

A Homogeneous Catalyst for Selective O₂ Oxidation at Ambient Temperature. Diversity-Based Discovery and Mechanistic Investigation of Thioether Oxidation by the Au(III)Cl₂NO₃(thioether)/O₂ System

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Abstract: A library of inorganic complexes with reversible redox chemistry and/or the ability to catalyze homogeneous oxidations by peroxides, including but not limited to combinations of polyoxometalate anions and redox-active cations, was constructed. Evaluation of library members for the ability to catalyze aerobic sulfoxidation (O₂ oxidation of the thioether, 2-chloroethyl ethyl sulfide, CEES) led to the discovery that a combination of HAuCl₄ and AgNO₃ forms a catalyst that is orders of magnitude faster than the previously most reactive such catalysts (Ru(II) and Ce(IV) complexes) and one effective at ambient temperature and 1 atm air or O₂. If no O₂ but high concentrations of thioether are present, the catalyst is inactivated by an irreversible formation of colloidal Au(0). However, this inactivation is minimal in the presence of O₂. The stoichiometry is R₂S + ¹/₂O₂ → R₂S(O), a 100% atom efficient oxygenation, and not oxidative dehydrogenation. However, isotope labeling studies with H₂¹⁸O indicate that H₂O and not O₂ or H₂O₂ is the source of oxygen in the sulfoxide product; H₂O is consumed and subsequently regenerated in the mechanism. The rate law evaluated for every species present in solution, including the products, and other kinetics data, indicate that the dominant active catalyst is Au(III)Cl₂NO₃(thioether) (**1**); the rate-limiting step involves oxidation of the substrate thioether (CEES) by Au(III); reoxidation of the resulting Au(I) to Au(III) by O₂ is a fast subsequent step. The rate of sulfoxidation as Cl is replaced by Br, the solvent kinetic isotope effect (*k*_{H₂O}/*k*_{D₂O} = 1.0), and multiparameter fitting of the kinetic data establish that the mechanism of the rate-limiting step involves a bimolecular attack of CEES on a Au(III)-bound halide and it does not involve H₂O. The reaction is mildly inhibited by H₂O and the CEESO product because these molecules compete with those needed for turnover (Cl[−], NO₃[−]) as ligands for the active Au(III). Kinetic studies using DMSO as a model for CEESO enabled inhibition by CEESO to be assessed.

Introduction

One of the ultimate challenges in catalysis is to formulate molecules or materials that catalyze useful (selective and reasonably rapid) reactions by O₂ or air under ambient conditions. Such catalysts would have a number of significant uses including the formulation of fabrics, coatings, and other materials that could catalytically clean the air and water by oxidative decontamination of the offending agents using solely the environment itself (air under ambient conditions). Unlike catalytic oxidation processes based on the second most economically and environmentally attractive oxidant, H₂O₂ (and related peroxides), there are almost no catalytic oxidation processes based on the most attractive oxidant, O₂/air, that proceed at a satisfactory rate at room temperature, thus certainly none that proceed rapidly, selectively, and without rapid catalyst decomposition under ambient conditions. Heterogeneous catalytic O₂-based oxidation processes typically are conducted between 270 and 720 °C (e.g., the O₂ oxidation of ethylene to ethylene oxide over Ag/Al₂O₃ is conducted at 270–290 °C and 30 bar O₂,¹ the oxidation of methanol to formaldehyde over 18–19 wt % Fe₂O₃ and 81–82 wt % MoO₃ is conducted at 350–450 °C with a large excess of O₂,¹ and the oxidation

of isobutyric acid to methacrylic acid over polyoxometalates (POMs) is conducted at elevated temperatures and O₂ pressures).^{2–5} The commercially important homogeneous catalytic O₂-based oxidations also all proceed well above ambient temperature. For example, the Mid-Century/Amoco process for oxidation of *p*-xylene to terephthalic acid catalyzed by cobalt and manganese acetates and bromides in acetic acid, the largest scale commercial homogeneous catalytic oxidation, is conducted at 225 °C using 15 atm O₂,^{6,7} and the Wacker oxidation of ethylene to acetaldehyde catalyzed by PdCl₂/CuCl₂ is conducted at 120–130 °C using 4 atm O₂.⁸

Selectivity is a general problem in both stoichiometric and catalytic strictly O₂-based oxidations because these reactions are nearly always dominated by radical chain autoxidation, a chemistry that is intrinsically nonselective and difficult to

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control.^{9–12} By “strictly” we mean that no reducing agent is present to convert O₂ to peroxo species or other forms of reduced oxygen that are both more reactive and more selective.⁹ Nearly all enzyme-catalyzed oxidation processes, including some dioxygenase-catalyzed reactions, involve consumption of a reducing agent (most frequently NADH, NADPH, ascorbate, and thiols). The few cases that involve homogeneous catalytic oxidations strictly by O₂ are noteworthy, but none of these proceed effectively under ambient conditions (rates, selectivities, and stabilities that approach viable ranges).^{10–15}

We report here the discovery of a Au(III)-based homogeneous catalyst for air (O₂)-based oxidations under ambient conditions that is significantly faster than any yet reported. We have chosen to probe this new catalyst using the O₂ oxidation of the thioether mustard analogue, 2-chloroethyl ethyl sulfide (CEES) to the corresponding sulfoxide, a reaction of current interest.^{16–25} Heterogeneous oxidation catalysts based on Au are under considerable scrutiny at present. For CO + 1/2 O₂ → CO₂ catalyzed by Au supported on TiO₂, the catalytic properties of Au depend on the support, the preparation method, and the size of the Au clusters. Optimal activity is achieved when the Au clusters are 3.0 nm in diameter, giving a turnover frequency (TOF) of 0.26 CO₂ (Au site)^{–1} s^{–1}.²⁶ In solution, the stoichiometric oxidation of thioethers by Au(III) is well known, dating from 1901^{27–32} to fairly thorough recent studies,^{32–36} but there is only one report of catalytic oxidation by Au compounds. The

oxidant in this case was nitrate not O₂.³⁷ Given the potential significance of a catalyst for selective and rapid oxidation by O₂ (or air) under ambient conditions, considerable effort has been expended to characterize this newly discovered Au-based catalyst despite its experimental sensitivity and complexity.

Experimental Section

Materials. HAuCl₄, AgNO₃, AgClO₄, TBANO₃, TBACl, TBAHSO₄, TMAOH, TBAClO₄, TBABPh₄, CH₃CO₂TBA, TBAH₂PO₄, anhydrous acetonitrile (CH₃CN), CEES, dimethyl sulfoxide (DMSO), 1,3-dichlorobenzene, and 95% labeled H₂¹⁸O were purchased from Aldrich. TBANO₂ was purchased from Fluka (TBA and TMA are abbreviations for tetra-*n*-butylammonium and tetramethylammonium cations, respectively). Diethyl ether and acetone were purchased from Aldrich and filtered through a column of neutral alumina before use. All POMs in the library (Figure 1) were synthesized from literature preparations.^{38–62} All reagents except for H₂¹⁸O (Aldrich) were dried in vacuo overnight. Stock solutions were prepared using anhydrous CH₃CN.

General Procedures. All gas chromatography analyses were performed on an HP5890 gas chromatograph equipped with a FID detector and a 5% phenyl methyl silicone capillary column. Mass abundance determinations were performed using a HP 5890 GC with a 5% phenyl methyl silicone capillary column and a 5971A mass selective detector. UV–visible spectra were run on a HP 8452A diode array spectrophotometer. The percentages of O₂ in or comprising the reaction atmosphere were varied using a series 810 Mass Trak flowmeter with dried argon as the other gas.

