

Supporting Information

Activation of platinum(IV) prodrugs by cytochrome *c* and characterization of the protein binding sites

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Figure SI1. ¹H,¹⁵N-HSQC spectra of ¹⁵N-labelled Pt(IV) complex treated with excess NADH at different concentrations of cyt *c*.

Figure SI2. ¹H NMR spectra of oxidized and reduced cyt *c*.

Figure SI3. HPLC total ion current chromatograms of tryptic digest of free cyt *c* and cyt *c* reacted with cisplatin.

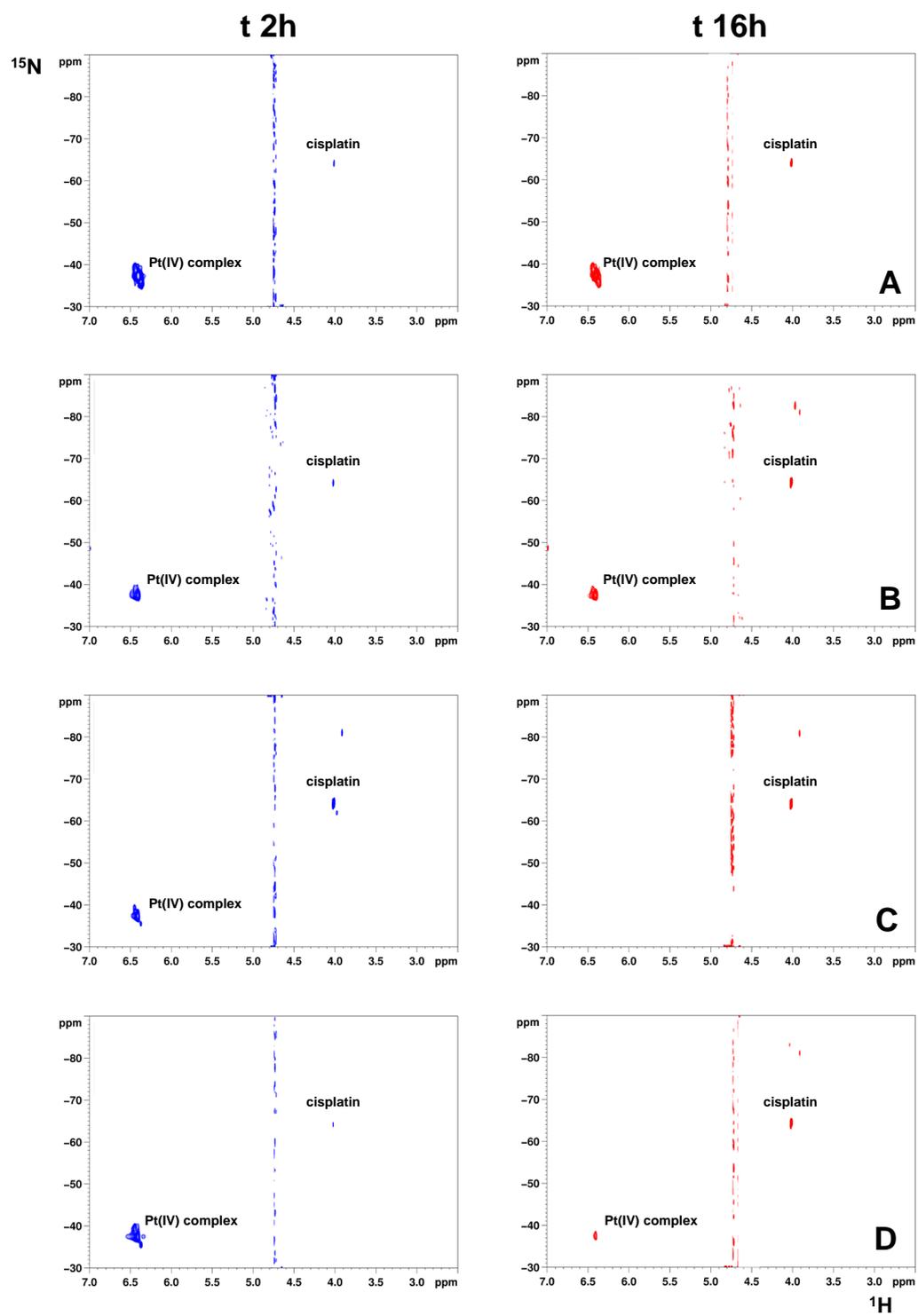


Figure S11. ^1H , ^{15}N -HSQC spectra of ^{15}N -labelled Pt(IV)-complex ($700\ \mu\text{M}$) treated with NADH ($7\ \text{mM}$) at pH 5.8 ($25\ \text{mM}$ phosphate buffer) and $37\ ^\circ\text{C}$ (A). Same composition of the previous solution but containing 700 (B), 70 (C) or $7\ \mu\text{M}$ cyt *c* (D). The spectra were recorded after 2 h and 16 h of incubation.

