# Supplementary material for EST2Prot: Mapping EST sequences to proteins

Paul Shafer<sup>1</sup>, David M. Lin<sup>2</sup> and Golan Yona<sup>1,\*</sup>

 $^{1}$  Department of Computer Science, Cornell University, Ithaca, NY

 $^2$  Department of Biomedical Sciences, Cornell University, Ithaca, NY

\*Corresponding author. Email: golan@cs.cornell.edu

# 1 Appendix - The EST2Prot webserver

The EST mapping system consists of 5 pages: the upload page, the summary page, the EST map page, the descriptor page, and the path page. The user starts with the upload page, which allows the user to submit ESTs for analysis. The user is then taken to the summary page, which summarizes each uploaded EST by displaying descriptors of proteins associated with it, and other pages with detailed information about the mapping. An overview of the webserver is given in Figure 1.



Figure 1: The Biozon EST2Prot webserver.

#### 1.1 The Upload Page

The upload page (Figure 2) allows the user to upload a list of ESTs for analysis. This is done by specifying either the Genbank accession number or Genbank GI number of each EST. The user may upload a file of identifiers or type their identifiers into a text box.

#### 1.2 The Summary Page

The *summary page* summarizes the possible functions of each uploaded EST (Figure 3). This page has four columns. The first column displays the Genbank identifier of the uploaded EST. If the identifier was found in Biozon's local copy of Genbank, the user may click on the identifier to view Biozon's record of the corresponding nucleic acid sequence.

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| Analyze EST dat              | ta                         |                                    |                                                        |
| Andry Lo Lo I da             |                            |                                    |                                                        |
| Before submitting you        | ır file, read carefully tl | ne <u>instructions</u> . Please no | ote: we currently analyze only mouse and               |
| human ESTs                   |                            |                                    |                                                        |
|                              |                            |                                    |                                                        |
| Upload your list of E9       | iT identifiers (use Gen    | Bank accession number              | <b>s or GI numbers)</b> (see <u>example file</u> )     |
|                              | Browse                     |                                    |                                                        |
| OR paste your list he        | re                         |                                    |                                                        |
| AI834965                     |                            |                                    |                                                        |
| AI834966                     |                            |                                    |                                                        |
| AI834969                     |                            |                                    |                                                        |
| AI834970                     |                            |                                    |                                                        |
| A1834973<br>AT834974         |                            |                                    |                                                        |
| A1834975                     |                            |                                    |                                                        |
| AI834976                     |                            |                                    |                                                        |
| A1834978                     |                            |                                    |                                                        |
| Union of Book of Association |                            | O                                  |                                                        |
| Have a list of target        | proteins :: (what is that) | Vyes (neuro-related)               | v no                                                   |
| Please note: uploadin        | g the file might take a    | minute or two                      |                                                        |
|                              |                            |                                    |                                                        |
|                              |                            |                                    |                                                        |

Figure 2: The Upload Page.

The second column displays definitions of proteins which are mapped to each EST. At most ten nonredundant definitions appear, and the number in parenthesis following each definition is the number of times that definition was observed. To facilitate the presentation of this information, we align the descriptions using a variation on a dynamic programming algorithm that considers the sentence structure as well as the actual descriptions when aligning descriptions (Yona & Leung, unpublished). Descriptions are then grouped based on their similarity scores. If "(sim)" follows a definition, then similarity data was used in the corresponding map.

The third column displays GO terms and Swiss-Prot keywords associated with the proteins mapped to each EST. Again, if "(sim)" follows a descriptor, then similarity data was used in the corresponding map.

In both the second and third columns, descriptors are displayed in order of map type. That is, descriptors of proteins mapped by type 1 paths appear first, type 2 paths appear second, and so on. Descriptors corresponding to similarity maps appear after direct maps and are also ordered by type.

The fourth column displays "yes" if the corresponding EST maps to a protein which is involved in an interaction and displays "no" otherwise. Similarly, if the proteins are on the list of target proteins then the corresponding column is marked.

#### 1.3 The EST Map page

The EST map page displays more detailed information on each of the proteins mapped to a particular EST (Figure 4). This page has six columns. The first column displays the mapped protein's NR identifier. The

user may click on the identifier to view Biozon's record of that protein, containing information on the broader biological context of the protein (such as the DNA sequences that encode the protein, the interactions it is involved with, the structures it is linked to and the other entities it is similar to).

