

ENTREZ DIRECT - QUICK TOUR

Entrez Direct (EDirect) is an advanced method for accessing the NCBI's set of interconnected data domains (publication, nucleotide, protein, structure, variation, expression, etc.) from a terminal window. Multi-step queries can be built incrementally, and functions take search terms from command-line arguments. Record retrieval and formatting are normally the final steps in the query.

ENTREZ DIRECT FUNCTION SUMMARY

EDirect consists of a set of Perl scripts that are downloaded to the user's computer. Operations can be grouped into several categories.

Navigation functions support exploration within the Entrez databases:

| | |
|----------------------|--|
| <code>esearch</code> | Performs a new Entrez search using terms in indexed fields. |
| <code>elink</code> | Looks up neighbors (within a database) or links (between databases). |
| <code>efilter</code> | Filters or restricts the results of a previous query. |

Retrieval functions return document summaries or downloaded records:

| | |
|-----------------------|---|
| <code>esummary</code> | Obtains document summaries for the query results in XML format. |
| <code>efetch</code> | Retrieves records or generates reports in a designated format. |

Desired fields from XML results can be extracted without writing a program:

| | |
|---------------------|---|
| <code>xtract</code> | Converts XML into a table of data values. |
|---------------------|---|

Several additional functions are also provided:

| | |
|---------------------|--|
| <code>einfo</code> | Obtains information on indexed fields in an Entrez database. |
| <code>epost</code> | Uploads lists of unique identifiers or sequence accession numbers. |
| <code>nquire</code> | Sends a URL request to a web page or CGI service. |

ENTERING QUERY COMMANDS

UNIX programs are run by typing the name of the program and then supplying any required or optional arguments on the command line. Argument names are letters or words that start with a dash ("-") character.

In order to begin an Entrez search, the user types "esearch" and then enters the required -db (database) and -query arguments. For example, the command:

```
esearch -db pubmed -query "capsaicin cancer pain management"
```

constructs the appropriate Entrez Utilities (EUtils) URL from the query terms and executes the search. EDirect handles many technical details behind the scenes (avoiding the learning curve normally required for EUtils programming), and saves the results on the Entrez history server.

CONSTRUCTING MULTI-STEP QUERIES

The vertical bar ("|") UNIX pipe character connects two programs, sending the standard output of the first to the standard input of the second. (The last program in the chain simply prints to the terminal.) Piping esearch to elink:

```
esearch -db pubmed -query "capsaicin cancer pain management" | elink -related
```

will look up related articles (precomputed PubMed neighbors) of the initial results. (Using "elink -target" retrieves associated records between databases.)

ENTERING QUERIES ON MULTIPLE LINES

The same pair of commands can be written on two lines with the backslash ("\") UNIX line continuation character, for improved readability and easier editing:

```
esearch -db pubmed -query "capsaicin cancer pain management" | \  
elink -related
```

RESTRICTING QUERY RESULTS

The current results can be refined by further term searching in Entrez (useful in sequence databases for limiting BLAST neighbors to a taxonomic subset):

```
esearch -db pubmed -query "capsaicin cancer pain management" | \  
elink -related | \  
efilter -query "conotoxin NOT morphine"
```

QUALIFYING QUERIES BY INDEXED FIELD

Query terms in esearch or efilter can be qualified by entering an indexed field abbreviation in brackets. Boolean operators and parentheses can also be used in the query expression for more complex searches.

Commonly-used fields for pubmed queries include:

| | | | | | |
|--------|--------------------|--------|------------------|--------|------------------|
| [AFFL] | Affiliation | [FILT] | Filter | [MESH] | MeSH Terms |
| [ALL] | All Fields | [JOUR] | Journal | [PTYP] | Publication Type |
| [AUTH] | Author | [LANG] | Language | [WORD] | Text Word |
| [FAUT] | Author - First | [MAJR] | MeSH Major Topic | [TITL] | Title |
| [LAUT] | Author - Last | [SUBH] | MeSH Subheading | [TIAB] | Title/Abstract |
| [PDAT] | Date - Publication | | | [UID] | UID |

and a qualified query looks like:

```
"Tager H [AUTH] AND glucagon [TIAB]"
```

Sequence databases are indexed with a different set of search fields, including:

| | | | | | |
|--------|--------------|--------|-------------------|--------|-----------------|
| [ACCN] | Accession | [GENE] | Gene Name | [PROT] | Protein Name |
| [ALL] | All Fields | [JOUR] | Journal | [SQID] | SeqID String |
| [AUTH] | Author | [KYWD] | Keyword | [SLEN] | Sequence Length |
| [GPRJ] | BioProject | [MLWT] | Molecular Weight | [SUBS] | Substance Name |
| [ECNO] | EC/RN Number | [ORGN] | Organism | [WORD] | Text Word |
| [FKEY] | Feature Key | [PACC] | Primary Accession | [TITL] | Title |
| [FILT] | Filter | [PROP] | Properties | [UID] | UID |

and a sample query in the protein database is:

```
"alcohol dehydrogenase [PROT] NOT (bacteria [ORGN] OR fungi [ORGN])"
```

The scripting examples at the end of this document will produce a report giving the entire set of indexed fields for every Entrez database.

