### This ESI replaces the version published on 16<sup>th</sup> May 2022 due to some errors, the scientific

content remains unchanged.

# Cisplatin and Oleanolic acid Co-loaded pH-Sensitive CaCO<sub>3</sub> Nanoparticles for Synergistic

### Chemotherapy

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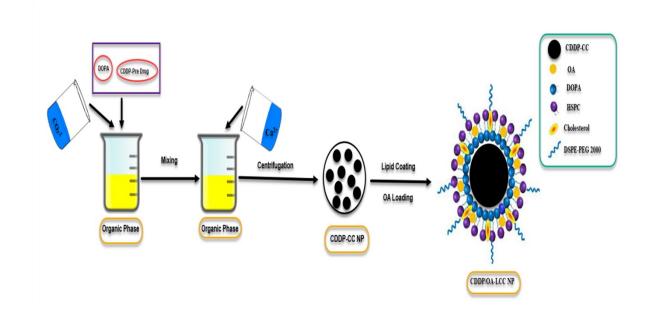
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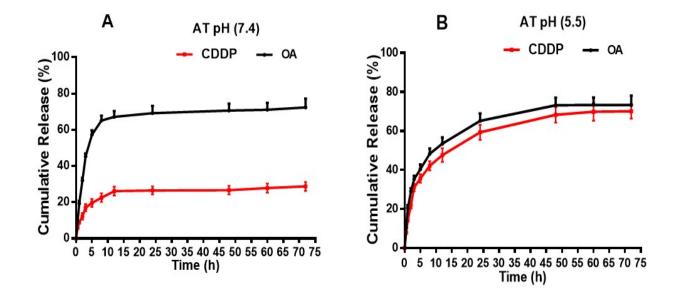
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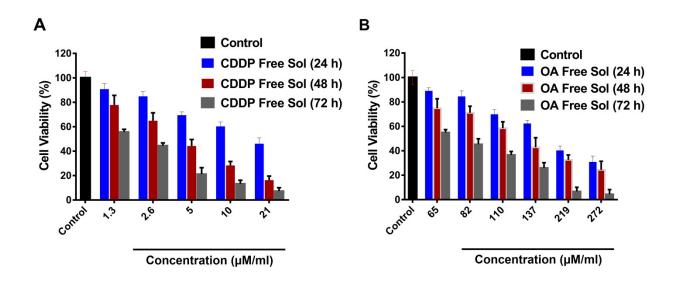
# **Supplementary Information**



**Supplementary Fig. S1.** Schematic illustration of the formulation of lipid coated cisplatin/oleanolic acid co-loaded calcium carbonate nanoparticles (CDDP/OA-LCC NPs).<sup>1</sup>

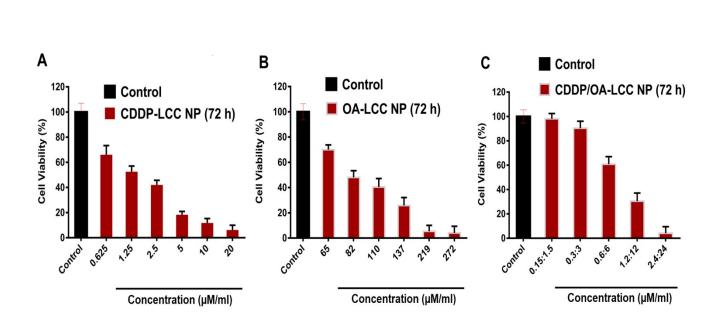


**Supplementary Fig. S2.** In vitro drug release profiles of CDDP and OA from the CDDP/OA-LCC NPs in PBS (72 h). (A) At pH 5.5; (B) At pH 7.4.<sup>1</sup>

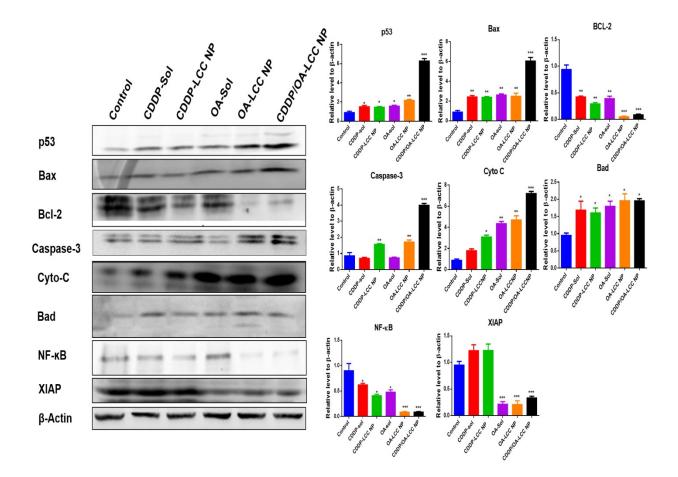


Supplementary Fig. S3. Cytotoxicity assay of free CDDP and free OA against HepG2 cells.

1



Supplementary Fig. S4. Cytotoxicity assay of the nanoparticles against HepG2 cells (72 h). (A) CDDP-LCC NPs MTT assay; (B) OA-LCC NPs MTT assay; (C) CDDP/OA-LCC NPs MTT assay with fixed ratios. Data presented as mean  $\pm$  SD, n=5.<sup>1</sup>



**Supplementary Fig. S5.** Western blot analysis of protein levels (p53, Bax, Bad, Cyto-C, caspase-3, NF-κB, Bcl-2 and XIAP) after treating HepG2 cells with CDDP-Sol, CDDP-LCC NP,

OA-Sol, OA-LCC NP, CDDP/OA-LCC NP *in vitro*.  $\beta$ -actin was used as a loading control. Quantification of protein level using Image J. Data presented as mean  $\pm$  S.D. (n=3).<sup>1</sup>

# **Reference**

M. W. Khan, P. Zhao, A. Khan, F. Raza, S. M. Raza, M. Sarfraz, Y. Chen, M. Li, T. Yang and X. Ma, *Int. J. Nanomed.*, 2019, 14, 3753-3771