The second column displays the protein's primary definition and the third column displays the protein's descriptors. Clicking the "see more" link in either of these columns takes the user to the *descriptor page* where the user finds a comprehensive list of the protein's definitions, GO terms, and Swiss-Prot keywords.

The fourth column indicates whether or not the protein is involved in an interaction. The fifth column displays the type of the corresponding map, and the sixth column contains a link to the *path page* where the user finds the details of the corresponding map between the EST and the protein.

#### 1.4 The Descriptor Page

The *descriptor page* displays all the definitions, GO terms, and Swiss-Prot keywords associated with a particular proteins (Figure 5). For each definition, the descriptor page displays the source database of that definition. The user may click on any of the displayed GO terms to view Biozon's record of the term and the corresponding graph. The descriptor page only displays GO terms actually assigned to the protein (not all ancestors of these GO terms). However, the parent GO terms can be viewed through the Biozon profile page of each GO term.

#### 1.5 The Map Page

The map page displays the details of every map from the chosen EST to the chosen protein (Figure 6). The maps are displayed in order of their type, with type 1 maps appearing first, type 2 maps appearing second, and so on. Maps which use similarity data appear after direct maps and are also ordered by type.

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## ESTs mapped to proteins

See <u>help</u> with output format

| Displa | aying Results | 1-10 S                                            | how 10 💟 results per page                 | <u>« prev 10</u>   <u>next 10 »</u> |
|--------|---------------|---------------------------------------------------|-------------------------------------------|-------------------------------------|
|        | ID            | Definition                                        | Descriptors                               | Interacts                           |
|        |               | Tubulin beta-4 chain (51)                         | chaperone activity                        |                                     |
|        |               | Class II beta-tubulin (313) (s                    | m) structural molecule activity           |                                     |
|        |               | beta 3 tubulin (73) (sim)                         | GTP binding                               |                                     |
|        |               | Tubulin (37) <i>(sim)</i>                         | microtubule                               |                                     |
|        |               | unnamed protein (17) (sim)                        | microtubule-based process                 |                                     |
| 1      | A1834965      | Olfactory enriched transcipt                      | microtubule-based movement                | yes                                 |
|        |               | similar to misato (2) (sim)                       | natural killer cell mediated<br>cytolysis |                                     |
|        |               | DJ20N2.2 (1) (sim)                                | MHC class I protein binding               |                                     |
|        |               | FtsZ (1) (sim)                                    | tubulin                                   |                                     |
|        |               | d.79.2.1 } (1) (sim)                              | Microtubules                              |                                     |
|        |               | View more                                         | <u>View more</u>                          |                                     |
|        |               | Adenylyltransferase thiF (1)                      |                                           |                                     |
|        |               | Ubiquitin-activating enzyme<br>(20)               | E1c<br>catalytic activity                 |                                     |
|        |               | UBA (6)                                           | thiamin biosynthesis                      |                                     |
|        |               | Mus musculus 12 days embry                        | <b>yo</b> transferase activity            |                                     |
|        |               | full-length enriched library,<br>clone:C530001N05 | nucleotidyltransferase activity           |                                     |
|        |               | product:MOP-4 homolog (1)                         | ubiquitin activating enzyme<br>activity   |                                     |
| 2      | AI834966      | A1s9Y protein (7) (sim)                           | protein modification                      | yes                                 |
|        |               | PP3895 (1) (sim)                                  | Transferase                               |                                     |
|        |               | Molybdopterin synthase<br>sulphurylase (2) (sim)  | Nucleotidyltransferase                    |                                     |
|        |               | HesA (2) (sim)                                    | Thiamine biosynthesis                     |                                     |
|        |               | Ydr540cp (1) (sim)                                | Complete proteome                         |                                     |
|        |               |                                                   |                                           |                                     |

Figure 3: The Summary Page.

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## AI834966 is mapped to the following proteins:

See help with output format

Mapping Modes: (<u>what is that?</u>)

