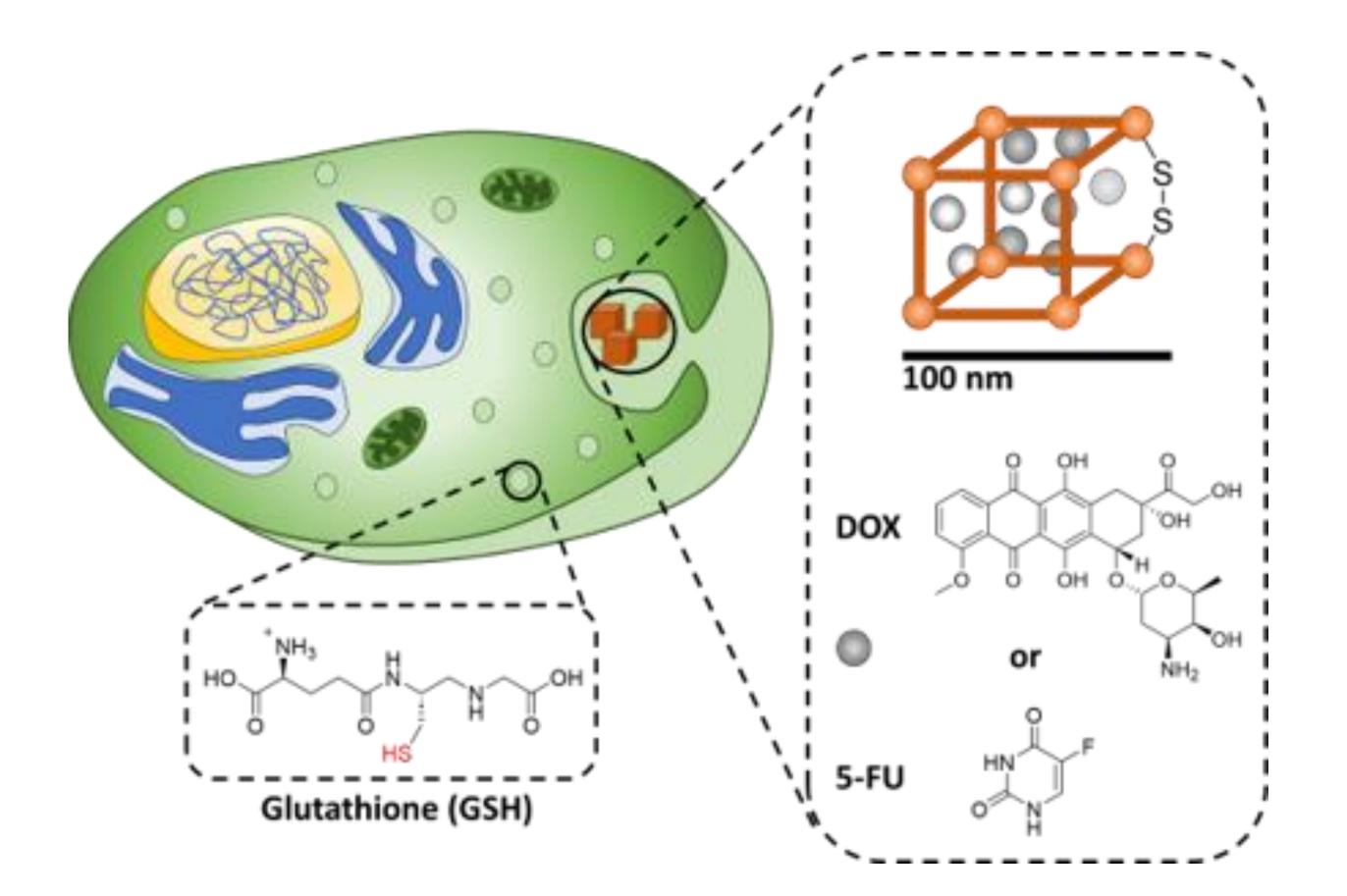
Glutathione-Sensitive Nanocarrier Metal Organic Frameworks for the Synergistic Release of Chemotherapeutics <u>Anthony Traficante, Pat Strobel, Nate Oldenhuis, Aylin Aykanat</u> Chemistry Department, University of New Hampshire, Durham, NH 03824

