Predicting Cell Types in Non-Human Single Cell RNA Sequencing Data

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1 Introduction

NeurIPS requires electronic submissions. The electronic submission site is

https://cmt.research.microsoft.com/NeurIPS2018/

Please read the instructions below carefully and follow them faithfully.

1.1 Style

Papers to be submitted to NeurIPS 2018 must be prepared according to the instructions presented here. Papers may only be up to eight pages long, including figures. Additional pages *containing only acknowledgments and/or cited references* are allowed. Papers that exceed eight pages of content (ignoring references) will not be reviewed, or in any other way considered for presentation at the conference.

The margins in 2018 are the same as since 2007, which allow for $\sim 15\%$ more words in the paper compared to earlier years.

Authors are required to use the NeurIPS IATEX style files obtainable at the NeurIPS website as indicated below. Please make sure you use the current files and not previous versions. Tweaking the style files may be grounds for rejection.

1.2 Retrieval of style files

The style files for NeurIPS and other conference information are available on the World Wide Web at

http://www.neurips.cc/

The file neurips_2018.pdf contains these instructions and illustrates the various formatting requirements your NeurIPS paper must satisfy.

The only supported style file for NeurIPS 2018 is neurips_2018.sty, rewritten for LaTeX 2ε . Previous style files for LaTeX 2.09, Microsoft Word, and RTF are no longer supported!

The LaTeX style file contains three optional arguments: final, which creates a camera-ready copy, preprint, which creates a preprint for submission to, e.g., arXiv, and nonatbib, which will not load the natbib package for you in case of package clash.

New preprint option for 2018 If you wish to post a preprint of your work online, e.g., on arXiv, using the NeurIPS style, please use the preprint option. This will create a nonanonymized version of your work with the text "Preprint. Work in progress." in the footer. This version may be distributed as you see fit. Please **do not** use the final option, which should **only** be used for papers accepted to NeurIPS.

32nd Conference on Neural Information Processing Systems (NeurIPS 2018), Montréal, Canada.

At submission time, please omit the final and preprint options. This will anonymize your submission and add line numbers to aid review. Please do *not* refer to these line numbers in your paper as they will be removed during generation of camera-ready copies.

The file neurips_2018.tex may be used as a "shell" for writing your paper. All you have to do is replace the author, title, abstract, and text of the paper with your own.

The formatting instructions contained in these style files are summarized in Sections 2, 3, and 4 below.

2 Related Work

The text must be confined within a rectangle 5.5 inches (33 picas) wide and 9 inches (54 picas) long. The left margin is 1.5 inch (9 picas). Use 10 point type with a vertical spacing (leading) of 11 points. Times New Roman is the preferred typeface throughout, and will be selected for you by default. Paragraphs are separated by $\frac{1}{2}$ line space (5.5 points), with no indentation.

The paper title should be 17 point, initial caps/lower case, bold, centered between two horizontal rules. The top rule should be 4 points thick and the bottom rule should be 1 point thick. Allow ¼ inch space above and below the title to rules. All pages should start at 1 inch (6 picas) from the top of the page.

For the final version, authors' names are set in boldface, and each name is centered above the corresponding address. The lead author's name is to be listed first (left-most), and the co-authors' names (if different address) are set to follow. If there is only one co-author, list both author and co-author side by side.

Please pay special attention to the instructions in Section 4 regarding figures, tables, acknowledgments, and references.

3 Methods

All headings should be lower case (except for first word and proper nouns), flush left, and bold. First-level headings should be in 12-point type.

3.1 Headings: second level

Second-level headings should be in 10-point type.

3.1.1 Headings: third level

Third-level headings should be in 10-point type.

Paragraphs There is also a \paragraph command available, which sets the heading in bold, flush left, and inline with the text, with the heading followed by 1 em of space.

4 Preliminary Results

These instructions apply to everyone.

4.1 Citations within the text

The natbib package will be loaded for you by default. Citations may be author/year or numeric, as long as you maintain internal consistency. As to the format of the references themselves, any style is acceptable as long as it is used consistently.

The documentation for natbib may be found at

http://mirrors.ctan.org/macros/latex/contrib/natbib/natnotes.pdf

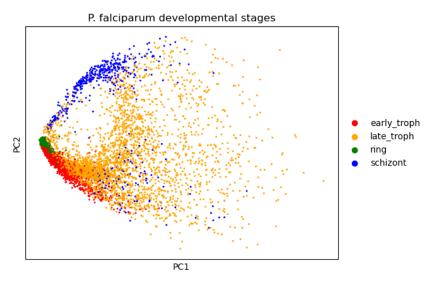


Figure 1: Principal components analysis of *P. falciparum* single cell RNA sequencing data, colored by lifecycle stage.

