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A distantly supervised dataset for automated data extraction from diagnostic studies

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Abstract
Systematic reviews are important in evidence based medicine, but are expensive to produce. Automating or semi-automating the data extraction of index test, target condition, and reference standard from articles has the potential to decrease the cost of conducting systematic reviews of diagnostic test accuracy, but relevant training data is not available. We create a distantly supervised dataset of approximately 90,000 sentences, and let two experts manually annotate a small subset of around 1,000 sentences for evaluation. We evaluate the performance of BioBERT and logistic regression for ranking the sentences, and compare the performance for distant and direct supervision. Our results suggest that distant supervision can work as well as, or better than direct supervision on this problem, and that distantly trained models can perform as well as, or better than human annotators.

1 Background
Evidence based medicine is founded on systematic reviews, which synthesize all published evidence addressing a given research question. By examining multiple studies, a systematic review can examine the variation between different studies, the discrepancies between them, as well as look at the quality of evidence across studies in a way that is difficult in a single trial. Since a systematic review needs to consider the entire body of published literature, producing a systematic review is expensive and labor-intensive process, often requiring months of manual work (O’Mara-Eves et al., 2015).

To ensure that the results of a systematic review are as comprehensive and unbiased as possible, their production follows a strict and systematic procedure. To catch and resolve disagreements, all steps of the process are performed in duplicate by at least two reviewers. There have recently been examples of systematic reviews using automation in a limited capacity (Bannach-Brown et al., 2019; Przybyla et al., 2018; Lerner et al., 2019), but the impact of automation on the reliability of systematic reviews is not yet fully understood. Automation is not part of accepted practice in current guidelines (De Vet et al., 2008).

After a set of potentially included studies have been identified, systematic reviewers complete a so-called data extraction form for each study. These forms comprise a semi-structured summary of the studies, identifying and extracting a consistent, pre-specified set of data items from abstracts or full-text articles in a coherent format (see the left part of Table 1 for sample excerpts). The coherent format allows the data from the studies to be synthesized qualitatively or quantitatively to address the research question of the review.

In this study we will focus on systematic reviews of diagnostic test accuracy (DTA), which examine the accuracy of tests and procedures for diagnosing medical conditions, and which have seen little attention in previous literature on automated data extraction. To compare and synthesize results across studies, reviewers extract diagnostic accuracy from each study, but also determine the index test (the specific diagnostic test or procedure that is being tested), what target condition the test seeks to diagnose, and the reference standard (the diagnostic test or procedure that is being used as the gold standard) (see Fig 1 for an example). These data must be determined for each study to know if the diagnostic accuracy in different studies can be compared.
Table 1: Examples of raw data from three data extractions forms in unstructured format (left) and a structured summary of the data intended for distant supervision by pattern matching (right).

![Table image](image_url)

Although typhoid fever is confirmed by culture of Salmonella enterica serotype Typhi, rapid and simple diagnostic serologic tests would be useful in developing countries. We examined the performance of Widal test in a community field site and compared it with Typhidot and Tubex tests for diagnosis of typhoid fever. [...] Sensitivity, specificity, positive predictive value (PPV), and negative predictive value (NPV) of the 3 serologic tests were calculated using culture-confirmed typhoid fever cases as "true positives" and paratyphoid fever and malaria cases as "true negatives". [...] The sensitivity, specificity, PPV, and NPV of Typhidot and Tubex were not better than Widal test.

There is a need for more efficient rapid diagnostic test for typhoid fever especially during the acute stage of the disease. Until then, culture remains the method of choice.

![Legend image](image_url)

1.1 BERT

BERT (Bidirectional Encoder Representations from Transformers) is a deep learning model that is unsupervisedly pretrained on a large general language corpus, then supervisedly fine-tuned on natural language processing tasks (Devlin et al., 2018). Despite being a general approach, with almost no task-specific modifications, BERT achieves state-of-the-art performance across a number of natural language processing tasks, including text classification, question answering, inference, and named entity recognition.

Pretrained models like BERT can be used directly for screening automation or automated data extraction. However, by default BERT is trained on a general language corpus, which differs radically in word choice and grammar from the special language found in biomedicine and related fields (Sager et al., 1980). Pretraining on biomedical corpora, rather than general corpora, has been demonstrated to improve performance on several biomedical natural language processing tasks (Lee et al., 2019; Beltagy et al., 2019; Si et al., 2019).

1.2 Objectives

In this study we seek to:

1. Construct a dataset for training machine learning models to identify and extract data from full-text articles on diagnostic test accuracy. We focus on the target condition, index test, and reference standard.

