Linguistic Analysis of the bioRxiv Preprint Landscape

This manuscript (<u>permalink</u>) was automatically generated from <u>greenelab/annorxiver manuscript@5624e41</u> on September 9, 2020.

Authors

• David N. Nicholson

Department of Systems Pharmacology and Translational Therapeutics, University of Pennsylvania \cdot Funded by GBMF4552; T32 HG00046

• Jane Roe

Department of Something, University of Whatever; Department of Whatever, University of Something

Abstract

Introduction

- 1. What is a preprint
- 2. Why are preprints important?
- 3. Mention how preprints are being integrated into scientist's everyday workflow
- 4. Talk about biorxiv and discuss how it is one of the repositories that maintains preprints along with citation of others such as arxiv/medrxiv etc.
- 5. Discuss works that analyze biorxiv from an audience perspective (quantifying tweets etc.)
- 6. Mention the gap which consists of analysing the language of biorxiv preprints (first to do this)
- 7. ^ Why is this important? What will this allow for future research projects?
- 8. Provide list of contributions within this manuscript

Methods

Datasets

BioRxiv

BioRxiv [1] is a repository of biological and biomedical research preprints. We downloaded an xml snapshot of this repository on February 3, 2020 from bioRxiv's Amazon S3 resource [2] that contained the full text and image content of 98,023 preprints. Preprints on bioRxiv are versioned, and in our snapshot 26,905 of 98,023 contained more than one version. When preprints had multiple versions, we used only the latest one. Preprints in this snapshot were grouped into one of twenty-nine different categories. Each preprint was also classified as a new result, confirmatory finding, or contradictory finding. Some preprints in this snapshot have been withdrawn from bioRxiv. When a preprint is withdrawn, its content is replaced with the reason for withdrawal. Because we used the latest version, withdrawn preprints in our analysis contained only statements indicating their removal.

PubMed Central

PubMed Central (PMC) [3] is a repository that contains free-to-read articles. PMC contains two types of contributions: closed access articles from research funded by the United States National Institutes of Health (NIH) appearing after an embargo period and articles published under Gold Open Access [4] publishing schemes. Paper availability within PMC is largely dependent on the journal's participation level [5]. Individual journals have can fully participate in submitting articles to PMC, selectively participate sending only a few few of papers to PMC, only submit papers according to NIH's public access policy [6], or not participate at all. As of September 2019, PMC had 5,725,819 articles available [7]. Out of these 5 million articles, about 3 million were open access and available for text processing systems [8,9]. We downloaded a snapshot of this open access subset on January 31, 2020. This snapshot contains papers such as literature reviews, book reviews, editorials, case reports, research articles and more; however, we used only the research articles.

Comparing Corpora

We used gensim [10] (version 3.8.1) to preprocess the bioRxiv and PubMed Central corpora. We removed the 337 gensim-provided stopwords. Throughout our analysis we encountered non-word symbols (e.g., \pm), so we refer to words and symbols as tokens to avoid confusion.

Following the cleaning process, we calculated the frequency of every token shared between both corpora. Because many tokens were unique to one set or the other and observed at low frequency, we used the union of the top 100 most frequent tokens from each corpus to compare them. We generated a contingency table and calculated the odds ratio for each token. Furthermore, we also calculated the 95% confidence interval for each odds ratio [11].

Visualizing the Preprint Landscape

Generate Document Representation

We used gensim [10] (version 3.8.1) to train a word2vec continuous bag of words (CBOW) [12] model over the bioRxiv corpus. Our neural network architecture had 300 hidden nodes, and we trained this model for 20 epochs. We set a fixed random seed and otherwise used gensim's default settings. Following training, we generated a document vector for every article within bioRxiv and PubMed Central. This document vector is calculated by taking the average of every token present within a given article, ignoring those that were absent from the word2vec model.

Dimensionality Reduction of Document Embeddings

We used principal component analysis (PCA) [13] to project bioRxiv document vectors into a low dimensional space. We trained this model using scikit-learn's [14] implementation of a randomized solver [15] with a random seed of 100, output of 50 principal components, and default settings for all other parameters. For each principal component we calculated its cosine similarity with all tokens in our word2vec model's vocabulary. We report the top 100 positive and negative scoring tokens in the form of word clouds, where the size of each word corresponds to the magnitude of similarity and color represents positive (blue) or negative (orange) association.

Discovering Unannotated Preprint-Publication Relationships

Automated procedures are in place to link preprints to peer reviewed versions and many journals require authors to update preprints with a link to the published version. However, automated procedures at bioRxiv are often based on exact matching of certain attributes and authors can forget to establish a link after publication. For example, authors can change the title between a preprint and published version (e.g., [16] and [17]), which prevents bioRxiv from automatically establishing a link. If the authors do not report the publication to bioRxiv, the preprint and the published version are treated as distinct entities despite representing the same underlying research. We recognized that the distance in embedding space could reveal preprint to published version links that were missed by existing automated processes. First, we used CrossRef [18] to identify bioRxiv preprints that were linked to a corresponding published article. We filtered out links that contained papers not in PubMed Central's Open Access corpus. Following the preprocessing step, we calculated the distribution of known preprint to published distances by taking the Euclidean distance between the preprint's embedding coordinates and the coordinates of the published version. We also calculated a background distribution, which consisted of the distance between each preprint with an annotated publication and a randomly selected article from the same journal the published version. Next, we calculated distances between preprints without a published version link with PubMed Central articles that weren't matched with a corresponding preprint. We filtered any potential links with distances that were greater than the minimum value of the background distribution to reduce the curation burden. Lastly, we binned the remaining pairs based on percentiles from the annotated pairs at the [0,25th percentile), [25th percentile, 50th percentile), [50th percentile, 75th percentile), and [75th percentile,

