proteins

```
CAH2 HUMAN STANDARD; PRT; 259 AA.

P009T8;

21-JUL-1986 (REL. 01, CREATED)

21-JUL-1986 (REL. 01, LAST SEQUENCE UPDATE)

01-MAR-1992 (REL. 21, LAST ANNOTATION UPDATE)

CARBONIC ANHYDRASE II (EC 4.2.1.1) (CARBONATE DEHYDRATASE II).
ID
AC
DT
DŤ
ĐŤ
DE
GN
              CA2
             HOMO SAPIENS (HUMAN).
EUKARYOTA; METAZOA; CHORDATA; VERTEBRATA; TETRAPODA; MAMMALIA;
EUTHERIA; PRIMATES.
OS
ÕČ
ÕČ
RN
             [1]
SEQUENCE FROM N.A.
RP
RM
              87231043
             NONTGOMERY J.C., VENTA P.J., TASHIAN R.E., HEWETT-EMMETT D.;
NUCLEIC ACIDS RES. 15:4687-4687(1987).
 RA
RL
DN
             [2]
SEQUENCE FROM N.A.
 RP
              88085190
 RM
             MURAKANI H., MARELICH G.P., GRUBB J.H., KYLE J.W., SLY W.S.;
GENOMICS 1:159-166(1987).
 RA
RL
             RN
 RP
 RM
              77006079
             HENDERSON L.E., HENRIKSSON D., NYMAN P.O.;
J. BIOL. CHEM. 251:5457-5463(1976).
RA
RL
RN
               [4]
 RP
              SEQUENCE.
             74143468
LIN K.-T.D., DEUTSCH H.F.;
J. BIOL. CHEM. 249:2329-2337(1974).
 RM
RA
 RL
               [5]
 RN
 RP
              SEQUENCE OF 1-76 FROM N.A.
DM
              86077780
              VENTA P.J., MONTGOMERY J.C., HEWETT-EMMETT D., TASHIAN R.E.;
BIOCHIM. BIOPHYS. ACTA 826:195-201(1985).
 RA
 RL
 RN
              X-RAY CRYSTALLOGRAPHY, 2.0 ANGSTROMS.
72111787
 RP
RM
              LILJAS A., KANNAN K.K., BERGSTEN P.-C., WAARA I., FRIDBORG K.,
STRANDBERG B., CARLBOM U., JARUP L., LOVGREN S., PETEF M.;
NATURE NEW BIOL. 235:131-137(1972).
 RA
 RA
 RL
 RN
               171
              X-RAY CRYSTALLOGRAPHY, 2.0 ANGSTROMS.
 RP
              89315726
ERIKSSON A.E., JONES T.A., LILJAS A.;
PROTEINS 4:274-282(1988).
 RM
 RA
 RI
 RN
               [8]
               X-RAY CRYSTALLOGRAPHY, 2.0 ANGSTROMS.
 RP
              RIKSSON A.E., KYLSTEN P.M., JONES T.A., LILJAS A.;
PROTEINS 4:283-293(1988).
 RM
 RA
 RL
 RN
               [9]
             VÁRIANT JOGJAKARTA.
83100296
 RP
RM
              JONES G.L., SOFRO A.S.M., SHAW D.C.;
BIOCHEM. GENET. 20:979-1000(1982).
 RA
 RL
               [10]
 RN
             VARIANT MELBOURNE.
83236368
 RP
 RM
             83236368
JONES G.L., SHAW D.C.;
HUM. GENET. 63:392-399(1983).
-!- FUNCTION: REVERSIBLE HYDRATATION OF CARBON DIOXIDE.
-!- CATALYTIC ACTIVITY: H(2)CO(3) = CO(2) + H(2)O.
-!- THERE ARE AT LEAST 6 ENZYMATIC FORMS OF CARBONIC ANHYDRASE: CA-I
(OR B), CA-II (OR C), CA-III (OR M), CA-IV, CA-V AND CA-VI.