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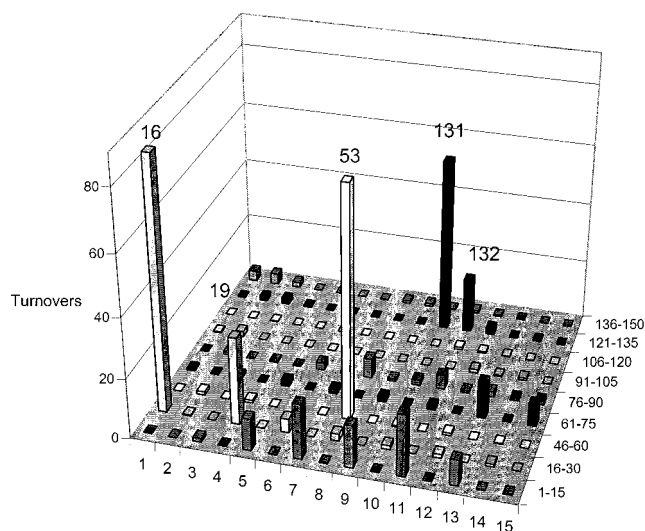


Figure 1. Combinatorial library of catalysts for the oxidation of CEES. Catalytic activity (turnovers of CEESO after 11 h) is depicted on the z axis. Each entry on the x,y plane (base plane) represents a distinct catalyst. Formulas of all the entries (primarily POM derivatives) are given in the Supporting Information (Table S1). Catalytic activity was assessed in an identical manner for all 150 members of the library (see text for details). Catalyst 131 was the Au/Ag/NO₃[−] combination (in less than ideal mole ratios) that led to discovery of the parent system (optimized mole ratios) that was subsequently investigated in depth.

Evaluation of the Library of Catalysts. A total of 150 combinations of substituted POM anions and counteranions were generated, and each was evaluated for the ability to catalyze the oxidation of CEES by O₂ (see Figure 1). For the 150 reactions, every other reaction involved mixing 1.0 equiv of a POM with 5 equiv of HAuCl₄ in a 20-mL vial. The other vials contained POM only. A few vials contained POM-free control compounds. The compositions of all entries in the library are summarized in Table S1 in the Supporting Information. Each vial was purged with O₂, and 1.0 mL of a solution containing 100 equiv of CEES and 50 equiv of 1,3-dichlorobenzene, the internal standard for gas chromatographic analysis, in CH₃CN was added via syringe. Each reaction was monitored at 4 and 11 h for turnovers of CEESO. Product distributions were quantified by GC.

General Procedure for Sample Preparation. Stock solutions of HAuCl₄, AgNO₃, AgClO₄, TBANO₃, TBACl, and TBAClO₄ were prepared in anhydrous CH₃CN and wrapped with aluminum foil to prevent light exposure. All reagents were dried in vacuo before they were dissolved in anhydrous CH₃CN. Appropriate amounts of each stock solution were added via syringe to a 20-mL glass vial fitted with a PTFE septum that was first purged with O₂. (The precise amounts of all reactants and other reagents and conditions are given in the figure captions.) The atmospheric pressure was adjusted to 1.0 atm by inserting a needle into the septum for several seconds. The reactions were carried out at 25 ± 1 °C.

Analysis of Precipitate Formed after CEES Addition. In a 20-mL vial purged with O₂, 0.10 mL (4.8 × 10^{−6} mol, 1 equiv) of HAuCl₄, 0.035 mL (3.6 × 10^{−6} mol, 0.75 equiv) of AgNO₃, 0.06 mL (6.0 × 10^{−6} mol, 1.25 equiv) of AgClO₄, and 0.083 mL (7.2 × 10^{−4} mol, 150 equiv) of 1,3-dichlorobenzene were added by syringe. The volume was adjusted to 0.96 mL with anhydrous CH₃CN. After the solution was stirred for 1 min, 0.042 mL of CEES (3.6 × 10^{−4} mol, 75 equiv) was added and the solution was allowed to stir for an additional 1 min. The vial was then centrifuged and the solution was filtered off leaving a white precipitate. The precipitate was washed with a small amount of CH₃CN and dried under reduced pressure overnight. Elemental analyses confirmed the precipitate was AgCl. Anal. Calcd for AgCl: Ag, 75.26; Cl, 24.74. Found: C, 0.03; H, 0.08; N, 0.03; Ag, 75.08; Cl, 24.89.

Substituting NO₃[−] with Other Anions. These experiments were conducted using protocols and solutions similar to those in the

experiments described above. Solutions of 0.095 mL (9.5 × 10^{−6} mol, 2 equiv) of AgClO₄, 0.10 mL (4.8 × 10^{−6} mol, 1 equiv) of HAuCl₄, and 0.083 mL (7.2 × 10^{−4} mol, 150 equiv) of 1,3-dichlorobenzene were added by syringe to a 20-mL vial that was purged with O₂. Solutions of the following in anhydrous CH₃CN (all 0.2 M) were prepared in 20-mL vials: TBANO₃, TBABPh₄, TBAHSO₄, TMAOH, TBAPF₆, CH₃CO₂TBA, and TBAH₂PO₄. Each of the aforementioned 0.2 M solutions (0.024 mL, 4.8 × 10^{−6} mol, 1 equiv) was added to vials containing the Au/Ag solution. The volume was adjusted to 0.96 mL with anhydrous CH₃CN. After stirring for 1 min, 0.042 mL of CEES was added by syringe. Aliquots were analyzed by GC every 15 min.

Assessment of Au(III) Concentration. For a blank, CH₃CN (2.00 mL) was added to a 1.0-cm quartz cuvette equipped with a stopcock sidearm, a standard taper 14/20 joint, a magnetic stirring bar, and a septum stopper. The background absorbance of this blank was measured on a HP 8452A diode array spectrophotometer. To this cuvette was added by syringe stock solutions of 0.089 mL (9.0 × 10^{−6} mol, 0.75 equiv) of AgNO₃, 0.15 mL (1.5 × 10^{−5} mol, 1.25 equiv) of AgClO₄, and 0.12 mL (1.2 × 10^{−6} mol, 1.0 equiv) of HAuCl₄. It was apparent from the spectrum of this solution, that the 350–450-nm range was the optimal wavelength for quantifying the concentration of Au(III). The solution was immediately transferred to a 20-mL vial that was purged with either 100% O₂ or a mixture of 50:50 O₂ and Ar. To this solution was added 0.020–0.155 mL (3.5 × 10^{−4} mol, 30 equiv) of CEES. After shaking for 1 min, the vial was centrifuged and the solution was transferred back to the cuvette previously purged with O₂ and spectra were collected during the reaction time every 5–10 min. Every 15 min, CEES consumption and CEESO formation were quantified by analyzing 1-μL aliquots by GC.

Establishing the Stoichiometry with Respect to O₂. O₂ consumption was determined by a volumetric method. The reaction vessel containing 1.36 mL of CH₃CN was purged by O₂, and solutions (all in CH₃CN) of 0.20 mL (5.0 × 10^{−6} mol) of HAuCl₄, 0.094 mL (5.0 × 10^{−6} mol) of AgNO₃, 0.094 mL (5.0 × 10^{−6} mol) of AgClO₄, and 0.166 mL (7.5 × 10^{−4} mol) of 1,3-dichlorobenzene (internal standard for GC) were added via syringe through a rubber septum stopper. The reaction was initiated by injection of 0.084 mL (3.8 × 10^{−4} mol) of CEES. The consumption of O₂ was recorded, and 1-μL aliquots were periodically taken and injected into the GC to quantify CEESO formation.