minimum background distance). We randomly sampled 50 articles from each bin for manual annotation. We shuffled these four sets to produce a list of 200 potential preprint-published pairs with a randomized order. We supplied these candidates to two scientists to manually determine if each link between a preprint and a putative matched version was correct or incorrect. After the curation process we encountered eight mismatches. These mismatches were supplied to a third scientist, who carefully reviewed each case and made a final determination. Lastly, we used this curated set to evaluate the extent to which distance in the embedding space revealed true but unannotated links between preprints and their published verisons.

Journal Recommendation

We aimed to predict the journal a paper would eventually be published in based on its embedding representation. We illustrate our recommendations as a short list along with a network visualization available at https://greenelab.github.io/annorxiver-journal-recommender/. Since we sought to examine if embeddings were related to publication venue, we used a simple k-nearest neighbors approach with Euclidean distance to recommend journals.

First, we filtered all journals that had fewer than 100 papers in the PMC dataset. A subset of our PMC corpus was directly linked to papers in bioRxiv as they had been published as open access articles. We held out this subset and treated it as our gold standard test set. We used the remainder of the PMC corpus for training and initial evaluation via cross validation. We considered a list of ten journal suggestions to be an appropriate number and we considered a prediction to be a true positive if the correct journal appeared within the ten closest neighbors of the query article.

Certain journals publish articles in a focused topic area, while others publish articles that cover many topics. Likewise, some journals have a publication rate of at most hundreds of papers per year while others publish at a rate of at least ten-thousand papers per year. Accounting for these characteristics, we designed two approaches for recommending journals.

The first approach is based on individual paper proximity, which enabled us to provide an example of the specific article or articles that led to the prediction. Conversely, predictions using this technique could be biased due to the overrepresentation of general topic journals. We call this approach the paper-based classifier. This classifier takes a query article that has been projected onto the embedding space trained on bioRxiv preprints as input and reports the journals of the top ten closest papers. The number of journals returned via this method could be less than ten as multiple papers in close proximity to query article may belong to the same journal.

The second approach is based on close proximity to a journal's centroid. This technique provides recommendations that are more focused on domain-specific publication venues. We call this approach the journal-based classifier. This classifier was trained by computing journal centroids via taking the average embedding of all papers published in each journal. Following the centroid calculation, this classifier takes a query article projected onto the same embedding space as above for input and reports the top ten nearest journals centroids. Both the paper-based classifier and the journal-based classifier were optimized via 10-fold cross validation. We evaluated performance of both classifiers on our gold standard test set of published preprints.

We used SAUCIE [19] to train a model that uses the latent space of a neural network to learn an embedding suitable for visualization. This model enabled us to visualized the PMC corpus and to efficiently embed new papers and preprints with the space. We trained this model using a learning rate of 0.001, lambda_b of 0, lambda_c of 0.001, and lambda_d of 0.001 for 2000 iterations. We used the fully trained model to project user-requested *bioRxiv* preprints onto the generated landscape to enable users to see where their preprint falls along the landscape.

Comparing bioRxiv to PubMed Central

bioRxiv Repository

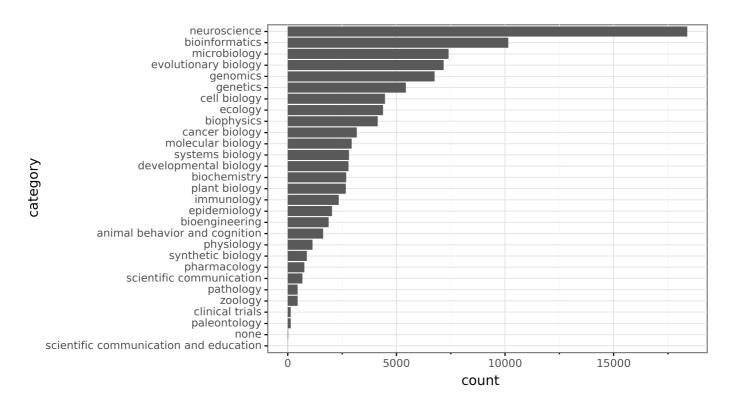


Figure 1: Neuroscience and bioinformatics are the two most common topics for preprints on bioRxiv. This bar chart depicts the number of preprints that fall into each author-selected topic area.