-!- DISEASE: DEFECTS IN CA2 ARE THE CAUSE OF OSTEOPETROSIS WITH RENAL
TUBULAR ACIDOSIS (MARBLE BRAIN DISEASE).
EMDI - VIOT3O. HSCA2
 RA
 RL
 CC
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ČČ
CC

        TUBLAR
        CLOSIS
        CAL ARE
        THE CAL

        TUBULAR
        ACIDOSIS
        (MARBLE BRAIN DISE/

        EMBL;
        Y00339; HSCA2.
        EMBL; X03251; HSCAII.

        EMBL;
        J03037; HSCAII.
        EMBL; X03251; HSCAII.

        EMBL;
        J03037; HSCAII.
        EMBL; X03251; HSCAIIA.

        PIR;
        A23202; A23202.
        PIR; A27175; A27175.

        PDB;
        1CA2; 15-JAN-90.
        PDB; 2CA2; 15-APR-90.

        PDB;
        2CA2; 15-APR-90.
        PMB; 2S9730; NINTH EDITION.

        PROSITE;
        PS00162; CARBONIC ANHYDRASE.
        LYASE; ACETYLATION; ZINC; 3D-STRUCTURE.

        INIT MET
        0
        0

        MOD RES
        1
        ACETYLATION; ZINC; 3D-STRUCTURE.

        INIT MET
        0
        0

        MCT_SITE
        63
        63

        ACT_SITE
        64
        66

        METAL
        93
        93
        ZINC (CATAI

DR
DR
D₽
DR
DR
DP
DR
DR
DR
DR
DR
KW
FT
 FŤ
                                                                                          ACETYLATION.
FT
FŤ
FT
                                                                                          ZINC (CATALYTIC).
                                              95
118
              METAL
                                                                  95
FT
                                                                                          ZINC (CATALYTIC).
ZINC (CATALYTIC).
                                                                 118
 FŤ
              METAL
                                             126
196
17
235
             ACT_SITE
ACT_SITE
VARTANT
FT
                                                                 126
                                                                 198
17
FT
                                                                                         K -> E (JOGJAKARTA).
P -> H (MELBOURNE).
 FŤ
                                                                235
FT
              VARIANT
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FT	VARIANT	251	251		N ->	D.		
FT	TURN	Ŕ	10	. · · ·				
FŤ	HELTX	12	18					
FT	HEIIX	20	23					
FŤ	THEN	25	24					
ĒŤ	STRAND	21	20					
ÉŤ	TIIDN	34	32	•				
FT	STRAND	79	20					
ET.	THON		27					
6T	STRAND	41	40					
6T	TIDA	40	47					
6T	STRAND	21	22					
ET.	STRAND	55	00					
ET	STRAND	0) 77	07					
51	JIKANU	//	00					
	CTDAND	01	82					
F	STRAND	100	90					
F 1	TUKN	100	101					
	STRAND	107	108					
<u></u>	IUKN	109	110					
<u></u>	STRAND	111	111					
	STRAND	115	123					
11	HELIX	124	126					
11	HELIX	129	132					
11	TURN	135	134					
11	TURN	136	137					
FI	STRAND	139	148					
FT	HELIX	153	165					
FI	TURN	168	169					
11	STRAND	1/1	1/3					
11	HELIX	1/9	182					
11	STRAND	189	194					
F1	TURN	199	200					
FI	STRAND	205	210					
FT	STRAND	214	216					
FT	HELIX	218	224					
FT	TURN	225	226					
FT	STRAND	228	228					
FT	TURN	232	233					
FT	STRAND	238	238					
FT	TURN	250	251					
FT	STRAND	255	256					
SQ	SEQUENCE	259 AA;	291	115 MW;	365	693 CN;		
	SHHWGYGKHN	GPEHWHK	DFP 1	AKGERÓS	PV D	IDTHTAKYD	PSLKPLSVSY	DOATSLRILN
	NGHAFNVEFD	DSQDKAV	LKG (PLDGTYR	LIQ	FHFHWGSLD	GQGSEHTVDK	KKYAAELHLV
	HWNTKYGDFG	KAVQQPD	GLA \	/LGIFLKV	GS AI	KPGLQKVVD	VLDSIKTKGK	SADETNEDPR
	GLLPESLDYW	TYPGSLT	TPP L	LECVTWI	VL KI	EPISVSSEQ	VLKFRKLNFN	GEGEPEELMV
	DNWRPAQPLK	NRQIKAS	FK					
11								

Figure 1. A sample entry from SWISS-PROT

- PIR, the protein sequence database of the Protein Identification Resource [2]
- EcoGene, from the EcoSeq/EcoMap integrated Escherichia coli database [6].
