Improving subcellular localization prediction using text classification and the gene ontology

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1 INTRODUCTION

There is a flood of biological data from laboratories around the world. To deal with this overload, there are now many computational methods that make it easier for fellow researchers to find relevant research results quickly and easily. We have developed a technique that uses the text of journal abstracts available through Swiss-Prot (Swiss-Prot, 2008) and PubMed (PubMed, 2008) in conjunction with the Gene Ontology (GO) (Gene Ontology Consortium, 2000) hierarchy to significantly improve text classification for biological journal abstracts.

In general, a text classifier maps each text document into one or more predefined labels. For example, text classification of a newspaper article attempts to predict the appropriate newspaper section label, based on the content of that article—whether it belongs in the sports or world news section. Our goal is to improve text classification in a predictor that maps each relevant PubMed abstract into a subcellular location (e.g. nucleus or mitochondrion), so that we can predict the location of a protein based on abstracts written about it, as recorded in a field within the protein’s Swiss-Prot entry.

Classification of biological abstracts is an interesting specialization of general text classification, since these abstracts contain scientific terminology that is often not understandable by non-scientists. They contain specialized terms and acronyms and terms that have many synonyms. For example, the ‘PAM complex’, which exists in the mitochondrion of a biological cell, is also referred to by the phrases ‘prerecognition translocon-associated import motor’ and ‘mitochondrial import motor’. This example also illustrates the fact that biological terms often span word boundaries, which means that their phrasal meaning would be lost if we tokenized the text into individual words.

Text classification has been used in the biological domain before, including its use for subcellular localization prediction. However, no researchers have used the hierarchical structure of the GO to improve text classification in the biological domain.

We want to learn these text classifiers. Many different learning algorithms have been explored for general text classification (Dumais et al., 1998). Support vector machines (SVMs) (Vapnik, 1995) were found to have the highest precision/recall break-even point (BEP, the point where precision equals recall). Joachims (1998) performed a very thorough evaluation of the suitability of SVMs for text, discovering that SVMs are perfect for textual data because textual data produces sparse training instances in very high dimensional space.

Soon after Joachims’ survey, researchers started using SVMs to classify biological journal abstracts. Stapley et al. (2002) used SVMs to improve the prediction of the subcellular localization of yeast proteins. They created a dataset by mining Medline for abstracts that contained a yeast gene name. Their text classifiers achieved -measure in the range 0.31–0.80, where p is precision, r is recall, TP is true positives, FP is false positives, FN is false negatives and PNP is number of prediction on positive proteins.

\[
F\text{-}measure = \frac{2 \times p \times r}{p + r}.
\]

They also built a subcellular predictor based on amino acid content that had an F-measure in the range 0.03–0.61. Using text and amino acid composition together yielded an F-measure in the range 0.33–0.82 with improvements of up to 0.05 over text alone. These results are modest, but prior to Stapley et al. (2002), most localization classification systems were built using text rules or were sequence based. This was one of the first applications of SVMs to biological journal abstracts and it showed that text and amino acid composition together yield better results than either alone.

In other research, text classification was used to augment subcellular localization predictors for animal and plant datasets.
We show that each of these techniques, abstract filtering and term augmentation (SR or TG), individually improves subcellular localization predictors.

Although, classification of biological journal abstracts is a challenging problem, a solution would yield important benefits. With sufficiently accurate text classifiers, the journal abstracts referenced from a protein database entry could be used to automatically annotate that protein. Here, we use these techniques to help a text classifier predict the subcellular localization of a protein. However, these ideas could also be used to help other text classifiers predict other annotations (e.g. general function, pathway participation, relation to disease). Biologists could use these prediction annotations to more efficiently identify proteins of interest. In addition, when biologists submit new research articles to databases like Swiss-Prot, text classifiers could be used to suggest annotations for approval by the submitting biologist. By spending less time sifting through unannotated proteins, researchers will be able to spend more time performing important experiments and uncovering novel knowledge.