Evaluation of the Reaction Kinetics. The kinetic data were evaluated and curves fit using the Solver subprogram of the Microsoft Excel. The sums of the squares of the difference between experimental and theoretical values were minimized. When fitting was performed varying two parameters at once, several minimums were found in some cases. In this event, the parameters giving the least deviation from experimental data were used.

Rate Law Determination for CEES Oxidation by O₂ Catalyzed by the HAuCl₄/AgNO₃/AgClO₄ System. Screening of a library followed by subsequent evaluation of various Ag(I) salt mixtures identified the most reactive O₂-based catalyst as one composed of AgNO₃, AgClO₄, and HAuCl₄; thus this system was chosen for further investigation. When 1.0 equiv of HAuCl₄ was used, the optimal reactivity was achieved with 0.75 and 1.25 equiv of Ag(I) and NO₃[−] respectively. The order of each component was determined by varying its concentration over the range given below while the concentrations of all other components were held constant. Unless otherwise noted, the initial concentration of HAuCl₄, [HAuCl₄]₀, used in each reaction was 4.8 mM. [CEES]₀ was varied from 0.15 to 3.1 M. [Ag(I)]₀ was varied from 0 to 15.2 mM, [HAuCl₄]₀ was varied from 0 to 11.9 mM, and [NO₃[−]]₀ was varied from 0 to 54 mM. To establish the order in the active multicomponent catalyst itself, the following ratio of concentrations of catalyst components [2Ag(I)/1Au(III)/0.75NO₃[−]] was varied from 1.2 to 7.4 mM (based on total Au concentration). [Cl[−]]₀ was varied from 0 to 6.3 mM, and [ClO₄[−]]₀ was varied from 0 to 0.1 M. The kinetic impact of the product sulfoxide was assessed using DMSO as a model for CEESO (CEESO itself could not, of course, be used because it is the actual product). [DMSO]₀ was varied from 0 to 0.36 M, [H₂O]₀ was varied from 0 to 0.91 M, and the percentage of O₂ in the gas phase (headspace over reaction) was varied from 0 to 100%. For rate order determinations, initial rate methods were employed as

per convention. Given the lack of reactivity of the CEESO product under the reaction conditions, the reactions were monitored until at least 10% of the starting CEES had been consumed.

H_2^{18}O Labeling Experiment. To determine the atomic origin of oxygen in the sulfoxide product (H_2O , H_2O_2 , or O_2), an experiment using H_2^{18}O was performed. For this experiment, a stock solution of 1.53 M H_2^{18}O in anhydrous CH_3CN was prepared using 95 atom % ^{18}O -labeled H_2O . All other stock solutions used were the same as those in the previous experiments. In a 20-mL vial purged with O_2 , 0.10 mL (4.8×10^{-6} mol, 1 equiv) of HAuCl_4 , 0.035 mL (3.6×10^{-6} mol, 0.75 equiv) of AgNO_3 , 0.06 mL (6.0×10^{-6} mol, 1.25 equiv) of AgClO_4 , and 0.083 mL (7.2×10^{-4} mol, 150 equiv) of 1,3-dichlorobenzene internal standard were added by syringe. The volume was adjusted to 0.82 mL with anhydrous CH_3CN . After the solution was stirred for 1 min, 0.042 mL of CEES (3.6×10^{-4} mol, 75 equiv) was added followed immediately by 0.140 mL of H_2^{18}O (3.6×10^{-4} mol, 75 equiv). The isotopic composition of the products was quantified by GC/MS.

Assessment of H_2O_2 -Based Sulfoxidation. To assess sulfoxidation arising from any H_2O_2 generated in situ, an experiment using H_2O_2 was performed. In a Schlenk flask, 2.0 mL of a 30% aqueous H_2O_2 solution was reduced in volume to 0.7 mL resulting in 85% v/v H_2O_2 . *Caution. The procedure of concentrating H_2O_2 or any other peroxide is potentially dangerous. There is some risk of explosion.* The concentrated H_2O_2 solution was immediately diluted with anhydrous CH_3CN and titrated iodometrically; final $[\text{H}_2\text{O}_2] = 1.2$ M in CH_3CN . The same concentrations of each component were used as in the H_2^{18}O experiment except that the H_2^{18}O was omitted. After allowing the solution to react for 2 h, 0.060 mL of the 1.2 M H_2O_2 solution (7.2×10^{-5} mol, 15.0 equiv) was added by syringe.

Solvent Kinetic Isotope Effect, $k_{\text{H}_2\text{O}}/k_{\text{D}_2\text{O}}$. In a 20-mL vial purged with O_2 , 0.10 mL (4.8×10^{-6} mol, 1 equiv) of HAuCl_4 , 0.035 mL (3.6×10^{-6} mol, 0.75 equiv) of AgNO_3 , 0.060 mL (6.0×10^{-6} mol, 1.25 equiv) of AgClO_4 , and 0.083 mL (7.2×10^{-4} mol, 150 equiv) of 1,3-dichlorobenzene were added by syringe. The volume was adjusted with 0.675 mL of anhydrous CH_3CN . To this solution were added, 0.042 mL (3.6×10^{-4} mol, 75 equiv) of CEES, immediately followed by 0.0065 mL (3.6×10^{-4} mol, 75 equiv) of D_2O . In a separate experiment, the same procedure was used except that H_2O was substituted for D_2O . The rate behaviors for the two systems (H_2O and D_2O) were compared.

Replacement of Cl^- Ligands with Br^- Ligands To Assess Atom Transfer in Rate-Limiting Step. In a 20-mL vial purged with O_2 , 0.10 mL (4.8×10^{-6} mol, 1 equiv) of HAuBr_4 , 0.035 mL (3.6×10^{-6} mol, 0.75 equiv) of AgNO_3 , 0.060 mL (6.0×10^{-6} mol, 1.25 equiv) of AgClO_4 , and 0.083 mL (7.2×10^{-4} mol, 150 equiv) of 1,3-dichlorobenzene were added by syringe. The volume was adjusted to 0.680 mL with anhydrous CH_3CN . To this solution, 0.041 mL of CEES (3.5×10^{-4} mol, 75 equiv) was added by syringe. The reaction products were quantified as a function of time by GC and GC/MS as described above. Another experiment was performed using the same procedure except that the volume was adjusted to 0.315 mL with CH_3CN and 0.406 mL of CEES was used.

Attempted Oxidation of Dimethyl Sulfoxide to Dimethyl Sulfone. In a 20-mL vial purged with O_2 , 0.10 mL (4.8×10^{-6} mol, 1 equiv) of HAuCl_4 , 0.035 mL (3.6×10^{-6} mol, 0.75 equiv) of AgNO_3 , 0.060 mL (6.0×10^{-6} mol, 1.25 equiv) of AgClO_4 , and 0.083 mL (7.2×10^{-4} mol, 150 equiv) of 1,3-dichlorobenzene were added by syringe. The volume was adjusted to 0.820 mL with anhydrous CH_3CN . To this solution, 0.025 mL (3.6×10^{-4} mol, 75 equiv) of DMSO was added by syringe. The reaction products were quantified as a function of time by GC and GC/MS as described above.