- FlyBase, the Drosophila Genetic database prepared by Michael Ashburner at the Department of Genetics, University of Cambridge.
- Gene-protein database of Escherichia coli K-12 (2D-gel spots) [7].
- HIV, the human retroviruses and AIDS Database [8]
- OMIM, the on-line version of the book 'Mendelian Inheritance in Man' [9]
- PROSITE, the Dictionary of Protein Sites and Patterns [10]
- REBASE, the restriction enzymes database [11] TFD, the transcription factors data bank [12]
- 11D, the transcription factors tata ballk [12]

Cross-references are provided in the form of pointers to information related to SWISS-PROT entries and found in data collections other than SWISS-PROT. They are implemented using a specific type of line, the 'DR' (for *D*ata bank *R*eference) line. For example the sample sequence shown in Figure 1 contains DR lines that point to EMBL, PIR, PDB, OMIM, and PROSITE. In that particular example it is therefore possible to retrieve the nucleic acid sequence(s) that encodes for that protein (EMBL), the X-ray crystallographic atomic coordinates (PDB), or the description of genetic disease(s) associated with that protein (OMIM).

CONTENT OF THE CURRENT RELEASE

Release 21.0 of SWISS-PROT (March 1992) contains 23742 sequence entries, comprising 7,866,596 amino acids abstracted from 23919 references. The data file (sequences and annotations) requires 40 Mb of disk storage space. The database is distributed with 15 documentation and index files (user's manual, release notes, list of organisms, citation index, keyword index, etc.) that require about 11 Mb of disk space.

DISTRIBUTION

SWISS-PROT is distributed on magnetic tape and on CD-ROM by the EMBL Data Library. The CD-ROM contains both SWISS-PROT and the EMBL Nucleotide Sequence Database as well as other data collections and some database query and retrieval software for MS-DOS PC compatible computers. For all enquiries regarding the subscription and distribution of SWISS-PROT one should contact:

EMBL Data Library European Molecular Biology Laboratory Postfach 10.2209, Meyerhofstrasse 1

6900 Heidelberg, Germany

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Telephone: (+49 6221) 387 258 Telefax : (+49 6221) 387 519 or 387 306 Electronic network address: datalib@EMBL-heidelberg.de

Individual sequence entries can be obtained from the EMBL File Server [13]. Detailed instructions on how to make best use of this service, and in particular on how to obtain protein sequences, can be obtained by sending to the network address netserv@EMBL-heidelberg.de the following message:

HELP HELP PROT

If you have access to a computer system linked to the Internet you can obtain SWISS-PROT using FTP (File Transfer Protocol), from the following file servers:

GenBank On-line Service [14] Internet address: genbank.bio.net (134.172.1.160)

NCBI (National Library of Medecine, NIH, Washington D.C., U.S.A.)

Internet address: ncbi.nlm.nih.gov (130.14.20.1)

ExPASy (Expert Protein Analysis System server, University of Geneva, Switzerland)

Internet address: expasy.hcuge.ch (129.195.254.61)

The present distribution frequency is four releases per year. No restrictions are placed on use or redistribution of the data.

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