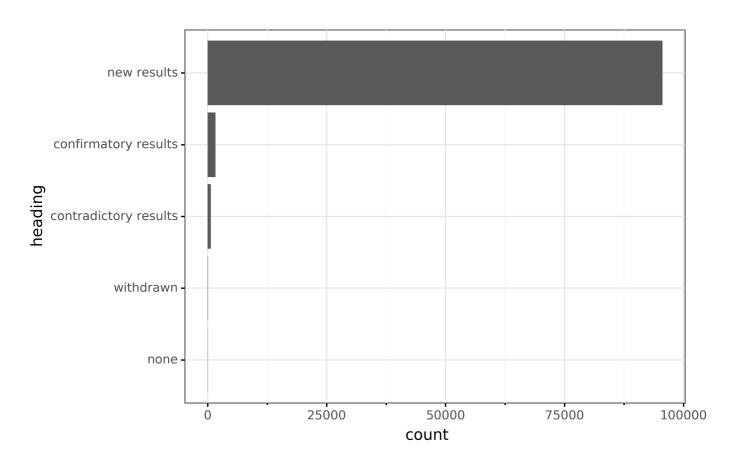


Figure 2: Most of bioRxiv's preprints report new research findings. This bar chart depicts the number of articles categorized based on author-selected article types.

Each preprint on bioRxiv has an author-selected topic area, and the predominant area in past reports has been neuroscience [20]. Our analysis of the full text release of bioRxiv confirms this previous finding (Figure 1). The author-selected topic area abundances that we found in the full text largely matched previous studies [20,21]. One exception was microbiology, which has a larger share of preprints than in a previous report from 2018 [20] (Figure 1). Authors also select from three article types when they upload their preprints. We found that most preprints are categorized as new results (Figure 2), which is consistent with previous findings [21].

Global Comparison of bioRxiv and PubMed Central

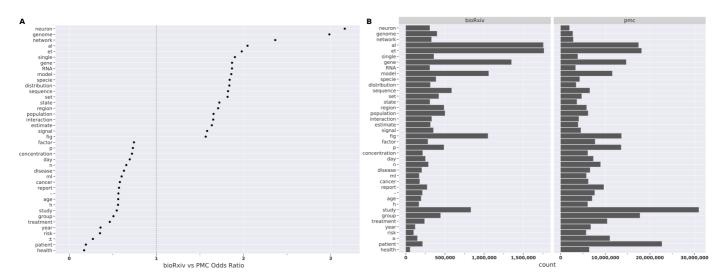


Figure 3: BioRxiv is more focused on biological discoveries rather than disease treatments and clinical trials. The plot on the left (A) is a point range plot of the odds ratio with respect to bioRxiv. Values greater than one indicate a high association with bioRxiv whereas values less than one indicate high association with PubMed Central. The dotted line provides a breaking point between both categories. The plot on the right (B) is a bar chart of token frequency appearing in bioRxiv and PMC respectively.

The linguistic style of the bioRxiv corpus differs from the PMC corpus. We compared preprints in bioRxiv with published manuscripts in PMC. We found that tokens such as "neuron", "genome", "RNA" and "network" had a high odds ratio, indicating enrichment in bioRxiv (Figure $\underline{3}$). Likewise, we found tokens such as "patient", "health", \pm , and "ml" to have a low odds ratio, indicating enrichment in PMC (Figure $\underline{3}$). This separation of tokens suggests a prevalence of articles focused on clinical trials and patient research within PMC compared to bioRxiv. Furthermore, bioRxiv has a predominance of neuroscience and bioinformatic topics. In regard to writing, citation styles diversify from the familiar "et al." form as preprints transition through the publication process. Additionally, published articles have an increase of typesetting (\pm) and measurement symbols ("ml", "age") compared to preprints.

Published Preprint Differences

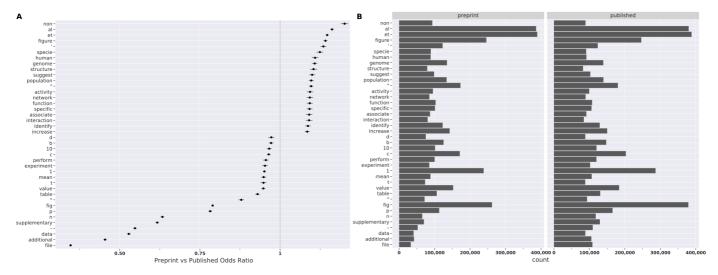


Figure 4: Top scoring tokens for preprints are focused on figure citations whereas their published versions are more focused on data availability. The plot on the left (A) is a point range plot of the odds ratio with respect to preprints. Values greater than one indicate a high association with preprints while values less than one indicate a high association with published articles. The dotted line provides a breaking point between both categories. The plot on the right (B) is a barchart of token frequency appearing in preprints and published versions of preprints respectively.

A preprint's linguistic style can change once a preprint has undergone the revision process prior to being published. We quantified this linguistic difference by calculating the odds ratio of tokens appearing in the union in bioRxiv preprints and their published counterparts within PMC. Tokens with an odds ratio greater than one are mainly centered on paper/figure references and research specific terms (Figure 4). Tokens with an odds ratio of less than one are focused on data availability, and research measurements such as number of cases and controls or significance testing (Figure 4).