Effect of DMSO (a CEESO Model) on the Rate. In a 20-mL vial purged with O_2 , 0.100 mL (4.77×10^{-6} mol, 1 equiv) of HAuCl_4 , 0.035 mL (3.58×10^{-6} mol, 0.75 equiv) of AgNO_3 , 0.060 mL (5.96×10^{-6} mol, 1.25 equiv) of AgClO_4 , and 0.083 mL (7.15×10^{-4} mol, 150 equiv) of 1,3-dichlorobenzene were added by syringe. The total volume, factoring in for added CEES and DMSO, was adjusted to 1.0 mL with anhydrous CH_3CN . To the vial, 0.045 mL (3.58×10^{-4} mol, 75 equiv) of CEES was added immediately followed by DMSO. The concentration of added DMSO was varied from 0 to 0.36 M.

Results

Identification of $\text{Au(III)Cl}_2\text{NO}_3(\text{thioether})/\text{O}_2$ Catalytic Oxidation System. A 150-member library was constructed, largely but not exclusively, by combining polyoxoanions (POMs for convenience) and selected cations. Each member of this library was evaluated for its ability to catalyze oxidation of the thioether (CEES) using only O_2 as the oxidant under ambient conditions (room temperature and atmospheric pressure). The POMs chosen were largely those known to have reversible redox chemistry and/or the ability to catalyze homogeneous oxidations by peroxides and other terminal oxidants under mild conditions. The cations chosen included redox-active d-block ions or their precursors including HAuCl_4 as well as the s-block and p-block (e.g., quaternary ammonium) cations conventionally used as the counterions in POMs. This library also contained POM-free control formulations such as the chloride, nitrate, or perchlorate salts of the same redox-active d-block cations used in combination with the POMs. Figure 1 (Experimental Section) summarizes the results of one such library (all members of this library on the x,y-base plane are given in Table S1 in the Supporting Information). While most showed little or no activity, three (entries 16, 53, and 131) exhibited considerable activity. The compositions of these three catalysts were $\text{CuPW}_{11}\text{O}_{39}^{5-}$, $\text{MnPW}_{11}\text{O}_{39}^{5-}$, and AgNO_3 , each with 5 equiv of HAuCl_4 . The rate of the third or POM-free system based on Au and Ag was increased severalfold by varying the ratios of HAuCl_4 and AgNO_3 (and subsequently the ratios of different Ag(I) salts) such that it became the fastest catalyst we evaluated and significantly faster than the two most reactive catalysts in the literature for the homogeneous O_2 -based oxidation of thioethers, $\text{RuCl}_2(\text{Me}_2\text{SO})_4$,⁶³ and $(\text{NH}_4)_2\text{Ce}(\text{NO}_3)_6$.⁶⁴ The former catalyst produces 3.4 turnovers h^{-1} of tetrahydrothiophene oxide (THTO) from tetrahydrothiophene (THT) at 110 °C and 110 psig of O_2 , while the latter catalyst produces 17.6 turnovers of THTO from THT after 30 min at 60 °C and 14 bar O_2 . *Neither $\text{RuCl}_2(\text{Me}_2\text{SO})_4$ nor $(\text{NH}_4)_2\text{Ce}(\text{NO}_3)_6$ has demonstrable activity at ambient temperature.* The unoptimized Au(III)/Ag(I) system arising from the library in Figure 1 was producing nearly comparable turnovers to the Ce(IV) system, the more reactive of the two literature catalysts, at ambient temperature and only 1 atm O_2 . Specifically, this initial system (1 AgNO_3 /5 HAuCl_4 only) gave 35.4 turnovers of CEESO after 4 h under these very mild conditions. Furthermore, CEES is significantly harder to oxidize than THT both thermodynamically and kinetically (it is less reducing and less nucleophilic).¹⁹ Given the remarkable reactivity of this mixture of Au(III) and Ag(I) salts, we chose to investigate this new catalyst in detail. The principal system used in this study comprises 1, 0.75, and 1.25 equiv of the HAuCl_4 , AgNO_3 , and AgClO_4 precursors respectively, based on rate and tractability (ability to vary the concentrations of other added species and maintain completely homogeneous reaction conditions at all times). We report here the general features of this catalytic system, the thioether oxidation reaction stoichiometry, the complex rate law and other kinetic features of this reaction, and the reaction mechanism.

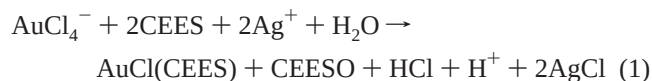
General Features of the $\text{Au(III)Cl}_2\text{NO}_3(\text{thioether})/\text{O}_2$ Catalytic Oxidation System. Extensive studies of *stoichiometric* thioether oxidation by Au(III) complexes in the literature have established five points: (1) HAuCl_4 does not oxidize thioethers at a measurable rate under ambient conditions; (2) halide and thioether ligands exchange in rapid preequilibria

(63) Riley, D. P. *Inorg. Chem.* **1983**, 22, 1965–1967.

(64) Riley, D. P.; Smith, M. R.; Correa, P. E. *J. Am. Chem. Soc.* **1988**, 110, 177–180.

before the rate-limiting redox step that involves thioether reduction of Au(III) forming Au(I) and sulfoxide; (3) the formation constant for the Au(III) complex with one thioether ligand (K_1) is greater than the formation constant for the Au(III) complex with two thioether ligands (K_2); (4) Au(III) complexes with three thioether ligands do not form (K_3 is too small to measure); and (5) the lability of the transient Au(III) complexes renders them nonisolable.^{35–37,65,66} The data given below indicate our Au(III)-based catalytic system for O₂ oxidation shares these features of the stoichiometric thioether–Au(III) studies. However, the catalytic system has additional components and is more mechanistically complex.

Like the Au catalyst precursor, HAuCl₄, by itself, and the Ag salt precursors, AgNO₃ and AgClO₄, by themselves, are both effectively inactive as catalysts for the air oxidation of thioethers at ambient temperature and pressure (HAuCl₄ and AgNO₃ are entries 1 and 130 in the library, Figure 1, respectively). The addition of a soluble Ag(I) salt (e.g., AgNO₃ or AgClO₄) to AuCl₄[–] does not result in the immediate precipitation of AgCl. However, the addition of CEES to the Au(III)/Ag(I) solution leads to the immediate disappearance of the yellow chromophore of Au(III) and the precipitation of AgCl (confirmed by elemental analysis after isolation). Quantification of CEES consumption and CEESO generation confirmed the stoichiometry in eq 1 for this catalyst preparation reaction.



Analysis of the soluble catalyst after removal of AgCl (by taking the reaction to dryness) indicated it contained Au and CEES and/or CEESO moieties but no Ag. Attempted recrystallization of this material produced only a catalytically inactive powder that analyzed for Au(0): Au, 99.83; H, 0.08; C, 0.02. If the catalyst is prepared using 5 mM Au, and excess CEES in the absence of O₂, the resulting Au(0) colloid is violet, suggesting the Au(0) clusters are ~80–100 nm. The attempted growth of crystals of the active catalyst involved much higher concentrations of Au and diffusion of ether into the CH₃CN solution. This procedure produced a colloid that was gold in color consistent with the presence of particles of <40 nm in diameter. (Au(0) colloids are blue/violet, red, or brown/gold for average particle sizes of 80–100, 40–80, and <40 nm, respectively.)⁶⁷ The likely concentrations of the Au(0) particles present during colloid formation are consistent with the different particle sizes and colors.⁶⁸

Once Au(0) forms in this system, it is not reoxidized, and thus this represents an irreversible inactivation of the catalyst. However, use of the ratios of reactants in the following experiments and in the presence of O₂, or even ambient air, produces soluble Au-based catalysts that remain in solution and retain activity for at least one week.⁶⁹

This catalyzed aerobic thioether (CEES) oxidation exhibits a short induction phase followed by the main reaction. This is illustrated in Figure 2. Control experiments establish that the byproducts (not shown) vinyl chloride and thioformaldehyde hydrate result exclusively from thermal decomposition of the

