

Supporting information for

The leaving group in Au(I)-phosphine compounds dictates cytotoxic pathways in CEM leukemia cells and reactivity towards a Cys₂His₂ model Zinc Finger

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Growth Inhibition Profiles

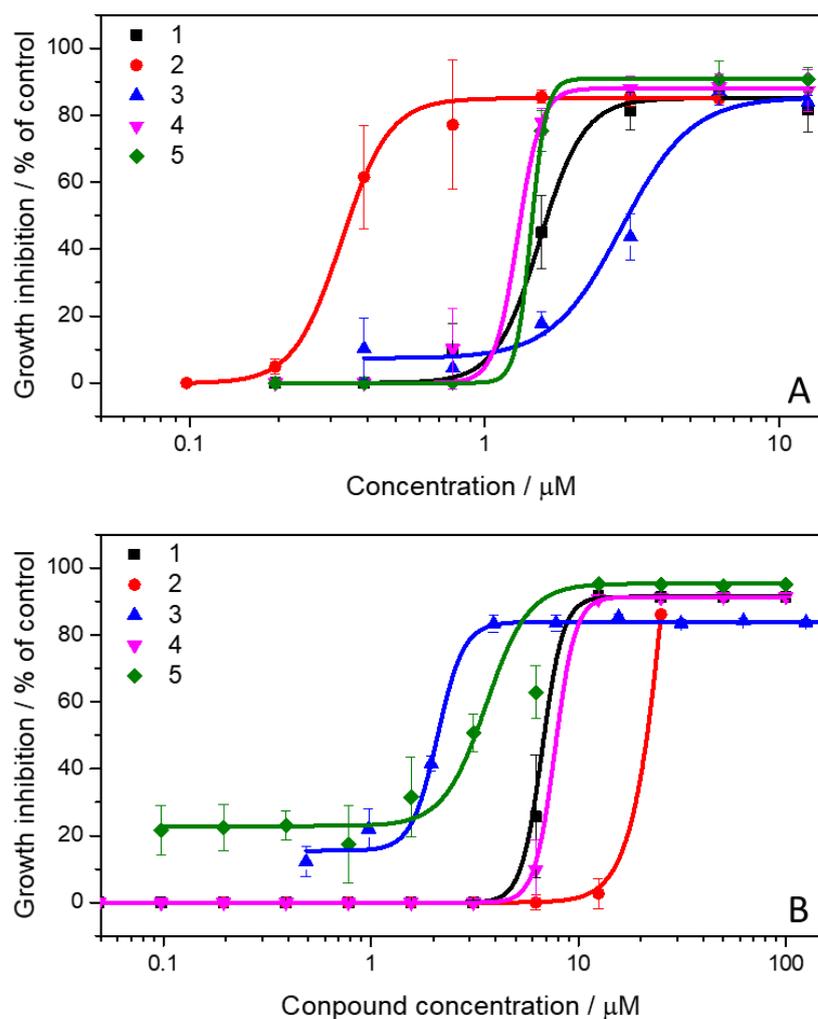


Figure S1. Growth inhibition profile of compounds of A. CEM and B. HUVEC cell lines exposed to compounds 1. $[\text{AuCl}(\text{Et}_3\text{P})]$, 2. $[\text{Au}(\text{dmap})(\text{Et}_3\text{P})]^+$, 3. Auranofin, 4. $[\text{AuCl}(\text{Cy}_3\text{P})]$ and 5. $[\text{Au}(\text{dmap})(\text{Cy}_3\text{P})]^+$.

