

## Researchers develop engineered Saccharomyces cerevisiae yeast strain for optimized production of carotenoids

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(A) PEST fusion to Gal80 for copper-induced GAL expression in YPD media. Copper sulfate ( $20 \mu M$ ) was used to induce the carotenogenic genes under the control of copper-inducible GAL system. (B) Carotenoid contents of engineered yeasts in YNBD and YPD media at 72 h. (C) Growth curves of JS-BE5 and JS-BE5-PEST cultivated in YNBD and YPD media. Strain JS-BE5-PEST is derived



from JS-BE5 with PEST-Gal80N. (D) Representative HPLC results showing the abundance of carotenoids produced by JS-BE5-PEST. P values were obtained from two-tailed unpaired Student's t test. \*\*\*P

More than 90% of the commercially available carotenoids are synthetically produced using chemicals. To meet the increasing demand for cost-effective natural compounds in carotenoid synthesis, researchers at Xiamen University, China, have developed an engineered S. cerevisiae yeast strain capable of selectively overproducing carotenoids.

They redesigned the genomic sequence and critical pathways to optimize <u>carotenoid</u> production. This novel and successful research approach can be extended to other model microorganisms in the future. The results of this work were published online in the <u>BioDesign Research</u> journal.

Carotenoids, classified as tetraterpenoids, feature a carbon-hydrogen backbone and are found in algae, plants, and some microorganisms. The <u>bright colors</u> in most vegetables and fruits are a result of these carotenoid pigments. Due to their excellent antioxidant properties, carotenoids are widely used in health care, medicine, and cosmetics. In recent years, these antioxidant molecules have gained popularity as a nutraceutical food supplement.

Most of the present-day, commercially available carotenoids are produced through <u>chemical synthesis</u>. The extraction and purification of carotenoids from <u>natural sources</u> is restricted by the costs and time-consuming nature of the processes involved. To circumvent these issues and meet the increasing demand for carotenoids, a group of researchers from China, led by Dr. Jifeng Yuan of Xiamen University, have successfully attempted to produce carotenoids by employing microbial synthesis techniques.

"Over the years, biotechnology has evolved drastically, allowing for specific modifications at the genetic level and regulating both, intracellular and intercellular dynamics. Our team has used multiple strategies on the model microorganism of Saccharomyces cerevisiae to improve the microbial synthesis of carotenoids," says Dr. Yuan, sharing the inspiration behind using microbial synthesis technology.



They initially re-engineered the <u>metabolic pathways</u> in S. cerevisiae <u>yeast</u> to increase the overall content of acetyl-CoA, which can be easily converted into carotenoids. Further, they suppressed the secondary metabolic pathway of squalene synthase-ergosterol. This ergosterol pathway reduces the overall availability of energy and raw materials needed for carotenoid production.

Also, as carotenoids are hydrophobic-resistant to water, their storage within yeast microorganisms was a challenge. The researchers introduced the human lipid binding/transfer protein saposin B (hSapB) genetic sequence to enhance the storage of carotenoids.

By changing the growth medium of yeasts to a low-cost, highly beneficial medium rich in yeast–peptone–dextrose and re-engineering a genetic expression system, the output of biomass increased by five-fold.

The S. cerevisiae yeast strain was selectively re-engineered to optimize the production of high content of beta-carotene, a pre-cursor molecule of carotenoids.

"We are very hopeful about the scalability and ease of implementing our novel carotenoid production technology and believe that in the near future such engineering strategies can be extended to other microorganisms as well," concludes Dr. Yuan.

Advancing bio-engineering to create novel model microorganisms holds promise for future industrial carotenoid synthesis, facilitating the production of essential health supplements. This approach can significantly contribute to mitigating vitamin deficiencies and addressing malnutrition on a global scale.

**More information:** Jian Fan et al, Multidimensional Optimization of Saccharomyces cerevisiae for Carotenoid Overproduction, *BioDesign Research* (2023). DOI: 10.34133/bdr.0026



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