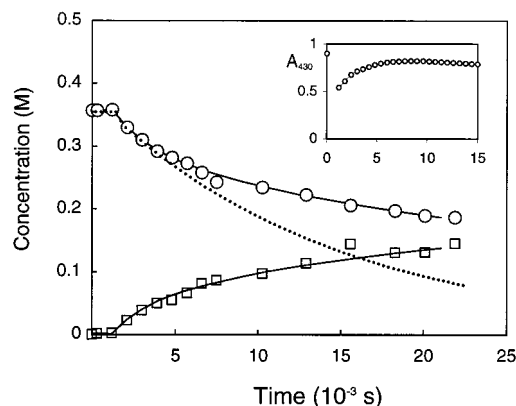


Figure 2. Kinetics of CEES oxidation by the HAuCl₄/0.75AgNO₃/1.25AgClO₄/O₂ system. CEES consumption and CEESO formation as a function of time are indicated by the symbols ○ and □, respectively. Inset: absorbance at 430 nm as a function of time (the absorbance remains constant after 15 000 s, ~4 h). Solid lines represent a fitting of CEES consumption and CEESO formation to the theoretical model in the text. The theoretical kinetic curve (simple-exponential decay with pseudo-first-order rate constant; $k_{\text{obs}} = 7 \times 10^{-5} \text{ s}^{-1}$) for CEES consumption, assuming no inhibition by CEESO is indicated by the dotted line. Conditions: 25 °C; 1 atm O₂; [HAuCl₄] = 4.8 mM; [AgNO₃] = 3.6 mM; [AgClO₄] = 6.0 mM.

CEESO product in the hot injector port of the gas chromatograph. When correction is made for this (or it is eliminated completely by using lower injector port temperatures), the selectivity for sulfoxide is $97 \pm 5\%$. This minimum error is dictated by the limits of the FID detector of the GC. While it is the main reaction itself and not the induction period that is of principal importance and interest, it is appropriate to examine the induction period because it sometimes provides information about the main reaction and because it formally constitutes a component of the overall reaction. After reduction of the starting Au(III) complex, eq 1, the system remains colorless and catalytically inactive during an induction period. During the induction period, no CEES is consumed and no CEESO is formed (Figure 2). At the end of the induction period, the solution rapidly becomes yellow, and the electronic absorption spectrum (350–450 nm) is very similar to that of conventional isolable square-planar Au(III) complexes.⁷⁰ During the main reaction (after the induction period), the concentration of Au(III) remains constant, CEES is consumed, and CEESO is formed (Figure 2, including inset). The dotted and solid lines in Figure 2 arise from fitting of experimental data to the proposed mechanism and are addressed in the Discussion section. While it may not be readily apparent from Figure 2, the kinetics of CEES consumption and CEESO formation in the main reaction do not obey any simple kinetic law; the reaction gradually slows down as a result of a mild inhibition by CEESO product (vide infra). The length of the induction period depends on concentrations of reagents used. The higher the concentrations of CEES, Au, O₂, and the lower the concentration of H₂O, the shorter the

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(66) Natile, G.; Bordignon, E.; Cattalini, L. *Inorg. Chem.* **1976**, 15, 246–248.

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(68) Weisser, H. B. *Inorganic Colloid Chemistry*; John Wiley & Sons: New York, 1933; Vol. 1, p 26.

(69) With the exception of [Au(CN)₂][–], Au(I) complexes disproportionate in H₂O and several disproportionation equilibria have been measured: (a) Schmid, G. M.; Curley-Fiorino, M. E. *Encyclopedia of Electrochemistry of the Elements*; M. Dekker: New York, 1975; Vol. 4. (b) Skibsted, L. H.; Bjerrum, J. *Acta Chim. Scand., Ser. A* **1977**, 31, 155–156. (c) Goolsby, D.; Sawyer, D. T. *Anal. Chem.* **1968**, 40, 1978. (d) Fenske, G. P.; Mason, W. R. *Inorg. Chem.* **1974**, 13, 1783–1786. Although disproportionation is less favorable kinetically in CH₃CN solution, it is possible that the catalyst inactivates by hydrolysis of the Au(I) intermediate followed by its reduction to form Au(0): Kissner, R.; Welti, G.; Geier, G. *J. Chem. Soc., Dalton Trans.* **1997**, 10, 1773–1777.

(70) Mason, W. R.; Gray, H. B. *Inorg. Chem.* **1968**, 7, 7.

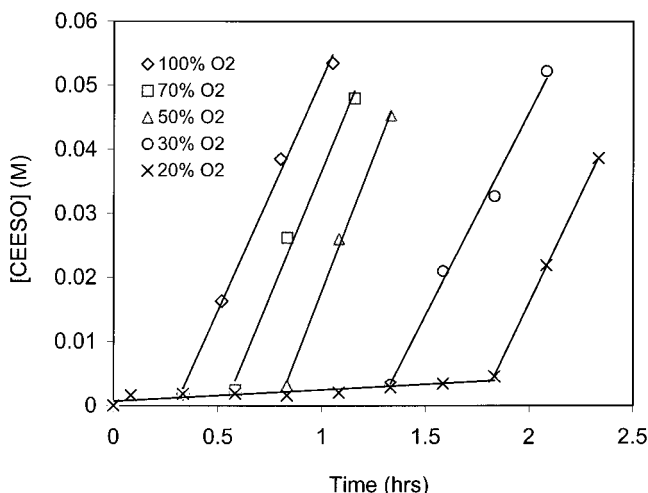
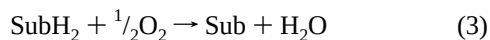
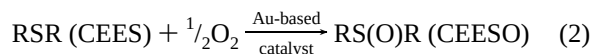


Figure 3. Induction period as a function of initial O_2 concentration in the $HAuCl_4/0.75AgNO_3/1.25AgClO_4/O_2$ system: 100% $O_2 = 1.0$ atm O_2 ; 70% $O_2 = 0.7$ atm O_2 ; 50% $O_2 = 0.5$ atm O_2 ; 30% $O_2 = 0.3$ atm O_2 ; 20% $O_2 = 0.2$ atm O_2 . In every case except 100% O_2 , the total gas pressure is adjusted to 1 atm using Ar. Conditions: 25 °C; $[HAuCl_4] = 4.8$ mM; $[AgNO_3] = 3.6$ mM; $[AgClO_4] = 6.0$ mM; $[CEES] = 0.15$ M.

induction period (Figure 3 shows the $[O_2]$ dependence of the induction period).

For clarification of the rate-limiting step in these Au-catalyzed O_2 -based oxidations, we sought to determine whether Au(I) and/or Au(III) was the dominant redox state of Au during conditions of catalytic turnover. This was assessed by examining the absorbance for Au(III) after the induction period as a function of the concentration of both CEES and O_2 with all other reactants and conditions kept constant in each case. The quantity of Au(III) was independent of both reactants ($[CEES]$ from 0.05 to 0.37 M and $[O_2]$ from 50 to 100%) within experimental error, and all the Au during turnover was effectively Au(III). Clearly, reduction of Au(III) species and not oxidation of Au(I) species is rate limiting in these aerobic oxidation reactions.

Reaction Stoichiometry. Once the catalyst preparation stoichiometry (eq 1) and the general features of the reaction including the induction period and main reaction were clear, then and only then could the stoichiometry of the main catalyzed reaction be determined clearly. By quantifying O_2 consumption as well as CEES consumption and CEESO formation, the stoichiometry of the reaction was established to be that in eq 2.



