# Mining Heterogenous Relationships from Pubmed Abstracts Using Weak Supervision

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### **Abstract**

This is a **rough draft** of a manscript on label function reuse for text mining heterogenous relationship from Pubmed Abstracts.

#Introduction Set introduction for paper here Talk about problem, goal, and significance of paper

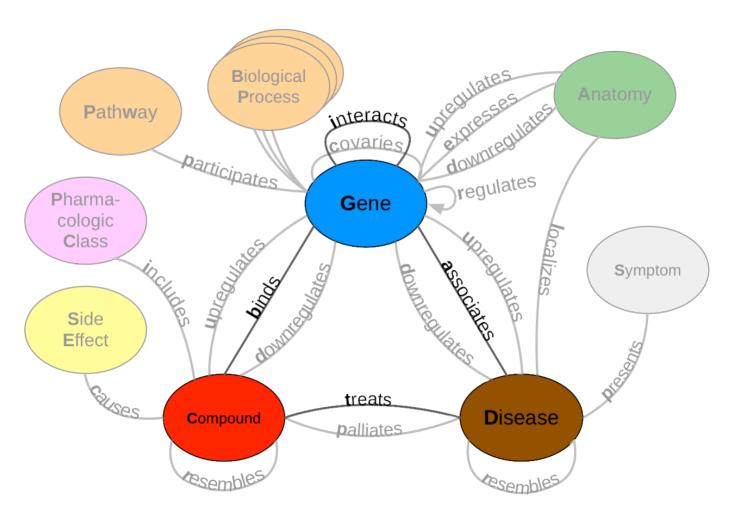
### **Recent Work**

Talk about what has been done in the field in regards to text mining and knowledge base integration

## **Materials and Methods**

### Hetionet

Hetionet [1] is a large heterogenous network that contains pharmacological and biological information. This network depicts information in the form of nodes and edges of different types: nodes that represent biological and pharmacological entities and edges which represent relationships between entities. Hetionet v1.0 contains 47,031 nodes with 11 different data types and 2,250,197 edges that represent 24 different relationship types (Figure 1). Edges in Hetionet were obtained from open databases, such as the GWAS Catalog [2] and DrugBank [2]. For this project, we analyzed performance over a subset of the Hetionet relationship types: disease associates with a gene (DaG), compound binds to a gene (CbG), gene interacts with gene (GiG) and compound treating a disease (CtD).



**Figure 1:** A metagraph (schema) of Hetionet where pharmacological, biological and disease entities are represented as nodes and the relationships between them are represented as edges. This project only focuses on the information shown in bold; however, we can extend this work to incorporate the faded out information as well.

### **Dataset**

We used PubTator [3] as input to our analysis. PubTator provides MEDLINE abstracts that have been annotated with well-established entity recognition tools including DNorm [4] for disease mentions, GeneTUKit [5] for gene mentions, Gnorm [6] for gene normalizations and a dictionary based look system for compound mentions [7]. We downloaded PubTator on June 30, 2017, at which point it contained 10,775,748 abstracts. Then we filtered out mention tags that were not contained in hetionet. We used the Stanford CoreNLP parser [8] to tag parts of speech and generate dependency trees. We extracted sentences with two or more mentions, termed candidate sentences. Each candidates sentence was stratified by co-mention pair to produce a training set, tuning set and a testing set (shown in Table  $\underline{1}$ ). Each unique co-mention pair is sorted into four categories: (1) in hetionet and has sentences, (2) in hetionet and doesn't have sentences, (3) not in hetionet and does have sentences and (4) not in hetionet and doesn't have sentences. Within these four categories each pair receives their own individual partition rank (continuous number between 0 and 1). Any rank lower than 0.7 is sorted into training set, while any rank greater than 0.7 and lower than 0.9 is assigned to tuning set. The rest of the pairs with a rank greater than or equal to 0.9 is assigned to the test set. Sentences that contain more than one co-mention pair are treated as multiple individual candidates. We hand labeled five hundred to a thousand candidate sentences of each relationship to obtain to obtain a ground truth set (Table 1, dataset).

**Table 1:** Statistics of Candidate Sentences. We sorted each candidate sentence into a training, tuning and testing set. Numbers in parentheses show the number of positives and negatives that resulted from the hand-labeling process.

Relationship	Train	Tune	Test
Disease Associates Gene	2.35 M	31K (397+, 603-)	313K (351+, 649-)
Compound Binds Gene	1.7M	468K (37+, 463-)	227k (31+, 469-)
Compound Treats Disease	1.013M	96K (96+, 404-)	32K (112+, 388-)
Gene Interacts Gene	12.6M	1.056M (60+, 440-)	257K (76+, 424-)

# **Label Functions for Annotating Sentences**

A common challenge in natural language processing is having too few ground truth annotations, even when textual data are abundant. Data programming circumvents this issue by quickly annotating large datasets by using multiple noisy signals emitted by label functions [9]. Label functions are simple pythonic functions that emit: a positive label (1), a negative label (-1) or abstain from emitting a label (0). We combine these functions using a generative model to output a single annotation, which is a consensus probability score bounded between 0 (low chance of mentioning a relationship) and 1 (high chance of mentioning a relationship). We used these annotations to train a discriminator model that makes the final classification step. Our label functions fall into three categories: databases, text patterns and domain heuristics. We provide examples for the categories, described below, using the following candidate sentence: "PTK6 may be a novel therapeutic target for pancreatic cancer."