EXAMINING INTERMEDIATE RESULTS

EDirect navigation functions produce a small structured XML object that is passed to the next command. Retrieval functions read the XML values and return the designated report from the history server.

The Count field in the XML intermediate gives the number of records returned by the previous step. A good measure of success is a reasonable (non-zero) value:

```
<ENTREZ_DIRECT>
  <Db>pubmed</Db>
  <WebEnv>NCID_1_4586177_172.16.22.25_5555_1366675055_388385336</WebEnv>
  <QueryKey>4</QueryKey>
  <Count>10</Count>
  <Step>3</Step>
</ENTREZ_DIRECT>
```

Checking the count before adding the next step helps avoid unsuccessful queries.

RETRIEVING PUBMED REPORTS

Piping the query results to efetch and specifying the "abstract" format:

```
esearch -db pubmed -query "capsaicin cancer pain management" | \
elink -related | \
efilter -query "conotoxin NOT morphine" | \
efetch -format abstract
```

returns a set of reports that can be read by a person:

1. PLoS One. 2013;8(3):e59293. doi: 10.1371/journal.pone.0059293. Epub ...

Expression and Pharmacology of Endogenous Cav Channels in SH-SY5Y Human Neuroblastoma Cells.

Sousa SR, Vetter I, Ragnarsson L, Lewis RJ.

Institute for Molecular Bioscience, The University of Queensland, St. Lucia, Australia.

SH-SY5Y human neuroblastoma cells provide a useful in vitro model to study the mechanisms underlying neurotransmission and nociception. These cells are derived ...

Using "efetch -format medline" instead produces a report that can be entered into common bibliographic management software packages:

```
PMID- 23536870
OWN - NLM
STAT- In-Data-Review
DA - 20130328
IS - 1932-6203 (Electronic)
IS - 1932-6203 (Linking)
VI - 8
IP - 3
DP - 2013
TI - Expression and Pharmacology of Endogenous Cav Channels in SH-SY5Y Human
    Neuroblastoma Cells.
PG - e59293
LID - 10.1371/journal.pone.0059293 [doi]
...
```

XML DOCUMENT SUMMARIES

EDirect also provides a document summary in structured XML format for every record in every database. Piping to esummary instead of efetch:

```

esearch -db pubmed -query "capsaicin cancer pain management" | \
elink -related | \
efilter -query "conotoxin NOT morphine" | \
esummary

```

will generate an XML document summary set:

```

<eSummaryResult>
  <DocumentSummarySet status="OK">
    <DocumentSummary uid="23536870">
      <Id>23536870</Id>
      <PubDate>2013</PubDate>
      <EPubDate>2013 Mar 25</EPubDate>
      <Source>PLOS One</Source>
      <Authors>
        <Author>
          <Name>Sousa SR</Name>
          <AuthType>Author</AuthType>
          <ClusterID>0</ClusterID>
        </Author>
        <Author>
          <Name>Vetter I</Name>
        </Author>
        ...
      </Authors>
    </DocumentSummary>
  </DocumentSummarySet>
</eSummaryResult>

```

The advantage of XML is that many pieces of information are in specific locations in a well-defined data hierarchy. Assembling individual units of data that are fielded by name, such as:

```

<PubDate>2013</PubDate>
<Source>PLOS One</Source>
<Volume>8</Volume>
<Issue>3</Issue>
<Pages>e59293</Pages>

```

requires matching the same general pattern, differing only by the element name. This is much easier than parsing the units from a long, complex string:

```
1. PLOS One. 2013;8(3):e59293 ...
```

The disadvantage of XML is that data extraction usually requires programming. But EDirect relies on the common pattern of XML value representation to provide a simplified approach to interpreting XML data, as discussed below.

CONVERSION OF XML DATA INTO TABULAR FORM

The xtract function uses command-line arguments to direct the selective conversion of XML data into a tab-delimited table. The -pattern argument divides the results into rows, while placement of data into columns is controlled by -element.

Additional arguments can limit data extraction to specified regions of the XML (-group, -block, and -subset), filter by data content (-match and -avoid) or element names (-present and -absent), and customize the table presentation.