This “dioxygenase” stoichiometry is the optimal one generally sought in air-based oxidations because it is 100% atom efficient (all of the oxidizing capacity of the oxidants and the atoms of the oxidant are accounted for in the desired product). However, other net reactions (stoichiometries) for O_2 /air oxidations are also possible, including the general and more common “oxidative dehydrogenation” process (eq 3 for substrate = $SubH_2$). While oxidative dehydrogenations are known in thioether oxidations,¹⁶ clearly this mechanism is more favorable for easily dehydrogenated substrates or those with no oxidizable lone pairs including aliphatic hydrocarbons and alcohols.

Determination of stoichiometry also involves the product selectivity. Sulfoxides, of course, can be and often are oxidized

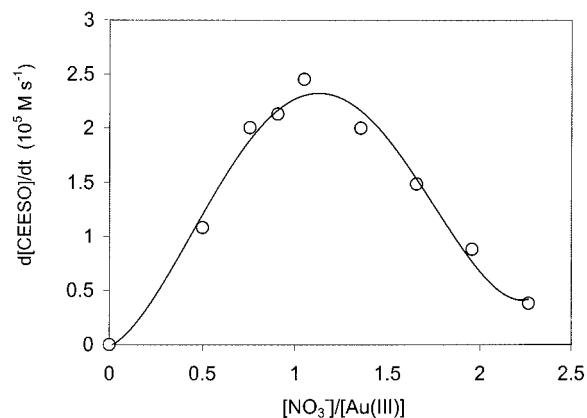


Figure 4. Rate of CEESO formation as a function of $[NO_3^-]/[Au(III)]$. Conditions: 25 °C; 1 atm O_2 ; $[HAuCl_4] = 4.8$ mM; $[CEES] = 0.37$ M.

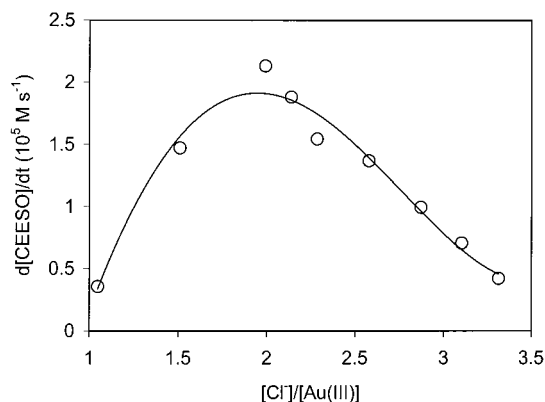


Figure 5. Rate of CEESO formation as a function of $[Cl^-]/[Au(III)]$. Conditions: 25 °C; 1 atm O_2 ; $[HAuCl_4] = 4.8$ mM; $[CEES] = 0.37$ M.

further to the corresponding sulfones. However, in our case, the selectivity for CEESO is very high ($97 \pm 5\%$).

Empirical Rate Law. The rate data are presented here and fully evaluated in context with mechanism elucidation in the Discussion section. The primary experimentally determined rate parameter was $+d[CEESO]/dt$, which is indicated henceforth as “rate”, but $-d[CEES]/dt$ was also evaluated in many cases. The rate for eq 2 was determined as a function of the concentrations not only of Au(III), Ag(I), CEES, and O_2 but also of NO_3^- , Cl^- , H_2O , $Au/Cl^-/NO_3^-$ together in a constant 1:2:1 mole ratio, and DMSO as a model for the CEESO product. A product model compound is helpful to assess product inhibition because its chemistry can be evaluated independently of the concentrations of all species in the main reaction itself. Determination of the experimental rate law for the reaction was not the usual straightforward process of evaluating the slopes of van’t Hoff $\ln$ – $\ln$ kinetics plots ($\ln$ observed rate versus $\ln$ of concentration of varied reactant), because linear plots were not obtained with respect to any reacting species.

Figures 4–8 give, respectively, the dependences of the rate of the main reaction, eq 2, on the concentrations of NO_3^- , Cl^- , CEES, H_2O , and total Au(III) ($Au(III)_T$). For the $Au(III)_T$ dependence, the 1:2:1 ratio of $Au/Cl^-/NO_3^-$ was kept constant. The fittings in Figures 6 and 8 ($[CEES]$ and $[Au(III)_T]$ dependences, respectively) are addressed in the Discussion section. Interestingly, both NO_3^- and Cl^- are critical for reaction, and the dominant transition-state complex appears to contain one NO_3^- and two Cl^- groups. Independently, it was established that replacement of NO_3^- by ClO_4^- , PF_6^- , BPh_4^- ,

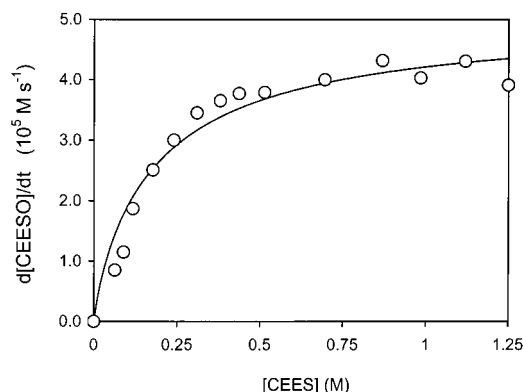


Figure 6. Rate of CEESO formation as a function of initial CEES concentration. The solid curve is a best least-squares fit to eq 19, namely, the bimolecular reaction. Conditions: 25 °C; 1 atm O₂; [HAuCl₄] = 4.8 mM; [AgNO₃] = 3.6 mM; [AgClO₄] = 6.0 mM.

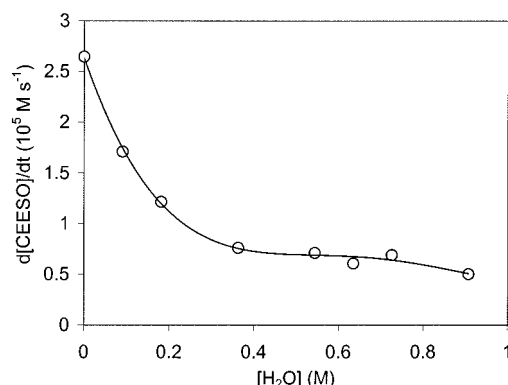


Figure 7. Rate of CEESO formation as a function of H₂O concentration. Conditions: 25 °C; 1 atm O₂; [HAuCl₄] = 4.8 mM; [AgNO₃] = 3.6 mM; [AgClO₄] = 6.0 mM; [CEES] = 0.37 M.

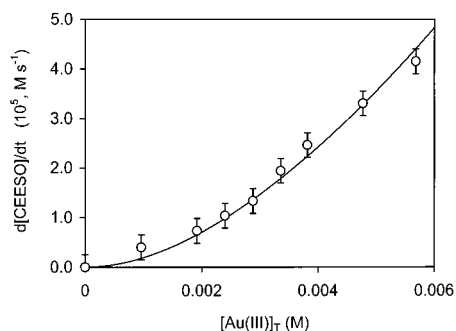


Figure 8. Rate of CEESO formation as a function of the concentration of total catalyst ($\text{Au(III)}_T = \text{Au/Cl/NO}_3$ in 1:2:1 ratio). The solid curve is a best least-squares fit to eq 19, namely, the bimolecular reaction. Conditions: 25 °C; 1 atm O₂; [CEES] = 0.37 M.

or HSO₄[−] generated species with little or no catalytic activity, confirming a significant and specific role for NO₃[−] in the active catalyst.

