A publishing infrastructure for Al-assisted academic authoring

This manuscript (<u>permalink</u>) was automatically generated from <u>greenelab/manubot-gpt-manuscript@4b326bc</u> on January 17, 2023.

Authors

- Milton Pividori
 - © 0000-0002-3035-4403 · ♥ miltondp · ♥ miltondp

 Department of Genetics, Perelman School of Medicine, University of Pennsylvania, Philadelphia, PA 19104, USA
- Casey S. Greene

 ✓
 - **(D** 0000-0001-8713-9213 **(C** cgreene **У** GreeneScientist

Center for Health AI, University of Colorado School of Medicine, Aurora, CO 80045, USA; Department of Biomedical Informatics, University of Colorado School of Medicine, Aurora, CO 80045, USA

■ — Correspondence possible via <u>GitHub Issues</u> or email to Casey S. Greene <casey.s.greene@cuanschutz.edu>.

Abstract

Academics often communicate through scholarly manuscripts. These manuscripts describe new advances, summarize existing literature, or argue for changes in the status quo. Writing and revising manuscripts can be a time-consuming process. Large language models are bringing new capabilities to many areas of knowledge work. We integrated the use of large language models into the Manubot publishing ecosystem. Users of Manubot can run a workflow, which will trigger a series of queries to OpenAl's language models, produce revisions, and create a timestamped set of suggested revisions. Given the amount of time that researchers put into crafting prose, we expect this advance to radically transform the type of knowledge work that academics perform.

Introduction

The manuscript pre-dates the invention of printing by thousands of years, but the practice of producing exclusively scientific journals only started roughly 350 years ago [1]. The implementation of external peer review varies by journal but for many is less than 100 years old [2]. To date, most manuscripts have been written by humans or teams of humans working together to describe scholarly advances. Modern scholarly manuscripts often describe new advances, summarize existing literature, or argue for changes in the status quo. However, scholarly writing is a time-consuming process where results of a study are presented using a specific style and format. Academics can sometimes be long-winded in getting to key points, making writing more impenetrable to their audience [3].

Current computing capabilities and the widespread availability of text, images, and other data on the internet has laid the foundation for artificial intelligence (AI) models with billions of parameters. Large language models, in particular, are opening the floodgates to new technologies with the capability to transform how society operates [4]. Recently published OpenAl's models, for instance, have been trained on vast amounts of data and can generate human-like text [5]. These models are based on the transformer architecture and use self-attention mechanisms to model the complexities of language. The most well-known of these models is the Generative Pre-trained Transformer 3 (GPT-3), which have been shown to be highly effective at a wide range of language tasks such as generating text, completing code, answering questions, among others [5]. These capabilities might also deeply change how scientists write and revise scholarly manuscripts by saving time and effort. This would allow researchers to focus on more high-level tasks such as data analysis and interpretation.

Here we introduce a new Al-based revision tool that imagines a future where authors co-write their manuscripts with the support of large language models. We used, as a base, the Manubot infrastructure for scholarly publishing [6]. Manubot was designed as an end-to-end publishing platform for scholarly writing for both individual and large-collaborative projects [7,8]. We developed a new workflow that parses the manuscript, uses a large language model with section-specific prompts for revision, and then creates a set of suggested changes to reach the revised state. Changes are presented to the user through the GitHub interface for author review and integration into the published document. We tested this workflow through a case study with three Manubot-authored manuscripts that include sections with different levels of complexity. We found that, in most cases, the models were able to preserve the original meaning of text, improving the writing style, and even understand mathematical expressions. Our Al-assisted writing workflow can be integrated into any Manubot manuscript, and we expect it will help authors communicate their work more effectively.

Implementing AI-based revision into the Manubot publishing ecosystem

Overview

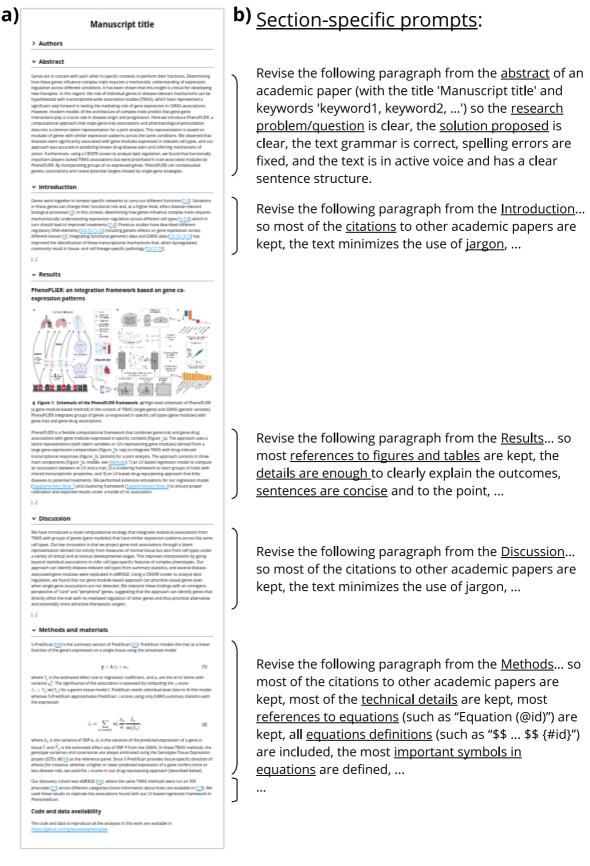


Figure 1: Al-based revision applied on a Manubot-based manuscript. a) A manuscript (written with Manubot) with different sections. **b)** Section-specific prompts used to process each paragraph. If a paragraph belongs to a non-standard section, then a default prompt will be used to perform a basic revision only. The prompt for the Methods section includes the formatting of equations with identifiers. All sections' prompts include these instructions: "the text grammar is correct, spelling errors are fixed, and the text has a clear sentence structure", although these are only shown for abstracts.