Topic Analysis of bioRxiv's Principal Components

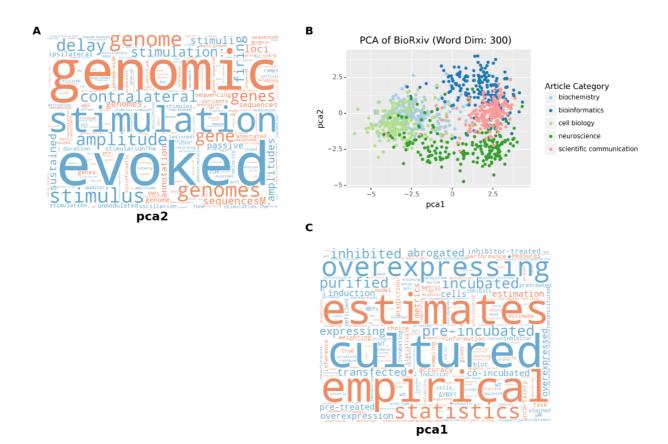


Figure 5: The top two principal components (PCs) appear to capture the concepts of molecular biology vs quantitative biology (PC1) and neuroscience vs bioinformatics (PC2). The word clouds (A, C) depict the cosine similarity score between tokens and the first two PCs. Tokens in orange are most similar to a PC's positive direction while tokens in blue

are most similar to a PC's negative direction. The size of each token indicates the magnitude of the similarity score. The scatter plot at the top right (B) is a visualization of documents being plotted along the PC directions. Article categories were hand-picked based on the concepts captured by each PC.

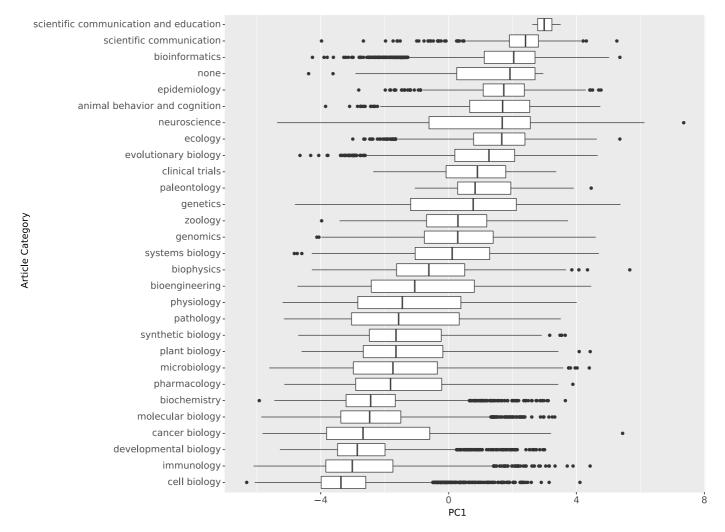


Figure 6: Preprint categories have a diverse spread of quantitative and molecular biology results. This is box plot shows preprints in each article category projected along the PC1 direction. Negative values indicate molecular biology concepts, while positive values indicate quantitative biology concepts.

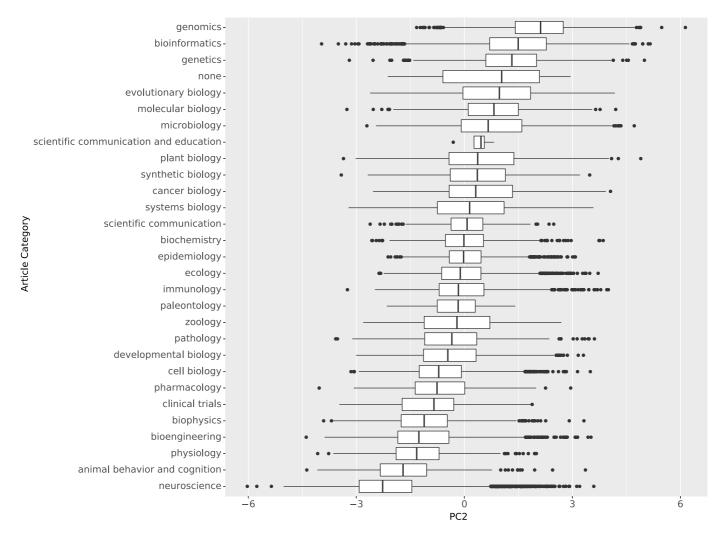


Figure 7: The second PC groups neuroscience related preprint categories and bioinformatics related preprint categories together. This is box plot shows preprints in each article category projected along the PC2 direction. Negative values indicate neuroscience concepts, while positive values indicate bioinformatic concepts.

Table 1: Top five and bottom five systems biology preprints projected onto the PC1 direction. These preprints contain quantitative and molecular biology concepts respectively.