Cellular morphology

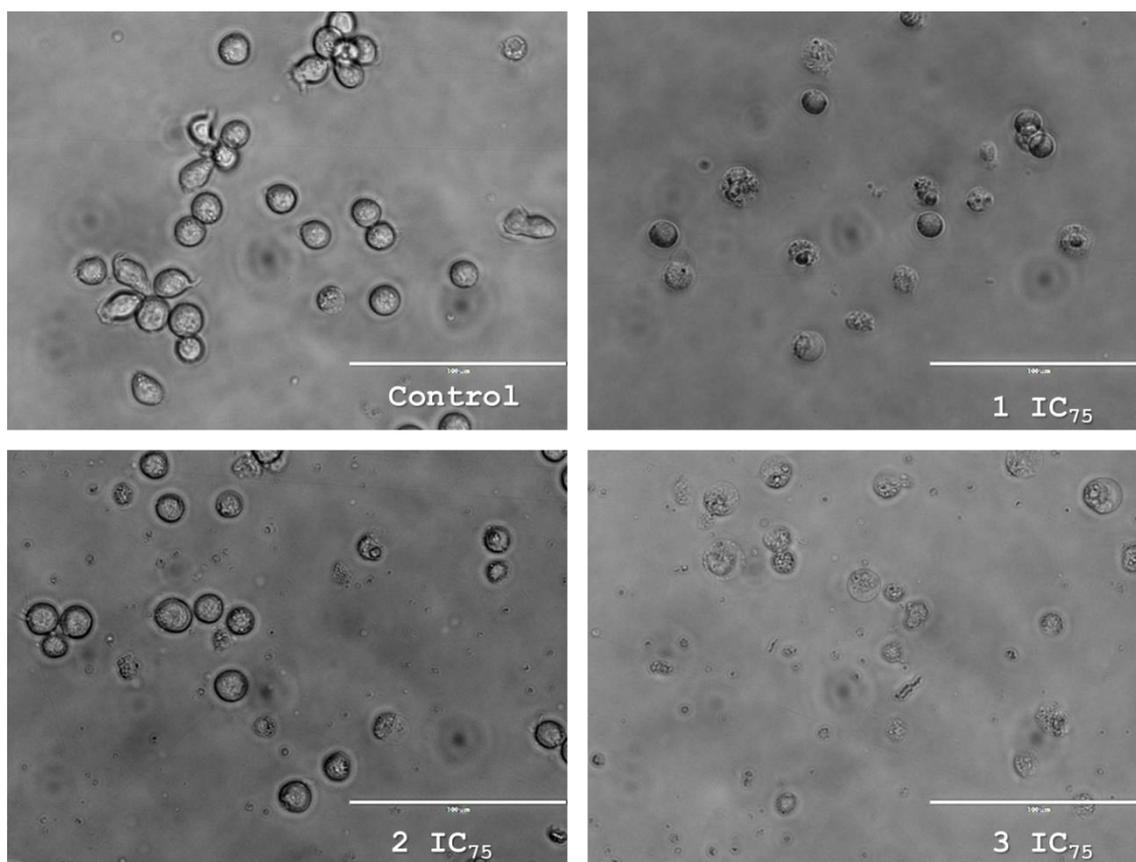


Figure S2. Morphology of CEM cell line untreated and treated with IC₇₅ concentrations of compounds [AuCl(Et₃P)] (1), [Au(dmap)(Et₃P)]⁺ (2) and auranofin (3).

Cell cycle analysis

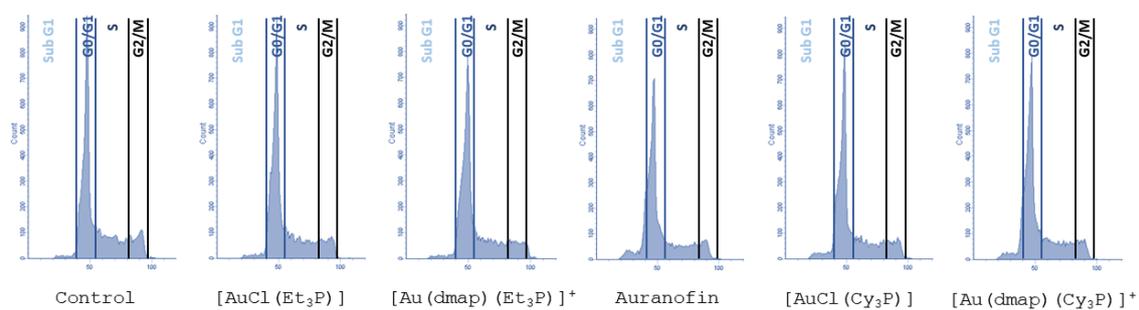


Figure S3. Flow cytometry profile of CEM cell treated with IC₂₅ concentrations for 6 hours.

