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Exploiting Multiply Annotated Corpora in Biomedical Information Extraction Tasks

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Abstract

This paper discusses the problem of utilising multiply annotated data in training biomedical information extraction systems. Two corpora, annotated with entities and relations, and containing a number of multiply annotated documents, are used to train named entity recognition and relation extraction systems. Several methods of automatically combining the multiple annotations to produce a single annotation are compared, but none produces better results than simply picking one of the annotated versions at random. It is also shown that adding extra singly annotated documents produces faster performance gains than adding extra multiply annotated documents.

1 Introduction

When annotating a corpus for an information extraction (IE) task, it is normal to annotate at least part of it multiple times in order to measure inter-annotator agreement (IAA) and thereby monitor the quality of the annotation and quantify the difficulty of the task. These multiply annotated documents can then be reconciled to produce a gold standard which is used to train and test a machine-learning based IE system. In our experience in the biomedical domain, however, this reconciliation process is very time-consuming, and therefore costly, with the reconciliation of a pair of annotated documents taking much longer than annotating the document for a third time. But if reconciliation is too expensive, and the corpus is to be used as training data, can one make better use of the multiply annotated data? This paper will set out to answer two questions with empirical evidence presented from biomedical IE tasks. The first question is how best to use the multiply annotated data in training machine learning systems; whether to try to automatically reconcile the documents using some type of combination algorithm, or whether to provide all copies to the machine learning algorithm and leave it to “reconcile” the documents. The second question concerns how best to spend annotation budget, on multiple annotation of documents already in the corpus, or on annotating new documents. To the best of our knowledge, these questions have not previously been addressed in the literature.

This paper is organised as follows: after reviewing related work in the following section, the corpora and experimental system are described in Section 3. An account of the experiments performed is provided in Section 4, with the results presented in Section 5 and discussed in Section 6.

2 Background

The experiments described in this papers make use of a text mining pipeline which was developed as part of the TXM project and integrated into a biomedical curation tool to enable assisted curation for biomedical literature (Alex et al., 2008). In order to train the machine learning components of the system, and to test the system, a collection of full-text biomedical articles were annotated with items of biomedical interest. The decision to annotate full-text articles instead of abstracts as training data for a biomedical text mining system is supported by previous work of Shah et al. (2003) and McIntosh and Curran (2007) who argue that paper abstracts do not always contain sufficient information.

In the pilot data annotation phase of TXM, multiply annotated papers were reconciled to create one gold standard annotation for each paper. However, this process was found to be very time-consuming, particularly as dealing with full-text articles. In order to obtain good machine learning results, a further decision was made to invest valuable annotation time to annotate more papers rather than reconcile multiple annotations. Consequently, the two corpora that were finally annotated as training material for the TXM pipeline were not reconciled and therefore contain multiple annotations for a proportion of papers (see Section 3). These multiple annotations were used to calculate IAA in order to monitor annotation consistency over time.

The collection of IAA during annotation has also been advocated by Lu et al. (2006) and Inderjeet et al. (2005). Furthermore, annotation consistency is an aspect that was highlighted as important in experiments carried out by Alex et al. (2006). Measures of IAA are also used to get a better understanding of how difficult it is to extract such information automatically.

3 Corpora and System

The two corpora used in the experiments in this paper address protein-protein interactions (the PPI corpus) and tissue expression (the TE corpus), and consist of full-text biomedical papers from PubMed and PubMedCentral. The PPI corpus contains approximately 75,000 sentences and the TE corpus around 63,000.

A total of nine qualified biologists were employed to annotate each corpus with named entities and relations (as shown in Tables 1 and 2), to map selected entity types to
identifiers in appropriate standard databases, and to enrich the relations with certain properties and attributes (Haddow and Matthews, 2007). Only the entities and relations will be considered in this paper. Each corpus was split into three sections (TRAIN, DEVTEST and TEST) with the intention that TRAIN and DEVTEST would be used to develop the NLP components, and TEST would only be used for testing the final system.

| Type         | Count |
|--------------|-------|
| CellLine     | 7,676 |
| Complex      | 7,668 |
| DrugCompound | 11,886|
| ExperimentalMethod | 15,311|
| Fragment     | 13,412|
| Fusion       | 4,344 |
| Modification | 6,706 |
| Mutant       | 4,829 |
| Protein      | 88,607|

Table 1: The entity types and occurrence counts for (a) PPI and (b) TE. Note that GOMOP stands for “Gene or mRNA:DNA or Protein” and was used when the annotators felt the author employed the term in an ambiguous way.

During the annotation of each corpus, a selection of papers were doubly or triply annotated, as shown in Table 3, and these multiply annotated papers were used to compute IAA. For each pair of corresponding annotations, the IAA is calculated by taking one of the annotators as the gold standard and scoring the other annotator against them by calculating precision, recall and $F_1$ in the usual way. The $F_1$ is not affected by the ordering of the annotators in this calculation (Brants, 2000). Entities are compared using exact match as in Tjong Kim Sang and De Meulder (2003), and relations are considered equal if they have the same type and arguments. When scoring relations, only those where both annotators agree on the entities were considered, in order for relation IAA not to be affected by entity IAA. A combined IAA is also included, where pairs of annotated documents were scored on all relations, not just those where the annotators agree on the entities. This combined score indicates the level of agreement on the whole task of relation markup.