(Höglund et al., 2006). These authors' text classifiers were based on the most distinguishing terms of documents, and they described the output of four protein sequence classifiers on their training data. They measured the performance of their classifier using recall (denoted sensitivity) and precision (denoted specificity). Their text-only classifier for the MultiLoc animal dataset had recall in the range [0.51–0.93] and precision in the range [0.32–0.91]. The MultiLocText classifiers, which include sequence-based features, have recall [0.82–0.93] and precision [0.55–0.95]. Their overall average accuracy increased by 16.2 and 9.0% to 86.4 and 94.5%, respectively, on the PLOC animal dataset when text was used to augment sequence-based features. Our technique (Fyshe and Szafron, 2006) is motivated by the improvements that Stapley et al. (2002) and Höglund et al. (2006) saw when they added text classification to other biological features. Our method uses only features extracted from the abstracts, augmented by features from the GO to improve text classification; it uses no information from the amino acid sequence. Since the text classifier described in this article has better performance than the text classifier of Höglund et al. (2008), and does not depend on other features, it can be used as a ‘plug-in’ replacement to augment predictors that also use non-textual features such as amino acid composition.

Our technique uses existing sources of well-organized data: Swiss-Prot, PubMed and the GO hierarchy. This research is a novel contribution to the area of text classification for biological journal abstracts as it identifies (i) a mechanism for determining which abstracts should be used (abstract filtering) and (ii) two techniques for employing the GO as a source of expert knowledge to add features for each protein (term augmentation).

In this article, we use the controlled vocabulary of biological terms represented by the GO to complement the information present in journal abstracts. Specifically, we improve text classification by effectively using:

- The GO as a thesaurus to identify synonyms [synonym resolution (SR)].
- The hierarchical structure of the GO to generalize specific terms into broad concepts [term generalization (TG)].

We show that each of these techniques, abstract filtering and term augmentation (SR or TG), individually improves subcellular localization predictors.

The first step in evaluating our abstract selection mechanism and the GO as a knowledge source for enhancing automatic subcellular localization annotation was to create protein datasets. We selected seven initial datasets (step A in Fig. 1). First, we included the MultiLoc animal and plant datasets (Höglund et al., 2006) to allow us to directly compare our results to the best natural language processing (NLP)-only subcellular location predictor we identified in the literature. Second, we used the latest versions of the five larger Proteome Analyst (PA) datasets (Lu et al., 2004) extracted from Swiss-Prot.

2 METHODS

Figure 1 presents an overview of the workflow for the experiments outlined in this article.

2.1 The datasets

(Höglund et al., 2006) created the MultiLoc dataset by searching for phrases in the Subcellular localization and Feature fields of Swiss-Prot version 42.0. The redundancy of the resulting dataset of 9761 proteins was reduced by removing sequences until no pair of sequences shared >80% similarity. Separate MultiLoc datasets were constructed from the remaining 5959 proteins. The plant dataset contains all proteins (not just plant proteins) that have subcell labels that occur in plants: chloroplast (ch), cytoplasm (cy), endoplasmic reticulum (er), extracellular space (ex), Golgi apparatus (go), mitochondrion (mi), nucleus (ni), peroxisome (pe), plasma membrane (pm) and vacuole (va). The animal classifier removes ch and replaces va with lysosomes (ly). Therefore, there is significant overlap between the proteins in the animal and plant datasets. For example, the protein dataset (step A in Fig. 1) has 5447 proteins for the MultiLoc animal dataset and 5856 proteins for the MultiLoc plant dataset from the total dataset of 5959 proteins.

Lu et al. (2004) created the PA datasets by collecting a group of proteins whose Swiss-Prot entries included a subcellular localization. PA parses the Swiss-Prot subcellular field to extract a short phrase that identifies the specific subcellular localization (e.g. inner membrane). Subcellular localization categories are organism specific. For example, a bacterial cell does not have a nucleus, and an animal cell does not have a periplasmic space. Therefore, PA creates five different datasets, one for each different category of biological cell: animal, green plant, fungi, gram-negative bacteria and gram-positive bacteria. In this article, we present results for the animal
dataset updated using Swiss-Prot version 51.3. The results for other PA datasets are available online (http://www.cs.uaic.ert/~bioinfo/nlp). PA's datasets have a separate 'binary' training file for each label so that a protein can appear with a positive label in more than one file. The PA dataset contains 39,385 labeled proteins, including 33,384 proteins that localize to more than one location. Unlike the MultiLoc datasets, these proteins were divided into five mutually exclusive datasets, based on the protein's organism. For example, the initial dataset shown in Figure 1 has 16,615 proteins for the PA animal dataset (including 2337 multilabeled proteins). We created a separate binary classifier for each label of each of the five organism categories, so there were actually nine binary classifiers used for animal classification, one for each location (the same locations as the MultiLoc animal dataset).