2. Train models to identify specific data items in full-text articles on diagnostic test accuracy.
One of the main aims of our study is to determine how such a dataset should be constructed to allow for training well performing models. In particular, do we need directly supervised data, or can we build reliable models with distantly supervised data? If we do need directly supervised data, how much is necessary?

2 Related Work

There have been attempts to extract several types of data relevant to systematic reviews, most notably extracting **PICO**\(^1\) statements from article text (Wallace et al., 2016; Kiritchenko et al., 2010; Kim et al., 2011; Nye et al., 2018). Other data items include background and study design (Kim et al., 2011), as well as automatically performing risk of bias assessments (Marshall et al., 2014). There is also a recent **TAC** track for data extraction in systematic reviews of environmental agents.\(^2\)

Similarly, previous work by Kiritchenko et al. (2010) aimed to extract 21 different kinds of data from articles, including treatment name, sample size, as well as the primary and secondary outcome from article text. Furthermore, the key criterion for extraction in a systematic review is not the actual data, but the context it appears in. For instance, both intervention studies and a diagnostic studies have target conditions, but these refer to different things: the intervention study seek to *treat* the condition while the diagnostic study seeks to *diagnose* it. As a consequence, in an intervention study the inclusion criterion often mentions the disease, while in a diagnostic study inclusion criteria may mention symptoms rather than the actual disease. This means that a data extraction system trained on interventions may not work as well (or at all) for systematic reviews of diagnostic test accuracy, even though it may seem that the same data is extracted in both. Furthermore, unlike the data required in diagnostic reviews, many previously considered data items are mentioned once in articles, often using formulaic expressions (e.g. sex, blinding, randomization).

Conventional methods for automated data extraction split articles into sentences and classify these using sequence tagging (e.g. **CRF, LSTM**) (Nye et al., 2018).

Despite the body of previous work on automation, many data items relevant to systematic reviews have been overlooked. A 2015 systematic review of data extraction found 26 articles describing the attempted extraction of 52 different data items, but almost all focused on interventions (Jonnalagadda et al., 2015). No study considered any data item specific to diagnostic studies, except for general data items common to both interventions and diagnostic studies, such as age, sex, blinding, or the generation of random allocation sequences. The likely reason for this is that traditional data extraction systems require bespoke training data for each particular data item to extract, which is generally only available through expensive, manual annotation by experts.

A cheaper way to construct datasets for data extraction is to use distant supervision, where the dataset is annotated per article or per review, rather than per sentence or per text span. Supervised methods are then trained on fuzzy annotations derived heuristically for each sentence. For instance, Wallace et al. (2016) used supervised distant supervision to learn to identify **PICO** statements in full text, and Marshall et al. (2014) used supervised distant learning with **SVMs** to identify risk of bias assessments.

There is likely a trade-off between quality and data size. All else being equal, direct supervision is generally better than distant supervision (distantly supervised training data adds a source of noise not present for direct supervision). At the same time, it may not be feasible for experts to annotate large amounts of data. Crowd-sourcing is sometimes used as an alternative to a group of known experts, but if a high degree of expertise is necessary to annotate, crowd-sourcing may not give sufficient guarantees about the expertise of the annotators.

3 Material

We used data from a previous dataset, the **LIMSI-Cochrane** dataset (Norman et al., 2018),\(^3\) to identify references included in previous systematic reviews of diagnostic test accuracy. The **LIMSI-Cochrane** dataset comprises 1,738 references to **DTA** studies from 63 **DTA** systematic reviews. The dataset includes the data extraction forms for each

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\(^1\)Population, intervention, control group, and outcome.


\(^3\)doi: 10.5281/zenodo.1303259
Table 2: The number of sentences in our dataset, broken into distantly annotated training and test sets, as well as a manually annotated subset. Distant annotations for each data type were not available for all studies, and the total number of labelled sentences are therefore different for each data type.

3.0.1 Distant annotation

The data forms from the systematic reviews were intended to be read by and be useful to the human systematic review authors. The contents are therefore usually semi-structured rather than structured, and will include different kinds of data depending on what is relevant to the systematic review (see Table 1).

We create a dataset of distant annotations from the LIMSI-Cochrane dataset by manually converting the semi-structured data into structured data items, and by ensuring that these items can be found in the corresponding article using pattern matching (see Table 1).