Metal Organic Frameworks (MOF)

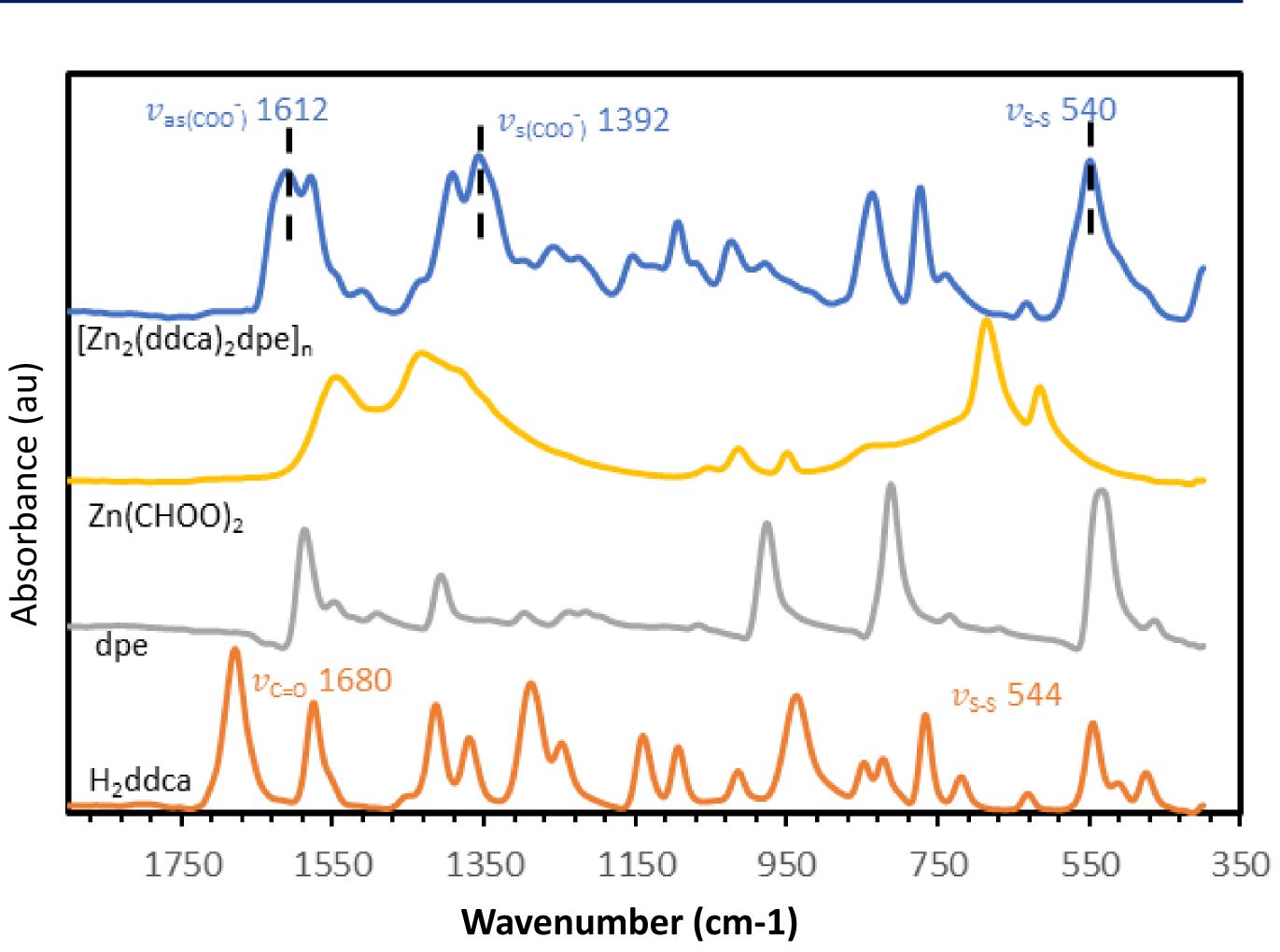
Metal Organic Frameworks, known as MOFs are porous, crystalline structures that are made up of metal centers bonded to organic linkers. Depending on the metal ion and linker, the size of the pores and geometry of the molecule can vary greatly. These pores can absorb and encapsulate other molecules, depending on both the size of the molecule and the pores, as well as potential intermolecular interactions between the molecule and framework. By changing the linkers, you can also alter the functionality of the MOF. This gives MOFs a wide variety of applications, such as filtration and drug delivery.