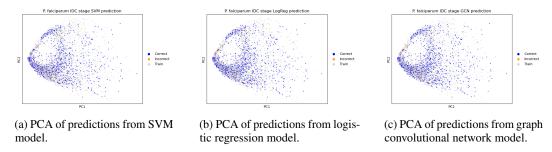


Figure 2: Model performance visualized on PCA plots.

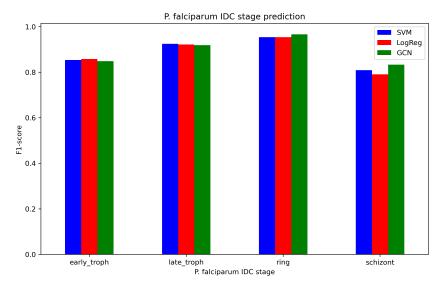


Figure 3: Prediction accuracy of models on P. falciparum single cell RNA sequencing data.

Table 1: Summary of datasets that will be used in this study.

Host	Source	Reference	Cells
Parasite	GitHub	Howick et al. 2019	6,737
Parasite	GSE146737	Wendt et al. 2020	43,642
Parasite	Zenodo	Briggs et al. 2021	8,599
Parasite	GitHub	Rezvani et al. 2022	8,719-12,910
Human	TENxPBMCData R package (Kasper et al. 2021)	10X Genomics	3,000
Human	scRNAseq package (Risso and Cole, 2021)	Lawlor et al. 2017	978
Human	GSE151530	Ma et al. 2021 and Ma et al. 2022	56,721
Mouse	scRNAseq package (Risso and Cole, 2021)	Zeisel et al. 2015	3,005
Zebrafish	GSE106587	Farrell et al. 2018	38,731
Non-human primate	GSE156755	Speranza et al. 2021	100,795

Of note is the command \citet, which produces citations appropriate for use in inline text. For example,

\citet{hasselmo} investigated\dots

produces

Hasselmo, et al. (1995) investigated...

If you wish to load the natbib package with options, you may add the following before loading the neurips_2018 package:

\PassOptionsToPackage{options}{natbib}

If natbib clashes with another package you load, you can add the optional argument nonatbib when loading the style file:

\usepackage[nonatbib] {neurips_2018}

As submission is double blind, refer to your own published work in the third person. That is, use "In the previous work of Jones et al. [4]," not "In our previous work [4]." If you cite your other papers that are not widely available (e.g., a journal paper under review), use anonymous author names in the citation, e.g., an author of the form "A. Anonymous."

4.2 Footnotes

Footnotes should be used sparingly. If you do require a footnote, indicate footnotes with a number in the text. Place the footnotes at the bottom of the page on which they appear. Precede the footnote with a horizontal rule of 2 inches (12 picas).

Note that footnotes are properly typeset *after* punctuation marks.²

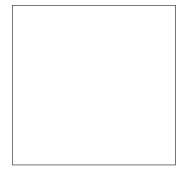


Table 2: Sample table title

Part		
Name	Description	Size (μm)
Dendrite Axon Soma	Input terminal Output terminal Cell body	$\begin{array}{c} \sim \! 100 \\ \sim \! 10 \\ \text{up to } 10^6 \end{array}$

4.3 Figures

All artwork must be neat, clean, and legible. Lines should be dark enough for purposes of reproduction. The figure number and caption always appear after the figure. Place one line space before the figure caption and one line space after the figure. The figure caption should be lower case (except for first word and proper nouns); figures are numbered consecutively.

You may use color figures. However, it is best for the figure captions and the paper body to be legible if the paper is printed in either black/white or in color.

4.4 Tables

All tables must be centered, neat, clean and legible. The table number and title always appear before the table. See Table 2.

Place one line space before the table title, one line space after the table title, and one line space after the table. The table title must be lower case (except for first word and proper nouns); tables are numbered consecutively.

Note that publication-quality tables *do not contain vertical rules*. We strongly suggest the use of the booktabs package, which allows for typesetting high-quality, professional tables:

https://www.ctan.org/pkg/booktabs

This package was used to typeset Table 2.

5 Future Plans

Please prepare submission files with paper size "US Letter," and not, for example, "A4."

Fonts were the main cause of problems in the past years. Your PDF file must only contain Type 1 or Embedded TrueType fonts. Here are a few instructions to achieve this.

• You should directly generate PDF files using pdflatex.

¹Sample of the first footnote.

²As in this example.

