

## Review of: "Microbiome Maps: Hilbert Curve Visualizations of Metagenomic Profiles"

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First, I would like to declare that I am not a computer scientist or a mathematician so I cannot comment on some of the algorithmic decisions made by the authors.

The authors describe a tool, which they call Jasper, that utilises Hilbert Curve to visualise genomes in metagenomic samples. Overall, the manuscript is very well written, easy to read, clear and to the point. I find that the advantages that the authors ascribe to their method to be genuinely interesting and of potential relevance and value to the community. I find myself interested in using the tool in the future and perhaps even recommending it. The website associated with the tool is well designed and has sufficient content.

## I have a few comments:

- · It is helpful to have line numbering for specific commenting
- Overall, the authors do not comment on how they consider abundance values reported from different metagenomic
  profiling tools. I would urge the authors to have a look at a recent review by Sun et al (2021) <sup>[1]</sup>. This review describes a
  somewhat concerning problem of reporting microbial abundances in metagenomics samples
- "While the Karken2 software does it for 16S datasets". This is not strictly true. Kraken2 is WGS metagenomic profiler that can also accommodate some 16S datasets. In fact the standard Kraken2 database is RefSeq (Bacteria, Archaea & Viruses). Refer to the publication [2] as well as the GitHub repo (https://github.com/DerrickWood/kraken2/wiki/Manual#standard-kraken-2-database).
- I am bit disappointed that the authors only provide a MacOS version of their tool (not clear in the manuscript). I would have been nice to have a Linux version or a webserver.
- "Note that unlike other HCV techniques, microbiome maps do not depict genomic positions....from a reference collection": not clear. Please elaborate.

## References

1. <sup>^</sup>Zheng Sun, Shi Huang, Meng Zhang, Qiyun Zhu, et al. (2021). <u>Challenges in benchmarking metagenomic profilers</u>.

Nat Methods, vol. 18 (6), 618-626. doi:10.1038/s41592-021-01141-3.



2. ^Derrick E. Wood, Jennifer Lu, Ben Langmead. (2019). Improved metagenomic analysis with Kraken 2. Genome Biol, vol. 20 (1). doi:10.1186/s13059-019-1891-0.