Redox-Responsive Nanoparticles



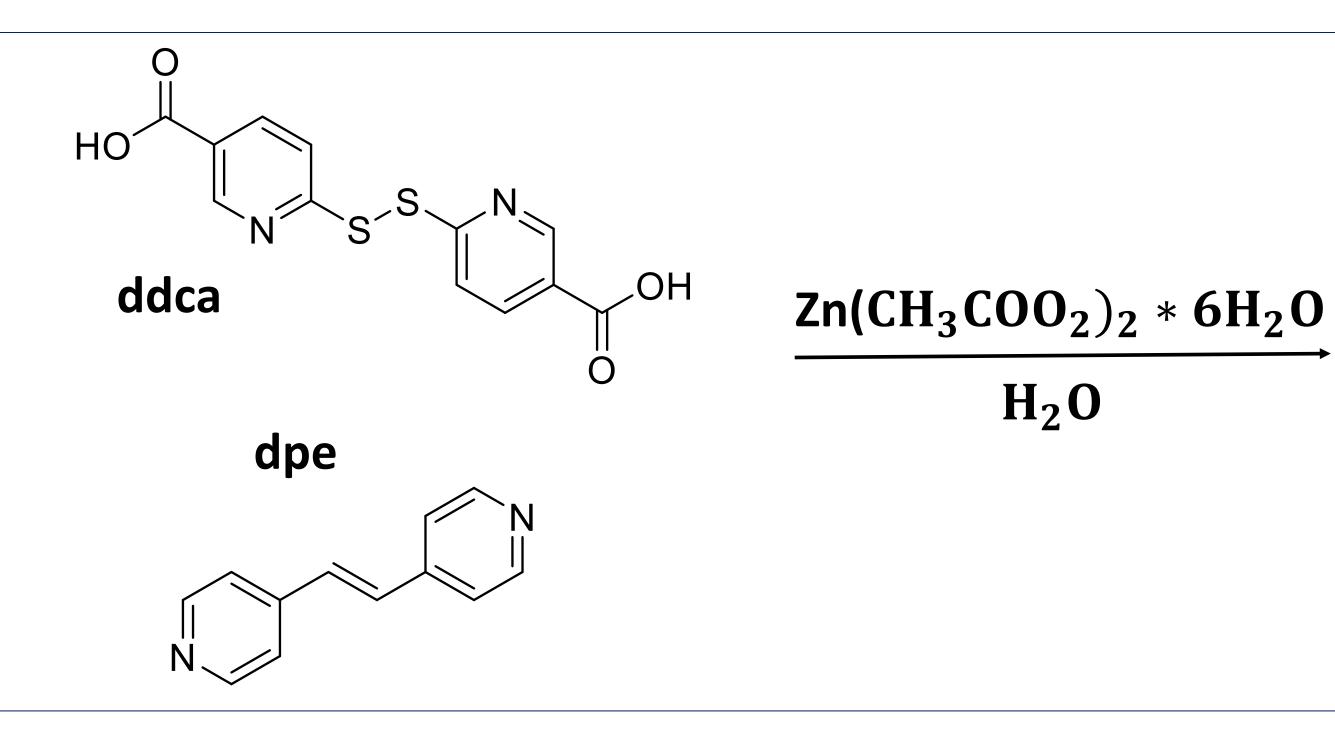
How MOF containing chemotherapeutics Doxorubicin and 5-flurorouracil is broken down inside of cells using glutathione, which is up to 1000x more present in cancer cells