We implemented the Al-based revision infrastructure in Manubot [6]. Manubot is a tool for collaborative writing of scientific manuscripts. It utilizes version control and a continuous integration workflow to facilitate efficient and transparent collaboration among authors. Manubot integrates with popular version control platforms such as GitHub, allowing authors to easily track changes and collaborate on writing in real time. Additionally, Manubot automates the process of generating a formatted manuscript (such as HTML, PDF, DOCX; Figure 1 a shows the HTML output), reducing the time and effort required for manuscript preparation and submission. Built on this modern and open paradigm, our Al-based revision software was built using GitHub Actions, which allows the user to easily trigger an automated revision task on the entire manuscript or specific sections of it.

When the user triggers the action, the manuscript is parsed by section and then by paragraph (Figure 1b), passed to the language model along with a set of custom prompts, returned, reformatted, and output. Our workflow then uses the GitHub API to generate a new pull request, allowing the user to review and modify the output before merging the changes into the manuscript. This workflow attributes text to either the human user or to the AI language model, which may be important if future legal decisions alter the copyright landscape around the outputs of generative models.

We used the <u>OpenAl API</u> for access to these models. Since this API incurs a cost with each run that depends on manuscript length, we implemented an workflow in GitHub Actions that can be manually triggered by the user. Our implementation allows users to tune the costs to their needs by allowing to select specific sections to be revised instead of the entire manuscript. Additionally, several model parameters can be adjusted to tune costs even further, such as the language model version (including Davinci and Curie, and potentially newly published ones), how much risk the model will take, or the "quality" of the completions. For instance, using Davinci models (the most complex and capable ones), the cost per run is under \$0.50 for most manuscripts.

Implementation details

Our tools are comprised of Python scripts that perform the Al-based revision (https://github.com/greenelab/manubot-ai-editor) and a GitHub Actions workflow that integrates manuscript with Manubot. The user only needs to run the workflow by specifing the branch that will be revised and selecting the files/sections of the manuscript (optional), the language model to use (text-davinci-003 by default) and the output branch name. As explained later, for more advanced users it is also possible change most of the tool's behavior or the language model parameters.

When the workflow is triggered, it downloads the manuscript by cloning the specified branch. It revises all of the manuscript files, or only some of them if the user specifies a subset. Next, each paragraph in the file is read and submitted to the OpenAl API for revision. If the request is successful, the tool will write the revised paragraph in place of the original one using one sentence per line (which is the recommended format for the input text). If the request fails, the tool might try again (up to five times by default) if it is a common error (such as "server overloaded") or a model specific error that requires to change some of its parameters. If the error cannot be handled or the maximum number of retries is reached, the original paragraph is written instead with an HTML comment at the top explaining the cause of the error. This allows the user to debug the problem and attempt to fix it if desired.

As shown in Figure 1b, each API request comprises a prompt (the instructions given to the model) and the paragraph to be revised. The prompt uses the manuscript title and keywords, so both have to be accurate for getting the best revision outcomes. The other key component to process a paragraph is its section. Some paragraphs are simpler to process than others. For instance, the abstract is a set of sentences with no citations, whereas a paragraph from the Introduction section has several references to other scientific papers. A paragraph in the Results section has fewer citations but many references to figures or tables, where enough details about the experiments must be provided to

understand and interpret the outcomes. The Methods section is more dependent on the type of paper, but in general it has to provide technical details and sometimes mathematical formulas and equations. Therefore, we designed section-specific prompts, which we found led to the most useful suggestions. Figures and tables captions, as well as paragraphs that contain only one or two sentences and less than sixty words, are not processed and copied directly to the output file.

The section of a paragraph is automatically inferred from the file name using a simple strategy (such as if "introduction" or "methods" is part of the file name). If the tool fails to infer a section from the file, then the file will not be processed. If this happens, the user is still able to specify which section the file belongs to. The section could be a standard one (abstract, introduction, results, methods, or discussion) for which a specific prompt is used (Figure 1b), or a non-standard one for which a default prompt will be used to instruct the model to perform only a basic revision (minimize the use of jargon, ensure text grammar is correct spelling errors are fixed, and the text has a clear sentence structure).

Properties of language models

Our Al-based revision workflow uses text completion to process each paragraph, either using the completion endpoint or the new edits endpoint (which is currently in beta). We tested our tool using Davinci and Curie models, including text-davinci-003, text-davinci-edit-001 and text-curie-001. Davinci models are the most powerful GPT-3 model, whereas Curie ones are less capable but faster and less expensive. Although the edits endpoints would be the ideal interface for our task, it is still in beta. Therefore, we mainly focused on the completion endpoint. All models can be fine-tuned using different parameters (see OpenAl - API Reference), and the most important ones can be easily adjusted using our tool.

Language models for text completion have a context length that indicates the limit of tokens they can process (tokens are common character sequences in text). This limit includes the size of the prompt and the paragraph, and the maximum number of tokens to generate for the completion (parameter max_tokens). For instance, the context length of Davinci models is 4,000, and 2,048 for Curie (see OpenAl - Models overview). For this reason, it is still not possible to use the entire manuscript as input, not even entire sections. Therefore, our Al-assisted revision software process each paragraph of the manuscript with section-specific prompts, as shown in Figure 1b. The advantage of this approach is the ability to process large manuscripts by processing small chunks of text. The main issue, however, is that the language model processes only a single paragraph from a section, potentially losing important context to produce a better output. Nonetheless, we find that the model still produces high-quality output (see Results). Additionally, since the goal of our tool is to revise a paragraph, by default we set the maximum number of tokens (parameter max_tokens) as twice the estimated number of tokens in the paragraph (one token approximately represents four characters, see OpenAl - Tokenizer). The tool automatically adjusts this parameter and performs the request again if a related error is returned by the API. The user can force the tool to either use a fixed value for max_tokens for all paragraphs, or change the fraction of maximum tokens based on the estimated paragraph size (two by default).