011130000015

(uba-1)

see more

| Displ | laying Results 1 - 1 | 0 Show 10 💟 results                                                                                                            | s per page                                           | <u>« pr</u> | <u>ev 10   n</u> | ext 10 :            |
|-------|----------------------|--------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------|-------------|------------------|---------------------|
|       | NR                   | Definition                                                                                                                     | Descriptors                                          | Interacts   | Mode             | Path                |
| 1     | 002510000138         | Adenylyltransferase thiF (EC 2.7.7).<br>see more                                                                               | catalytic activity<br>nucleotidyl<br><u>see more</u> | no          | 3                | <u>View</u><br>path |
| 2     | 004620000351         | Ubiquitin-activating enzyme E1c<br>(Nedd8-activating enzyme E1c)<br>Ubiquitin-activating enzyme 3 homolog).<br><u>see more</u> | catalytic activity<br>protein mod<br><u>see more</u> | no          | 3                | <u>View</u><br>path |
| 3     | <u>010240000010</u>  | Ubiquitin-activating enzyme E1 1.<br>see more                                                                                  | catalytic activity<br>cytoplasm I<br><u>see more</u> | yes         | 3                | <u>View</u><br>path |
| 4     | 010580000011         | Ubiquitin-activating enzyme E1 1.<br><u>see more</u>                                                                           | catalytic activity<br>ligase acti<br><u>see more</u> | no          | 3                | <u>View</u><br>path |
| 5     | <u>010580000028</u>  | Ubiquitin-activating enzyme E1 (A1S9<br>protein).<br><u>see more</u>                                                           | DNA replication<br>catalytic acti<br><u>see more</u> | no          | 3                | <u>View</u><br>path |
| 6     | 010770000014         | Ubiquitin activating enzyme 2.<br><u>see more</u>                                                                              | catalytic activity<br>ligase acti<br><u>see more</u> | no          | 3                | <u>View</u><br>path |

Mus musculus 12 days embryo spinal cord cDNA, RIKEN full-length enriched library, catalytic activity

UBA (human ubiquitin) related; UBiquitin

Activating enzme related (124.1 kD)

Figure 4: The Map Page.

catalytic activity

embryonic d...

see more

View

path

3

yes

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## Definitions associated with 010580000011

| Num. | Source     | Definition                                       |
|------|------------|--------------------------------------------------|
| 1    | Genpept    | Ube1x protein [Mus musculus]                     |
| 2    | Genpept    | ubiquitin activating enzyme E1 [Mus musculus]    |
| 3    | Genpept    | unnamed protein [Mus musculus]                   |
| 4    | PIR        | ubiquitinprotein ligase (EC 6.3.2.19) E1 - mouse |
| 5    | SWISS-PROT | Ubiquitin-activating enzyme E1 1.                |

#### GO terms and keywords associated with 010580000011

| Num. | Source | Definition                           |
|------|--------|--------------------------------------|
| 1    | GO     | catalytic activity                   |
| 2    | GO     | ubiguitin activating enzyme activity |
| 3    | GO     | protein modification                 |
| 4    | GO     | ubiquitin cycle                      |
| 5    | GO     | ligase activity                      |
| 6    | КW     | Ubl conjugation pathway              |
| 7    | КW     | Ligase                               |
| 8    | КW     | Multigene family                     |
| 9    | КW     | Repeat.                              |

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Figure 5: The Descriptor Page.

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|--------------|------------|--------------------------|-----------------|-----------|-----------|
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|              | New Search | Saved Searches           | Instructions    | Tools     | & Files   |

## Paths from EST AI834966 to protein 010580000011

| Мар                                                                                                                                 |
|-------------------------------------------------------------------------------------------------------------------------------------|
| DNA A1834966 in UniGene cluster Mm.34012 related to protein 010580000011                                                            |
| DNA AI834966 in UniGene cluster Mm.34012 related to protein 004620000351 similar to 010580000011 eval 1e-24                         |
| DNA A1834966 in UniGene cluster Mm.34012 related to protein 010240000010 similar to 010580000011 eval 0                             |
| DNA A1834966 in UniGene cluster Mm.34012 related to protein 010580000028 similar to 010580000011 eval 0                             |
| DNA A1834966 in UniGene cluster Mm.34012 related to protein 010770000014 similar to 010580000011 eval 0                             |
| DNA <u>AI834966</u> in UniGene cluster <u>Mm.34012</u> related to protein <u>011130000015</u> similar to <u>010580000011</u> eval 0 |
| DNA A1834966 in UniGene cluster Mm.34012 related to protein 002510000138 similar to 010580000011 eval 4e-12                         |
| DNA AI834966 in UniGene cluster Mm.34012 related to protein 002510000138 similar to 010580000011 eval 8e-10                         |
| DNA A1834966 in UniGene cluster Mm.34012 related to protein 004620000351 similar to 010580000011 eval 1e-5                          |
| DNA AI834966 in UniGene cluster Mm.34012 related to protein 004620000351 similar to 010580000011 eval 4.5e-1                        |

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Figure 6: The Path Page.