We split each of the XML documents into sentences using the nltk sentence splitter. The sentences are then divided into positive and negative depending on whether the relevant data items occur as a partial match in the sentence. Partial matches were calculated using tf-idf cosine similarity between the data item and the sentence, where we took the 20 top ranking sentences for each pair of data item and article, with a similarity score of 0.1 or higher. We chose 20 as a target number of sentences since we felt this was a reasonable upper limit on the number of relevant sentences in a single article. We added an absolute threshold of 0.1 to keep the system from annotating obviously non-relevant sentences (scores close to zero) when no matches could be found in the article. For articles that have multiple data items we used the concatenation of all data items. For example, in Table 1, the data items for ‘Schwartz 1997b’ would be: target condition: ‘Group A streptococcus; Group A streptococcal infection’, index test: ‘QuickVue In-Line Strep A; EIA; ELISA Immunoassays’, and reference standard ‘Microbial culture; Bacterial culture’.

We excluded all articles where the data items were not provided in the data form (because the reviewers did not extract this data), or where data forms were missing from the systematic review. Since we do not know which sentences were relevant or not in these articles we did not use these

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4. https://endnote.com/

5. https://www.nltk.org/
articles as either positive or negative data. As a consequence the total amount of sentences differ for the target condition, index test and reference standard.

We repeated the matching procedure for the target condition, the index test and the reference standard, resulting in three distinct datasets.

3.0.2 Expert annotation

We randomly split the evaluation set into three sets of five systematic reviews. Two experts (ML and RS) on systematic reviews of diagnostic test accuracy manually annotated the 15 articles by highlighting all sentences in the text that 1) mentions the target condition, index test, and reference standard 2) makes it clear that these are the target condition, index test and reference standard, and 3) do not simply mention these same items in an unrelated context. The annotation instructions were written and adjusted twice to remove ambiguity, and the reasons for disagreement were discussed and resolved after two rounds of annotation. As a compromise between getting more data and being able to use the agreement between the experts as baseline for the performance, one expert annotated the first five studies, the second expert annotated the next five studies, and both annotated the last five studies.

4 Method

We construct three pipelines, one for each of the target condition, index test, and reference standard, and we train and evaluate these separately.

We varied our experiments in three dimensions: We tried A) two machine learning algorithms, B) two levels of preprocessing, and C) distantly supervised training data versus directly supervised training data. The directly and distantly supervised models were evaluated on the same data.

4.0.1 A1: BioBERT

We here used a pointwise learning-to-rank approach, where we trained a sentence ranking model by using BioBERT, a version of BERT pretrained on PubMed and PMC (Lee et al., 2019), and fine-tuned the model by training it to regress probability scores. This model was thus trained to map sentences to relevance scores.

To train and evaluate, we used the default BERT setup for the GLUE datasets, modified to output a relevance score rather than a binary value. We used default parameters.

4.0.2 A2: Logistic Regression

We here used a pairwise learning-to-rank approach, where we trained a logistic regression model using stochastic gradient descent (sklearn). As features we used 1) lowercased, \textit{tf-idf} weighted word \textit{n}-grams, 2) lowercased, binary word \textit{n}-grams, 3) lowercased, \textit{tf-idf} weighted, stemmed word \textit{n}-grams, 4) lowercased, stemmed, binary word \textit{n}-grams, as well as i) lowercased, \textit{tf-idf} weighted character \textit{n}-grams, and ii) non-lowercased, \textit{tf-idf} weighted character \textit{n}-grams. We used word \textit{n}-grams up to length 3, and character \textit{n}-grams up to length 6. The first set of features is intended to capture contextual information (‘for the diagnosis of ...’); the second set of features is intended to capture medical technical terms, which are often distinctive at the morpheme level (e.g. ‘ischemia’, ‘anemia’). We deliberately did not use stop-words, since doing so would discard almost all the contextual information. This results in a sparse feature matrix consisting of approximately 1.8 million features for the distantly supervised experiments, and approximately 300,000 features for the directly supervised experiments.

We handled class imbalance by setting the weight for the positive class to 80. This was previously determined to be a reasonable weight in experiments on screening automation in diagnostic test accuracy systematic reviews, a problem with similar class imbalance.

4.0.3 B1: Raw Sentences

Here we used the sentences as they appear in the articles.

4.0.4 B2: Sentences with UMLS Concepts

In this setup we used the Unified Medical Language System, a large ontology of medical concepts maintained by the National Library of Medicine (Bodenreider, 2004; Lindberg et al., 1993). We used MetaMap\(^7\) to locate concept mentions in the sentences, and to replace these with their corresponding UMLS semantic types. For instance the sentence ‘Typhoid fever is a febrile and often serious systemic illness caused by Salmonella enterica serotype Typhi’ was transformed into ‘DSYN is a FNDG and TMCO serious DSYN caused by BACT enterica BACT’.