The rate dependence on the concentration of the substrate, CEES, shows saturation kinetics consistent with one or more intermediates in the rate-limiting step(s) containing varying numbers of coordinated CEES molecules. Not surprisingly, the [H₂O] and [DMSO] dependencies (Figure 7 and Table 1, respectively) are hyperbolic (largely inverse), consistent with both H₂O and DMSO competing with Cl[−], NO₃[−], and CEES for open coordination sites on the catalytic Au center and thus decreasing the concentration of the Au complexes required for CEESO production.

Table 1. Rate of CEESO Formation as a Function of [DMSO]^a

[DMSO], M	0	0.022	0.029	0.036	0.043	0.072	0.14	0.22	0.36
rate × 10 ⁵ , M s ^{−1}	2.9	1.84	1.40	1.33	1.01	0.69	0.49	0.42	0.24

^a Conditions: 25 °C; 1 atm O₂; [HAuCl₄] = 4.8 mM; [AgNO₃] = 3.6 mM; [AgClO₄] = 6.0 mM; [CEES] = 0.37 M.

The rate is independent of O₂ concentration. This is consistent with the Au oxidation state during turnover (*vide supra*) and implicates that Au(III) reduction and not Au(I) reoxidation is rate limiting. Interestingly, in the one other documented case of thioether oxidation catalyzed by soluble Au complexes, the oxidation of thioethers to sulfoxides by HNO₃ reported by Gasparrini et al., the rate-limiting step is reoxidation of Au(I) to Au(III).³⁷

One experiment to assess the mechanism of the slow thioether oxidation–Au(III) reduction step involved evaluation of the overall reaction rate when Cl[−] was replaced by Br[−] under otherwise identical conditions. At 0.3 M CEES, the initial rates for CEESO production for the Cl[−]-containing and Br[−]-containing systems are 2.0×10^{-5} and 3.4×10^{-5} M s^{−1}, respectively. At 3.0 M CEES, these initial rates are 3.9×10^{-5} and 7.3×10^{-5} M s^{−1}, respectively. These and other results clarify the nature of this step (see Discussion section).

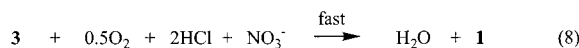
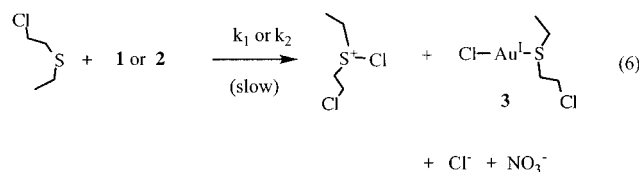
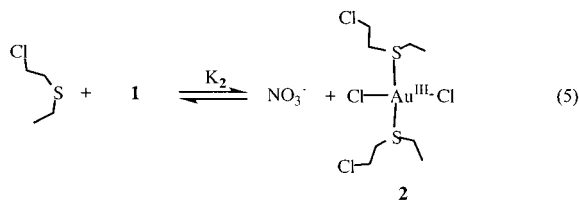
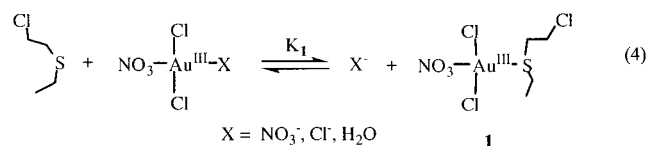
The temperature dependence (activation parameters) of the reaction was purposefully not assessed because reactions with several preequilibrium steps, such as the one in this study (see Discussion section), would yield minimally interpretable, and possibly misleading data.

H₂O₂ can oxidize CEES to CEESO in CH₃CN, when a strong acid is present. Furthermore, HCl from catalyst generation, eq 1, and the reactant HAuCl₄ itself are acidic. Since H₂O₂ could be formed in situ during Au(I) reoxidation by O₂, we assessed the possibility of CEES oxidation by intermediate H₂O₂ using an isotope labeling study. A reaction with H₂¹⁸O in place of regular H₂¹⁶O was conducted to provide this information and also to ascertain whether the oxygen in the product sulfoxide, CEESO, derives from H₂O, H₂O₂, or O₂. The CEESO produced in the early part of the reaction (up to 10 turnovers) is ~80% ¹⁸O and ~20% ¹⁶O. The somewhat lower percent ¹⁸O in the CEESO than in the H₂¹⁸O used derives in part from water molecules associated with HAuCl₄, a hygroscopic compound that is hard to dry. The percentage of ¹⁸O in the CEESO decreases with reaction time and turnovers. After 60 turnovers, the ¹⁸O in the CEESO decreases to ~60 atom %. These data indicate that CEESO is produced by a process involving only H₂O and not O₂ or H₂O₂. If H₂O₂ alone were involved, the ¹⁸O/¹⁶O ratio would not exceed 50%. The time dependence of the label is consistent with a consumption of labeled water which results in a dilution of the H₂¹⁸O pool with H₂¹⁶O (while no net H₂O is formed in the overall reaction, eq 2, H₂O is consumed and then regenerated in the mechanism, *vide infra*).

The rates of CEESO production, eq 2, in both 100% H₂O (usual conditions) and in ~100% D₂O were determined. The rates were the same within the experimental error (±5%).

Discussion

A mechanism for the selective aerobic oxidation of thioethers to sulfoxides catalyzed by Au(III) complexes that is compatible with all the data and the literature investigations (stoichiometric thioether–Au(III) reactions) is given in Scheme 1. While NO₃[−] is capable of reoxidizing the product Au(I) back to Au(III) as demonstrated by Natile and co-workers,³⁷ O₂ and not NO₃[−] is the terminal oxidant in our system. This is established because

Scheme 1. Mechanism for O₂-Based Oxidation of Thioethers Catalyzed by Au(III)Cl₂NO₃ (thioether) (**1**)

there is only 1 equiv of NO₃⁻ in our reactions, yet most of them have been taken to 30 or more turnovers and in one case 200 turnovers (200 equiv of CEESO product per equivalent of Au and equivalent of NO₃⁻). Figures 4 and 5 indicate that the kinetically dominant species leading to CEESO contains Au, Cl⁻, and NO₃⁻ in a 1:2:1 ratio, and Figure 6 indicates that this species also contains CEES (Figure 6 is addressed further below). Dimeric or oligomeric d⁸ square-planar Au(III) are not particularly common,³⁰ and a monomeric Au(III) complex with two Cl⁻ ligands, one NO₃⁻ ligand, and at least one CEES ligand is likely. While 5-coordinate Au(III) complexes cannot be rigorously ruled out, they are not likely to be significant.⁷¹ All the information on oxidation of thioethers by Au(III) complexes is consistent with the rapid exchange of all ligands on Au(III) prior to thioether oxidation.^{32–36} For the present catalytic system, these preequilibria are summarized by eqs 4 and 5 in Scheme 1. Explicitly, Cl⁻, NO₃⁻, and CEES lead to formation of the Au(III) complexes required for CEESO formation: a complex with one CEES ligand (henceforth **1**), with formation constant *K*₁, and a complex with two CEES ligands (henceforth **2**), with formation constant *K*₂. These ligands drive eq 4 to the right. H₂O (data in Figure 7), and the CEESO product, modeled by another sulfoxide, DMSO (data in Table 1), both inhibit the rate of reaction by decreasing the concentrations of the Au(III) complexes required for CEESO formation. These ligands drive eq 4 to the left. For completeness, CH₃CN may also competitively bind to Au(III), but this cannot be assessed since it is the solvent.

