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Editorial Reflections on BioEnergy: Perspectives from 2024

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1 Editorial Reflections on BioEnergy: Perspectives from 2024

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12 For millennia fermentations of grain and fruit — such as those used in beer and wine — provided valuable nutrition
13 for communities worldwide. In 2024, this biobased energy has taken on a very different context: advancements in
14 our ability to isolate organisms, manipulate DNA, and optimize bioproduction on different feedstocks have brought
15 about more efficient fermentations; the energy derived from cultivating organisms not only fuels our food but also
16 drives our cars and lifts our airplanes. This section of Current Opinion in Biotechnology: BioEnergy offers a
17 snapshot of the global effort through collaborations across academia, government, and industry, to optimize
18 microbial strains, expand feedstock variety, and streamline strain development time and cost.

19

20 Although historically focused on the production of bioethanol, the field as a whole is also exploring a host of
21 advanced biofuels and small molecule products. Chainani et al. discuss how coupling chemistry and biology for the
22 synthesis of advanced bioproducts integrates synthetic biology with chemical catalysis enabling the production of
23 novel biofuels and valuable bioproducts. This multidisciplinary approach allows for precision in designing microbes
24 that can produce a wider range of compounds, including advanced biofuels like isobutanol, higher alcohols, and
25 alkanes. By leveraging both chemical and biological systems, it becomes possible to tap into new molecules and
26 pathways that would have been out of reach using either system alone.

27

28 Meanwhile, nature has long been a source of inspiration and raw material for biotechnological innovation. Li et al.
29 discuss how tapping into these natural resources holds enormous potential for biofuel production. Bioprospecting
30 and metagenomics allow researchers to explore previously inaccessible microbial communities, such as those in
31 extreme environments, for novel enzymes and metabolic pathways. These enzymes and pathways could lead to the
32 production of new biofuels and bioproducts, potentially reducing the need for petroleum-derived materials and
33 chemicals.

34

35 The tools for metabolic engineering have evolved rapidly, but modeling these complex systems remains a challenge.
36 Han et al. discuss how advances in genome-scale metabolic models of industrially important fungi are helping
37 address this challenge by offering powerful computational tools to optimize strain performance. Such models allow
38 researchers to predict the effects of genetic modifications, optimize metabolic pathways, and ultimately improve the
39 yield and efficiency of biofuel production. For fungi, which are crucial players in industrial biotechnology for their
40 ability to produce bioethanol and other products from lignocellulosic biomass, these advances represent a significant
41 step forward in accelerating research
42 and reducing the trial-and-error approach that has historically dominated strain development. Designing and
43 optimizing microbes at the genomic level require innovative approaches to enzyme engineering. Generative models,
44 such as those using machine learning, allow researchers to design enzymes that are optimized for specific reactions,
45 including those involved in biofuel production. Barghout et al. discuss how, by training these models on vast
46 datasets of enzyme structures and functions, researchers can now create enzymes with higher specificity and
47 activity, increasing the efficiency of biofuel production processes. This approach not only accelerates enzyme
48 discovery but also opens the door to entirely new catalytic functions that might not exist in nature.

49

50 In the pursuit of biofuels, Hu et al. discuss how traditional model organisms like *Saccharomyces cerevisiae* and
51 *Escherichia coli* are being joined by emerging oleaginous yeasts, algae, bacteria, and fungi each with unique
52 metabolic capabilities that can be harnessed for next-generation biofuels. These

53 organisms can often metabolize a wider range of feedstocks, including those that are less amenable to traditional
54 microbial fermentation, and can produce biofuels that are more energy-dense than ethanol. Martinez-Garcia and
55 Lorenzo offer an in-depth review of *Pseudomonas putida*, while Zhang
56 et al. discuss how extremophiles like *Halomonas* spp. offer opportunities in nonsterile bioproduction. Although the
57 majority of this issue is focused on microbial bioproduction, Morgan et al. offer a review of plant synthetic biology
58 and how it offers promise for sustainability and space exploration. As researchers continue to explore these
59 nonmodel systems, they may uncover new opportunities to improve biofuel yields and expand the range of biofuels
60 produced.

61
62 Another area of biofuel research is improving microbial bioproduction under low-oxygen conditions. Many
63 industrial bioprocesses occur in conditions where oxygen is limited, such as in large-scale fermenters or in the
64 production of biofuels from anaerobic digestion. Microbes that can function efficiently under low-oxygen conditions
65 are therefore essential for these processes. Kulakowski et al. discuss how researchers are developing strains with
66 enhanced tolerance to low oxygen and engineering metabolic pathways
67 that can operate under these constraints, thereby improving the efficiency and robustness of biofuel production
68 systems.

69
70 Finally, the issue of feedstock availability and suitability is central to the future of biofuels. As biofuel production
71 expands, reliance on traditional feedstocks like corn and sugarcane becomes less sustainable. Scown et al. discuss
72 the growing need to explore nontraditional feedstocks, including agricultural residues, municipal waste, and
73 lignocellulosic biomass. Liu et al. offer a focus on bioproduction from lignin; Tan et al. discuss how methanotrophs
74 are being used to harness methane for bioproduction. Matching these diverse feedstocks with the right microbial or
75 chemical conversion processes is key to
76 making biofuels more economically viable and environmentally sustainable. This approach not only diversifies the
77 inputs into the biofuel production pipeline but also reduces competition with food crops, a crucial consideration as
78 global populations and food demands rise.

79
80 This special issue has touched on some of the leading research on the frontiers of biofuel research, bringing together
81 advances in microbial engineering, enzyme design, metabolic modeling, and feedstock conversion. As the world
82 continues to shift towards renewable energy, the role of biofuels and bioproducts will become increasingly critical in
83 mitigating climate change, reducing reliance on fossil fuels, and fostering sustainable economic growth. By
84 exploring both established and emerging systems, as well as leveraging cutting-edge techniques in synthetic biology,
85 machine learning, and bioprospecting, the biofuel field is well-positioned to continue advancing toward a more
86 sustainable future.