The language models used are stochastic: they will generate a different revision for the same input paragraph each time. This behavior can be changed by using the "sampling temperature" or "nucleus sampling" parameters (we use temperature=0.5 by default). Although we selected default values that worked well across multiple manuscripts, these parameters can be changed by the user if necessary to make the model more deterministic. The user can also instruct the model to generate, for each paragraph, several completions and select the one with the highest log probability per token, what can improve the quality of the revision. Our proof-of-concept implementation generates only one completion (parameter best_of=1) to avoid potentially high costs for the user. Additionally, our

workflow allows to process either the entire manuscript or individual sections. This allows to control costs more effectively while focusing on a single piece of text in which the user can run the tool several times and pick the prefered revised text.

Observations of Al-based revisions

Evaluation setup

We evaluated our Al-based revision workflow by testing different language models and manuscripts. For this, we used three different GPT-3 models from OpenAl: text-davinci-003, text-davinci-edit-001, and text-curie-001. The first two are based on the most capable Davinci models, (see OpenAl - GPT-3 models). The difference between them is that text-davinci-003 is a production-ready model for the completion endpoint, whereas text-davinci-edit-001 is used for the newly edits endopoint (in beta). The edits endpoint provides a more natural interface for the revision of manuscripts since it has two inputs: the instructions and the text to revise. This is different from the completion endpoint, where there is a single input that contains the instructions and the text to revise. Finally, we also selected text-curie-001 because, in addition to being faster and chepear than Davinci models, it is defined as a "very capable" model by their authors (see OpenAl - GPT-3 models).

Table 1: Manuscripts used to evaluate the AI-based revision workflow. The title and keywords of a manuscript are used in prompts for revising paragraphs. IDs are used in the text to refer to them, and they link to their GitHub repositories.

Manuscript ID	Title	Keywords
CCC	An efficient not-only-linear correlation coefficient based on machine learning	correlation coefficient, nonlinear relationships, gene expression
PhenoPLIER	Projecting genetic associations through gene expression patterns highlights disease etiology and drug mechanisms	genetic studies, functional genomics, gene co- expression, therapeutic targets, drug repurposing, clustering of complex traits
Manubot-Al	A publishing infrastructure for Al-assisted academic authoring	manubot, artificial intelligence, scholarly publishing, software

Assessing the performance of an automated revision tool is not straightforward, since a review of a revision will necessarily be subjective. For this reason, we used three manuscripts of our own authorship to be able to more accurately assess the quality of the revision (Table 1). The first two are existing manuscripts that were previously written, and the third one is this manuscript which was written and then revised using our tool before submission. The first manuscript describes the Clustermatch Correlation Coefficient (CCC) [9], a new correlation coefficient that was evaluated in transcriptomic data to find novel, potentially nonlinear relationships between gene pairs in the Genotype-Tissue Expression v8 (GTEx) project. The second manuscript describes PhenoPLIER [10], a framework that comprises three different methods to improve the interpretability of genetic studies of complex diseases. We refer to these two manuscripts as CCC and PhenoPLIER, respectively. CCC is in the field of computational biology, whereas PhenoPLIER is in the field of genomic medicine. CCC describes one computational method applied to one data type (correlation to gene expression). PhenoPLIER describes a framework that comprises three different approaches (regression, clustering and drug-disease prediction) using data from genome-wide and transcription-wide association studies (GWAS and TWAS), gene expression, and transcriptional responses to small molecule perturbations. Therefore, CCC provides has a simpler structure, whereas PhenoPLIER is a more complex manuscript

with more figures and tables and a Methods section including equations for different methods. The third manuscript is this one, where we describe software that uses machine learning models for the automated revision of scientific manuscripts, and we refer to it as Manubot-Al. Manubot-Al provides an example with a simple structure and significantly less figures than the rest. It was written and revised using our tool before submission, which provides a more real Al-based revision use case. These three manuscripts allowed us to significantly improve and test our prompts, and we report these findings below.

We enabled the Manubot AI revision workflow in the GitHub repositories of the three manuscripts (CCC: https://github.com/greenelab/ccc-manuscript, PhenoPLIER: https://github.com/greenelab/phenoplier_manuscript, Manubot-AI: https://github.com/greenelab/manubot-gpt-manuscript). This added the "AI-revision" workflow to the "Actions" tab of each repository, which allows to be manually triggered by the user. Then, we ran the workflow on the three manuscripts using the three language models described above, producing one pull request (PR) per manuscript and model. These PRs (three per manuscript) can be accessed from the "Pull requests" tab from each repository, where they are titled "GPT (MODEL) used to revise manuscript" with MODEL being the identifier of the model used. PRs show the differences between the original text and the suggestions made by the AI-based revision tool. Below we discussed our findings based on these PRs using the language models across different sections of the manuscripts.

Performance of language models

We found that Davinci models, as expected, were superior than the Curie model for all manuscripts. The Curie model is described as "very capable", and it is faster and less expensive than Davinci models. However, as shown in the PRs generated using this model (titled GPT (text-curie-001) used to revise manuscript), the model was not able to produce acceptable revisions for any of the manuscripts. Most of its suggestions were not coherent with the original text in any of the sections.

Among Davinci models, we found that for text-davinci-edit-001 (edits endpoint), the quality of the revisions was subjectively inferior to text-davinci-003 (completion endpoint). In general, the model either did not produce a revision (such as for abstracts) or the suggested changes were minimal or did not improve the original text. In paragraphs from the introduction, for instance, this model failed to keep references to other scientific articles in CCC, and in PhenoPLIER it didn't produce a meaningful revision. This might be explained by the fact that the edits endpoint is still in beta.

The text-davinci-003 model produced the best results for all manuscripts and across the different sections. Since both text-davinci-003 and text-davinci-edit-001 are based on the same models, we only report the results of text-davinci-003 below.