**Databases**: These label functions incorporate existing databases to generate a signal, as seen in distant supervision [10]. These functions detect if a candidate sentence's co-mention pair is present in a given database. If the pair is present, emit a positive label and abstain otherwise. If the pair isn't present in any existing database, then a separate label function will emit a negative label. We use a separate label function to prevent the label imbalance problem. This problem occurs when

candidates, that scarcely appear in databases, are drowned out by negative labels. The multitude of negative labels increases the likelihood of misclassification when training the generative model.

$$\Lambda_{DB}( extstylde{D}, extstylde{G}) = egin{cases} 1 & ( extstylde{D}, extstylde{G}) \in DB \ 0 & otherwise \end{cases}$$

$$\Lambda_{
eg DB}( extbf{ extit{D}}, extbf{ extit{G}}) = egin{cases} -1 & ( extbf{ extit{D}}, extbf{ extit{G}}) 
otherwise \end{cases}$$

**Text Patterns**: These label functions are designed to use keywords and sentence context to generate a signal. For example, a label function could focus on the number of words between two mentions or focus on the grammatical structure of a sentence. These functions emit a positive or negative label depending on the situation. In general, those focused on keywords emit positives and those focused on negation emit negatives.

$$\Lambda_{TP}( extstylde{D}, extstylde{G}) = \left\{egin{array}{ll} 1 & "target" \in Candidate\ Sentence\ 0 & otherwise \end{array}
ight.$$

$$\Lambda_{TP}( extbf{ extit{D}}, extbf{ extit{G}}) = egin{cases} -1 & "VB" 
otherwise \ 0 & otherwise \end{cases}$$

**Domain Heuristics**: These label functions use the other experiment results to generate a signal. For this category, we used dependency path cluster themes generated by Percha et al [11]. If a candidate sentence's dependency path belongs to a previously generated cluster, then the label function will emit a positive label and abstain otherwise.

$$\Lambda_{DH}( extstylde{D}, extstylde{G}) = \left\{egin{array}{ll} 1 & Candidate \, Sentence \in Cluster \, Theme \ 0 & otherwise \end{array}
ight.$$

Roughly half of our label functions are based on text patterns, while the others are distributed across the databases and domain heuristics (Table  $\underline{2}$ ).

**Table 2:** The distribution of each label function per relationship.

Relationship	Databases (DB)	Text Patterns (TP)	Domain Heuristics (DH)
Disease associates Gene (DaG)	7	20	10
Compound treats Disease (CtD)	3	15	7
Compound binds Gene (CbG)	9	13	7
Gene interacts Gene (GiG)	9	20	8

# **Training Models**

### **Generative Model**

talk about generative model and how it works ### Word Embeddings Word embeddings are representations that map individual words to real valued vectors of user-specified dimensions. These embeddings have been shown to capture the semantic and syntatic information between words [12]. Using all candidate sentences for each individual relationship pair, we trained facebook's fastText [13] to generate word embeddings. The fastText model uses a skipgram model [14] that aims to predict

the context given a candidate word and pairs the model with a novel scoring function that treats each word as a bag of character n-grams. We trained this model for 20 epochs using a window size of 2 and generated 300-dimensional word embeddings. We use the optimized word embeddings to train a discriminative model.

### **Discriminator Model**

talk about the discriminator model and how it works ### Discriminator Model Calibration talk about calibrating deep learning models with temperature smoothing

### **Experimental Design**

talk about sampling experiment

### **Results**

### **Random Sampling of Generative Model**

place the grid aurocs here for generative model

### **Discriminator Model Builds Off Generative Model**

place the grid of aurocs here for discriminator model

### **Random Noise Generative Model**

place the results of random label function experiment

### **Reconstructing Hetionet**

place figure of number of new edges that can be added to hetionet as well as edges we can reconstruct using this method

## **Discussion**

Here mention why performnace increases in the beginning for the generative model then decreases

Discuss discriminator model performance given generative model

Mention Take home messages

1. have a centralized set of negative label functions and focus more on contstructing positive label functions

# **Conclusion and Future Direction**

Recap the original problem - takes a long time to create useful label function

Proposed solution - reuse label functions

Mention incorporating more relationships Mention creating a centralized multitask text extractor using this method.