IR Spectroscopy



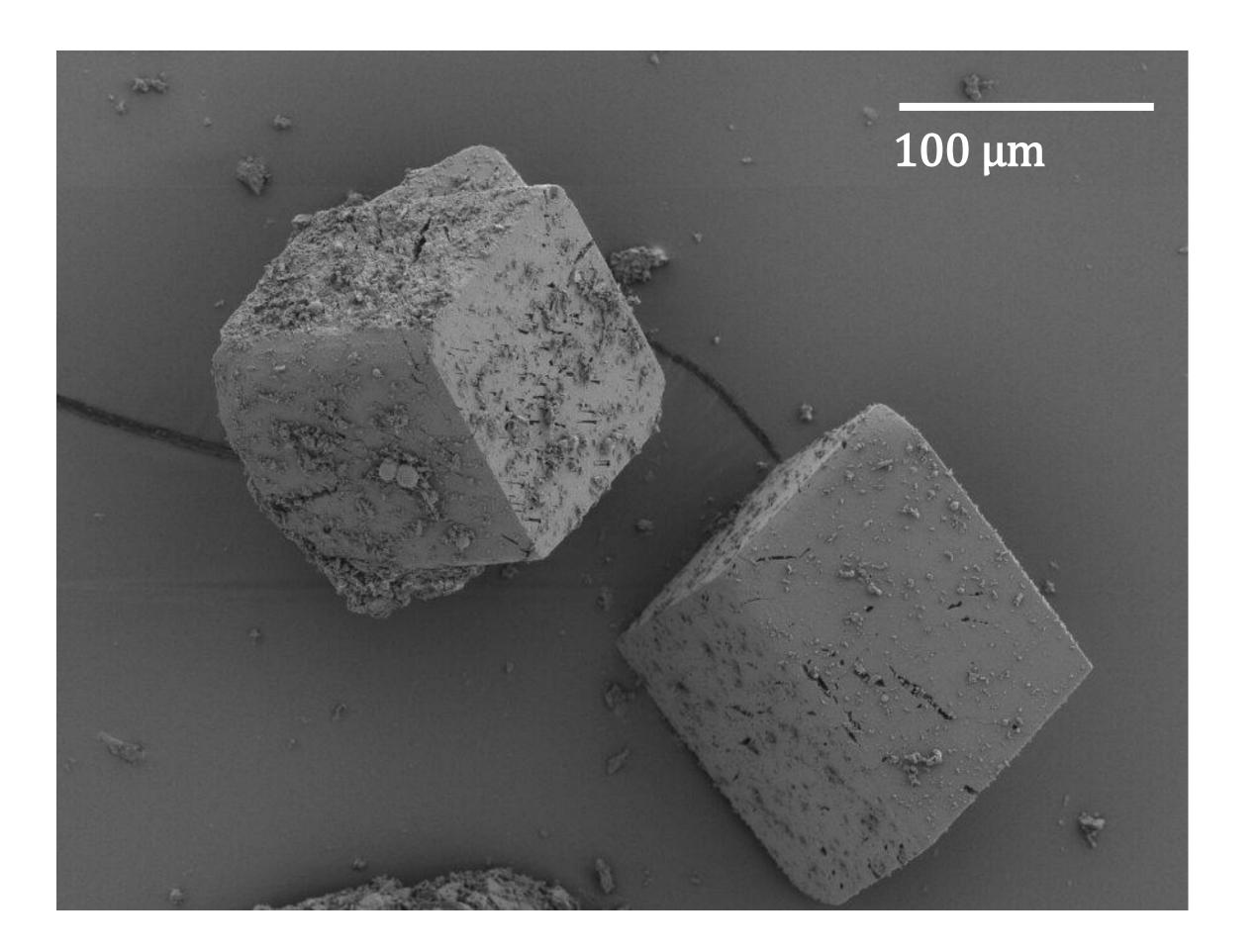
FTIR spectra of differed reaction times showing the presence of carboxylate and disulfide in our MOF