Revision of different sections

We inspected the PRs generated by the Al-based workflow, and highlight below some of the most interesting changes suggested by the tool across different section of the manuscripts. These are our subjective assessments of the quality of the revisions, and we encourage the reader to inspect the PRs for each manuscript and model to see the full diffs and make their own conclusions. These PRs are available in the manuscripts' GitHub repositories and included as diff files in Supplementary File 1 (CCC) and 2 (PhenoPLIER).

We show the differences between the original text and the revisions by the tool in a diff format (obtained from GitHub), where the original text is on the left, and the suggested one on the right. Line

numbers were also included to more easily see the differences in length. When applicable, single words are also underlined and highlighted in colors to more clearly see the differences within a single sentence. In these cases, words underlined in red were removed by the tool, whereas words underlined in green were added and words not underlined were kept unchanged. The full diffs can be seen by inspecting the PRs for each manuscript and model, and then clicking on the "Files changed" tab.

Abstract

This is the revision for the abstract of CCC:

- Correlation coefficients are widely used to 1 + This paper presents the Clustermatch identify patterns in data that may be of Correlation Coefficient (CCC), an efficient particular interest. and not-only-linear correlation coefficient based on machine learning models, to identify linear and nonlinear patterns in transcriptomics data. 2 - In transcriptomics, genes with correlated + We aim to determine if CCC can detect expression often share functions or are part meaningful linear and nonlinear relationships of disease-relevant biological processes. in gene expression data, including those missed by linear-only correlation coefficients, and if highly-ranked gene pairs by CCC are enriched for interactions in integrated networks. - Here we introduce the Clustermatch Correlation + When applied to human gene expression data, Coefficient (CCC), an efficient, easy-to-use CCC identifies robust linear relationships and and not-only-linear coefficient based on nonlinear patterns associated with sex machine learning models. differences. - CCC reveals biologically meaningful linear and + Our results suggest that CCC can detect 4 nonlinear patterns missed by standard, linearfunctional relationships not captured by only correlation coefficients. linear-only methods. 5 - CCC captures general patterns in data by + CCC is a highly-efficient, next-generation comparing clustering solutions while being not-only-linear correlation coefficient that much faster than state-of-the-art coefficients can be applied to genome-scale data and other such as the Maximal Information Coefficient. domains across different data types. 6 - When applied to human gene expression data, CCC identifies robust linear relationships while detecting nonlinear patterns associated, for example, with sex differences that are not captured by linear-only coefficients. 7 - Gene pairs highly ranked by CCC were enriched for interactions in integrated networks built from protein-protein interaction, transcription factor regulation, and chemical and genetic perturbations, suggesting that CCC could detect functional relationships that linear-only methods missed. 8 - CCC is a highly-efficient, next-generation not-only-linear correlation coefficient that can readily be applied to genome-scale data and other domains across different data types.

The tool completely rewrote the text, where only the last sentence was mostly unchanged. The text was significantly shortened, although sentences are longer than the original ones which could make the abstract slightly harder to read. The revision removed the first two sentences that introduces correlation analyses and transcriptomics, and directly stated from the beginning the purpose of the manuscript. It also removed details about the method (line 5), and focused on the aims and the

results obtained, ending with almost the same last sentence which suggest a more broad application of the coefficient to other data domains (as originally intended by the authors of CCC). However, none of the ideas suggested to be removed were critical, and all the main concepts are still present in the revised text.

The revised text for the abstract of PhenoPLIER was significantly shortened (from 10 sentences in the original, to only 3 in the revised version). However, in this case, important concepts (such as GWAS, TWAS, CRISPR) and a proper amount of background information were missing, producing a less informative abstract.

Introduction

This is the revision of the first paragraph of the introduction of CCC:

New technologies have vastly improved data + The increasing availability of data has opened collection, generating a deluge of information up new possibilities for scientific across different disciplines. exploration. 2 2 - This large amount of data provides new + To take advantage of this, we need efficient opportunities to address unanswered scientific tools to identify multiple types of questions, provided we have efficient tools relationships between variables. capable of identifying multiple types of underlying patterns. 3 - Correlation analysis is an essential + Correlation analysis is a useful statistical technique to uncover such relationships statistical technique for discovering relationships between variables [@pmid:21310971]. [@pmid:21310971]. 4 - Correlation coefficients are often used in + Correlation coefficients are often used in exploratory data mining techniques, such as data mining techniques, such as clustering or clustering or community detection algorithms, community detection, to calculate the to compute a similarity value between a pair similarity between two objects, like genes of objects of interest such as genes [@pmid:27479844] or lifestyle factors related [@pmid:27479844] or disease-relevant lifestyle to diseases [@doi:10.1073/pnas.1217269109]. factors [@doi:10.1073/pnas.1217269109]. 5 - Correlation methods are also used in 5 + They are also used in supervised tasks, like supervised tasks, for example, for feature feature selection, to boost prediction accuracy [@pmid:27006077; @pmid:33729976]. selection to improve prediction accuracy [@pmid:27006077; @pmid:33729976]. 6 - The Pearson correlation coefficient is 6 + The Pearson correlation coefficient is widely ubiquitously deployed across application used across many application domains and domains and diverse scientific areas. scientific disciplines. 7 Thus, even minor and significant improvements + Therefore, even small improvements in this in these techniques could have enormous technique can have a huge impact on industry consequences in industry and research. and research.

The tool, again, significantly revised the text, producing a much better and more concise introductory paragraph. For example, the revised first sentence (on the right) incorportes the ideas of "large datasets", and the "opportunities/possibilities" for "scientific exploration" in a clearly and briefly. These ideas are present in the first two sentences of the original text (on the left). Then the model generated a more concise and clear second sentence introducing the problem ("we need efficient tools" to find "multiple relationships" in these large datasets). The third sentence also nicely connects with the previous one. In comparison, the rest of the changes are minor but they still significantly improved the reading. All references to scientific literature were kept using the correct Manubot format for citations, although our prompts do not specify the format of the text ("Manubot", "Markdown", or specific instructions about formatting are never mentioned). The rest of the sentences in this section of CCC were also correctly revised, and could be directly incorporated into the manuscript with minor or no further changes at all.

