Increasing 5-aminolevulinic acid production in engineered Corynebacterium glutamicum via metabolic flux perturbation

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5-aminolevulinic acid (ALA) is an important metabolite for various biological processes and has been increasingly used in agricultural and medical fields. By considering the innate capability to overproduce glutamate, Corynebacterium glutamicum was chosen as a host for the biological production of ALA. For this purpose, mutated HemA (glutamyl t-RNA reductase) from Salmonella typhimurium was expressed to confer ALA production in C5 pathway hence utilizing the endogenous glutamate production route. Cultivation of the recombinant strain produced about 204 mg/L of ALA after 48 hours. Co-expression of HemL (glutamate-1-semialdehyde aminotransferase) further increased ALA concentration up to 457 mg/L which denoted a 25.9-fold increase over the control strain (pMT-Trc vector, 17 mg/L of ALA). Effects of metabolic perturbation on ALA production were performed of which addition of iron-chelating 2,2'-dipyridyl led to 529 mg/L of ALA being produced. Addition of glutamate-enhancing penicillin G gave out the highest ALA yield of about 584 mg/L of ALA. The results obtained from this study thus demonstrated the potential of developing C. glutamicum strain for the biological production of ALA via metabolic engineering.