2.2 Retriving and filtering abstracts

We obtained a protein's abstracts (step B in Fig. 1) using the PubMed identifiers recorded in its Swiss-Prot database entry. For the MultiLoc dataset we used Swiss-Prot version 42.0, the same version used by Höglund et al. (2006), so that our results could be directly compared to theirs. For the PA datasets we used Swiss-Prot version 51.3, the newest when the experiments were conducted. We did not include full article text, since it can be difficult to obtain automatically and since previous research has shown that using full text rather than only abstracts does not significantly improve the performance of text classification (Sinclair and Webber, 2004). A protein may have zero or more abstracts and many proteins in Swiss-Prot can refer to the same abstract. All proteins with no abstracts were removed from the datasets (step C in Fig. 1). For example, of the 5447 proteins in the MultiLoc animal dataset, 5137 have at least one abstract, so we eliminated the other 310 from the dataset. Of the 16,615 proteins in the PA animal dataset, 12,261 have at least one abstract so the 4354 with no abstracts were eliminated.

Since different abstracts may have different utilities in predicting subcellular localization, we experimented with abstract filtering mechanisms that remove abstracts that might not contribute to good label prediction. For example, an abstract that describes the sequencing of an entire organism is probably not useful, since it is referenced by proteins with different subcellular localizations. An ambiguous abstract is one whose removal from a training set results in a predictor whose F-measure is not lower than the F-measure of a predictor that uses that abstract. We investigated techniques for ambiguous abstract removal.

The complete set is the set of all abstracts referred to by any protein in the protein dataset. The exclusion set is the set of abstracts that are each referenced by any group of proteins that have no subcellular labels in common. For example, suppose proteins p_1 and p_2 each reference the same abstract, a_12, and p_1 is labeled [cy, ex] and p_2 is labeled [nu]. Then a_12 would be included in the exclusion set, since a_12 cannot help to differentiate between the labels. However, if instead, p_2 was labeled [nu, ex], then a_12 would not be included in the exclusion set, since a_12 could be describing aspects of proteins related to the common label, ex. We compared the approaches of using only the filtered set = complete set — exclusion set to using the complete set. For example, the MultiLoc animal dataset has 211 exclusion set abstracts and 12,549 filtered set abstracts, while the PA animal dataset has 122 exclusion set abstracts and 25,547 filtered set abstracts.

As described later, each predictor uses as features for a given protein, tokens that are extracted from the abstracts that it references. In the filtered set domain, any protein that has only exclusion set abstracts will have no remaining abstracts and therefore no features. For example, the MultiLoc animal dataset has 287 proteins that have only exclusion set abstracts and the PA animal dataset has 244 proteins with only exclusion set abstracts. There are two prediction choices available when a protein has no features. One is to make the prediction based on the prior probabilities — predict the class label with the highest probability. For the MultiLoc animal dataset this class is cy and for the PA animal dataset this class is extracellular. The other choice is to make no prediction, which decreases recall, but does not affect precision. We tried both approaches and discovered that F-measure is higher if we make no prediction. We explored whether using a filtered set instead of a complete set improves accuracy enough on the whole set of proteins to offset the fact that some proteins will have no predictions.

2.3 Processing abstracts

To use a classifier, the relevant abstracts for each protein must be assigned a feature vector that can be used by the classifier (step D in Fig. 1). We first create a set of tokens that represent individual words or phrases in an abstract. We use white space or hyphens to determine token boundaries, and then strip all leading and trailing punctuation marks from the tokens. Finally, tokens are stemmed using Porter’s stemming algorithm (Porter, 1980) to strip suffixes. In our baseline (BASE) classifiers, the feature vector has one component (feature) for each unique token that appears in an abstract referenced by any protein in the filtered set. For example, there are 46,772 unique tokens for the PA animal classifier. We later describe two other classifiers that use additional tokens: SR and TG. Our feature vector has one component for each unique token found in any abstract for any protein in the training dataset.

The simplest approach is to use a binary feature component for a protein with value 1 if the token appears in one of the abstracts referenced by that protein or 0 if it does not appear. However, it seems more appropriate to use a count of the number of times that the token appears in the selected abstracts as the feature component. In fact, there are more sophisticated NLP techniques called importance measures that assign a value to each token that can be used as components in our feature vector.

