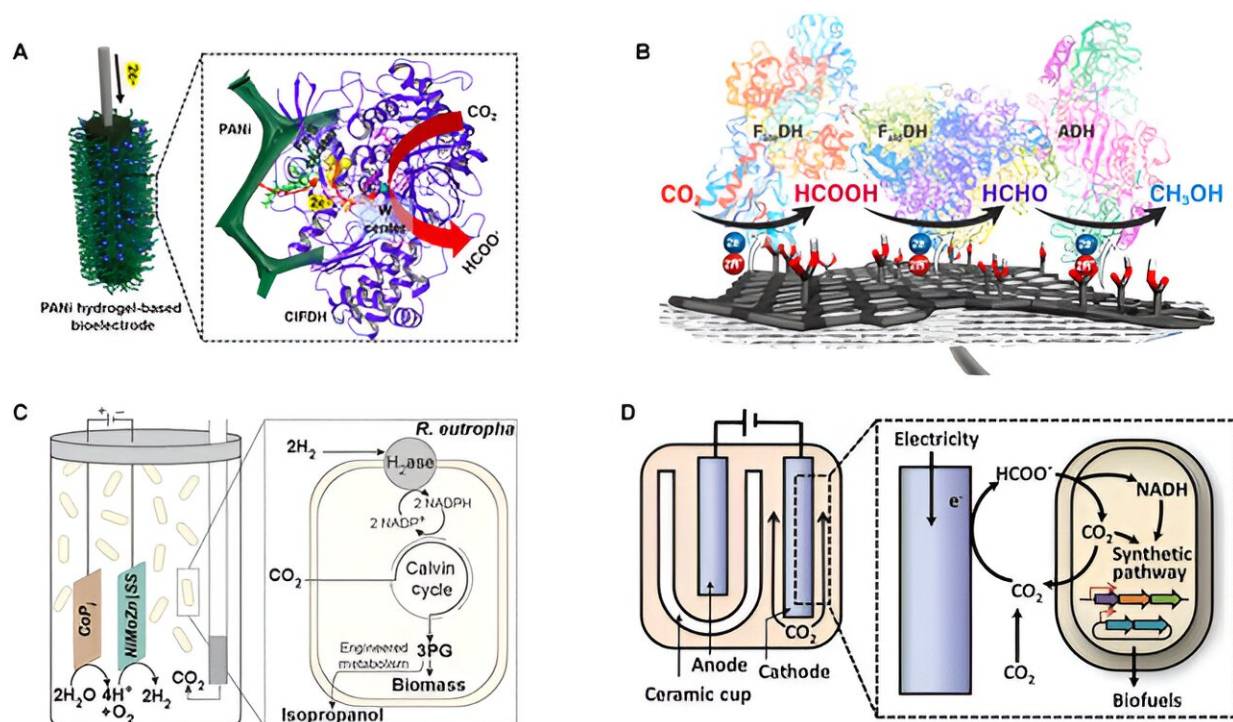


Revolutionizing biorefineries: Advancing toward sustainable third-generation technologies in CO₂ utilization

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Schematic illustration of electrical-bio system for CO₂ fixation. (A) Direct electron transfer from the conductive PANi hydrogel electrode to immobilized C1FDH for electroenzymatic CO₂ reduction to formate. (B) Direct electron transfer from the functionalized graphene to the immobilized tandem enzymatic cascade for CO₂ reduction to methanol. (C) Culturing of *C. necator* in bioelectrochemical cell with CoPi anode and NiMoZn/SS cathode to convert H₂ and CO₂ into 3PG that is subsequently converted into biomass or isopropanol in engineered Re2133-pEG12. (D) Electrochemical reduction of CO₂ to formate

that is converted to isobutanes and 3MB by the engineered R. eutrophic. Credit: *BioDesign Research* (2023). DOI: 10.34133/bdr.0021

The evolution of biorefineries, shifting from sugar-based and biomass feedstocks to third-generation (3G) technologies, marks significant progress toward sustainable development. 3G biorefineries use microbial cell factories or enzymatic systems to convert one-carbon (C1) sources such as CO₂ into value-added chemicals, powered by renewable energies.

Despite the potential of native C1 assimilating microbes, challenges like low carbon fixation efficiency and limited product scope hinder their scalability. Heterotrophic microorganisms, engineered through [synthetic biology](#) and computational tools, offer a promising solution to these challenges. The current research focuses on enhancing the efficiency of C1 fixation and productivity of desired compounds, with chemo-bio hybrid systems leveraging electricity and light as emerging strategies.

In October 2023, three researchers from Hangzhou, China, published a review article titled "[Design and Construction of Artificial Biological Systems for One-Carbon Utilization](#)" in *BioDesign Research*. In this review, significant advancements over the past decade in the development of third-generation (3G) biorefineries are discussed.

These refineries focus on using one-carbon (C1) sources such as CO₂, methanol, and formate, harnessing artificial autotrophic microorganisms, tandem enzymatic systems, and chemo-bio hybrid systems. This approach could revolutionize biotechnology, offering sustainable alternative strategies for industrial production.

Central to these developments are natural CO₂ fixation pathways, which

have been instrumental in engineering artificial systems for heterotrophic microorganisms such as *E. coli* and *Pichia pastoris*. Despite this progress, challenges such as energy imbalances, low carbon fixation efficiency, and the absence of native methanol assimilation pathways in certain heterotrophs still remain.

Chemo-bio hybrid systems, which combine electrocatalysis and biocatalysis, show promise for efficient CO₂ conversion. However, issues such as maintaining metabolite stability and [enzymatic activity](#) still need addressing.

Overcoming these challenges is essential for the success of these artificial biological systems in C1 utilization, with potential transformative impacts on various industries, including pharmaceuticals, agriculture, and food production.

More information: Wei Zhong et al, Design and Construction of Artificial Biological Systems for One-Carbon Utilization, *BioDesign Research* (2023). [DOI: 10.34133/bdr.0021](https://doi.org/10.34133/bdr.0021)

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