Bioengineering Seminar 生物工学セミナー

17th Lecture on Molecular Engine 第17回発動分子科学セミナー 3rd WRHI SatelliteLab Seminar 第3回WRHIサテライトラボセミナー

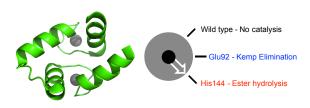
Prof. Ivan V. Korendovych

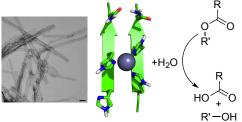
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De Novo Design of Protein and Peptide Catalysts.

Abstract: Design of a novel catalytic function in proteins and peptides, apart from its inherent practical value, is important for fundamental understanding of origins of enzymatic activity. Two applications of a minimalistic approach to design of artificial catalysts will be presented.





- •Strategic introduction of single mutations is sufficient to confer catalytic activities (Kemp elimination, ester hydrolysis, retroaldol reaction) onto calmodulin, a non-enzymatic protein. The catalytic efficiencies of the resulting allosterically regulated catalysts are on par with those of the best computational approaches. Directed evolution allowed for further improvement of catalysts' efficiency.
- •We designed a series of 7-residue peptides that self-assemble into amyloid-like fibrils to act as metal-dependent esterases and oxidases. Metal ions, help stabilize the fibril formation, while also acting as cofactors to catalyze chemical reactions. The resulting catalytic amyloids show efficiency that rivals that of some enzymes by weight. These results indicate that amyloid fibrils are able to not only catalyze their own formation they also can catalyze chemical reactions. Thus, amyloids might have served as intermediates in the evolution of modern-day enzymes.

References

Nature Chem. **2014**, 6, 303-309; ACIE **2013**, 52, 6246-6249; JACS **2015**, 137, 14905, J. Biol. Inorg. Chem. **2013**, 18, 411-418; PNAS **2011**, 108, 6823-6827; Protein Sci. **2015**, 24, 561-570; PNAS **2017**, 114, 6191-6196; ACS Cat. **2018**, 8, 59-62.

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Hub Initiative

