

## Nanozyme-enabled nanodecoys: A new strategy for fighting urinary tract infections

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## Urinary tract region

Urinary tract infections (UTIs), common bacterial infections in communities and medical facilities, are mainly mediated by FimH. The glycan sites of the



uromodulin protein play a crucial role in protecting against UTIs by interacting with FimH. A bioinspired approach using glycan-FimH interactions may effectively reduce bacteria through an antiadhesive mechanism, thereby curbing bacterial resistance. However, typical antiadhesive therapy alone fails to address the excessive reactive oxygen species and inflammatory response during UTIs. To bridge this gap, antioxidant nanozymes with antiadhesive ability were developed as nanodecoys to counter bacteria and inflammation. Specifically, ultrasmall dextran-coated ceria (DEC) was engineered to address UTIs, with dextran blocking FimH adhesion and ceria exhibiting anti-inflammatory properties. DECs, metabolizable by the kidneys, reduced bacterial content in the urinary tract, mitigating inflammation and tissue damage. In murine models, DECs successfully treated acute UTIs, repeated infections, and catheter-related UTIs. This dual approach not only highlights the potential of nanozymes for UTIs but also suggests applicability to other FimH-induced infections in the lungs and bowels, marking a significant advancement in nanozyme-based clinical approaches. Credit: ACS Nano (2024). DOI: 10.1021/acsnano.3c12783

Urinary tract infections (UTIs), affecting millions worldwide, are predominantly caused by uropathogenic Escherichia coli (UPEC). These infections are characterized by bacterial adhesion and colonization in the urinary tract, evading host immune responses. Researchers from Nanjing University have recently reported a new approach to combating UTIs through the development of bioinspired nanozymes acting as nanodecoys.

The nanozymes, designed to mimic the function of uromodulin (UMOD), a natural defense mechanism against bacterial invasion, offer a promising solution. By incorporating dextran onto ceria nanoparticles, the nanozymes simulate UMOD's glycans, effectively obstructing the adhesion of UPECs.

The bioinspired nanozymes exhibit multifaceted functionalities,



including anti-inflammatory and anti-adhesive properties, which position them as potential game-changers in UTI treatment. Through their unique design, the nanozymes can scavenge reactive nitrogen and oxygen species (RNOS) generated during infection-induced inflammation.

Additionally, they prevent <u>bacterial adhesion</u> to host cells and abiotic surfaces, reducing bacterial colonization. The nanozymes' ability to alleviate inflammation and mitigate <u>tissue damage</u> offers a comprehensive approach to UTI management.

In <u>preclinical studies</u> utilizing mouse models, the nanozymes demonstrated promising results in treating acute UTIs, repeated infections, and catheter-associated UTIs. The bioinspired nanozymes hold significant potential for clinical translation by effectively reducing bacterial colonization and inflammation in the urinary tract.

This innovative approach not only addresses the challenges posed by traditional antibiotic therapies but also offers a safer and more comprehensive strategy for managing UTIs and related complications.

The paper is **<u>published</u>** in the journal ACS Nano.

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