Piping the esummary output to:

```
xtract -outline
```

will give an indented overview of the XML structure hierarchy:

```

DocumentSummarySet
  DocumentSummary
    Id
    PubDate
    EPubDate
    Source
    Authors
      Author
        Name
        AuthType
        ClusterID
    ...

```

The outline can help in deciding what arguments to send to xtract. Thus:

```

esearch -db pubmed -query "capsaicin cancer pain management" | \
elink -related | \
efilter -query "conotoxin NOT morphine" | \
esummary | \
xtract -pattern DocumentSummary -element Id SortFirstAuthor Title

```

returns the PubMed identifier (PMID), first author name, and article title:

```

23536870  Sousa SR  Expression and Pharmacology of Endogenous Cav Channels ...
22410003  Vetter I   Characterisation of Na(v) types endogenously expressed ...
18956616  Fürst Z    [Central and peripheral mechanisms in antinociception: ...
12566085  Lo YK      Effect of arvanil (N-arachidonoyl-vanillyl-amine), a n ...
...

```

INTERACTION WITH UNIX UTILITIES

A tab-delimited table can be processed by many UNIX utilities. For example:

```

esearch -db pubmed -query "capsaicin cancer pain management" | \
elink -related | \
efilter -query "conotoxin NOT morphine" | \
esummary | \
xtract -pattern DocumentSummary -element Id SortFirstAuthor Title | \
sort -t '$\t' -k 2,2f -k 3,3f

```

takes the table from the previous example and sorts first by author name and then (if there are multiple publications by the same author) alphabetically by title:

```

11000661  Chiang JS  New developments in cancer pain therapy.
18956616  Fürst Z    [Central and peripheral mechanisms in antinociception ...
10864900  Jerman JC  Characterization using FLIPR of rat vanilloid recepto ...
12566085  Lo YK      Effect of arvanil (N-arachidonoyl-vanillyl-amine), a ...
...

```

Alternatively, a frequency table of title words is easily generated by passing the titles to a different set of UNIX commands:

```

esearch -db pubmed -query "capsaicin cancer pain management" | \
elink -related | \
efilter -query "conotoxin NOT morphine" | \
esummary | \
xtract -pattern DocumentSummary -element Title | \
sed 's/[^a-zA-Z0-9]/ /g' | tr 'A-Z' 'a-z' | xargs -n 1 | \
sort | uniq -c | sort -k 1,1nr -k 2,2f

```

The "sed" (stream editor) command takes the article titles and removes punctuation and accented characters, replacing them with a space and thus creating separate words. Piping to "tr" (translate) converts capital letters to lower case. The "xargs -n 1" command places each resulting word on a separate line. The words are then sorted alphabetically (by "sort"), and occurrence counts for each unique word are calculated (by "uniq -c"). Finally, those results are sorted by frequency:

```
8 of
7 in
4 cells
4 human
4 neuroblastoma
4 sh
4 sy5y
...
```

PUBMED ARTICLE XML RECORDS

The PubmedArticle object has a more detailed structure than the DocumentSummary, and is only available for records in the pubmed database:

```
esearch -db pubmed -query "capsaicin cancer pain management" | \
elink -related | \
efilter -query "conotoxin NOT morphine" | \
efetch -format xml | \
xtract -outline
```

More information is fielded, including author names and the abstract:

```
PubmedArticleSet
  PubmedArticle
    MedlineCitation
      PMID
      DateCreated
      Year
      Month
      Day
      Article
        Journal
          ISSN
          JournalIssue
          Volume
          Issue
          PubDate
          Year
          Title
          ISOAbbreviation
        ArticleTitle
        Pagination
          MedlinePgn
        ELocationID
        Abstract
          AbstractText
        Affiliation
        AuthorList
          Author
            LastName
            ForeName
            Initials
          ...
```

Using this information to craft a new xtract statement:

```
esearch -db pubmed -query "capsaicin cancer pain management" | \
elink -related | \
efilter -query "conotoxin NOT morphine" | \
efetch -format xml | \
xtract -pattern PubmedArticle -element "MedlineCitation/PMID" LastName
```

results in a table of all authors for each record:

```
23536870 Sousa Vetter Ragnarsson Lewis
22410003 Vetter Mozar Durek Wingerd Alewood Christie Lewis
18956616 Fürst
12566085 Lo Chiang Wu
...
```

(Note that -element "MedlineCitation/PMID" uses the parent/child construct to prevent picking up PMIDs that occur later in CommentsCorrections objects. The -first or -last arguments can be used instead of -element, if appropriate.)