Synthesis of Zinc-based MOF: $[Zn_2(ddca)_2(dpe)]_n * nH_2O$

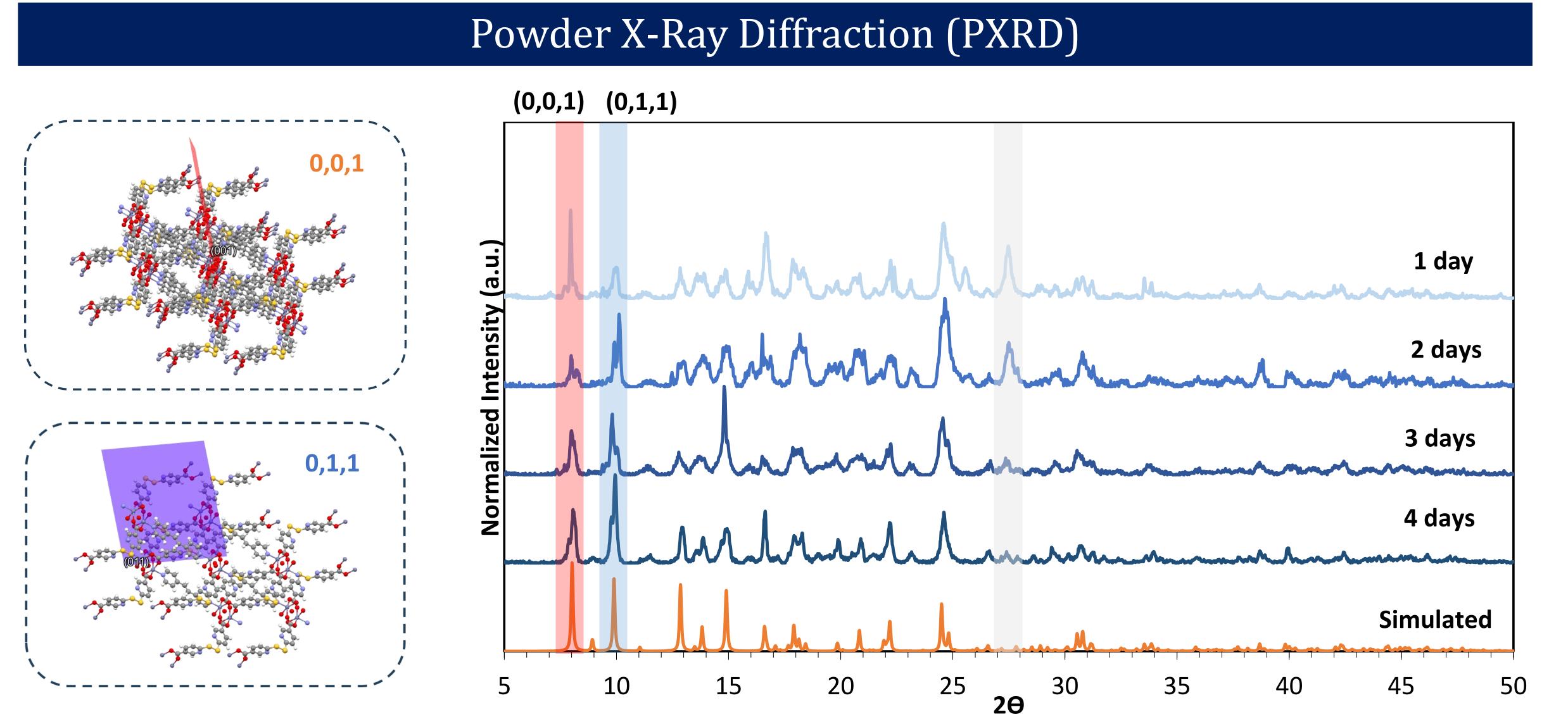


 H_2O

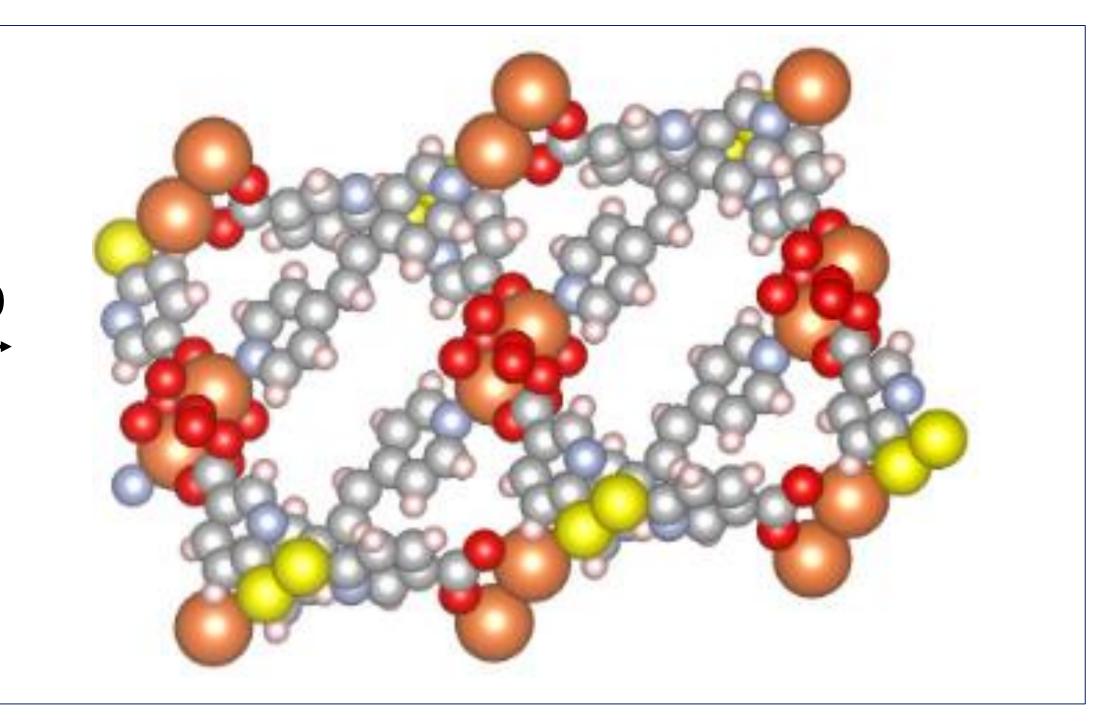
SEM Images

