



**Anti-Cancer N-Heterocyclic Carbene Complexes of
Gold(III), Gold(I) and Platinum(II): Thiol
"Switch-on" Fluorescent Probes, Thioredoxin
Reductase ... Reticulum Targeting Agents
(Springer Theses)**

Taotao Zou

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This thesis focuses on the development of gold- and non-classical platinum-based anti-cancer agents that display distinctively different anti-cancer mechanisms compared to the commonly used cisplatin. These metal complexes contain N-heterocyclic carbene (NHC) ligands which are able to form strong M-C(NHC) bonds, conferring high stability and favorable lipophilicity, reactivity and binding specificity of metal complexes on biomolecules. The author demonstrates significant advances made in anti-cancer gold(III), gold(I) and platinum(II) complexes. Detailed chemical synthesis, in vitro and/or in vivo anti-cancer activities are clearly presented including: (i) a class of Au(III) complexes containing a highly fluorescent N^NN ligand and NHC ligand that simultaneously act as fluorescent thiol "switch-on" probes and anti-cancer agents; (ii) a dinuclear gold(I) complex with a mixed diphosphine and bis(NHC) ligand displaying favorable stability and showing significant inhibition of tumor growth in two independent mice models with no observable side effects; and (iii) a panel of stable luminescent cyclometalated platinum(II) complexes exhibiting high specificity to localize to the endoplasmic reticulum (ER) domain, inducing ER stress and cell apoptosis. These works highlight the clinical potential that gold and platinum complexes offer for cancer treatment.

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