EXPLORATION OF XML SETS

Individual PubmedArticle objects can be retrieved directly by efetch:

```
efetch -db pubmed -id 1937004 -format xml
```

The resulting XML may contain a list of Medical Subject Headings (MeSH Terms):

```
...
<MeshHeadingList>
  <MeshHeading>
    <DescriptorName>Adenosine Triphosphatases</DescriptorName>
  </MeshHeading>
  <MeshHeading>
    <DescriptorName>Amino Acid Sequence</DescriptorName>
  </MeshHeading>
  <MeshHeading>
    <DescriptorName>Base Sequence</DescriptorName>
  </MeshHeading>
  ...

```

A -block statement visits each individual MeSH object. Customizing the output format (by changing the tab between fields to a newline character):

```
efetch -db pubmed -id 1937004 -format xml | \
xtract -pattern PubmedArticle -tab "\n" -element "MedlineCitation/PMID" \
-block MeshHeading -tab "\n" -element DescriptorName
```

produces a list of MeSH terms, one per line, following the PubMed ID:

```
1937004
Adenosine Triphosphatases
Amino Acid Sequence
Base Sequence
...
Recombination, Genetic
Saccharomyces cerevisiae
Saccharomyces cerevisiae Proteins
```

NESTED EXPLORATION OF SUBSETS WITHIN XML SETS

Some MeSH terms in a record may be assigned subheadings:

```
...
<MeshHeading>
  <DescriptorName>Recombination, Genetic</DescriptorName>
  <QualifierName>genetics</QualifierName>
</MeshHeading>
<MeshHeading>
  <DescriptorName>Saccharomyces cerevisiae</DescriptorName>
  <QualifierName>genetics</QualifierName>
  <QualifierName>radiation effects</QualifierName>
</MeshHeading>
<MeshHeading>
  <DescriptorName>Saccharomyces cerevisiae Proteins</DescriptorName>
</MeshHeading>
</MeshHeadingList>
...
```

Adding a `-subset` statement within the `-block` allows nested exploration of the subheadings for the current MeSH term:

```
efetch -db pubmed -id 1937004 -format xml | \
xtract -pattern PubmedArticle -tab "" -element "MedlineCitation/PMID" \
  -block MeshHeading -pfx "\n" -tab "" -element DescriptorName \
    -subset QualifierName -pfx "/" -tab "" -element QualifierName
```

and results in a list of MeSH terms and associated subheadings:

```
1937004
Adenosine Triphosphatases
Amino Acid Sequence
Base Sequence
...
Recombination, Genetic/genetics
Saccharomyces cerevisiae/genetics/radiation effects
Saccharomyces cerevisiae Proteins
```

EXPLORING SEPARATE XML REGIONS

Multiple `-block` statements can be used in a single `xtract` to explore different areas of the XML. This limits element extraction to the desired subregion, and allows disambiguation of fields with identical names.

Combining fields with commas allows them to be treated as sets. The tab that normally separates these can be replaced with a `-sep` argument:

```
efetch -db pubmed -id 781293,2678811,6301692,8332518 -format xml | \
xtract -pattern PubmedArticle -element "MedlineCitation/PMID" \
  -block AuthorList -sep "/" -element LastName "#Author" \
  -block PubDate -sep " " -element Year,Month MedlineDate \
  -block DateCreated -sep "-" -element Year,Month,Day | \
sort -t '$\t' -k 3,3n | \
column -s '$\t' -t
```

produces a table that allows easy parsing of author names, counts the number of authors present, and prints the date each record was published and the date it was entered into PubMed, sorting the results by the computed author count:

| | | | | |
|---------|----------------------------------|---|--------------|------------|
| 781293 | Casadaban | 1 | 1976 Jul | 1976-10-02 |
| 6301692 | Krasnow/Cozzarelli | 2 | 1983 Apr | 1983-06-17 |
| 8332518 | Benson/Lipman/Ostell | 3 | 1993 Jul | 1993-08-17 |
| 2678811 | Mortimer/Schild/Contopoulou/Kans | 4 | 1989 Sep-Oct | 1989-11-22 |

(The PubDate object can exist either in structured or string form, but would not contain a mixture of both types, so "-element Year,Month MedlineDate" will only contribute a single column to the output.)

RECORDING VALUES IN VARIABLES

A value can be recorded in a variable and then displayed multiple times as needed. Variables are indicated by a hyphen followed by a string of capital letters or digits. The variable "-PMID" is referred to as "&PMID" in an -element argument.