To produce an average IAA for a particular relation or entity type, the $F_1$ scores on each pair of corresponding annotated papers were micro-averaged, i.e. each relation or entity instance was given equal weight in computing the overall scores shown in Table 4.

| Corpus    | Annotated item | IAA |
|-----------|----------------|-----|
| PPI       | entities       | 84.9|
| PPI       | relations      | 76.1|
| PPI       | combined       | 59.7|
| TE        | entities       | 83.8|
| TE        | relations      | 74.1|
| PPI       | combined       | 55.7|

Table 4: Inter-annotator agreement for entities, relations, and the combined entity/relation annotation task.

Both named entity recognition (NER) and relation extraction (RE) were implemented using maximum entropy based machine learning approaches as part of an information extraction pipeline (Alex et al., 2008). The pipeline contains a pre-processing component which performs tokenisation and sentence splitting, linguistic analysis (such as part-of-speech tagging and chunking) and adds some biological markup (indicating species words and abbreviation definitions). The NER component uses the Curran and Clark (2003) named entity tagger augmented with extra orthographic and gazetteer features tailored to the domain, and more fully described in Alex et al. (2007). The RE component uses maximum entropy to classify candidate relations, based on features derived from the context of the entities in the candidate, the text in between, the part-of-speech and chunk tags, the entities themselves, and features derived from indicator words. The latter are either interaction words or expression level words collected from the training data which were marked up by the annotators in addition to the entity and relation annotations discussed earlier. An earlier version of the RE component was described by Nielsen (2006), but has since been extended to address further relation types, and some inter-sentential relations.

### 4 Experiments

To assess how different methods of combining multiply annotated documents affect performance, models were trained and tested using each of the combination methods listed below. Tests were conducted for both NER and RE, on the two corpora, giving four different testing configurations. Performance measurements are for the DEVTEST and TEST sections of each corpus, which include multiply annotated documents. It was thought fairest to leave the multiply annotated documents in the evaluation set, in order to ensure that it was as large as possible, and so that the system would obtain partial credit for predicting an entity or relation chosen by at least one annotator, even though this means that the system cannot obtain 100% accuracy. The following methods of combining multiply annotated training data were used:
| Corpus | Type | Count | Description |
|--------|------|-------|-------------|
| PPI    | PPI  | 11,523| Indicates that the text is referring to an interaction between Proteins, Fragments, Mutants, Complexes or Fusions. |
| PPI    | FRAG | 16,002| Connects Fragment or Mutant to its parent Protein. |
| TE     | TE   | 12,426| Links a gene or gene product to a Tissue, indicating that the text is stating that the gene or gene product is expressed in that Tissue. |
| TE     | FRAG | 4,735 | Plays the same role as in PPI. |

Table 2: The relation types in the corpora

| Annotations | PPI | TE |
|-------------|-----|----|
|             | TRAIN | DEVTEST | TEST | TRAIN | DEVTEST | TEST |
| 1           | 65   | 25    | 35   | 82    | 34     | 34   |
| 2           | 48   | 9     | 8    | 68    | 7      | 11   |
| 3           | 20   | 5     | 2    | 1     | 0      | 1    |
| Total documents | 133  | 39    | 45   | 151   | 41     | 46   |
| Total annotations | 221  | 58    | 57   | 221   | 48     | 59   |

Table 3: Counts of numbers of papers with 1, 2 or 3 annotations in each section of each corpus.

**all** Multiply annotated versions of the same documents are not combined at all, but all versions are included in the training set. The machine learner is, in effect, used to reconcile the different versions on its own.

**union** Each set of annotations on the same document is combined by including all entities and relations marked by each annotator. For entity annotation there is a problem combining annotations in this way if entities cross (in other words, overlap but do not nest in the sense of Alex et al. (2007)), since crossing entities cannot be represented in the NER system. Crossing entities are resolved by removing the one with the later starting point.

**intersection** Annotations are combined by only choosing the entities and relations which are common to all annotated versions of a multiply annotated document.

**one-random** For each multiply-annotated document, one annotated version is chosen at random and the rest are discarded.

**best-ner** An NER model is trained using the all configuration. For each multiply annotated document in the training set, this NER model is then applied to all annotated versions in TRAIN and the version on which the model achieves the highest score is chosen. The rationale behind this method is that the system will do better on annotated versions which are more consistent with the rest of the corpus, so the combined corpus will be more consistent, and therefore provide better training data.

**best-re** The same as best-ner, except the annotated version is chosen by considering performance on relation extraction.

**consistent** An arbitrary order of preference is applied to the annotators, and for each multiply annotated document, the annotation corresponding to the annotator which came highest in the order of preference was selected. This combination method may create a more consistent corpus by favouring certain annotators.

