

Reviewer Report

Title: The state of Medusozoa genomics: current evidence and future challenges

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Reviewer name: Sheila Kitchen

Reviewer Comments to Author:

Santander et al. review the state of genome assemblies and cytogenetics of Medusozoa. This review captures the progression of the sequencing efforts in the past decade and how the field is moving with new technological advances. From their assessment of the literature and unpublished data, they found that a weakness in their community is a general lack of standardization in analysis and limited availability of intermediate assembly components, such as the repeat libraries, and associated metadata. In the end they provide recommendations for standards to be applied to ongoing and future genomic projects.

I felt that these recommendations fell short of extending beyond basic requirements of publishing genomes today. While these recommendations are in line with recommendations of other genomic consortia (Vertebrate Genomes Project [Rhie et al. 2021, Nature], Sanger/Moore Aquatic Symbiosis Genomics, etc.) and most publishers including GigaScience (deposit data, reproducible methods, code availability statements, etc), they are quite general. I was left wondering if this was a commentary on the whole field of genomics. To that end, are there specific recommendations regarding medusozoans that would enhance data usage community wide that could be stated here? Are there established assembly pipelines (i.e. tools that provide the highest quality assemblies from various species) or types of sequencing effort (i.e. long read + HiC maps, transcriptome-informed gene annotation) that should be endorsed as part of your assessment? Are there specific taxonomic gaps that should be prioritized (starting Line 238)?

The majority of the resources you identified only have short-read Illumina data which inevitably means that chromosome-scale assemblies are not possible yet. However, these assemblies are sufficient for gene model comparisons across species (starting on Line 187). Is there a way to standardize gene prediction for cases where short reads may be all that is available? Re-analysis of gene predictions with different tools may lead to varying estimates and can lead to erroneous orthology assignments (see <https://doi.org/10.1111/jpy.12947>, <https://doi.org/10.1371/journal.pbio.3000862>, and <https://www.biorxiv.org/content/10.1101/2022.01.13.476251v1>). Re-analysis of *Rhopilema* gene content using different tools increases gene predictions closer to the median gene count you've found. Regarding the recommendation for depositing intermediates into repositories (#3), is there one established for the community or are you referring to more general ones like Dryad, FigShare, Rebase, etc.? Providing an example genome project or two that shares these associated files might be helpful. There can be cost associated with hosting these resources. Do you see that as a barrier to researchers providing this sort of data?

A recommendation that is provided earlier in the paper is the call for lineage-specific single copy ortholog sets (Line 228). Should this be re-stated in the final recommendations as well?

Minor Comments:

Line 31-33: This sentence seems to be constructed of two thoughts but missing a connector between them.

Line 98: ... assembly statistics using the statswrapper.sh script ...

Line 169: ... [55], and the ...

Line 315: Remove "of" between reusing and previously.

Line 337: "reran" should be "rerun".

Line 389: Typo, "projects"

Figures: The resolution of the figures provided made it difficult to review. Specifically Figure 3 was quite pixelated.

Methods

Are the methods appropriate to the aims of the study, are they well described, and are necessary controls included? Choose an item.

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Are the conclusions adequately supported by the data shown? Choose an item.

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