For this study, we evaluated two importance measures to produce feature vector components, term frequency—inverse document frequency (tfidf) and redundancy, given in Equations (2) and (3), respectively. Note that f(t_i) is the number of times token k appears in an abstract, f(t_i, d_i) is the number of times token k appears in abstract i, N is the total number of abstracts, d(t_k) is the number of abstracts that contain token k and r(t_k) defined in Equation (4), is called empirical entropy.

\[
\text{tfidf}(t_i) = f(t_i) \times \log \frac{N}{d(t_k)}
\]

(2)

\[
\text{redundancy}(t_i) = f(t_i) \times r(t_k)
\]

(3)

\[
r(t_k) = \log N + \sum_{i=1}^{N} \frac{f(t_i, d_i)}{f(t_i)} \log \frac{f(t_i, d_i)}{f(t_i)}
\]

(4)

2.4 Synonym resolution

The GO hierarchy can be used as a thesaurus for biological words and phrases. For example GO encodes the fact that ‘metal binding’ is a synonym for ‘metal ion binding’ (Fig. 2). SR uses the GO’s ‘exact synonym’ field to find synonyms and incorporates synonym information into the feature vector obtained using the BASE processing technique. Specifically, we searched stemmed versions of the abstracts for matches to stemmed GO node names or synonyms. If a match was found, the GO node name (deemed the canonical representative for its set of synonyms) was added as a token for the protein. However, this node name was prefixed with the string ‘go_’, so that the SVM classifier could differentiate between the case where a GO node name appears exactly in text and the case where a GO node name is added by SR.

For example, in Fig. 3, the phrase ‘metal binding’ appears in the text. The GO hierarchy indicates that this is a synonym of the GO node ‘metal ion binding’. Therefore the term ‘metal ion binding’ is added as a ‘go’ token. This approach combines the weight of several synonyms into one representative, allowing the classifier to more accurately model the author’s intent, and to identify multiword phrases that would be otherwise lost during tokenization. Comparing column 5 to column 3 or column 6 to column 4 in Table 1 shows an increase of 3.5% in the number of feature components and the average number of feature vector components per positive training instance for the classifiers constructed from the PA animal dataset. We will show that

\[\text{We assume this was done in the Höglund et al. (2006) study but were unable to verify it.}\]
We studied the effect of p123 on metal binding, specifically on transition metal ions.

Fig. 2. A subgraph of the GO hierarchy. GO nodes are shown as ovals, and synonyms are shown as rectangles.

Fig. 3. A sentence that illustrates abstract processing (step D in Fig. 1). Text is white-space tokenized and stemmed. Stop words are removed to create token set used in the BASE classifier. SR adds a canonical representative for each synonym group. TG adds GO hierarchy node names that are parents of nodes identified using SR. The term ‘metal ion binding’ is the canonical synonym of ‘metal binding’ so SR adds the GO term ‘metal ion binding’ as a token. TG also adds the names of its parent nodes as tokens.

in some cases, this increase in information translates to improved classifier accuracy.

2.5 Term generalization

In order to express the relationships between terms, the GO hierarchy is organized in a directed acyclic graph. For example, ‘cation binding’ is a type of ‘metal binding’, which is a specific type of ‘binding’. This ‘is a’ relationship is expressed as a series of parent–child relationships (Fig. 2).

In TG, we first use the GO for SR, but we also use the GO’s hierarchical structure to generalize specific terms into broader concepts. For TG, if a GO node name (or synonym) is found in an abstract, then in addition to adding a feature for the canonical synonym, we add a feature for the name of each ancestor (TG in Fig. 3). Just as in SR, the additional features are prefixed with the string ‘go_’. TG further increases the number of feature components and the average number of feature vector components per training instance by another 6.9%, as shown in Table 1.

Table 1. PA filtered dataset for the Animal classifiers, where Train Ins is the number of positive training instances, Feature is the number of feature vector components, F/Ins is the average number of feature vector components per positive training instance.

