

CHEMOENZYMATIC SYNTHESIS AND BIOLOGICAL FUNCTIONS OF COMPLEX GLYCANS AND GLYCOCONJUGATES

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Carbohydrates are involved in a wide range of physiological and pathological processes. Due to the complexity and diversity of carbohydrates, it is difficult to isolate a large number of structurally well-defined glycans and glycoconjugates in high purity and sufficient quantities from natural sources, which results in poor understanding of their biological roles and structure-function relationships. To address the issue, we have developed an innovative diversity-oriented chemoenzymatic platform by integrating chemical synthesis and enzyme-catalyzed diversification. This platform enables the efficient synthesis of a wide variety of complex carbohydrates, including sulfated ganglioside glycans [1], glycosphingolipids [2,3], and *O*-glycopeptides [4,5]. Furthermore, we have employed high-throughput glycan microarray and surface plasmon resonance technologies to investigate the structure-binding relationships of various carbohydrates. Our research has unveiled the specific recognition modes of different glycan sequences towards disease-related proteins, thereby providing new opportunities for biomedical discovery and the development of carbohydrate-based therapeutics.

References:

- 1. Z. Xu, Y. Liu, J. Liu, W. Ma, Z. Zhang, D. G. Chapla, L. Wen, K. W. Moremen, W. Yi, T. Li, *Nat. Chem.* **2024**, *16*, 881-892
- 2. Y. Ma, Y. Liu, C. Cao, J. Peng, Y. Jiang, T. Li, J. Am. Chem. Soc. 2025, 147, 6974-6982.
- 3. Y. Liu, M. Yan, M. Wang, S. Luo, S. Wang, Y. Luo, Z. Xu, W. Ma, L. Wen, T. Li, ACS Cent. Sci. 2024, 10, 417-425
- 4. W. Ma, Y. Deng, Z. Xu, X. Liu, D. G. Chapla, K. W. Moremen, L. Wen, T. Li, *J. Am. Chem. Soc.* **2022**, *144*, 9057-9065
- 5. W. Ma, J. Luo, H. Liu, Q. Du, T. Hao, Y. Jiang, Z. Huang, L. Lan, Z. Li, T. Li, *Angew. Chem. Int. Ed.* **2025**, *64*, e202424312.