\(^6\) https://github.com/google-research/bert

\(^7\) https://metamap.nlm.nih.gov/
Table 3: Agreement in terms of recall where columns are considered ground truth, e.g. annotator RS chose 62% of ML’s annotations for the target condition.

<table>
<thead>
<tr>
<th></th>
<th>Auto</th>
<th>ML</th>
<th>RS</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Target condition</strong></td>
<td>Auto</td>
<td>1.00 0.07 0.04</td>
<td></td>
</tr>
<tr>
<td></td>
<td>ML</td>
<td>0.90 1.00 0.38</td>
<td></td>
</tr>
<tr>
<td></td>
<td>RS</td>
<td>1.00 0.62 1.00</td>
<td></td>
</tr>
<tr>
<td><strong>Index test</strong></td>
<td>Auto</td>
<td>1.00 0.09 0.07</td>
<td></td>
</tr>
<tr>
<td></td>
<td>ML</td>
<td>1.00 1.00 0.61</td>
<td></td>
</tr>
<tr>
<td></td>
<td>RS</td>
<td>0.93 0.70 1.00</td>
<td></td>
</tr>
<tr>
<td><strong>Reference standard</strong></td>
<td>Auto</td>
<td>1.00 0.01 0.03</td>
<td></td>
</tr>
<tr>
<td></td>
<td>ML</td>
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<td></td>
</tr>
<tr>
<td></td>
<td>RS</td>
<td>1.00 0.40 1.00</td>
<td></td>
</tr>
</tbody>
</table>

4.0.5 **C1: Directly Supervised Training**

We here trained and evaluated on the articles manually annotated by our two experts (ML and RS), using leave-one-out cross-validation. In other words, to evaluate on each of the ten articles annotated by each annotator we used the remaining 9 articles annotated by the same expert as training data. This was done separately for each expert, and the annotations from the other expert was not used.

4.0.6 **C2: Distantly Supervised Training**

We here trained on the distant annotations from the 48 systematic reviews in the training set, and evaluated on the 15 manually annotated articles in the evaluation set, where each annotator provided annotation data for 10 articles (with a 5 article overlap). The articles used for evaluation were the same as in C1.

4.1 Evaluation

Since our model output ranked sentences, rather than a binary classification, we evaluated all experiments in terms of average precision.

As a comparison, we also evaluated the average precision using the ranking given by the other annotator. In plain language, we tried to evaluate how useful it would have been for the expert to highlight sentences for each other. The expert annotations were binary (Yes/No), rather than a ranking score, so we calculated the average precision by interpolating ties in the ranking.

5 Results

Out of the 1,738 references in the LiMST-Cochrane dataset, 1152 had either a PMID or DOI assigned. EndNote was able to retrieve PDF articles for 666 of these references. A total of 90,996 sentences were distantly labeled for target condition, 94,290 sentences were distantly labeled for index test, and 79,504 sentences were distantly labeled for reference standard. The first annotator (ML) annotated 981 sentences and the second annotator (RS) annotated 1,031 sentences (Table 2).

We present the results of our algorithm evaluated on the annotations by ML in Table 4, and evaluated on the annotations by RS in Table 5.

The ranking performance exhibited large variations. Neither BioBERT or logistic regression were consistently better than the other, neither distant supervision or direct supervision were consistently better than the other, and neither raw sentence nor sentences augmented with UMLS concepts were consistently better than the other. For the target condition, the best performance was achieved by logistic regression on raw sentences using either distant or direct supervision, with a maximum at 0.412 compared to human performance at 0.376 and 0.386 respectively. For the index test, the performance fell within the range 0.344–0.468 compared to human performance at 0.525 and 0.516 respectively. For the reference standard, BioBERT exhibited substantially inferior results on the reference standard compared to logistic regression, while logistic regression performance fell within the range 0.345–0.467, compared to human performance at 0.267 and 0.381 respectively.

The performance also varied between systematic reviews, with consistently close to perfect performance on a few reviews (CD007394 and CD0008782), and consistently very low performance on a few (CD009647 and CD010339). These also correspond to the articles with the highest and lowest inter-annotator agreement. The consensus of the two experts is that CD010339 is not a diagnostic test accuracy study.