<table>
<thead>
<tr>
<th>Class</th>
<th>Train Ins</th>
<th>BASE Feature</th>
<th>F/Ins</th>
<th>SR Feature</th>
<th>F/Ins</th>
<th>TG Feature</th>
<th>F/Ins</th>
</tr>
</thead>
<tbody>
<tr>
<td>cy</td>
<td>2350</td>
<td>21 1576</td>
<td>9.18</td>
<td>23 991</td>
<td>10.21</td>
<td>26 559</td>
<td>11.30</td>
</tr>
<tr>
<td>er</td>
<td>108</td>
<td>34 754</td>
<td>32.18</td>
<td>37 980</td>
<td>35.17</td>
<td>43 500</td>
<td>40.28</td>
</tr>
<tr>
<td>ex</td>
<td>5289</td>
<td>25 355</td>
<td>4.79</td>
<td>27 175</td>
<td>5.14</td>
<td>29 114</td>
<td>5.50</td>
</tr>
<tr>
<td>go</td>
<td>28</td>
<td>16 866</td>
<td>60.21</td>
<td>19 971</td>
<td>70.39</td>
<td>23 757</td>
<td>84.82</td>
</tr>
<tr>
<td>ly</td>
<td>141</td>
<td>58 922</td>
<td>41.79</td>
<td>63 955</td>
<td>45.35</td>
<td>70 599</td>
<td>50.06</td>
</tr>
<tr>
<td>mi</td>
<td>889</td>
<td>10 508</td>
<td>11.82</td>
<td>11 559</td>
<td>13.00</td>
<td>12 918</td>
<td>14.53</td>
</tr>
<tr>
<td>nu</td>
<td>3618</td>
<td>23 192</td>
<td>6.41</td>
<td>25 513</td>
<td>7.05</td>
<td>27 708</td>
<td>7.66</td>
</tr>
<tr>
<td>pe</td>
<td>108</td>
<td>36 036</td>
<td>33.36</td>
<td>39 492</td>
<td>36.50</td>
<td>44 666</td>
<td>41.35</td>
</tr>
<tr>
<td>pm</td>
<td>206</td>
<td>57 487</td>
<td>27.90</td>
<td>63 735</td>
<td>30.95</td>
<td>72 377</td>
<td>35.13</td>
</tr>
<tr>
<td>All</td>
<td>12 261</td>
<td>45 197</td>
<td>3.69</td>
<td>48 904</td>
<td>3.99</td>
<td>52 157</td>
<td>4.25</td>
</tr>
</tbody>
</table>

The compartments are cy, er, ex, go, ly, mi, nu, pe and pm. The 12 261 proteins in the dataset include 468 multi-labeled proteins.

TG gives the SVM algorithm an opportunity to learn correlations that exist between general terms and subcellular localization even if the general term itself never appears in an abstract as only the names of its more specific children occur. Without TG, the SVM has neither concept of the relationship between child and parent terms, nor between sibling terms. For some localization categories more general terms may be the most informative and in other cases specific terms may be best. Because our technique adds features to training instances and never removes any, the SVM can assign lower weights to the generalized terms in cases where the localization category is not well characterized by more general terms.

3 RESULTS AND DISCUSSION

Our BASE, SR and TG classifiers were trained and evaluated using cross validation on seven separate datasets: MultiLoc animal, MultiLoc plant, PA animal, PA plant, PA fungi, PA gram negative bacteria and PA gram-positive bacteria. For brevity, this article contains F-measure scores for the MultiLoc animal and PA animal datasets using tfidf. Results using the other datasets (including precision and recall scores) and using redundancy as well as tfidf are available online (http://www.cs.ualberta.ca/~bioinfo/nilp/). For the MultiLoc datasets, we used 5-fold cross validation so the results could be compared directly to Höglund et al. (2006). For the PA datasets we used 10-fold cross validation.

Table 2 shows the F-measure scores for the MultiLoc animal dataset and Swiss-Prot version 42.0, including the text-only classifier (denoted TEXT) reported by Höglund et al. (2006). The best score is bolded in each row. The BASE-complete predictor is comparable to the TEXT classifier. The F-measures differ in the range -15.1% (mi) to +20.6% (ly) with an overall difference of +0.05%. This overall score is not the simple average of the individual scores. Instead, it is the mean of the five aggregate F-measures, each taken over the single confusion matrix constructed for all predictions. For the TEXT predictor, we did not have fold data so the overall score is the aggregate F-measure over the single confusion matrix for all predictions.
We used a one-sample t-test to compare the mean F-measure of the five overall BASE-complete scores (df = 4) to the overall TEXT score. The t-score was 0.3465 (p = 0.3732), indicating that we only have 62.68% confidence that the F-measure of the BASE-complete classifier is higher than the F-measure of the TEXT classifier, so we have established that it is comparable—not better. This indicates that our BASE predictor is not a ‘straw man’, since it is as good as the best-known predictors appearing in the literature that are based on abstracts alone. The key differences between BASE-complete and TEXT are:

