

# Polydopamine-coated selenium nanoparticles as a stable and reusable catalyst for tuneable nitric oxide generation

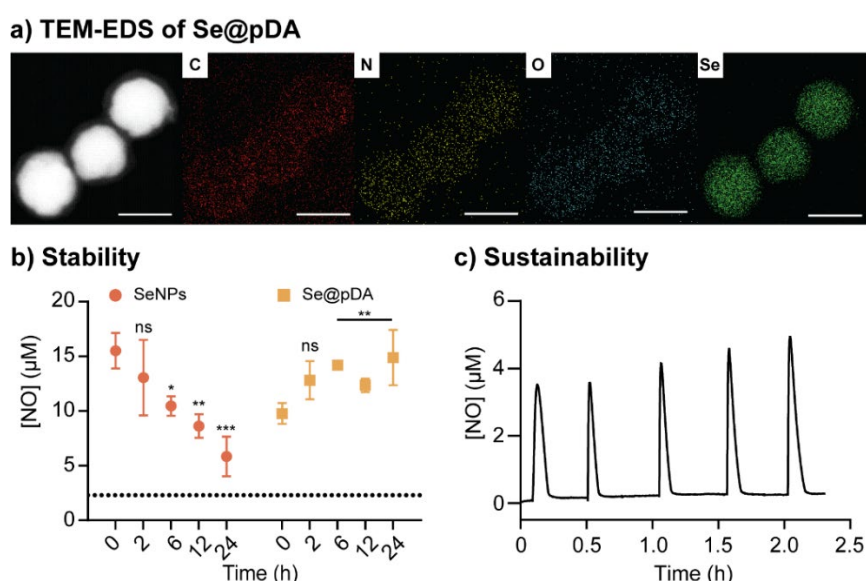
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Nitric oxide (NO) is a signaling molecule involved in a wide range of physiological and pathological processes. Its effects are concentration dependent, promoting cell survival and proliferation at low concentrations (nM), or exhibiting antibacterial/anticancer properties at higher concentrations ( $\mu\text{M}$ ).<sup>1</sup> The significant role of NO in neuroscience, physiology, and immunology has promoted the development of NO delivery strategies. However, a key limitation of NO delivery is its short half-life (5s), which restricts its diffusion distance in human tissue. To address this issue, catalytic approaches have emerged, employing nanomaterials as catalysts to decompose endogenous NO donor S-nitrosoglutathione (GSNO) to generate NO *in-situ*.

Selenium, an essential element in various enzymes/proteins, has been found to catalytically generate NO from NO donors.<sup>2,3</sup> Our previous study established that selenium nanoparticles (SeNPs) are highly efficient NO-generating catalysts with low cytotoxicity. However, the stability of SeNPs under physiological conditions requires improvement. To address this issue, we modified SeNPs with a polydopamine (pDA) coating, creating core-shell nanoparticles known as Se@pDA NPs. These Se@pDA NPs exhibit increased stability compared to SeNPs in PBS buffer for up to 24 hours. Furthermore, the pDA coating overcomes the pH-dependent NO generation limitation of SeNPs, enabling NO delivery across physiological conditions ranging from pH 5.5 to 8.5, thus broadening its potential therapeutic applications. Considering the concentration-dependent nature of NO's effect, achieving tunable NO generation is crucial. This can be accomplished by varying the thickness of the pDA coating. Additionally, our study investigated the long-term stability and sustainability of Se@pDA NPs, demonstrating their potential as a stable and controlled platform for NO-induced therapeutic applications.



**Figure 1:** a) TEM-EDS of Se@pDA NPs. b) Cumulative NO generation catalysed by SeNPs or Se@pDA NPs, particles were incubated in PBS 7.4 for 0, 2, 6, 12, and 24 hours followed by the addition of GSNO (50  $\mu\text{M}$ ) (dashed line represents GSNO control). c) Sustainable NO generation catalysed by Se@pDA NPs in the presence of GSH (1 mM).

## References

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