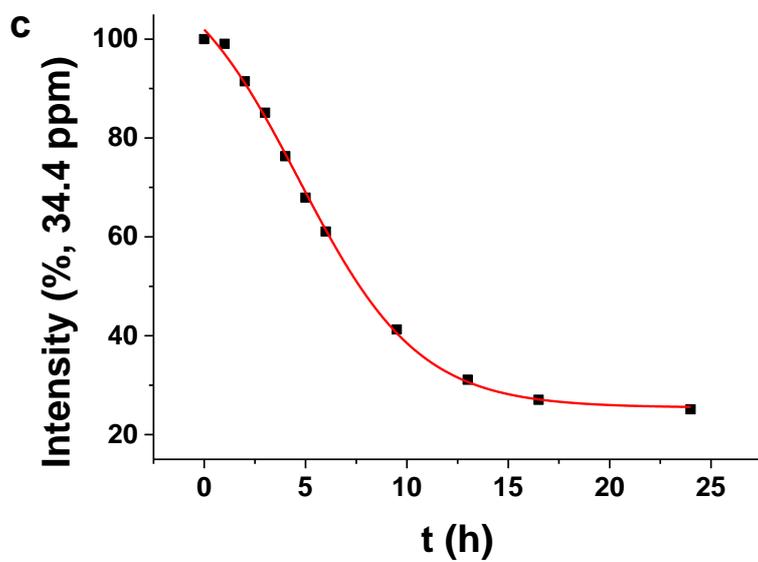
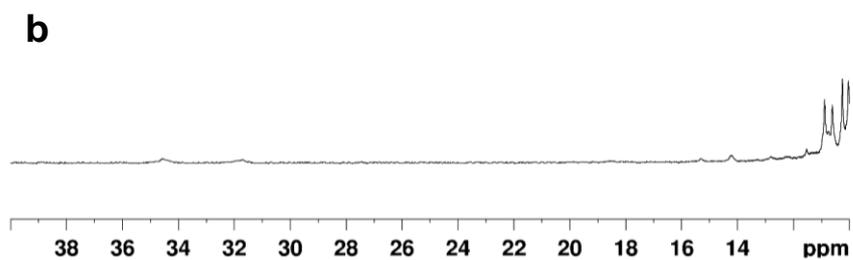
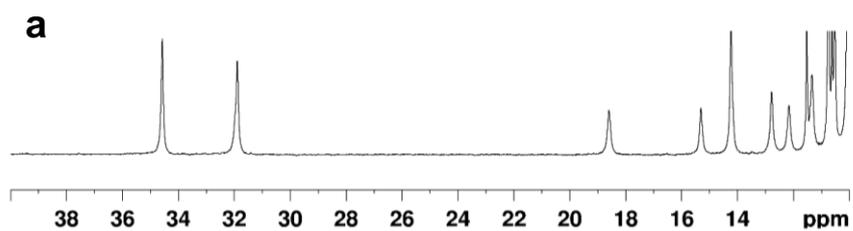


Figure SI2. ^1H NMR spectrum of oxidized (a) and reduced cyt *c* (b) in the region ranging from 10 to 40 ppm. Kinetics of cyt *c* reduction by NADH obtained by plotting the intensity of the ^1H NMR peak at 34.4 ppm as a function of time (c).

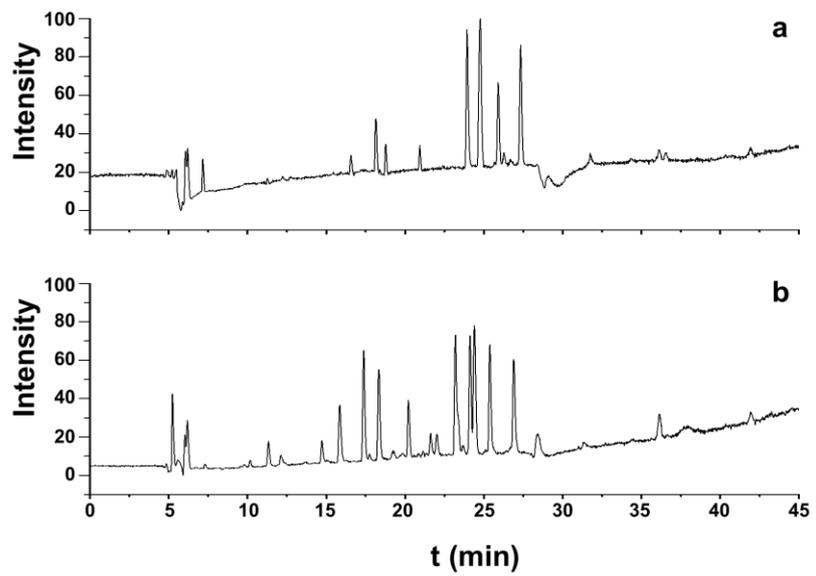


Figure SI3. HPLC total ion current chromatograms of tryptic digest of free *cyt c* (a) and *cyt c* reacted with a 10-fold excess of cisplatin (b).