1. After white space tokenization and stemming, TEXT uses only a subset of machine-learned distinguishing terms (~800 terms for 10,000 abstracts). BASE uses all terms (stemmed).
2. TEXT uses a probabilistic term weighting scheme, where the weight of a distinguishing term is assigned the ratio of its occurrence count in the abstract to the sum of the occurrence counts of all distinguished terms in abstracts referenced by the protein. We use standard NLP algorithms: tfidf and redundancy.
3. TEXT uses a single multilabel classifier that predicts a single label. We use a set of binary classifiers, one for each label in the ontology of the category (animal or plant). This allows us to make correct predictions for proteins that localize to more than one location.

Our goal is to evaluate the use of abstract filtered sets as an ambiguous abstract removal technique and the use of GO terms to add effective features. First, we show that ambiguous abstract removal using a filtered set almost always produces a predictor with higher F-measure than a complete set predictor that removes no abstracts. However, we discuss a specific situation where this assertion fails, meaning that it is better to use the complete set of abstracts rather than the filtered set. Table 2 shows that a filtered set improves the F-measure score of all BASE, SR, and TG predictors except for the nu class. The overall improvements are 6.9, 6.4 and 5.3%, respectively. We used a one sample t-test to test whether the overall F-measure for the filtered set are at least 5% higher than the overall F-measure for the complete set across the 5-folds and three types of classifiers (BASE, SR and TG—fifteen data points in total) by comparing ‘filtered set score minus complete set score minus 0.05’ to 0. This test establishes, with 99.8% confidence (p = 0.001989, df = 14, t = 3.44) that the filtered set F-measure is at least 5% better than the complete set F-measure. A test for predicting when to use a complete set for a particular class instead of using the filtered set is given later in this article.

Table 2 also shows that using the GO for SR or SR plus TG improves predictions over BASE when the abstract set is fixed in almost all cases. Out of 18 cases (nine complete and nine filtered), only two resulted in lower F-measures: SR-complete for er (0.570–0.554) and TG-filtered for mi (0.746–0.727). In all other cases, using the GO improves performance. The overall F-measure increases by 1.6% (0.731–0.747) for SR-complete, 2.6% (0.731–0.757) for TG-complete, 1.1% (0.800–0.811) for SR-filtered and 1.0% (0.800–0.810) for TG-filtered. We used a one sample t-test to test whether the overall F-measure for the SR and TG predictors are at least 1% higher than the overall F-measure for the BASE predictor across the 5-folds and two dataset abstractions (complete set and filtered set—10 data points each for SR and TG) by comparing ‘SR or TG score—BASE score—0.01’ to 0. For SR, we measured 92% confidence (p = 0.08207, df = 9, t = 1.5148) that the SR F-measure is at least 1% better than the BASE F-measure. For TG, we measured 95% confidence (p = 0.04599, df = 9, t = 1.8855) that the TG F-measure is at least 1% better than the BASE F-measure. The difference between overall F-measure scores for the SR and TG predictors was not statistically significant.

Combining the benefits of filtered sets and the GO results in an overall 8.0% improvement in F-measure, when SR is used and 7.9% when TG is used. A two-sample t-test shows that SR-filtered has 4% higher F-measure than BASE-complete with confidence 96% (p = 0.03913, df = 7.98, t = 2.019), TG-filtered has 4% higher F-measure than BASE-complete with confidence 96% (p = 0.04091, df = 7.98, t = 1.9903). These results indicate that, in general, if a predictor uses NLP techniques on biological abstracts then filtered sets, and using the GO are effective in improving F-measure.

The results of similar experiments with the PA dataset using Swiss-Prot version 51.3 and the tfidf importance measure are shown in Table 3 (without an entry for TEXT). However, there are now 10-folds so the number of data points is doubled. Using the filtered set of abstracts rather than the complete set results in a 4% higher overall F-measure score (4.7% for BASE, 5.0% for SR and 4.8% for TG) with confidence 99% (p = 0.0026, df = 29, t = 3.0274). Of the 27 comparisons, only one has a complete set F-measure higher than filtered set F-measures (SR for ly) it is not statistically significant.

