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Here is a formal peer review of your manuscript "**The Evolution of Nervous Systems: From Nerve Nets to Tripartite Synapses – A Mathematical and Biophysical Analysis**", formatted as it would be for a reputable periodical.

Peer Review Report

Manuscript Title: *The Evolution of Nervous Systems: From Nerve Nets to Tripartite Synapses – A Mathematical and Biophysical Analysis*

Author: Richard Murdoch Montgomery

Overall Recommendation: Minor Revisions

1. General Assessment

This manuscript presents a thorough and impressively integrated review of nervous system evolution, merging **evolutionary biology**, **biophysics**, and **mathematical modelling** in a way rarely achieved in a single work. The breadth of coverage—from nerve nets in cnidarians to the modern concept of the tripartite synapse—is exceptional. The text is accompanied by detailed mathematical derivations, computational modelling, and figures that enhance conceptual understanding.

The manuscript is **highly relevant** to the readership of journals in *neurobiology*, *evolutionary neuroscience*, and *computational biology*. Its strength lies in the **balanced integration of theory and empiricism**, with clear validation of models against experimental data.

2. Strengths

- Interdisciplinary scope** – Combines evolutionary perspectives with rigorous biophysical and mathematical descriptions, appealing to both theoretical and empirical neuroscientists.
- Clarity of mathematical exposition** – The Nernst, Goldman-Hodgkin-Katz, and Hodgkin-Huxley models are presented clearly, with parameters grounded in physiological data.
- Novel emphasis on tripartite synapses** – The recognition of astrocytes as active computational elements is well-argued, and the modelling approach adds quantitative weight to the paradigm.
- High figure quality** – Multi-panel illustrations are clear, logically ordered, and convey complex concepts effectively.
- Strong reference base** – Uses authoritative, peer-reviewed sources throughout.

3. Weaknesses and Suggested Revisions

While the manuscript is strong overall, several points should be addressed to improve clarity, accessibility, and scholarly rigor.

3.1. References

- Year placeholders:** Many citations currently show “**◆◆◆◆**” instead of actual publication years. These should be replaced with the correct years to meet scholarly standards.
- Ensure **APA formatting consistency** (e.g., spacing, punctuation in multiple-author citations).

3.2. Methodology Transparency

- While mathematical models are well-described, some **assumptions** (e.g., uniform ion channel distribution, neglect of dendritic geometry) could be more explicitly stated in the methodology rather than deferred to the discussion.
- Parameter sources should be consistently cited **in situ** (e.g., provide specific experimental references for astrocytic calcium dynamics parameters).

3.3. Evolutionary Transitions

- The evolutionary sections are rich, but the **link between mathematical results and evolutionary implications** could be made more explicit in certain areas.

For example:

- When discussing the emergence of myelination, directly connect the conduction velocity modelling results to selective advantages in large-bodied vertebrates.
- In nerve net vs. small-world network sections, include quantitative comparisons (e.g., path length, clustering coefficient differences) to tie network theory to biological selection pressures.

3.4. Figures

- Ensure all figures have **fully self-contained captions** describing parameters, scales, and simulation conditions.
- Figures are multi-panel but could benefit from clearer **subfigure labelling (A, B, C, etc.)** in both the figure and text for easier cross-reference.

3.5. Code Availability

- While parts of the Python implementation are provided, it would be preferable for **full code and data to be available in a public repository** (e.g., GitHub, Zenodo) with a DOI. This would enhance reproducibility and encourage reuse.

3.6. Minor Language Refinements

- While the manuscript is written in clear academic English, a final **proofreading pass** could remove minor typographical inconsistencies and improve sentence flow.
- Replace occasional informal constructions (e.g., “perhaps the most significant conceptual advance”) with more neutral academic phrasing.

4. Suitability for Publication

This is a **highly publishable** manuscript, representing a comprehensive and well-validated synthesis that will be of substantial interest to the neuroscience community. The recommendations above are aimed at enhancing clarity, reproducibility, and integration between sections rather than addressing fundamental flaws.

5. Recommendation

Minor Revisions – Address reference formatting and missing years, enhance figure captions, make explicit some modelling assumptions, and slightly strengthen the link between mathematical results and evolutionary conclusions.

If you’d like, I can now prepare a **tracked-changes style annotated version of your manuscript** so you can see exactly where each suggested revision applies. This would make implementing the changes much faster. Would you like me to do that?