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
Conditional Robust Calibration (CRC): a new computational Bayesian methodology for model parameters estimation and identifiability analysis [22]	4.70055 4908074 704	Non e	h t t p s : / w w w

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
			i o r x i v . o r g /
			c o n t e n t / b i o
			r x i v / e a r !
			/ 2 0 1 7 / 1 0 / 0 2 /
			/ 1 9 7 4 0 / F 1

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
			. I a r g e . j p g
Machine learning of stochastic gene network phenotypes [23]	4.41066 0604449 826	CC- BY- NC- ND	https://www.biorxiv.org/content/biorxiv/ea

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
			r y/2019/10/31/825943/F5.large.jpg
Notions of similarity for computational biology models [24]	4.35529 5926618 207	CC- BY- NC- ND	h t t p s : / w w . b i o r x i

Title [citation]	PC_1	Lice nse	i
			/ () () () () () () () () () () () () ()

Title [citation]	PC_1	Lice nse	F i g u r e L i n
			e j p
GpABC: a Julia package for approximate Bayesian computation with Gaussian process emulation [25]	4.35151 7618262 304	CC- BY- NC- ND	https://www.biorxiv.org/content/biorxiv/early/2

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
			0 1 9 / 0 9 / 1 8 / 7 6 9 2 9 9 / F 1 I a r g e j p g e j p g
SBpipe: a collection of pipelines for automating repetitive simulation and analysis tasks [26]	4.32184 7854182 741	CC- BY- NC- ND	h t t p s : / / w w . b i o r x i v . o r g

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
			/ c o n t e n t / b
			i o r x i v / e a r
			J y / 2 0 1 7 / 0
			/ 0 9 / 1 0 7 2 5
			/ F 1 I a r g e
			j p g

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
Spatiotemporal proteomics uncovers cathepsin-dependent host cell death during bacterial infection [22]	-4.26396 4235099 807	CC- BY- ND	https://www.biorxiv.org/content/biorxiv/early/2018/1

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
Systems analysis by mass cytometry identifies susceptibility of latent HIV-infected T cells to	-4.27901	CC-	1 / 0 7 / 4 5 5 0 4 8 / F 1 . l a r g e . j p g h
targeting of p38 and mTOR pathways [28]	6673409	BY- NC- ND	t t p s : / / w w w . b i o r x i v . o r g / c o n t

Title [citation]	PC_1	Lice nse	i g u r e L i n k
			e n
			t /
			b i
			o r
			i
			/ /
			e a
			r I
			у / 2
			0
			8
			0
			1
			9
			3 7
			1
			2
			/ F
			1
			l a
			r g
			e
			j p g
ADPH consumption by L-cystine reduction creates a metabolic vulnerability upon glucose eprivation [29]	-4.59220 9988884 236	Non e	h t t

Title [citation]	PC_1	Lice nse

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
	4.70564		7 3 3 1 6 2 / F 1 . I a r g e . j p g
Inhibition of Bruton's tyrosine kinase reduces NF-kB and NLRP3 inflammasome activity preventing insulin resistance and microvascular disease [30]	-4.73661 3689905 791	Non e	https://www.biorxiv.org/content/b

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
			i o r x i v / e a r l y / 2 0 1 9 / 0 8 / 2 8 / 7 4 5 9 4 3 / F 1 . l a r g e . j p g
AKT but not MYC promotes reactive oxygen species-mediated cell death in oxidative culture [31]	-4.82679 3742506 695	Non e	h t p s : / w

Title [citation]	PC_1	Lice nse	e L i n k
			w b i o r x i v
			org/conten
			t / b i o r x i v /
			e a r l y / 2 0 1 9
			y / 2 0 1 9 / 0 9 / 0 1 / 7 5 4 5 7

Title [citation]	PC_1	Lice nse	F i g u r e L i n k
			2 / F
			1
			I
			a r
			g
			e
			j
			p
			g

We explored the primary differences between the full text of bioRxiv preprints by performing principal components analysis on generated document embeddings. We visualized the correspondence between tokens and the loadings for each principal component (Figure {5})A,C). We also visualized documents projected on selected principal components (Figure {5})B). The first principal component separates bioRxiv preprints that encompass molecular biology results with preprints that contain quantitative biology results (Figure {5})C). This highlights the bisection of biomedical research where majority of results can be categorized under the molecular biology category or the quantitative biology category. Furthermore, this bisecting trend is evident across individual preprint categories as most categories lie on either side of the first principal component (Figure 6). We also provide example preprints from the systems biology category to reinforce this concept (Table 1).

The second principal component represents the concept of neuroscience vs bioinformatics (Figure {5})A). This principal component suggests that the bulk of preprints within bioRxiv are largely focused around neuroscience and bioinformatic concepts. This split is evident in Figure 7 as enriched categories along this principal component are quite related to neuroscience (negative end) or bioinformatics (positive end). As with the first principal component we provide example preprints from the systems biology category to reinforce this concept (Supplemental Table 2). More principal component word clouds can be found on our journal recommender website (greenelab.github.io/annorxiver-journal-recommender) and within our online repository (see Data Availability).