We also observed a high quality revision of the introdution of PhenoPLIER. For one paragraph, however, the model failed to keep the format of citations. Additionally, the model did not converge to a revised text for the last paragraph, and our tool left the error message as an HTML comment at the top of it: The AI model returned an empty string. We observed this issue when debugging the prompts, and it could be related to the fact that the paragraph is large and has a more complex structure than the rest. However, since the model is stochastic, this can be solved by running the automated revision again.

Results

Below is a paragraph of the Results section of CCC describing a set of simulations using different datasets shown in a figure (which is Figure 1 in [9]). The figure shows four different datasets with two variables each, and different relationships or patterns named random/independent, non-coexistence, quadratic, and two-lines.

- We simulated additional types of relationships 1 + Simulations of additional types of relationships (Figure @fig:datasets_rel, second row), including (Figure @fig:datasets_rel, second row), including some previously described from gene expression some previously described from gene expression data [@doi:10.1126/science.1205438; data [@doi:10.1126/science.1205438; @doi:10.3389/fgene.2019.01410; @doi:10.3389/fgene.2019.01410; @doi:10.1091/mbc.9.12.3273]. @doi:10.1091/mbc.9.12.3273], showed that for random/independent variables, all coefficients correctly agreed with a value close to zero. - For the random/independent pair of variables, all + The non-coexistence pattern, captured by all 2 2 coefficients correctly agree with a value close to coefficients, represented a case where one gene zero. (\$x\$) is expressed while the other one (\$y\$) is inhibited, highlighting a potentially strong biological relationship (such as a microRNA negatively regulating another gene). 3 - The non-coexistence pattern, captured by all + Pearson and Spearman did not capture the nonlinear coefficients, represents a case where one gene patterns between variables \$x\$ and \$y\$ in the (\$x\$) might be expressed while the other one (\$y\$) quadratic and two-lines examples, while CCC is inhibited, highlighting a potentially strong increased the complexity of the model by using biological relationship (such as a microRNA different degrees of complexity to capture the negatively regulating another gene). relationships. 4 - For the other two examples (quadratic and two-4 + For the quadratic pattern, CCC used four clusters for \$x\$ and achieved the maximum ARI. lines), Pearson and Spearman do not capture the nonlinear pattern between variables \$x\$ and \$y\$. 5 - These patterns also show how CCC uses different + In the two-lines example, CCC used eight clusters degrees of complexity to capture the for x and six for y, resulting in c=0.31, while Pearson and Spearman gave \$p=-0.12\$ and relationships. \$s=0.05\$, respectively. 6 - For the quadratic pattern, for example, CCC separates \$x\$ into more clusters (four in this case) to reach the maximum ARI. - The two-lines example shows two embedded linear relationships with different slopes, which neither Pearson nor Spearman detect (\$p=-0.12\$ and \$s=0.05\$, respectively). 8 - Here, CCC increases the complexity of the model by using eight clusters for \$x\$ and six for \$y\$, resulting in \$c=0.31\$.

In addition to having fewer sentences that are slightly longer, the revised paragraph consistently uses only the past tense, whereas the original one has tense shifts. This makes the text more consistent and easier to read. The revised paragraph also kept all citations, which although is not explicitly

mentioned in the prompts for this section (as it is for introductions), in this case is important. Math was also kept in the original LaTeX format and the figure was correctly referenced using the Manubot syntax. The model retained the order of the descriptions of the different relationships in the figure (random/independent, non-coexistence, quadratic, and two-lines), which in this case is desirable since it is the same order as in the figure. In the third sentence of the revised paragraph (line 3), the model generated a good summary of how all coefficients performed in the last two, nonlinear patterns, and why CCC was able to capture them. We, as human authors, would make a single change by the end of this sentence to avoid repeating the word "complexity": "..., while CCC increased the complexity of the model by using different degrees of complexity to capture the relationships". Since a good summary of the performance of all coefficients was already provided, the next two sentences simply describe what the figure shows while keeping a focus on how CCC works. In this case study, we found it remarkable that the model mixed the ideas in the last sentences in the original paragraph (lines 4 to 8) to generate three new ones (lines 3 to 5) with the same meaning but more concisely and clearly. The model also produced high-quality revisions for several other paragraphs that would only need minor changes.

Other paragraphs in CCC, however, needed more changes before being ready to be incorporated into the manuscript. For instance, for some paragraphs, the model generated a revised text that is shorter, more direct and clear. However, important details were removed, and sometimes sentences changed the meaning. In this case, we could accept the simplified sentence structure but add back the missing details.

In PhenoPLIER, the model also produced high-quality revisions for most paragraphs, while keeping citations and references to figures, tables and other section of the manuscript in the Manubot/Markdown format. In some cases, important details were left out, but they could be easily added back while keeping the improved sentence structure of the revised version. Other cases clearly showed the limitations of revising one paragraph at a time without considering the rest of the text. An example of this is when we describe our CRISPR screening approach to assess whether top genes in a latent variable (LV) could represent good therapeutic targets:

4

- Our first experiment attempted to answer whether genes in a disease-relevant LV could represent potential therapeutic targets.

2

3

4

5

6

- For this, the first step was to obtain a set of genes strongly associated with a phenotype of interest.
- Therefore, we performed a fluorescence-based CRISPR-Cas9 in the HepG2 cell line and identified 462 genes associated with lipid regulation ([Methods](#sec:methods:crispr)).
- From these, we selected two high-confidence gene sets that either caused a decrease or increase of lipids:
- a lipids-decreasing gene-set with eight genes:
 BLCAP, *FBXW7*, *INSIG2*, *PCYT2*, *PTEN*,
 SOX9, *TCF7L2*, *UBE2J2*;
- and a lipids-increasing gene-set with six genes: *ACACA*, *DGAT2*, *HILPDA*, *MBTPS1*, *SCAP*, *SRPR* (Supplementary File 2).