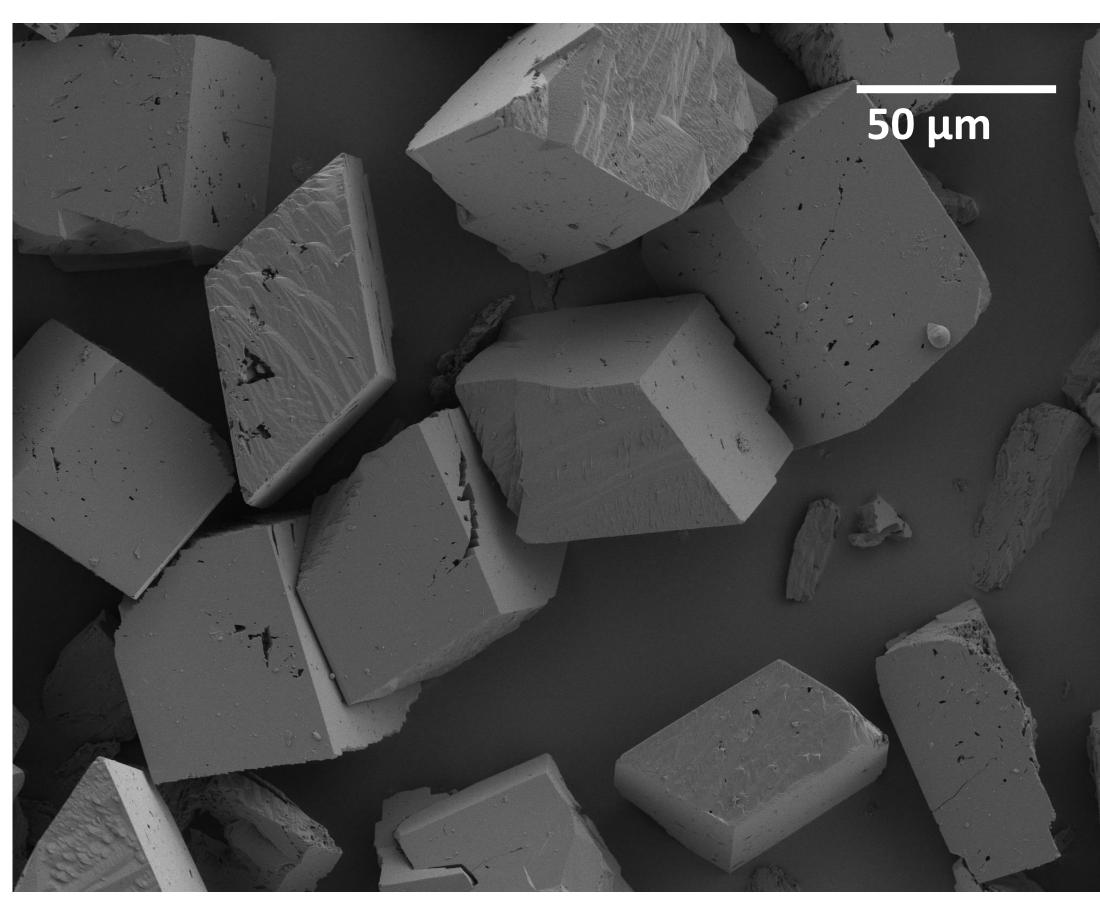


Scanning electron microscope (SEM) images of $[Zn_2(ddca)_2(dpe)]_n^* nH_2O$ before and after DTT degradation studies