Notice that intersection and union create a new version of the annotated document by combining annotations from each version, whereas one-random, best-ner, best-re and consistent simply choose one of the annotated versions, based on some criteria. We considered using a voted strategy, but since most of the multiply annotated documents only have two annotations, voted would just collapse to intersection or union, depending on how ties were resolved. Note that the variation in the training corpora produced by each combination strategy will be higher for relations than for entities, since the IAA (Table 4) is much lower for relations. The combined IAA score is relevant here, as it shows the overall difference between different annotators’ views of the relations in the same document.

To compare the effect of adding extra multiply annotated data versus extra single annotated data, learning curves for each of these two configurations were produced. Firstly, all but one version of each multiply annotated document was removed from the original training set (DEVTEST and TRAIN combined), and a matching number of singly annotated documents was also removed. A model was then trained on the remaining training documents, and tested on TEST. Learning curves for singly and multiply annotated documents, respectively, were produced by re-adding the removed documents in 10 separate batches, training and testing after adding each batch. The whole process was repeated 20 times, averaging the results, with the documents placed in a different random order each time.
5 Results

The results for the different combination types on the NER task are shown in Table 5. Each combination type was tested in two different configurations; training on TRAIN and testing on DEVTEST, and training on TRAIN and DEVTEST combined then testing on TEST. The $F_1$ scores for each combination were micro-averaged across entity classes to give an overall $F_1$ score.

| Method        | PPI | TE    | PPI | TE    |
|---------------|-----|-------|-----|-------|
|               | DEVTEST | TEST | DEVTEST | TEST |
| all           | 75.1 | 72.5  | 65.4 | 63.3  |
| union         | 74.8 | 72.1  | 65.1 | 63.7  |
| intersection  | 74.8 | 72.0  | 63.9 | 62.8  |
| one-random    | 75.0 | 71.9  | 65.2 | 63.7  |
| best-ner      | 74.7 | 72.1  | 64.9 | 63.5  |
| best-re       | 75.1 | 72.1  | 65.1 | 63.6  |
| consistent    | 74.7 | 72.4  | 65.0 | 63.6  |

Table 5: Comparison of performance of NER, training models on different versions of the training set produced by different combination methods. Performance is given on DEVTEST and TEST, and measured using $F_1$.

The differences in scores between each combination type appear small, and do not follow a consistent pattern. A Friedman rank sum test was performed separately on the PPI and TE TEST results, comparing the $F_1$ scores on each file, in each combination type. For both domains the test gave $p = 0.38$, indicating that there is no evidence of a significant difference between the methods. The corresponding results for RE are show in Table 6, again comparing the performance of each combination method on DEVTEST and TEST, for both domains.

| Method        | PPI | TE    | PPI | TE    |
|---------------|-----|-------|-----|-------|
|               | DEVTEST | TEST | DEVTEST | TEST |
| all           | 58.9 | 58.7  | 58.3 | 53.3  |
| union         | 56.4 | 58.0  | 56.4 | 53.1  |
| intersection  | 56.9 | 58.3  | 54.6 | 53.1  |
| one-random    | 57.5 | 58.6  | 57.2 | 53.6  |
| best-ner      | 57.9 | 58.5  | 57.0 | 53.9  |
| best-re       | 57.7 | 58.6  | 57.0 | 52.8  |
| consistent    | 57.8 | 58.6  | 57.0 | 53.4  |

Table 6: Comparison of performance of RE, training models on different versions of the training set produced by different combination methods. Performance is given on DEVTEST and TEST, and measured using $F_1$.

Testing with the Friedman rank sum test as for NER gives $p = 0.72$ for PPI and $p = 0.23$ for TE, again indicating that the effect of the different combination methods is not significant.

The effect of adding extra multiply annotated data to the training set, compared with adding extra singly annotated data, is shown in Figures 1 and 2. In each graph, the curve labelled “single” shows the performance of the system as more singly annotated data is added, and the “multiple” curve shows how performance changes as more multiply annotated data is added. The systems are trained on subsets of TRAIN and DEVTEST combined, and tested on TEST.

6 Discussion and Conclusion

The comparisons between methods shown in Tables 5 and 6 indicate that there is very little difference between the different combination methods. For RE, there is a slight preference for the all combination method, but this preference is not significant, and could have arisen on the DEVTEST set because the system was optimised with the all combination method on that test set. The union and intersection combination strategies cause a change in the precision/recall balance (data not shown), since the former tends to increase the number of relations/entities in the annotated corpus, and the latter tends to reduce them.

It is perhaps surprising that using all the annotated versions of each document does not produce better performance than simply picking one of the versions at random. This is especially true for RE, where the IAA of below 60 $F_1$ means that there is a significant amount of extra information in the other annotated versions of a document. It is possible that because of the large training corpus, adding this extra data did not really offer anything extra, or that the inconsistency in the different versions of the annotated data just confused the machine learner. Maybe with a smaller training corpus the differences between different combination methods would be more marked.

The learning curves in Figure 2 suggest that adding extra multiple annotation can offer small improvements in NER...
and RE performance, but annotations of new documents will offer a faster rate of improvement. Whilst multiply annotated versions of documents may be required to measure IAA and to monitor annotation quality, it is better to keep the multiple annotation to the minimum required for these purposes, in order to maximise the budget available for extra singly annotated documents.

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