Journal Recommendations/Audience Associations

- 1. Title will change once analysis is finished
- 2. Provide key figure for this section and take-home message

Discussion

Conclusion

Acknowledgements



References

1. bioRxiv: the preprint server for biology

Richard Sever, Ted Roeder, Samantha Hindle, Linda Sussman, Kevin-John Black, Janet Argentine, Wayne Manos, John R. Inglis

bioRxiv (2019-11-06) https://doi.org/ggc46z

DOI: 10.1101/833400

2. Machine access and text/data mining resources | bioRxivhttps://www.biorxiv.org/tdm

3. PubMed Central: The GenBank of the published literature

R. J. Roberts

Proceedings of the National Academy of Sciences (2001-01-16) https://doi.org/bbn9k8

DOI: 10.1073/pnas.98.2.381 · PMID: 11209037 · PMCID: PMC33354

4. Gold open access: the best of both worlds

M. A. G. van der Heyden, T. A. B. van Veen

Netherlands Heart Journal (2017-12-01) https://doi.org/ggzfr9

DOI: 10.1007/s12471-017-1064-2 · PMID: 29196877 · PMCID: PMC5758455

5. How Papers Get Into PMChttps://www.ncbi.nlm.nih.gov/pmc/about/submission-methods/

6. 8.2.2 NIH Public Access

Policyhttps://grants.nih.gov/grants/policy/nihgps/html5/section-8/8.2.2_nih_public_access_policy.html

7. PMC Overviewhttps://www.ncbi.nlm.nih.gov/pmc/about/intro/

8. PMC text mining subset in BioC: about three million full-text articles and growing

Donald C Comeau, Chih-Hsuan Wei, Rezarta Islamaj Doğan, Zhiyong Lu *Bioinformatics* (2019-09-15) https://doi.org/ggzfsb

DOI: 10.1093/bioinformatics/btz070 · PMID: 30715220 · PMCID: PMC6748740

9. PubTator central: automated concept annotation for biomedical full text articles

Chih-Hsuan Wei, Alexis Allot, Robert Leaman, Zhiyong Lu

Nucleic Acids Research (2019-07-02) https://doi.org/ggzfsc

DOI: 10.1093/nar/gkz389 · PMID: 31114887 · PMCID: PMC6602571

10. Software Framework for Topic Modelling with Large Corpora

Radim Řehůřek, Petr Sojka

Proceedings of the LREC 2010 Workshop on New Challenges for NLP Frameworks (2010-05-22)

11. Odds Ratio (OR)

Steven Tenny, Mary R. Hoffman

StatPearls (2020) http://www.ncbi.nlm.nih.gov/books/NBK431098/

12. Efficient Estimation of Word Representations in Vector Space

Tomas Mikolov, Kai Chen, Greg Corrado, Jeffrey Dean *arXiv* (2013-09-10) https://arxiv.org/abs/1301.3781

13. Probabilistic Principal Component Analysis

Michael E. Tipping, Christopher M. Bishop

Journal of the Royal Statistical Society: Series B (Statistical Methodology) (1999-08)

https://doi.org/b3hjwt

DOI: <u>10.1111/1467-9868.00196</u>

14. Scikit-learn: Machine learning in Python

F. Pedregosa, G. Varoquaux, A. Gramfort, V. Michel, B. Thirion, O. Grisel, M. Blondel, P. Prettenhofer, R. Weiss, V. Dubourg, ... E. Duchesnay *Journal of Machine Learning Research* (2011)

15. Finding structure with randomness: Probabilistic algorithms for constructing approximate matrix decompositions

Nathan Halko, Per-Gunnar Martinsson, Joel A. Tropp *arXiv* (2014-04-29) https://arxiv.org/abs/0909.4061

16. The *Drosophila* Cortactin Binding Protein 2 homolog, Nausicaa, regulates lamellipodial actin dynamics in a Cortactin-dependent manner

Meghan E. O'Connell, Divya Sridharan, Tristan Driscoll, Ipsita Krishnamurthy, Wick G. Perry, Derek A. Applewhite

bioRxiv (2018-07-24) https://doi.org/gg4hp7

DOI: <u>10.1101/376665</u>

17. The *Drosophila* protein, Nausicaa, regulates lamellipodial actin dynamics in a Cortactindependent manner

Meghan E. O'Connell, Divya Sridharan, Tristan Driscoll, Ipsita Krishnamurthy, Wick G. Perry, Derek A. Applewhite

Biology Open (2019-06-15) https://doi.org/gg4hp8

DOI: <u>10.1242/bio.038232</u> · PMID: <u>31164339</u> · PMCID: <u>PMC6602326</u>

18. CrossRef Text and Data Mining Services

Rachael Lammey

Insights the UKSG journal (2015-07-07) https://doi.org/gg4hp9

DOI: <u>10.1629/uksg.233</u>

19. Assessing the Heterogeneity of Cardiac Non-myocytes and the Effect of Cell Culture with Integrative Single Cell Analysis

Brian S. Iskra, Logan Davis, Henry E. Miller, Yu-Chiao Chiu, Alexander R. Bishop, Yidong Chen, Gregory J. Aune

bioRxiv (2020-03-05) https://doi.org/gg9353

DOI: <u>10.1101/2020.03.04.975177</u>

20. Tracking the popularity and outcomes of all bioRxiv preprints

Richard J Abdill, Ran Blekhman

eLife (2019-04-24) https://doi.org/gf2str

DOI: 10.7554/elife.45133 · PMID: 31017570 · PMCID: PMC6510536

21. Altmetric Scores, Citations, and Publication of Studies Posted as Preprints

Stylianos Serghiou, John P. A. Ioannidis

JAMA (2018-01-23) https://doi.org/gftc69

DOI: 10.1001/jama.2017.21168 · PMID: 29362788 · PMCID: PMC5833561

22. Conditional Robust Calibration (CRC): a new computational Bayesian methodology for model parameters estimation and identifiability analysis

