

Efficient synthetic system for production of microbial drugs

Ya-Ping Xue, Yu-Guo Zheng

Key Laboratory of Bioorganic Synthesis of Zhejiang Province, College of Biotechnology and Bioengineering, Zhejiang University of Technology, Hangzhou 310014, China.

Engineering Research Center of Bioconversion and Biopurification of Ministry of Education, Zhejiang University of Technology, Hangzhou 310014, China.

National and Local Joint Engineering Research Center for Biomanufacturing of Chiral Chemicals, Zhejiang University of Technology, Hangzhou 310014, P. R. China

Microbial drugs have been widely used to control bacterial or fungal infections, tumors, diabetes, immune diseases, and et al. They play an important role in safeguarding human health and improving the quality of life. *Actinomycetes*, *fungi*, *Pseudomonas*, and *Bacillus* are the main drug-producing microorganisms. However, the yield is still too low for a variety of important microbial drugs. There are several bottlenecks such as multi-component coexistence, low matrix utilization, and low yield, which restrict the production of microbial drugs. We have studied the high-yield mechanism of several microbial drugs such as acarbose, daptomycin, fidaxomicin, doxorubicin, amphotericin B, echinomycin and et al, successfully optimized the metabolic networks and developed highly efficient production techniques. In the case of acarbose, a highly effective treatment for type II diabetes, due to the complex biosynthetic pathway and massive structural analogs of acarbose, it is difficult to develop a high-efficiency, low-cost technology for production of acarbose with high-quality. After years of research, we have invented a high-throughput screening method combined with efficient breeding technology to improve the acarbose-producing strain. By investigating acarbose metabolic pathway, and anabolic regulatory genes, the key regulatory factors which affect acarbose synthesis was discovered. Based on these results, a highly efficient synthesis technology was

developed for acarbose production by system optimization and the first industrial production line of acarbose in China was built.



Xue, Ya-Ping

Dr. Xue Ya-Ping received his Ph.D. in biochemical engineering. He is Professor at the College of Biotechnology and Bioengineering in Zhejiang University of Technology. His research interests are synthetic biology, biochemical engineering, metabolic engineering, and biotransformation and biocatalysis. He has published about 70 papers and filed more than 100 patents. He has developed several industrial biotechnologies for production of pharmaceuticals and fine chemicals. In 2018, he was selected into the National "Ten Thousand Talents Program".