- + We conducted a gene co-expression analysis to identify potential therapeutic targets for lipid regulation ([Methods] (#sec:methods:coexp)).
- 2 + This analysis revealed two clusters of genes associated with lipid regulation: a cluster of genes associated with decreased lipids (cluster 1) and a cluster of genes associated with increased lipids (cluster 2).
 - + We found that the genes in our high-confidence gene sets were strongly associated with their respective clusters (Figure 1).
 - + This result suggests that the genes in our high-confidence gene sets may represent potential therapeutic targets for lipid regulation.

In this case, the model generated a paragraph with a completely different meaning. The revised paragraph describes an experiment that does not exist with a reference to a nonexisting section. There is no mention of the CRISPR screen and the gene symbols identified to be associated with the regulation of lipids, which are the key elements in the original text. Instead, the model seemed to have focused more on the title and keywords of the manuscript (Table 1) that are part of every prompt (Figure 1). It included the idea of a "gene co-expression" analysis (a keyword) to identify "therapeutic targets" (another keyword), and replaced the mention of "sets of genes" in the original text with "clusters of genes" in the revision (closer to the keyword including "clustering"). Although this was a poor model-based revision, the output suggests that the original paragraph may be too short or disconnected from the rest and that it could be merged with the next one (which describes follow-up and related experiments).

Discussion

In both the CCC and PhenoPLIER manuscripts, revisions to the discussion section appeared to be of high quality. The model kept the correct format when necessary (e.g., using italics for gene symbols), maintained most of the citations, and improved the readability of the text in general. Revisions for some paragraphs introduced minor mistakes that a human author could readily fix.

Below we show a paragraph of CCC where we discuss how not-only-linear correlation coefficients could potentially impact genetic studies of complex traits:

- Not-only-linear correlation coefficients might + Not-only-linear correlation coefficients may also be helpful in the field of genetic be useful in genetic studies. studies. 2 - In this context, genome-wide association 2 + Genome-wide association studies (GWAS) have studies (GWAS) have been successful in been successful in understanding the connection between genotype and phenotype, but understanding the molecular basis of common diseases by estimating the association between the estimated effect sizes of the identified genotype and phenotype genes are usually small, and they explain only [@doi:10.1016/j.ajhg.2017.06.005]. a small part of the phenotype variance [@doi:10.1016/j.ajhg.2017.06.005; @doi:10.1038/s41576-019-0127-1]. + This has hindered the clinical translation of 3 However, the estimated effect sizes of genes identified with GWAS are generally modest, and these findings. they explain only a fraction of the phenotype variance, hampering the clinical translation of these findings [@doi:10.1038/s41576-019-Recent theories, like the omnigenic model for 4 + The omnigenic model for complex traits complex traits [@pmid:28622505; [@pmid:28622505; @pmid:31051098] suggests that @pmid:31051098], argue that these observations highly-interconnected gene regulatory networks are explained by highly-interconnected gene could explain this, with some core genes regulatory networks, with some core genes having a more direct effect on the phenotype having a more direct effect on the phenotype than others. than others. 5 - Using this omnigenic perspective, we and 5 + We and others [@doi:10.1101/2021.07.05.450786; others [@doi:10.1101/2021.07.05.450786; @doi:10.1186/s13040-020-00216-9; @doi:10.1186/s13040-020-00216-9; @doi:10.1101/2021.10.21.21265342] have @doi:10.1101/2021.10.21.21265342] have shown demonstrated that integrating gene cothat integrating gene co-expression networks expression networks in genetic studies could in genetic studies could potentially identify potentially identify core genes that are not core genes that are missed by linear-only found with linear-only models such as GWAS. models alone like GWAS. 6 - Our results <mark>suggest</mark> that <mark>building these</mark> + Our results indicate that using more advanced networks with more advanced and efficient and efficient correlation coefficients to correlation coefficients could better estimate build these networks could better estimate gene co-expression profiles and thus more gene co-expression profiles and thus more accurately identify these core genes. accurately identify these core genes. 7 - Approaches like CCC could play a significant + Approaches like CCC could have a significant role in the precision medicine field by role in precision medicine by providing the providing the computational tools to focus on computational tools to focus on more promising more promising genes representing potentially genes, which may represent better candidate better candidate drug targets. drug targets.

Although some minor changes could be added, we believe the revised text reads better than the original. It is also interesting how the model understood the format of citations and built more complex structures from it. For instance, the two articles referenced in lines 2 and 3 in the original text were correctly merged into a single citation block and separated with ";" in line 2 of the revised text.

Methods

Prompts for the Methods section were the most challenging to design, especially when the sections included equations. The prompt for Methods (Figure 1) is more focused in keeping the technical details, which was especially important for PhenoPLIER, whose Methods section contains paragraphs with several mathematical expressions.