Collision Induced Dissociation

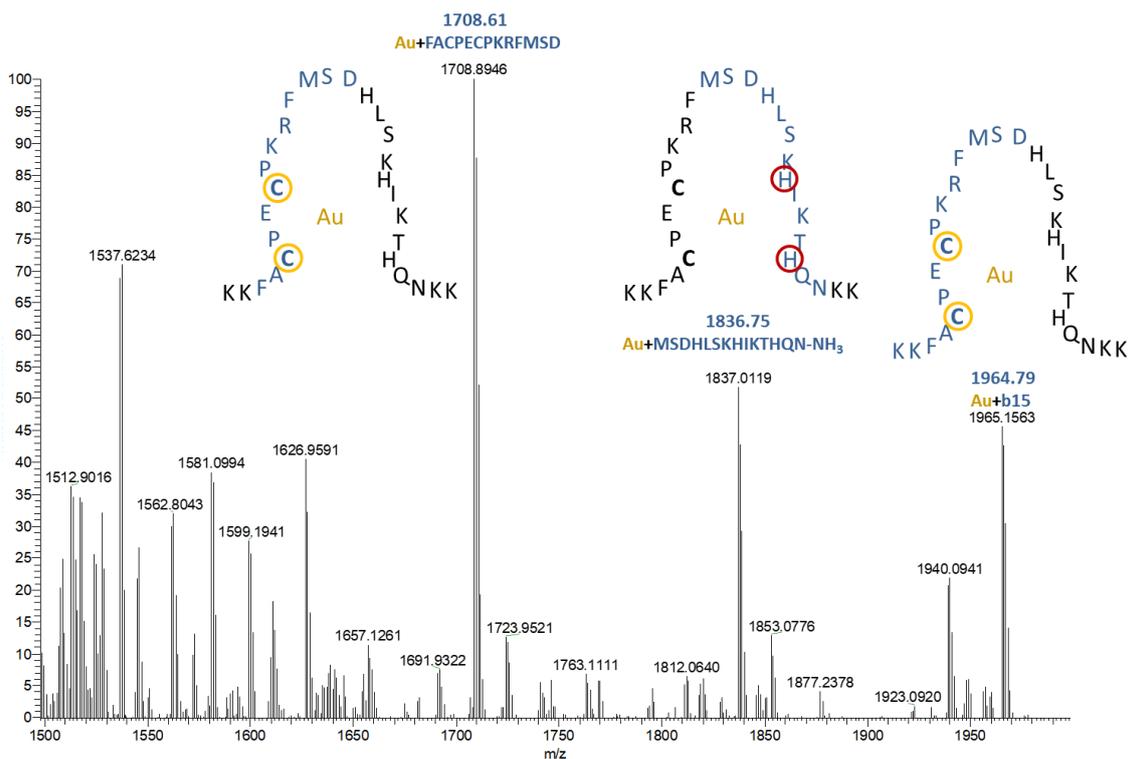


Figure S4. Collision-Induced Dissociation (CID) spectra obtained for the AuF⁴⁺ parent ion 891.9 m/z (3+), product of the reaction between [Au(dmap)(Et₃P)]⁺ and Sp1 ZnF3. Species diagnostic for Cys binding and for His binding are observed. Either one of the Cys residues marked in yellow, and either of the His residues marked in red are expected to bind to Au(I). Theoretical m/z values for each assigned peak are also given. The coordination sphere of the AuF obtained is also Cys(S)-Au-N(His).

