Peer Review

Review of: "Mechanisms of Glycolysis and Fermentation: A Non-Equilibrium

Thermodynamics Perspective"

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This review article presents a highly original and thought-provoking reinterpretation of glycolysis and

fermentation through the lens of non-equilibrium thermodynamics. The authors challenge conventional

biochemical paradigms, offering an integrative perspective that attempts to reconcile classical models

with physical laws, particularly focusing on proton flow (H+) as a central energy entity. The manuscript

demonstrates deep historical knowledge, referencing foundational work by pioneers such as Meyerhof,

Krebs, and Mitchell. Its ambition to unify biochemical pathways with thermodynamic principles is

commendable and could stimulate significant academic discussion across disciplines, including

biochemistry, physics, and systems biology.

However, the manuscript is excessively dense, highly speculative in several areas, and often strays into

philosophical and theoretical assertions without sufficient empirical grounding. The language is

frequently convoluted, with repetitive arguments and unclear terminology that may confuse readers

unfamiliar with both advanced thermodynamics and metabolic biochemistry. Furthermore, while

historical references are abundant, there is limited engagement with contemporary empirical research

that could substantiate the proposed models. The figures, though conceptually interesting, lack clarity

and are not adequately explained. There are also concerns regarding overstatements, particularly

accusations of "scientific misconduct" against prevailing biochemical models, which are inappropriate

for scientific discourse.

Declarations

Potential competing interests: No potential competing interests to declare.