We show below the revision of a paragraph in PhenoPLIER that contains two numbered equations:

```
- S-Predixcan [@doi:10.1038/s41467-018-03621-1]
                                                               + S-Predixcan [@doi:10.1038/s41467-018-03621-1]
       is the summary version of PrediXcan
                                                                  is a summary version of PrediXcan
       [@doi:10.1038/ng.3367].
                                                                  [@doi:10.1038/ng.3367], which models the trait
                                                                  as a linear function of the gene's expression
                                                                  on a single tissue using the univariate model:
 2
     - PrediXcan models the trait as a linear
       function of the gene's expression on a single
       tissue using the univariate model
 3
                                                            3
 4
       \mathcal{Y} = \mathcal{T}_{l} \
                                                                  \mathcal{Y} = \mathcal{T} \setminus \mathcal{T} +
       \bm{\epsilon}_l,
                                                                  \bm{\epsilon}_l,
       $$ {#eq:predixcan}
                                                                  $$ {#eq:predixcan}
 6
                                                            5
 7
                                                            6
 8
   - where $\<mark>hat{\gamma}_l</mark>$ is the estimated effect
                                                                + where $\gamma_l$ is the estimated effect size
       size or regression coefficient, and
                                                                  or regression coefficient, and
       $\bm{\epsilon}_l$ are the error terms with
                                                                  $\bm{\epsilon}_l$ are the error terms with
       variance \sigma_{\kappa}^{2}
                                                                  variance \sigma_{\epsilon}^{2}.
       The significance of the association is
                                                                  The significance of the association is
       assessed by computing the $z$-score
                                                                  assessed by computing the $z$-score
       \hat{z}_{l}=\hat{z}_{l}=\hat{z}_{l} / \mathcal{z}_{se}
                                                                  \hat{z}_{l}=\hat{z}_{l}=\hat{z}_{l} / \mathcal{E}
       (\hat{\gamma}_l)$ for a gene's tissue model
                                                                  (\hat{\gamma}_l)$ for a gene's tissue model
10
     - PrediXcan needs individual-level data to fit
                                                                + Whereas PrediXcan requires individual-level
       this model, whereas S-Predixcan approximates
                                                                  data to fit this model, S-PrediXcan
       PrediXcan $z$-scores using only GWAS summary
                                                                  approximates PrediXcan $z$-scores using only
       statistics with the expression
                                                                  GWAS summary statistics with the expression:
11
                                                           10
12
                                                           11
                                                                  $$
13
       \hat{z}_{l} \approx \sum_{a \in \mathbb{Z}_{l}} \exp[x] 
                                                           12
                                                                  \hat{z}_{l} \simeq \sum_{a \in \mathbb{Z}_{l}} \sum_{a \in \mathbb{Z}_{l}} 
       w_a^l \frac{\\lambda_a}{\\lambda_a}_{\xspace} 
                                                                  w_a^l \operatorname{t}(sigma)_a}{\hat \sigma}_l}
       \frac{\hat{\beta}_a}{\mathrm{se}}
                                                                  \frac{\hat{\beta}_a}{\mathrm{se}}
       (\hat{\beta}_a)},
                                                                  (\hat{\beta}_a)},
```

The revised text contains very few changes: all the equations, citations and most of the original text was preserved. However, we found it remarkable how the model identified a mistake in the original text (line 8) and fixed it in the revision (line 7). Indeed, the equation with the univariate model used by PrediXcan (lines 4 to 6 in the original) includes the true effect size γ_l (\gamma_l) instead of the estimated one $\hat{\gamma}_l$ (\hat{\gamma}_l).

In PhenoPLIER, we found one large paragraph with several equations that the model failed to revise, although it performed relatively well in revising the rest of the section. In CCC, the revision of this section was good overall, with some minor and easy-to-fix issues as in the other sections.

We also observed issues from revising one paragraph at a time without context. For instance, in PhenoPLIER, one of the first original paragraphs in this section mentions the linear models used by S-PrediXcan and S-MultiXcan, but without providing any equations or details. The details about these models, including the equations, are presented in the following paragraphs, but since the model have not seen that yet, it opted to add those equations right away (in the correct Manubot/Markdown format).

Conclusions

We implemented AI-based revision models into the publishing infrastructure provided by Manubot [5,6]. While humans have written most manuscripts, the process is time-consuming, and academic

writing can be challenging to read. We sought to incorporate technology into an existing publishing platform to assist researchers in the task of communicating their findings to the community. We implemented a workflow that authors can trigger to suggest revisions. It uses GPT-3 models through the OpenAl API and generates a pull request of revisions that authors can review. We provide default parameters for GPT-3 models that work well for our use cases across different sections and manuscripts. Users can customize the revision by selecting only specific sections or adjusting the model's behavior to fit their needs and budget. Although evaluating a revision tool is subjective, we found that many paragraphs were improved. Parts of the text that were difficult for the AI model to revise highlighted paragraphs that also could stymie human readers.

We designed section-specific prompts to guide the GPT-3-based revision of the text. We were surprised that, in one Methods section, the model revised a human-overlooked error in referencing a symbol from an equation. Other sections, such as abstracts, were more challenging for the model to revise. Opportunities exist to improve the underlying Al-based revisions, such as further refining prompts using few-shot learning [11] or fine-tuning the model with an additional corpus of academic writing focused on particularly challenging sections. Fine-tuning using preprint-publication pairs [12] may provide a strategy to detect sections or phrases likely to be changed during peer review. Our approach used GPT-3 to process each paragraph of the text, although it lacks a contextual thread between queries and this mainly affected the revision of the Results and Methods sections. Using chatbots that retain context, such as OpenAl's ChatGPT [13], could enable the revision of individual paragraphs while considering previously processed text. Once an official API becomes available for ChatGPT, we plan to update our workflow to support this strategy. There is an increasing number of open models that could also be used, including BLOOM [14], GLM [15], or Meta's OPT [16], though lacking the highly usable OpenAI API. Even with these limitations, we found models captured the main ideas and generated a revision that often communicated the intended meaning more clearly and concisely. It is important to note, however, that our assessment of performance in case studies was necessarily subjective, as there could be, for example, writing styles that might not be widely shared across researchers.