For example:

```
efetch -db pubmed -id 1937004 -format xml | \
xtract -pattern PubmedArticle -PMID "MedlineCitation/PMID" \
-block MeshHeading -tab "\n" -element "&PMID",DescriptorName
```

produces a list of MeSH terms, with the PMID in the first column of each row:

```
1937004 Adenosine Triphosphatases
1937004 Amino Acid Sequence
1937004 Base Sequence
...
1937004 Recombination, Genetic
1937004 Saccharomyces cerevisiae
1937004 Saccharomyces cerevisiae Proteins
```

CONDITIONAL PROCESSING

Xtract provides arguments that filter by data content (-match and -avoid) or element names (-present and -absent). Parallel statements are used to handle alternative conditions. For example:

```
efetch -db pubmed -id 1937004 -format xml | \
xtract -pattern PubmedArticle -PMID "MedlineCitation/PMID" \
-group MeshHeading \
-block MeshHeading -present QualifierName \
-subset DescriptorName -TERM "DescriptorName" \
-subset QualifierName -tab "\n" -element "&PMID","&TERM",QualifierName \
-block MeshHeading -absent QualifierName \
-subset DescriptorName -tab "\n" -element "&PMID",DescriptorName
```

has separate sections for MeSH terms with and without subheadings, and produces:

```
1937004 Adenosine Triphosphatases
1937004 Amino Acid Sequence
1937004 Base Sequence
...
1937004 Recombination, Genetic          genetics
1937004 Saccharomyces cerevisiae        genetics
1937004 Saccharomyces cerevisiae        radiation effects
1937004 Saccharomyces cerevisiae Proteins
```

SELECTING XML ATTRIBUTES

The MeSH term and subheading fields actually have major topic attributes:

```

<MeshHeading>
  <DescriptorName MajorTopicYN="N">Saccharomyces cerevisiae</DescriptorName>
  <QualifierName MajorTopicYN="Y">genetics</QualifierName>
  <QualifierName MajorTopicYN="N">radiation effects</QualifierName>
</MeshHeading>

```

that can be selected as "DescriptorName@MajorTopicYN" or simply "@MajorTopicYN":

```

efetch -db pubmed -id 1937004 -format xml | \
xtract -pattern PubmedArticle -PMID "MedlineCitation/PMID" \
-group MeshHeading \
-block MeshHeading -present QualifierName \
-subset DescriptorName -TERM "DescriptorName" -MAJR "@MajorTopicYN" \
-subset QualifierName -tab "\n" \
-element "&PMID","&TERM","&MAJR",QualifierName,"@MajorTopicYN" \
-block MeshHeading -absent QualifierName \
-subset DescriptorName -tab "\n" \
-element "&PMID",DescriptorName,"@MajorTopicYN"

```

The results are suitable for importing into a database or spreadsheet program:

| | | | | |
|---------|-----------------------------------|---|-------------------|---|
| 1937004 | Adenosine Triphosphatases | N | | |
| 1937004 | Amino Acid Sequence | N | | |
| 1937004 | Base Sequence | N | | |
| ... | | | | |
| 1937004 | Recombination, Genetic | N | genetics | N |
| 1937004 | Saccharomyces cerevisiae | N | genetics | Y |
| 1937004 | Saccharomyces cerevisiae | N | radiation effects | N |
| 1937004 | Saccharomyces cerevisiae Proteins | Y | | |

STORING COMMON PHRASES IN ALIAS FILES

Long or complicated search phrases can be saved in a file to avoid having to retype (or copy and paste) the full text for each query. Each line of the file has a shortcut keyword, a tab character, and the expanded search term. Shortcuts are referenced by placing them in parentheses and prefixing with a pound ("#") sign.

For example, given a file named "query_aliases" containing:

```

jour_filt      [MULT] AND ncbijournals [FILT]
trans_imm      (transposition OR target) immunity

```

the esearch line in:

```

esearch -alias query_aliases -db nlmcatalog -query "Science (#jour_filt)" | \
esummary | \
xtract -pattern DocumentSummary -element ISOAbbreviation \
-subset ISSNInfo -sep "|" -element issn,issntype | \
column -s $'\t' -t

```

will be expanded to:

```

esearch -db nlmcatalog -query "Science [MULT] AND ncbijournals [FILT]"

```

with the query producing:

| | | |
|------------------------|-----------------|--------------------------|
| J. Zhejiang Univ. Sci. | 1009-3095 Print | 1009-3095 Linking |
| Science (80-) | 0193-4511 Print | 0193-4511 Linking |
| Science | 0036-8075 Print | 1095-9203 Electronic ... |

LINKING TO SEQUENCE DATABASES

Using elink with the -target argument will link to associated records in the destination database:

```
esearch -db pubmed -query "conotoxin AND nicotinic acetylcholine receptor" | \
elink -target protein
```

Filtering can then be done using indexed fields in the protein database:

```
esearch -db pubmed -query "conotoxin AND nicotinic acetylcholine receptor" | \
elink -target protein | \
efilter -query "mat_peptide [FKEY]"
```

