

Solvent effects on reactions of singlet molecular oxygen, $O_2(^1g)$, with antimalarial drugs

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Detection of $O_2(^1g)$ emission, $\lambda_{max}=1270$ nm, following laser excitation and steady-state methods were employed to measure total reaction rate constants, k_T , for the reaction between singlet oxygen and the antimalarial drugs quinine (QU), quinacrine (QC), chloroquine (CQ) and amodiaquine (AQ) in several solvents. Values for k_T range from $0.45 \pm 0.03 \times 10^7$ M⁻¹ s⁻¹ for AQ in benzene to $25.1 \pm 0.88 \times 10^7$ M⁻¹ s⁻¹ for CQ in N, N -dimethylformamide. Analysis of solvent effect on k_T for QU, QC, and CQ by using the LSER formalism indicates that singlet oxygen deactivation by these drugs is accelerated by solvents with large π^* values and hydrogen bond acceptor (HBA) properties and is inhibited by hydrogen bond donors (HBD) solvents. This result support the formation of an exciplex intermediate of charge transfer character, as proposed for reactions of tertiary amines with singlet oxygen, process largely governed by physical quenching. AQ behaves in a different manner. The LSER equation for this drug