6 Discussion

Raw sentences worked consistently better for logistic regression on the target condition (8/8), and worked better than UMLS concepts as a general trend (20/24). While general concepts could theoretically improve performance by help-
Table 4: Average precision results for the 8 different machine learning models on the data annotated by the first annotator (ML), compared to the performance of an independent human expert (annotator RS). The ‘Raw’ columns denote results for models trained and evaluated on raw sentences. The ‘UMLS’ columns denote results for models trained and evaluated on sentences where the concept mentions have been replaced with their corresponding UMLS semantic types. The ‘n pos’ column denotes the number of positive sentences labeled by ML for each article. Rows were omitted for which no sentences were labeled positive. In the baseline results, cells are marked ‘—’ if the article was not annotated by the other expert (RS).

### Target condition

<table>
<thead>
<tr>
<th>n pos</th>
<th>BioBERT</th>
<th>Logistic Regression</th>
<th>As ranked by the other expert (RS)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Distant</td>
<td>Supervised</td>
<td>Distant</td>
</tr>
<tr>
<td></td>
<td>Raw UMLS</td>
<td></td>
<td>Raw UMLS</td>
</tr>
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<td>CD007394</td>
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<td>CD007427</td>
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<td>CD008054</td>
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<td>0.353</td>
</tr>
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<td>CD009372</td>
<td>29</td>
<td>0.354</td>
<td>0.225</td>
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<tr>
<td>CD010339</td>
<td>16</td>
<td>0.388</td>
<td>0.305</td>
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<tr>
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<td>0.269</td>
<td>0.351</td>
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<tr>
<td>CD011420</td>
<td>6</td>
<td>0.251</td>
<td>0.342</td>
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<tr>
<td>mean:</td>
<td>0.336</td>
<td>0.311</td>
<td>0.304</td>
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</table>

### Index test

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<tr>
<th>n pos</th>
<th>BioBERT</th>
<th>Logistic Regression</th>
<th>As ranked by the other expert (RS)</th>
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</thead>
<tbody>
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<td></td>
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<td>Supervised</td>
<td>Distant</td>
</tr>
<tr>
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<td>Raw UMLS</td>
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<td>Raw UMLS</td>
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<tr>
<td>mean:</td>
<td>0.450</td>
<td>0.468</td>
<td>0.344</td>
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</table>

### Reference standard

<table>
<thead>
<tr>
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<th>Logistic Regression</th>
<th>As ranked by the other expert (RS)</th>
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<td>0.075</td>
<td>0.291</td>
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</table>

The models generalize, this may also remove important semantic information from the sentences, keeping the models from ranking accurately. We also note that BioBERT already encodes a language model (similar to word embeddings), and concepts may therefore be unhelpful for the model.

BioBERT performed consistently better than logistic regression on the index test when using distant supervision (4/4), but not when using direct supervision (0/4). Logistic regression performed consistently better than BioBERT on both the target condition and the reference standard (16/16). On the reference standard the difference in performance is substantial, with BioBERT scoring very poorly, and logistic regression performing much better than human performance. The reason for BioBERT’s poor performance on the reference standard may be due to the relative sparsity of the annotations for this subtask (see Table 2).

Distant supervision was consistently on par with or better than direct supervision. The top performing models also outperformed the human annotators on the target condition and the reference standard, and came comparatively close on the index test (0.468 versus 0.525 and 0.444 versus 0.516).

### 6.1 Limitations

We only manually annotated a small sample of the dataset. The small size is further compounded by problems with converting PDF to text, which may also bias the training and evaluation in favor of articles where the conversion works better (mainly articles from big publishers).

The dataset was constructed from articles included in previous systematic reviews of diagnostic test accuracy. These include articles that con-
### 7 Conclusions

Our results suggest that distant supervision is sufficient to train models to identify target condition, index test, and reference standard in diagnostic articles. Our results also suggest that such models can perform on par with human annotators.

We constructed a dataset of full-text articles of diagnostic test accuracy studies, with distant annotations for target condition, index test and reference standard, that can be used to train machine learning models. We also provide a subset of the data manually annotated by experts for evaluation. Our dataset cannot be publicly distributed due to copyright restrictions, but will be available upon request. We also plan to distribute the code for the distant annotations and data preprocessing, as well as the cleaned data extraction forms.

#### 7.1 Future Work

The dataset is being updated, and we plan to increase the amount of manually annotated data to improve the statistical reliability of the experiments. We also plan to let all experts annotate the same articles to simplify the comparisons.

### Acknowledgments

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**Table 5:** Average precision results for the 8 different machine learning models on the data annotated by the second annotator (RS), compared to the performance of an independent human expert (annotator ML). Abbreviations are the same as in Table 4. In the baseline results, cells are marked ‘—’ if the article was not annotated by the other expert (ML).
References


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