

## **Acids Unlocking CO<sub>2</sub>-Based Biomanufacturing: Multi-Omics-Driven Optimization of Gas Fermentation Coupled to Acetate-Based Production of Proteins and Omega-3 Fatty Acids**

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Industrial carbon emissions remain a major barrier to achieving climate neutrality, yet CO<sub>2</sub>-derived feedstocks offer new avenues for sustainable biomanufacturing. As part of the Research Council of Norway-funded Centre for Research and Innovation “SFI Industrial Biotechnology” and the collaborative research project “CO<sub>2</sub>Value”, we present an integrated value chain that transforms industrial CO<sub>2</sub> into high-value proteins and omega-3 fatty acids through gas fermentation and acetate-driven microbial bioprocesses.

At the core of this work is a thermophilic gas-fermenting bacterium capable of converting CO<sub>2</sub> and H<sub>2</sub> into acetate with high carbon efficiency. We combine a curated genome-scale metabolic model with quantitative multi-omics (proteomics and metabolomics), to unravel metabolic regulation under autotrophic and mixotrophic growth. This systems-level understanding provides the foundation for rational strain improvement and process optimisation, identifying metabolic bottlenecks and energy-flux constraints in thermophilic CO<sub>2</sub> fixation.

The acetate generated from gas fermentation is concentrated and directly deployed as the sole carbon source in two parallel biomanufacturing routes. First, filamentous fungi are used to produce protein-rich biomass for sustainable feed applications, achieving high yields and favourable amino-acid profiles. Second, marine protists are cultivated to synthesize omega-3 fatty acids, reaching lipid contents of up to 80% of dry cell weight under optimised fed-batch conditions. In both pathways, we demonstrate complete process chains from CO<sub>2</sub> to final product, including fermentation optimisation and downstream processing.

Together, these results highlight a scalable and circular CCU concept that couples thermophilic gas fermentation with acetate-based microbial production platforms. By integrating system biology, bioprocess engineering, and downstream valorisation, this work advances the technological readiness of CO<sub>2</sub>-derived proteins and lipids for feed and food sectors, therefore contributing to a more resilient and sustainable bio-based economy.