Fortunato Bianconi, Chiara Antonini, Lorenzo Tomassoni, Paolo Valigi

bioRxiv (2017-10-02) https://doi.org/gg9393

DOI: <u>10.1101/197400</u>

23. Machine learning of stochastic gene network phenotypes

Kyemyung Park, Thorsten Prüstel, Yong Lu, John S. Tsang

bioRxiv (2019-10-31) https://doi.org/gg94bm

DOI: <u>10.1101/825943</u>

24. Notions of similarity for computational biology models

Ron Henkel, Robert Hoehndorf, Tim Kacprowski, Christian Knüpfer, Wolfram Liebermeister, Dagmar Waltemath

bioRxiv (2016-03-21) https://doi.org/gg939z

DOI: 10.1101/044818

25. **GpABC:** a Julia package for approximate Bayesian computation with Gaussian process emulation

Evgeny Tankhilevich, Jonathan Ish-Horowicz, Tara Hameed, Elisabeth Roesch, Istvan Kleijn, Michael PH Stumpf, Fei He

bioRxiv (2019-09-18) https://doi.org/gg94bj

DOI: 10.1101/769299

26. SBpipe: a collection of pipelines for automating repetitive simulation and analysis tasks

Piero Dalle Pezze, Nicolas Le Novère

bioRxiv (2017-02-09) https://doi.org/gg9392

DOI: <u>10.1101/107250</u>

27. Spatiotemporal proteomics uncovers cathepsin-dependent host cell death during bacterial infection

Joel Selkrig, Nan Li, Jacob Bobonis, Annika Hausmann, Anna Sueki, Haruna Imamura, Bachir El Debs, Gianluca Sigismondo, Bogdan I. Florea, Herman S. Overkleeft, ... Athanasios Typas bioRxiv (2018-11-07) https://doi.org/gg94bc

DOI: <u>10.1101/455048</u>

28. Systems analysis by mass cytometry identifies susceptibility of latent HIV-infected T cells to targeting of p38 and mTOR pathways

Linda E. Fong, Victor L. Bass, Serena Spudich, Kathryn Miller-Jensen

bioRxiv (2018-07-19) https://doi.org/gg9398

DOI: 10.1101/371922

29. NADPH consumption by L-cystine reduction creates a metabolic vulnerability upon glucose deprivation

James H. Joly, Alireza Delfarah, Philip S. Phung, Sydney Parrish, Nicholas A. Graham *bioRxiv* (2019-08-13) https://doi.org/gg94bf

DOI: <u>10.1101/733162</u>

30. Inhibition of Bruton's tyrosine kinase reduces NF-kB and NLRP3 inflammasome activity preventing insulin resistance and microvascular disease

Gareth S. D. Purvis, Massimo Collino, Haidee M. A. Tavio, Fausto Chiazza, Caroline E. O'Riodan, Lynda Zeboudj, Nick Guisot, Peter Bunyard, David R. Greaves, Christoph Thiemermann bioRxiv (2019-08-28) https://doi.org/gg94bg

DOI: 10.1101/745943

31. AKT but not MYC promotes reactive oxygen species-mediated cell death in oxidative culture

Dongqing Zheng, Jonathan H. Sussman, Matthew P. Jeon, Sydney T. Parrish, Alireza Delfarah, Nicholas A. Graham

bioRxiv (2019-09-01) https://doi.org/gg94bh

DOI: <u>10.1101/754572</u>

32. Pangenome Analysis of Enterobacteria Reveals Richness of Secondary Metabolite Gene Clusters and their Associated Gene Sets

Omkar S. Mohite, Colton J. Lloyd, Jonathan M. Monk, Tilmann Weber, Bernhard O. Palsson *bioRxiv* (2019-09-25) https://doi.org/gg94bk

DOI: <u>10.1101/781328</u>

33. QTG-Finder: a machine-learning based algorithm to prioritize causal genes of quantitative trait loci

Fan Lin, Jue Fan, Seung Y. Rhee

bioRxiv (2019-04-29) https://doi.org/gg94bd

DOI: <u>10.1101/484204</u>

34. Identification of candidate genes underlying nodulation-specific phenotypes in *Medicago* truncotula through integration of genome-wide association studies and co-expression networks