RETRIEVING SEQUENCE REPORTS

Nucleotide and protein sequences can be retrieved in FASTA format:

```
efetch -db protein -id 26418308 -format fasta
```

which is the input format for many sequence analysis programs:

```
>gi|26418308|gb|AAN78128.1| alpha-conotoxin ImI precursor [Conus imperialis]
IVRRGCCSDPRCAWRCG
```

Sequence records can also be retrieved in GenBank or GenPept flatfile formats:

```
efetch -db protein -id 26418308 -format gp
```

which has features annotating particular regions of the sequence:

```
LOCUS      AAN78128                      17 aa          linear   INV 03-JAN-2003
DEFINITION alpha-conotoxin ImI precursor, partial [Conus imperialis].
ACCESSION  AAN78128
VERSION    AAN78128.1  GI:26418308
DBSOURCE   accession AY159318.1
KEYWORDS   .
SOURCE     Conus imperialis
  ORGANISM Conus imperialis
            Eukaryota; Metazoa; Lophotrochozoa; Mollusca; Gastropoda;
            Caenogastropoda; Hypsogastropoda; Neogastropoda; Conoidea; Conidae;
            Conus.
```

...

```
FEATURES             Location/Qualifiers
    source             1..17
                        /organism="Conus imperialis"
                        /db_xref="taxon:35631"
    Protein            <1..17
                        /product="alpha-conotoxin ImI precursor"
    mat_peptide        5..16
                        /product="alpha-conotoxin ImI"
                        /note="the C-terminal glycine of the precursor is post
                        translationally removed"
    CDS                1..17
                        /coded_by="AY159318.1:<1..54"
                        /note="nAChR antagonist"
```

```
ORIGIN
      1 ivrrgccsdprcawrcg
```

```
//
```

SEQUENCE RECORDS IN INSDSEQ XML

Sequence records can be retrieved in an XML version of the GenBank or GenPept flatfile. The query:

```
efetch -db protein -id 26418308 -format gpc -mode xml
```

returns INSDSeq XML, which includes structured feature table information:

```
...
<INSDFeature>
  <INSDFeature_key>mat_peptide</INSDFeature_key>
  <INSDFeature_location>5..16</INSDFeature_location>
  <INSDFeature_intervals>
    <INSDInterval>
      <INSDInterval_from>5</INSDInterval_from>
      <INSDInterval_to>16</INSDInterval_to>
      <INSDInterval_accession>AAN78128.1</INSDInterval_accession>
    </INSDInterval>
  </INSDFeature_intervals>
  <INSDFeature_qual>
    <INSDQualifier>
      <INSDQualifier_name>product</INSDQualifier_name>
      <INSDQualifier_value>alpha-conotoxin ImI</INSDQualifier_value>
    </INSDQualifier>
    <INSDQualifier>
      <INSDQualifier_name>note</INSDQualifier_name>
      <INSDQualifier_value>the C-terminal glycine of the precursor is
        post translationally removed</INSDQualifier_value>
    </INSDQualifier>
    <INSDQualifier>
      <INSDQualifier_name>calculated_mol_wt</INSDQualifier_name>
      <INSDQualifier_value>1357</INSDQualifier_value>
    </INSDQualifier>
    <INSDQualifier>
      <INSDQualifier_name>peptide</INSDQualifier_name>
      <INSDQualifier_value>GCCSDPRCAWRC</INSDQualifier_value>
    </INSDQualifier>
  </INSDFeature_qual>
</INSDFeature>
...
```

Feature and qualifier names are data values, not XML element tags, and require -match to select the desired object. A query for snail venom mature peptides:

```
esearch -db pubmed -query "conotoxin AND nicotinic acetylcholine receptor" | \
elink -target protein | \
efilter -query "mat_peptide [FKEY]" | \
efetch -format gpc -mode xml | \
xtract -pattern INSDSeq -ACCN INSDSeq_accession-version \
-group INSDFeature -match ">mat_peptide<" \
-avoid "<INSDFeature_partial" -pfx "\n" -element "&ACCN" \
-block INSDQualifier -match ">peptide<" -element "%INSDQualifier_value" \
-block INSDQualifier -match ">product<" -element INSDQualifier_value \
-block INSDQualifier -match ">peptide<" -element INSDQualifier_value | \
grep conotoxin | \
sort -t '$\t' -u -k 3,4 | \
sort -t '$\t' -k 2,2n -k 3,3f | \
column -s '$\t' -t
```

calculates the length of each mature peptide, and prints the product name and peptide sequence, removing redundant entries and sorting by peptide length:

```
AAN78128.1 12 alpha-conotoxin ImI GCCSDPRCAWRC
AAN78127.1 12 alpha-conotoxin ImII ACCSDRRRCRWRC
AAO33169.1 16 alpha-conotoxin GIC GCCSHPACAGNNQHIC
ABE27010.1 25 conotoxin fel4.1 SPGSTICKMACRTGNGHKYPFCNCR
ABE27011.1 25 conotoxin fel4.2 SSGSTVCKMMCRLLGYGHLYPSCGCR
...
```

(The sequence accession is captured in a variable for use with each `mat_peptide` in the record. Prefix substitution ensures that every peptide is shown on a separate line. This also works for multi-product precursor proteins such as proinsulin.)