For SR, we measured 99.98% confidence (p = 0.000159, df = 19, t = 4.383) that the SR F-measure is better than the BASE F-measure (0.4% for a complete set predictor and 0.7% for a filtered set predictor). For TG, we measured 99% confidence (p = 0.0009, df = 19, t = 3.6344) that the TG F-measure is better than the BASE F-measure (0.6% for a complete set predictor and 0.7% for a filtered set predictor). Combining the benefits of filtered sets and the GO results in an overall 5.4% improvement in F-measure, when either TG or SR is used. A two sample t-test shows that SR-filtered has 3% higher F-measure than BASE-complete with confidence 99% (p = 0.01498, df = 17.79, t = 2.3592), TG-filtered has 3% higher F-measure than BASE-complete with confidence 99% (p = 0.01370,
The complete (Comp) and filtered (Filtered) abstract sets for the BASE, SR, and TG predictors. The Improve column (IMPR) indicates the amount by which TG filtered improved over BASE complete. The table shows that cy, er, go, ly, mi, nu, pe and pm. The bolded value in each line is the highest F-measure for that cellular compartment.

Table 4. Numbers of proteins with only exclusion set abstracts, number of abstracts in the exclusion set and the ratio of these two numbers in the MultiLoc animal dataset and the PA animal dataset

<table>
<thead>
<tr>
<th>Class</th>
<th>MultiLoc animal</th>
<th>PA animal</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Proteins</td>
<td>Abstracts</td>
</tr>
<tr>
<td>cy</td>
<td>33</td>
<td>156</td>
</tr>
<tr>
<td>er</td>
<td>22</td>
<td>50</td>
</tr>
<tr>
<td>ex</td>
<td>0</td>
<td>46</td>
</tr>
<tr>
<td>go</td>
<td>9</td>
<td>33</td>
</tr>
<tr>
<td>ly</td>
<td>0</td>
<td>17</td>
</tr>
<tr>
<td>mi</td>
<td>56</td>
<td>94</td>
</tr>
<tr>
<td>nu</td>
<td>116</td>
<td>76</td>
</tr>
<tr>
<td>pe</td>
<td>10</td>
<td>29</td>
</tr>
<tr>
<td>pm</td>
<td>41</td>
<td>102</td>
</tr>
<tr>
<td>Overall</td>
<td>287</td>
<td>211</td>
</tr>
</tbody>
</table>

The compartments are cy, er, go, ly, mi, nu, pe and pm. df = 17.79, \( t = 2.405 \). These results confirm that filtered sets, and using GO provide a significant improvement when using NLP techniques based on biological abstracts.

To discover the reason why the complete set of abstracts was better than the filtered set for the nu class in the MultiLoc animal dataset, we examined many statistics based on the number of abstracts in the exclusion set for each class and the number of proteins with labels in each class that yield no features when the abstract filtered set is used instead of the abstract complete set. Table 4 shows that nu (MultiLoc animal) is the only class in which the ratio of proteins removed to abstracts removed is larger than the overall ratio for the filtered set. Therefore, we conjecture that a class ratio that is higher than the overall ratio should be used as the criteria for determining whether to make an exception to using the filtered set instead of the complete set. We then computed the ratios for the PA animal dataset that are shown in Table 4. The ratios suggest that no PA animal complete set classifier should have a (statistically significant) higher F-measure than the PA animal filtered set classifier for the same class. Table 3 provides evidence for our conjecture.

The NLPl-based classifiers described in this article can be used to augment other non-NLP predictors. We are currently using our new NLP techniques to improve the coverage and accuracy of the PA family of predictors (Lu et al., 2005; Szafron et al., 2004). There are several ways that classifiers can be combined into an ensemble predictor and it is not clear yet which technique will provide the best overall win in terms of increased coverage and accuracy.

In summary, using the GO for SR and TG is a useful mechanism for improving text classification using biological abstracts. In addition, we recommend ambiguous abstract removal using a filtered set for those predictors whose ratio of empty feature vectors per abstract removed is lower than the overall ratio for all of the predictors.

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REFERENCES