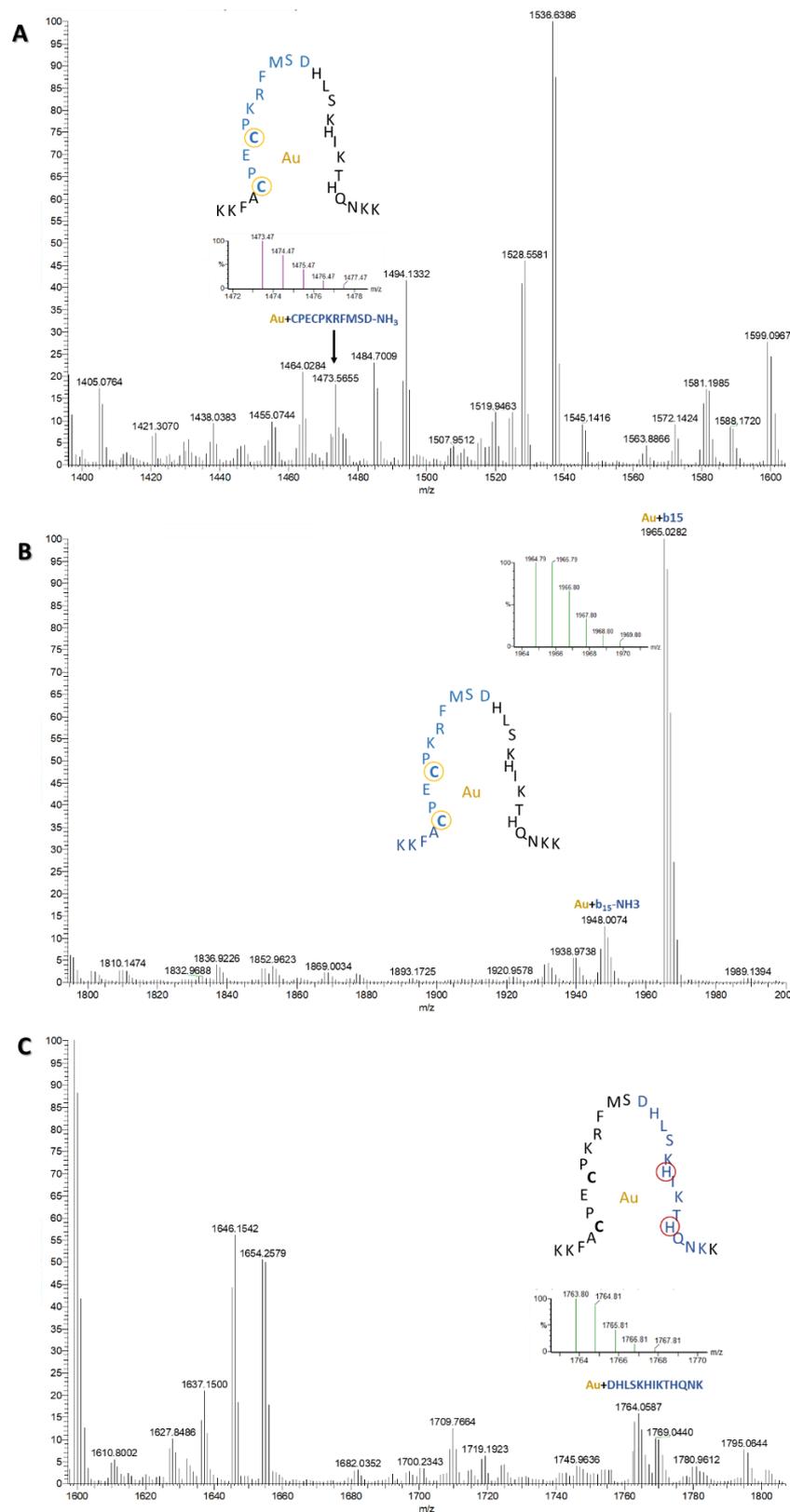


Figure S5. Collision-Induced Dissociation (CID) spectra obtained for the AuF^{3+} parent ion 1188.9 m/z (3+), product of the reaction between $[\text{AuCl}(\text{Et}_3\text{P})]$ and Sp1 ZnF3. A and B show species diagnostic for Cys binding (either one of the Cys residues marked in yellow are expected to bind Au(I)). C shows a fragment diagnostic for His binding, where either of the His residues marked in red are expected to bind Au(I). Theoretical isotope patterns for each assigned peak are given as insets. The coordination sphere of the AuF obtained is Cys(S)-Au-N(His).

Mechanism auranofin-induced apoptosis on CEM cells

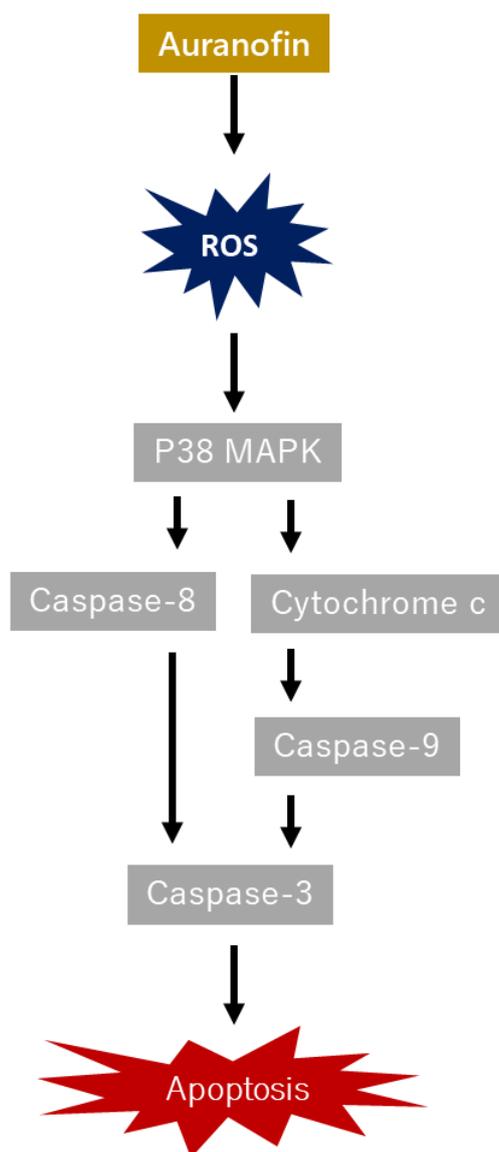


Figure S6. Apoptotic mechanism caused by auranofin on CEM cells. The mechanism is consistent with previous results on HL-60 leukemia cells treated with auranofin.¹

References

- 1 S.-J. Park and I.-S. Kim, *Br. J. Pharmacol.*, 2005, **146**, 506–13.