(Incorporating fragments of the surrounding XML tags in `-match` arguments prevents coincidental matches to free text in note qualifiers.)

Multiple `-avoid` or `-match` conditions are specified with `-and` and `-or` commands:

```
-group INSDFeature -avoid ">proprotein<" -and ">sig_peptide<" \
-block INSDQualifier -match ">calculated_mol_wt<" -or ">peptide<" \
```

EXTENSIVE MODIFICATION OF TABULAR OUTPUT

The normal column-oriented output can be extensively modified to produce a custom report. A query on a gene that undergoes messenger RNA splicing:

```
efetch -db nuccore -id "GQ370762.1" -format gbc -mode xml | \
xtract -pattern INSDSeq -pfx ">Feature " -sfx "\n" -tab "" -first INSDSeqid \
-group INSDFeature -avoid "INSDFeature_key>source<" -FKEY INSDFeature_key \
-block INSDInterval -element INSDInterval_from INSDInterval_to \
INSDInterval_point INSDInterval_point "&FKEY" -FKEY "" -tab "" -lbl "\n" \
-block INSDQualifier -avoid ">translation<" -and ">peptide<" -pfx "\t\t\t" \
-sfx "\n" -tab "" -element INSDQualifier_name,INSDQualifier_value
```

clears the feature key variable after its first use to reproduce the 5-column feature table format used for GenBank submissions:

```
>Feature gb|GQ370762.1|
51      1474      gene
                    gene          HBB
51      142      mRNA
273     495
1346    1474
                    gene          HBB
                    product       beta-globin
51      142      CDS
273     495
1346    1474
                    gene          HBB
                    codon_start    1
                    transl_table    1
                    product       beta-globin
                    protein_id     ACU56984.1
                    db_xref        GI:256028940
```

(A location interval will have either an `INSDInterval_from` and `INSDInterval_to` pair or a single `INSDInterval_point`. Additional work would be required to support the 5' and 3' partial flags for features with incomplete locations.)

The actual 5-column feature table representation of any sequence record can be obtained directly by using "efetch -format ft".

AUTOMATION WITH SCRIPTS

Taking an adventurous plunge into the world of programming, a shell script can be written when each output line of one step needs to be processed independently, instead of output being piped in its entirety to the next command. (The simplest shell script can be merely a copy of the set of commands that are typed into the terminal for execution.)

In shell scripts, variables can be set to the results of a command by enclosing the statements in backtick ("`) characters. The variable name is prefixed by a dollar sign ("\$") to use its value as an argument in another command. Comments start with a pound sign ("#") and are ignored. Quotation marks within quoted strings are "escaped" with a backslash ("\"). Subroutines can be used to collect common code or simplify the organization of the script.

For example:

```
#!/bin/bash -norc

parse_fields() {
    echo "$1" | \
    xtract -pattern Field -element Name FullName Description | \
    sort -t '$\t' -k 2,2f | \
    perl -pe 's/(.+?)\t(.+?)\t(.+)/\[ $1\]\t$2\t$3/' | \
    column -s '$\t' -t
}

dbs=`einfo -dbs | xtract -pattern DbName -element DbName | sort`

for db in $dbs; do
    eix=`einfo -db $db`

    echo "$db"
    echo ""
    flds=`parse_fields "$eix"`
    echo "$flds"
    echo ""

    sleep 1
done
```

will obtain the list of Entrez databases, and then return the abbreviations, names, and descriptions of indexed search fields, for each individual database:

```
...
epigenomics

[ACCN]  Accession      Accession number of sequence
[ALL]   All Fields     All terms from all searchable fields
[AUTH]  Author         Author
[CDAT]  Create Date    CreateDate
[DOCT]  Document Type  DocType
[FILT]  Filter         Limits the records
[KYWD]  Keyword        Keyword
...
```

AUTOMATION WITHOUT SCRIPTS

Creative use of the "sh" and "xargs" commands can obtain the same behavior from the command line, without the need to write separate script files. Within the "sh" command string, the database argument passed by xargs is substituted at the "\$0" variable. All of the sh commands are run separately on each database.