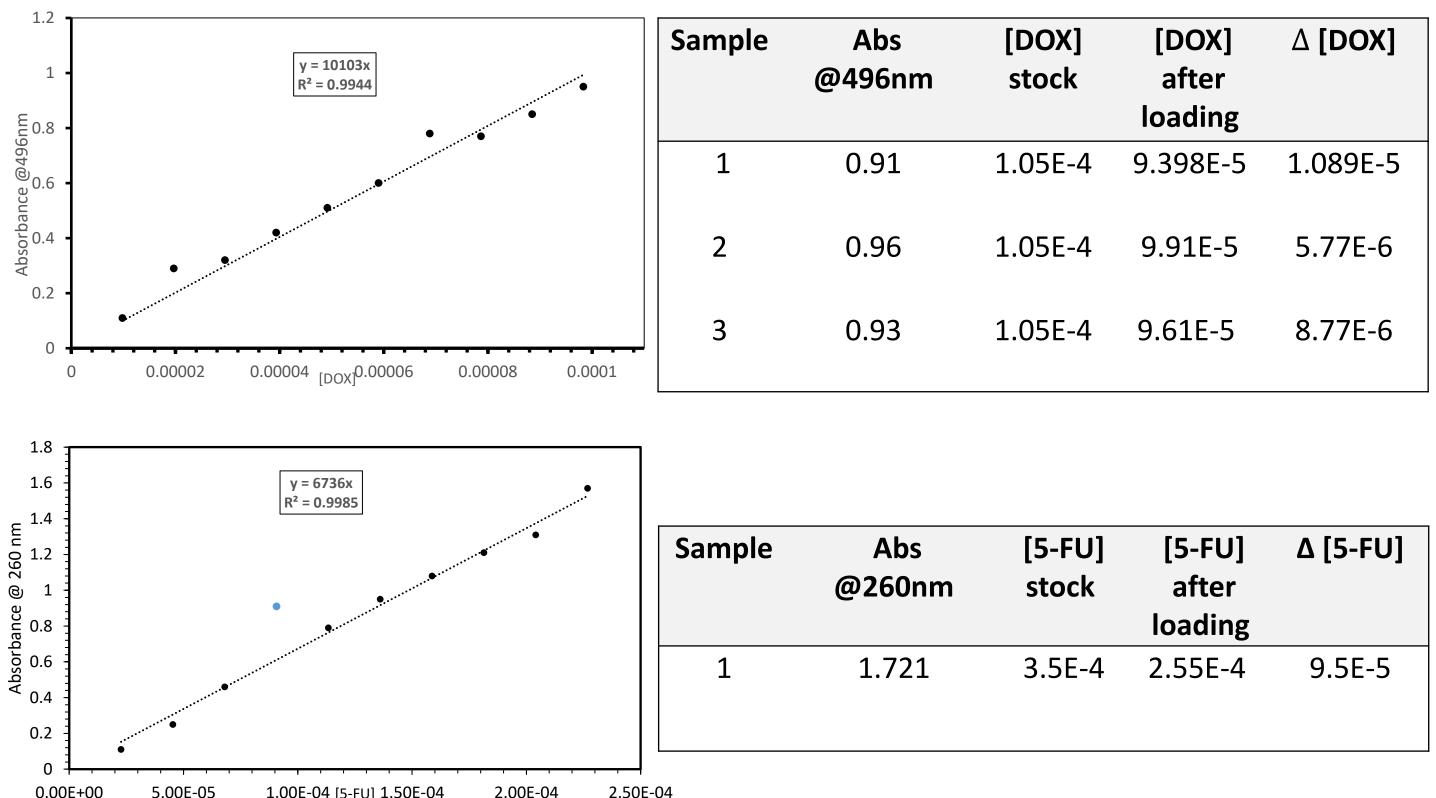


PXRD of $[Zn_2(ddca)_2(dpe)]_n^* nH_2O$ showing planes in the molecule and the corresponding peak values as well as evidence of longer synthetic timing correlating to loss of starting material into product





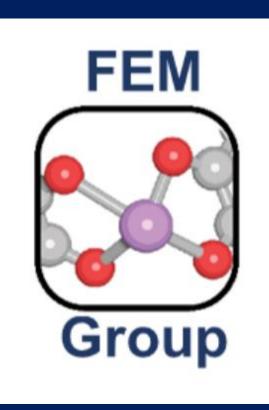
Drug Loading of Doxorubicin into $[Zn_2(ddca)_2(dpe)]_n * nH_2O$; the DOX solution lightens up as the drug is loaded into MOF because of a decrease in concentration of doxorubicin in the solution



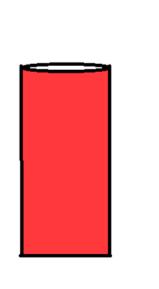


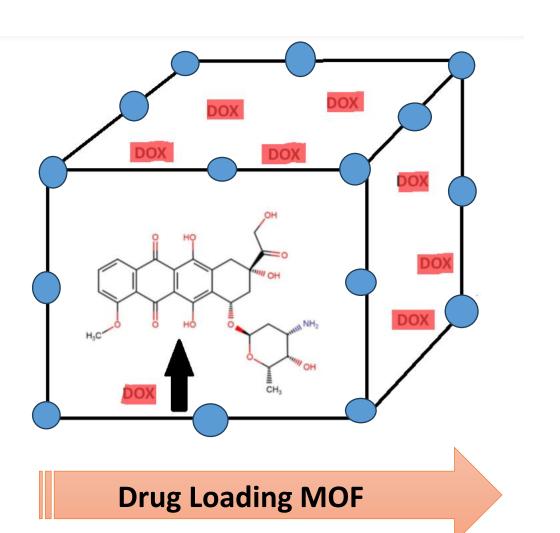
Special thanks to Prof Aylin Aykanat, Prof Nate Oldenhuis, Pat Strobel, Andrew Jenkins, Forde Kates, Carmelina Minico, Lucas Laventure, Elisabeth Gotschlich, and all others that have helped in this project





Chemotherapeutic Loading







stock solution

Dox solution after add Zn-MOF to load drug

Sample	Abs @260nm	[5-FU] stock	[5-FU] after loading	Δ [5-FU]
1	1.721	3.5E-4	2.55E-4	9.5E-5

Calibration curve of doxorubicin hydrochloride and 5-fluorouracil stock solution concentration and comparing before and after concentration

of stock solution with the addition of $[Zn_2(ddca)_2(dpe)]_n * nH_2O$ to load pores with chemotherapeutic

Acknowledgements

References

(1)Chen, X.; Qiao, Y.; Gao, L.; Cui, H.; Zhang, M.; Lv, J.; Hou, X. Synthesis, Structure, and Characterization of Two Zn/Cd Coordination Polymers with Flexible Disulfide Carboxylate Derivatives. Zeitschrift anorg allge chemie 2013, 639 (2), 403-408. <u>https://doi.org/10.1002/zaac.201200416</u>.

(2) Kirlikovali, K. O.; Hanna, S. L.; Son, F. A.; Farha, O. K. Back to the Basics: Developing Advanced Metal–Organic Frameworks Using Fundamental Chemistry Concepts. ACS Nanosci. Au **2023**, 3 (1), 37–45. https://doi.org/10.1021/acsnanoscienceau.2c00046.