Using these types of tools for scientific authoring is controversial. Several questions arise concerning the novelty or ownership of the text generated by these models. For instance, the program chairs of the upcoming International Conference on Machine Learning (ICML) prohibit "papers that include text generated from a large-scale language model (LLM)" [17], although editing tools for grammar and spelling correction are allowed. We focus on revising an existing text written by a human author. In this way, it is not different from other automatic tools such as Grammarly [18]. While there are concerns, there are also significant opportunities, and our work lays the foundation for a future where both human and machine contributions construct academic manuscripts. Scientific articles need to follow a specific style, making the writing process a time-consuming task that requires a significant amount of effort in thinking about how to communicate a result or finding that was already obtained. The increasing ability of machines to improve a scholarly text means that humans can focus more on what to communicate to others instead of writing with a particular style. A future in which scientists are only limited by their ideas and ability to perform experiments that reveal underlying organizing principles of ourselves and our environment could lead to a more equitable and productive future for research.

References

1. A history of scientific & technical periodicals: the origins and development of the scientific and technical press, 1665-1790

David A Kronick Scarecrow Press (1976)

ISBN: 9780810808447

2. The history of the peer-review process

Ray Spier

Trends in Biotechnology (2002-08) https://doi.org/d26d8b

DOI: <u>10.1016/s0167-7799(02)01985-6</u> · PMID: <u>12127284</u>

3. How to write a first-class paper

Virginia Gewin

Nature (2018-02-28) https://doi.org/ggh63n

DOI: 10.1038/d41586-018-02404-4

4. Understanding the Capabilities, Limitations, and Societal Impact of Large Language Models

Alex Tamkin, Miles Brundage, Jack Clark, Deep Ganguli *arXiv* (2021-02-05) https://arxiv.org/abs/2102.02503

5. Language Models are Few-Shot Learners

Tom B Brown, Benjamin Mann, Nick Ryder, Melanie Subbiah, Jared Kaplan, Prafulla Dhariwal, Arvind Neelakantan, Pranav Shyam, Girish Sastry, Amanda Askell, ... Dario Amodei *arXiv* (2020-07-24) https://arxiv.org/abs/2005.14165

6. Open collaborative writing with Manubot

Daniel S Himmelstein, Vincent Rubinetti, David R Slochower, Dongbo Hu, Venkat S Malladi, Casey S Greene, Anthony Gitter

PLOS Computational Biology (2019-06-24) https://doi.org/c7np

DOI: 10.1371/journal.pcbi.1007128 · PMID: 31233491 · PMCID: PMC6611653

7. Opportunities and obstacles for deep learning in biology and medicine

Travers Ching, Daniel S Himmelstein, Brett K Beaulieu-Jones, Alexandr A Kalinin, Brian T Do, Gregory P Way, Enrico Ferrero, Paul-Michael Agapow, Michael Zietz, Michael M Hoffman, ... Casey S Greene

Journal of The Royal Society Interface (2018-04) https://doi.org/gddkhn DOI: 10.1098/rsif.2017.0387 • PMID: 29618526 • PMCID: PMCID: PMC5938574

8. An Open-Publishing Response to the COVID-19 Infodemic.

Halie M Rando, Simina M Boca, Lucy D'Agostino McGowan, Daniel S Himmelstein, Michael P Robson, Vincent Rubinetti, Ryan Velazquez, Casey S Greene, Anthony Gitter *ArXiv* (2021-09-17) https://www.ncbi.nlm.nih.gov/pubmed/34545336

PMID: 34545336 • PMCID: PMC8452106

9. An efficient not-only-linear correlation coefficient based on machine learning

Milton Pividori, Marylyn D Ritchie, Diego H Milone, Casey S Greene *Cold Spring Harbor Laboratory* (2022-06-17) https://doi.org/gqcvbw

DOI: 10.1101/2022.06.15.496326

10. Projecting genetic associations through gene expression patterns highlights disease etiology and drug mechanisms

Milton Pividori, Sumei Lu, Binglan Li, Chun Su, Matthew E Johnson, Wei-Qi Wei, Qiping Feng, Bahram Namjou, Krzysztof Kiryluk, Iftikhar Kullo, ... Casey S Greene Cold Spring Harbor Laboratory (2021-07-06) https://doi.org/gk9g25

DOI: 10.1101/2021.07.05.450786

11. Generalizing from a Few Examples

Yaqing Wang, Quanming Yao, James T Kwok, Lionel M Ni *ACM Computing Surveys* (2020-06-12) https://doi.org/gg37m2

DOI: 10.1145/3386252

12. Examining linguistic shifts between preprints and publications

David N Nicholson, Vincent Rubinetti, Dongbo Hu, Marvin Thielk, Lawrence E Hunter, Casey S Greene

PLOS Biology (2022-02-01) https://doi.org/gggzn2

DOI: 10.1371/journal.pbio.3001470 · PMID: 35104289 · PMCID: PMC8806061

13. https://openai.com/blog/chatgpt

14. BLOOM: A 176B-Parameter Open-Access Multilingual Language Model

BigScience Workshop, :, Teven Le Scao, Angela Fan, Christopher Akiki, Ellie Pavlick, Suzana Ilić, Daniel Hesslow, Roman Castagné, Alexandra Sasha Luccioni, ... Thomas Wolf arXiv (2022-12-13) https://arxiv.org/abs/2211.05100

15. **GLM-130B: An Open Bilingual Pre-trained Model**

Aohan Zeng, Xiao Liu, Zhengxiao Du, Zihan Wang, Hanyu Lai, Ming Ding, Zhuoyi Yang, Yifan Xu, Wendi Zheng, Xiao Xia, ... Jie Tang arXiv (2022-10-06) https://arxiv.org/abs/2210.02414

16. **OPT: Open Pre-trained Transformer Language Models**

Susan Zhang, Stephen Roller, Naman Goyal, Mikel Artetxe, Moya Chen, Shuohui Chen, Christopher Dewan, Mona Diab, Xian Li, Xi Victoria Lin, ... Luke Zettlemoyer arXiv (2022-06-22) https://arxiv.org/abs/2205.01068

- 17. ICML 2023 https://icml.cc/Conferences/2023/llm-policy
- 18. Write your best with Grammarly. https://www.grammarly.com/