Because the sh instructions are enclosed between single quote ("' ") characters (apostrophes), in-line perl commands cannot be used, nor can sort be told to use tabs (instead of tabs or spaces) to delimit columns. But a combination of xtract customization and the trick of temporarily replacing spaces with asterisks:

```
einfo -dbs | xtract -pattern DbName -element DbName | sort | \
xargs -n 1 sh -c 'einfo -db "$0" | \
  xtract -pattern DbInfo -tab "\n\n" -element DbName \
    -block Field -pfx "[" -sep "]" -tab "\n" -element Name,FullName | \
  sed "s/ / */g" | sort -k 2,2f | sed "s/* / /g" | expand'
```

allows the desired sorting by full field name and produces results that are alphabetized for each database:

```
...
pubmed
[AFFL] Affiliation
[ALL] All Fields
[AUTH] Author
[COLN] Author - Corporate
[FAUT] Author - First
[FULL] Author - Full
[LAUT] Author - Last
[AUCL] Author Cluster ID
[BOOK] Book
[CDAT] Date - Completion
[CRDT] Date - Create
...
```

APPENDICES

Common command-line arguments for the EDirect functions are shown below:

esearch

-db -query

elink

-db -id -target -name -related -batch

efilter

-query

esummary

-db -id

efetch

-db -id -format -mode
-seq_start -seq_stop -strand -complexity

xtract

```

-pattern  -division  -group    -branch
-block    -section   -subset    -unit
-match    -avoid     -present   -absent  -and  -or
-element  -first     -last
-pfx      -sfx       -sep       -tab     -ret  -lbl
-outline  -synopsis

```

Efetch -format and -mode values for each database are shown below:

| -db | -format | -mode | Record Type |
|------------|--|--------------|--|
| (all) | full uid url xml | | Same as native except for mesh Unique Identifier List Entrez URL Converted to -format full -mode xml |
| bioproject | native native | xml | BioProject Report RecordSet XML |
| biosample | native native | xml | BioSample Report BioSampleSet XML |
| biosystems | native | xml | Sys-set XML |
| gds | native summary | xml | RecordSet XML Summary |
| gene | gene_table native native native | asn.1 xml | Gene Table Gene Report Entrezgene ASN.1 Entrezgene-Set XML |
| homologene | alignmentscores fasta homologene native native native | asn.1 xml | Alignment Scores FASTA Homologene Report Homologene List HG-Entry ASN.1 Entrez-Homologene-Set XML |
| mesh | full native native | xml | Full Record RecordSet XML MeSH Report |
| nlmcatalog | native native | xml | Full Record NLMCatalogRecordSet XML |

| | | | |
|-------------|--------------|-------|-------------------------------|
| pmc | medline | | MEDLINE |
| | native | xml | pmc-articleSet XML |
| pubmed | abstract | | Abstract |
| | medline | | MEDLINE |
| | native | asn.1 | Pubmed-entry ASN.1 |
| | native | xml | PubmedArticleSet XML |
| (sequences) | acc | | Accession Number |
| | est | | EST Report |
| | fasta | | FASTA |
| | fasta | xml | TinySeq XML |
| | fasta_cds_aa | | FASTA of CDS Products |
| | fasta_cds_na | | FASTA of Coding Regions |
| | ft | | Feature Table |
| | gb | | GenBank Flatfile |
| | gb | xml | GBSet XML |
| | gbc | xml | INSDSet XML |
| | gbwithparts | | GenBank with Contig Sequences |
| | gp | | GenPept Flatfile |
| | gp | xml | GBSet XML |
| | gpc | xml | INSDSet XML |
| | gss | | GSS Report |
| | native | asn.1 | Seq-entry ASN.1 |
| | native | xml | Bioseq-set XML |
| | seqid | | Seq-id ASN.1 |
| snp | chr | | Chromosome Report |
| | docset | | Summary |
| | fasta | | FASTA |
| | flt | | Flat File |
| | native | asn.1 | Rs ASN.1 |
| | native | xml | ExchangeSet XML |
| | rsr | | RS Cluster Report |
| | ssexemplar | | SS Exemplar List |
| sra | native | xml | EXPERIMENT_PACKAGE_SET XML |
| | | | |
| structure | mmdb | | Ncbi-mime-asn1 strucseq ASN.1 |
| | native | | MMDB Report |
| | native | xml | RecordSet XML |
| taxonomy | native | | Taxonomy List |
| | native | xml | TaxaSet XML |

FOR MORE INFORMATION

The comprehensive user GUIDE contains the definitive documentation of EDirect functionality.