Jean-Michel Michno, Liana T. Burghardt, Junqi Liu, Joseph R. Jeffers, Peter Tiffin, Robert M. Stupar, Chad L. Myers

bioRxiv (2018-08-16) https://doi.org/gg94bb

DOI: <u>10.1101/392779</u>

35. Raw sequence to target gene prediction: An integrated inference pipeline for ChIP-seq and RNA-seq datasets

Nisar Wani, Khalid Raza

bioRxiv (2017-11-16) https://doi.org/gg9394

DOI: <u>10.1101/220152</u>

36. The y-ome defines the thirty-four percent of *Escherichia coli* genes that lack experimental evidence of function

Sankha Ghatak, Zachary A. King, Anand Sastry, Bernhard O. Palsson

bioRxiv (2018-12-03) https://doi.org/gg9396

DOI: <u>10.1101/328591</u>

37. The effects of time-varying temperature on delays in genetic networks

Marcella M Gomez, Richard M Murray, Matthew R Bennett

bioRxiv (2015-09-24) https://doi.org/gg939x

DOI: <u>10.1101/019687</u>

38. An analog to digital converter creates nuclear localization pulses in yeast calcium signaling

lan S Hsu, Bob Strome, Sergey Plotnikov, Alan M Moses

bioRxiv (2018-06-28) https://doi.org/gg9397

DOI: <u>10.1101/357939</u>

39. Nicotinic modulation of hierarchal inhibitory control over prefrontal cortex resting state dynamics: modeling of genetic modification and schizophreniarelated pathology

Marie Rooy, Fani Koukouli, Uwe Maskos, Boris Gutkin

bioRxiv (2018-04-13) https://doi.org/gg9395

DOI: <u>10.1101/301051</u>

40. Electrical propagation of vasodilatory signals in capillary networks

Pilhwa Lee

bioRxiv (2019-11-13) https://doi.org/gg94bn

DOI: <u>10.1101/840280</u>

41. Dendritic spine geometry and spine apparatus organization govern the spatiotemporal dynamics of calcium

Miriam Bell, Tom Bartol, Terrence Sejnowski, Padmini Rangamani

bioRxiv (2019-05-29) https://doi.org/gg9399

DOI: 10.1101/386367

Supplemental Tables

Table 2: Top five and bottom five systems biology preprints projected onto the PC2 direction. These preprints contain bioinformatis and neuroscience concepts respectively.

Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
and their Associated Gene Sets [32]	3.5865 702659 438883	CC- BY- ND	https://www.biorxiv.org/content/bi

Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
			orxiv/early/2019/09/25/781328/F1.large.jpg
QTG-Finder: a machine-learning based algorithm to prioritize causal genes of quantitative trait loci [33]	3.4703 883830 23157	No ne	h t p s : / w w

Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
			W.biorxiv.org/content/biorxiv/early/2019/04/29/484204

Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
			/ F 1
Identification of candidate genes underlying nodulation-specific phenotypes in Medicago trunco through integration of genome-wide association studies and co-expression networks [34]	otula 3.3814 906334 073953	CC- BY- NC- ND	https://www.biorxiv.org/content/biorxiv

Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
			/ e a r l y / 2 0 1 8 / 0 8 / 1 6 / 3 9 2 7 7 9 / F 1 . l a r g e . j p g
Raw sequence to target gene prediction: An integrated inference pipeline for ChIP-seq and RNA-seq datasets [35]	3.3632 576028 389742	No ne	h t t p s : // w w w b i

x i v v	Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
				r x i v . o r g / c o n t e n t / b i o r x i v / e a r l y / 2 0 1 7 / 1 1 / 1 6 / 2 2 0

Title [citation]	PC_2	Lic ens e	
			; ; ; ;
he y-ome defines the thirty-four percent of Escherichia coli genes that lack experimental evidence if function [36]	3.2874 278664 1417	CC- BY	

Title [citation]	PC_2	Lic ens e	L i n k
			<pre>y/2018/12/03/328591/F1.large.jpg</pre>
The effects of time-varying temperature on delays in genetic networks [37]	-2.7047 102478 958056	No ne	h t t p s : / / w w . b i o r x i v .

Title [citation]	PC_2	Lic ens e

Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
			p g
An analog to digital converter creates nuclear localization pulses in yeast calcium signaling [38]	-2.7757 450002 60895	No ne	https://www.biorxiv.org/content/biorxiv/early/201

Title [citation]	PC_2	Lic ens e	F i g u r e L i n k
			8 / 0 6 / 2 8 / 3 5 7 9 3 9 / F 1 . l a r g e . j p g
Nicotinic modulation of hierarchal inhibitory control over prefrontal cortex resting state dynamics: modeling of genetic modification and schizophreniarelated pathology [39]	-3.0473 423827 98414	No ne	https://www.biorxiv.org/c

Title [citation]	PC_2	Lic ens e	e L i n
			ontent/biorxiv/early/2018/04/13/301051/F1.large.jpg
Electrical propagation of vasodilatory signals in capillary networks [40]	-3.1077	CC-	h

	155787 93087	BY- NC- ND	F i g u
Title [citation]	PC_2	Lic ens e	
			t t p s
			/ / w w
			b i o r x i
			v o r g /
			c o n t e n
			t / b i o r
			x i v / e a
			r I y / 2
			0 1 9 / 1

Title [citation]	PC_2	Lic ens e	L i
Dendritic spine geometry and spine apparatus organization govern the spatiotemporal dynamic calcium [41]	s of -3.2153 349907 2831	CC- BY- NC- ND	

Title [citation]	PC_2	Lic ens e	L i n k
			t / b i o r x i v / e a r r
			I y / 2 0 1 9 / 0 5 /
			2 9 / 3 8 6 3 6 7 / F 1
			. l a r g e . j p g