- You can check which fonts a PDF files uses. In Acrobat Reader, select the menu Files>Document Properties>Fonts and select Show All Fonts. You can also use the program pdffonts which comes with xpdf and is available out-of-the-box on most Linux machines.
- The IEEE has recommendations for generating PDF files whose fonts are also acceptable for NeurIPS. Please see http://www.emfield.org/icuwb2010/downloads/IEEE-PDF-SpecV32.pdf
- xfig "patterned" shapes are implemented with bitmap fonts. Use "solid" shapes instead.
- The \bbold package almost always uses bitmap fonts. You should use the equivalent AMS Fonts:

```
\usepackage{amsfonts}
```

followed by, e.g., \mathbb{R} , \mathbb{R} , \mathbb{R} , or \mathbb{C} . You can also use the following workaround for reals, natural and complex:

```
\newcommand{\RR}{I\!\!R} %real numbers
\newcommand{\Nat}{I\!\!N} %natural numbers
\newcommand{\CC}{I\!\!!\!C} %complex numbers
```

Note that amsfonts is automatically loaded by the amssymb package.

If your file contains type 3 fonts or non embedded TrueType fonts, we will ask you to fix it.

5.1 Margins in LaTeX

Most of the margin problems come from figures positioned by hand using \special or other commands. We suggest using the command \includegraphics from the graphicx package. Always specify the figure width as a multiple of the line width as in the example below:

```
\usepackage[pdftex]{graphicx} ...
\includegraphics[width=0.8\linewidth]{myfile.pdf}
```

See Section 4.4 in the graphics bundle documentation (http://mirrors.ctan.org/macros/latex/required/graphics/grfguide.pdf)

A number of width problems arise when LATEX cannot properly hyphenate a line. Please give LaTeX hyphenation hints using the \- command when necessary.

Acknowledgments

Use unnumbered third level headings for the acknowledgments. All acknowledgments go at the end of the paper. Do not include acknowledgments in the anonymized submission, only in the final paper.

References

References follow the acknowledgments. Use unnumbered first-level heading for the references. Any choice of citation style is acceptable as long as you are consistent. It is permissible to reduce the font size to small (9 point) when listing the references. Remember that you can use more than eight pages as long as the additional pages contain *only* cited references.

- 1. Heumos, L., et al. (2023). Best practices for single-cell analysis across modalities. *Nat Rev Genet*, 24, 550–572.
- 2. Yang, F., et al. (2022). scBERT as a large-scale pretrained deep language model for cell type annotation of single-cell RNA-seq data. *Nat Mach Intell*, 4, 852–866.
- 3. Chen, J., et al. (2023). Transformer for one stop interpretable cell type annotation. *Nat Commun*, 14, 223.
- 4. Howick, V. M., et al. (2019). The Malaria Cell Atlas: Single parasite transcriptomes across the complete *Plasmodium* life cycle. *Science*, 365, eaaw2619.
- Kipf, T. N., & Welling, M. (2016). Semi-Supervised Classification with Graph Convolutional Networks. arXiv:1609.02907.

- Pedregosa, F., et al. (2012). Scikit-learn: Machine Learning in Python. J Mach Learn Res, 12, 2825–30.
- 7. Spektral. https://github.com/danielegrattarola/spektral
- 8. F1 Score. https://towardsdatascience.com/the-f1-score-bec2bbc38aa6
- TENxPBMCData. https://bioconductor.org/packages/release/data/experiment/ html/TENxPBMCData.html
- scRNAseq. https://bioconductor.org/packages/release/data/experiment/manuals/ scRNAseq/man/scRNAseq.pdf
- 11. Lawlor, N., et al. (2017). Single-cell transcriptomes identify human islet cell signatures and reveal cell-type-specific expression changes in type 2 diabetes. *Genome Res*, 27(2), 208-222.
- 12. Zeisel, A., et al. (2015). Cell types in the mouse cortex and hippocampus revealed by single-cell RNA-seq. *Science*, 347, 1138-1142.
- 13. Ma, L., et al. (2021). Single-cell atlas of tumor cell evolution in response to therapy in hepatocellular carcinoma and intrahepatic cholangiocarcinoma. *J Hepatol*, 75(6), 1397-1408.
- 14. Ma, L., et al. (2022). Multiregional single-cell dissection of tumor and immune cells reveals stable lock-and-key features in liver cancer. *Nat Commun*, 13(1), 7533.
- 15. Farrell, J. A., et al. (2018). Single-cell reconstruction of developmental trajectories during zebrafish embryogenesis. *Science*, 360(6392).
- Speranza, E., et al. (2021). Single-cell RNA sequencing reveals SARS-CoV-2 infection dynamics in lungs of African green monkeys. Sci Transl Med, 13(578).
- 17. Wendt, G., et al. (2020). A single-cell RNA-seq atlas of Schistosoma mansoni identifies a key regulator of blood feeding. *Science*, 369(6511), 1644-1649.
- 18. Briggs, E. M., et al. (2021). Single-cell transcriptomic analysis of bloodstream Trypanosoma brucei reconstructs cell cycle progression and developmental quorum sensing. *Nat Commun*, 12, 5268.
- 19. Rezvani, Y., et al. (2022). Comparative single-cell transcriptional atlases of Babesia species reveal conserved and species-specific expression profiles. *PLOS Biol*, 20(9), e3001816.