### 1. Systematic integration of biomedical knowledge prioritizes drugs for repurposing

Daniel Scott Himmelstein, Antoine Lizee, Christine Hessler, Leo Brueggeman, Sabrina L Chen, Dexter Hadley, Ari Green, Pouya Khankhanian, Sergio E Baranzini

eLife (2017-09-22) <a href="https://doi.org/cdfk">https://doi.org/cdfk</a>

DOI: 10.7554/elife.26726 · PMID: 28936969 · PMCID: PMC5640425

### 2. The new NHGRI-EBI Catalog of published genome-wide association studies (GWAS Catalog)

Jacqueline MacArthur, Emily Bowler, Maria Cerezo, Laurent Gil, Peggy Hall, Emma Hastings, Heather Junkins, Aoife McMahon, Annalisa Milano, Joannella Morales, ... Helen Parkinson

Nucleic Acids Research (2016-11-29) https://doi.org/f9v7cp

DOI: <u>10.1093/nar/gkw1133</u> · PMID: <u>27899670</u> · PMCID: <u>PMC5210590</u>

### 3. PubTator: a web-based text mining tool for assisting biocuration

Chih-Hsuan Wei, Hung-Yu Kao, Zhiyong Lu

Nucleic Acids Research (2013-05-22) https://doi.org/f475th

DOI: <u>10.1093/nar/gkt441</u> · PMID: <u>23703206</u> · PMCID: <u>PMC3692066</u>

### 4. DNorm: disease name normalization with pairwise learning to rank

R. Leaman, R. Islamaj Dogan, Z. Lu

Bioinformatics (2013-08-21) https://doi.org/f5gj9n

DOI: 10.1093/bioinformatics/btt474 · PMID: 23969135 · PMCID: PMC3810844

### 5. GeneTUKit: a software for document-level gene normalization

M. Huang, J. Liu, X. Zhu

Bioinformatics (2011-02-08) https://doi.org/dng2cb

DOI: 10.1093/bioinformatics/btr042 · PMID: 21303863 · PMCID: PMC3065680

### 6. Cross-species gene normalization by species inference

Chih-Hsuan Wei, Hung-Yu Kao

BMC Bioinformatics (2011-10-03) https://doi.org/dnmvds

DOI: <u>10.1186/1471-2105-12-s8-s5</u> · PMID: <u>22151999</u> · PMCID: <u>PMC3269940</u>

# 7. Collaborative biocuration-text-mining development task for document prioritization for curation

T. C. Wiegers, A. P. Davis, C. J. Mattingly

Database (2012-11-22) https://doi.org/gbb3zw

DOI: 10.1093/database/bas037 · PMID: 23180769 · PMCID: PMC3504477

### 8. The Stanford CoreNLP Natural Language Processing Toolkit

Christopher Manning, Mihai Surdeanu, John Bauer, Jenny Finkel, Steven Bethard, David McClosky *Proceedings of 52nd Annual Meeting of the Association for Computational Linguistics: System Demonstrations* (2014) <a href="https://doi.org/gf3xhp">https://doi.org/gf3xhp</a>

DOI: 10.3115/v1/p14-5010

### 9. Data Programming: Creating Large Training Sets, Quickly

Alexander Ratner, Christopher De Sa, Sen Wu, Daniel Selsam, Christopher Ré *arXiv* (2016-05-25) <a href="https://arxiv.org/abs/1605.07723v3">https://arxiv.org/abs/1605.07723v3</a>

### 10. Distant supervision for relation extraction without labeled data

Mike Mintz, Steven Bills, Rion Snow, Dan Jurafsky

Proceedings of the Joint Conference of the 47th Annual Meeting of the ACL and the 4th International Joint Conference on Natural Language Processing of the AFNLP: Volume 2 - ACL-IJCNLP '09 (2009)

https://doi.org/fg9q43

DOI: 10.3115/1690219.1690287

### 11. A global network of biomedical relationships derived from text

Bethany Percha, Russ B Altman

Bioinformatics (2018-02-27) https://doi.org/gc3ndk

DOI: <u>10.1093/bioinformatics/bty114</u> · PMID: <u>29490008</u> · PMCID: <u>PMC6061699</u>

### 12. Distributed Representations of Words and Phrases and their Compositionality

Tomas Mikolov, Ilya Sutskever, Kai Chen, Greg Corrado, Jeffrey Dean *arXiv* (2013-10-16) <a href="https://arxiv.org/abs/1310.4546v1">https://arxiv.org/abs/1310.4546v1</a>

### 13. Enriching Word Vectors with Subword Information

Piotr Bojanowski, Edouard Grave, Armand Joulin, Tomas Mikolov *arXiv* (2016-07-15) <a href="https://arxiv.org/abs/1607.04606v2">https://arxiv.org/abs/1607.04606v2</a>

### 14. Efficient Estimation of Word Representations in Vector Space

Tomas Mikolov, Kai Chen, Greg Corrado, Jeffrey Dean *arXiv* (2013-01-16) <a href="https://arxiv.org/abs/1301.3781v3">https://arxiv.org/abs/1301.3781v3</a>