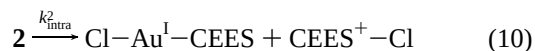
(71) The only 5-coordinate Au(III) complexes with appreciable stability contain chelating ligands such as bromodicyano(1,10-phenanthroline)-gold(III) isolated from dimethylformamide: Marangoni, G.; Pitteri, B.; Bertolasi, Y.; Gilli, G.; Ferretti, V. *J. Chem. Soc., Dalton Trans.* **1986**, 1941. If there is an interaction between a positive Au(III) center and a negative axial ligand counterion, this ligand would likely be Cl⁻, because NO₃⁻ associates very weakly with Au(III) (and not at all with Au(I)).

Literature studies of thioether-Au(III) reactions establish that Au(III) complexes with one thioether ligand, such as **1**, are considerably more abundant in solutions of excess thioether than complexes with two thioether ligands, such as **2** (*K*₁ ≫ *K*₂).^{35,36} No Au(III) complex with three thioether ligands is given in Scheme 1 because there are no data indicating these species form in measurable quantities (i.e., *K*₂ ≫ *K*₃; *K*₃ ~ 0). Evaluation of the kinetics data below also indicates that **1** and not **2** is the kinetically dominant Au(III) complex for catalytic aerobic oxidation of CEES to CEESO.

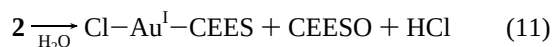
Three lines of evidence establish that the redox step, oxidation of thioether/reduction of Au(III), eq 6 in Scheme 1, is rate limiting in the catalytic O₂-based oxidation of thioethers: First, all the rate information (Figures 4–8) are consistent with this; second, the rate is independent of O₂ pressure, and third, Au(III), not Au(I), is the dominant form of Au during catalytic turnover. The rate constants *k*₁ and *k*₂ in eq 6 are for **1** and **2**, respectively; however, eq 6 has been written in terms of **1** and its redox products, (CEES)Au^ICl (**3**), Cl⁻, and NO₃⁻ (versus **2** and its products), because most of the observed chemistry derives from **1**. Because eq 6 is clearly rate determining, the subsequent steps, eqs 7 and 8, are not.

There are several possible mechanisms for the rate-limiting redox step, eq 6, but one is more consistent with the data, analysis of the data below, and the stoichiometric Au(III)–thioether studies in the literature. First, a mechanism involving rate-limiting formation of the required Au(III) complex (primarily **1** in our case) prior to redox chemistry is not compatible with any of the literature studies, nor is it compatible with the kinetic requirement of two molecules of thioether in the slow step. The detailed equilibrium and kinetics studies (extensive data and fitting) of two groups establish that two molecules of thioether are involved in the slow step: Natile and co-workers³⁶ on the oxidation of a Me₂S, Et₂S, and *n*-Pr₂S by Au(III)(Cl)_x(RS)_{4–x} and Elding and co-workers³⁵ on the oxidation of a Me₂S by *trans*-Au(III)(CN)₂(Cl)_{2–x}(RS)_{2–x}. While the Natile system and ours differ in three ways, theirs does not undergo catalytic turnover, does not contain NO₃⁻, and is in 95:5 CH₃OH/H₂O (versus CH₃CN in our case), the Au(III)–thioether coordination and redox chemistry in the two systems are otherwise very similar. Elding's complex, while not as close to ours as that of Natile, nonetheless shares structural and redox chemistry very similar to ours as well.

A second possible mechanism is an intramolecular decomposition of the required Au(III) complex, eqs 9 and 10.



This is effectively a rate-limiting reductive elimination to form a thioxonium salt (whose fate, eq 7 in Scheme 1, is addressed below) and a Au(I) complex. While this mechanism has been proposed for some thioether oxidations by Au(III),^{32,34,35} it is likely not operable in our catalytic aerobic oxidation chemistry. First, eq 9 is not compatible with the finding that two molecules of thioether are involved in the rate-limiting redox step (vide supra).^{35,36} However, this argument does not eliminate eq 10, which involves two thioether molecules. The conversion of **2** to **3** + CEESO can, in principle, be assisted by H₂O, eq 11,³⁵



but the nonexistent solvent effect ($k_{\text{H}_2\text{O}}/k_{\text{D}_2\text{O}} = 1.0$) argues against eq 11 being rate limiting. Again, the kinetics data are in accord with H₂O functioning as an inhibitor via competition for the pool of Au(III) complexes present during catalytic turnover, i.e., driving eq 4, Scheme 1, to the left.

Additional evidence against the unimolecular collapse of **1** and **2** (eqs 9 and 10) derives from analysis of the reaction kinetics, which we now address. If eqs 9 and 10 were rate limiting, the overall rate would be eq 12. It is reasonable to

$$\text{rate} = k_{\text{intra}}^1 [\mathbf{1}] + k_{\text{intra}}^2 [\mathbf{2}] \quad (12)$$

assume, that under our experimental conditions, $[\text{AuCl}_2\text{NO}_3(\text{CH}_3\text{CN})] \ll [\mathbf{1}] + [\mathbf{2}]$, since K_1 is big and $[\text{CEES}] \gg [\text{Cl}^-] + [\text{NO}_3^-]$. The equilibrium expressions from eqs 4 and 5, Scheme 1, and reaction mass balance expressions afford eqs 13 and 14, from which we can write eqs 15 and 16. Expressions for $[\mathbf{1}]$ and $[\mathbf{2}]$, eqs 17 and 18, are obtained by

$$K_2[\mathbf{1}][\text{CEES}] = [\mathbf{2}][\text{NO}_3^-] = [\mathbf{2}]^2 \quad (\text{because } [\mathbf{2}] = [\text{NO}_3^-]) \quad (13)$$

$$[\mathbf{1}] + [\mathbf{2}] = [\text{Au(III)}]_{\text{T}} \quad (\text{since } [\text{Au(I)}] \ll [\text{Au(III)}]) \quad (14)$$

$$[\mathbf{2}] = (K_2[\mathbf{1}][\text{CEES}])^{0.5} \quad (15)$$

$$[\mathbf{1}] + (K_2[\mathbf{1}][\text{CEES}])^{0.5} = [\text{Au(III)}]_{\text{T}} \quad (16)$$

solving eqs 15 and 16:

$$[\mathbf{1}] = (1/4)(\sqrt{K_2[\text{CEES}] + 4[\text{Au(III)}]_{\text{T}}} - \sqrt{K_2[\text{CEES}]})^2 \quad (17)$$

$$[\mathbf{2}] = [\text{Au(III)}]_{\text{T}} - (1/4)(\sqrt{K_2[\text{CEES}] + 4[\text{Au(III)}]_{\text{T}}} - \sqrt{K_2[\text{CEES}]})^2 \quad (18)$$

Combining eqs 12, 17, and 18 gives eqs 19a and 19b for the reaction rate.

$$\text{rate} = k_{\text{intra}}^2 [\text{Au(III)}]_{\text{T}} + (k_{\text{intra}}^1 - k_{\text{intra}}^2)[\mathbf{1}] \quad (19a)$$

$$\text{rate} = k_{\text{intra}}^2 [\text{Au(III)}]_{\text{T}} + (1/4)(k_{\text{intra}}^1 - k_{\text{intra}}^2) \times (\sqrt{K_2[\text{CEES}] + 4[\text{Au(III)}]_{\text{T}}} - \sqrt{K_2[\text{CEES}]})^2 \quad (19b)$$

This rate law indicates that reaction rate orders depend on whether **1** or **2** is more reactive. If $k_{\text{intra}}^1 > k_{\text{intra}}^2$, then the rate will increase quadratically with $[\text{Au(III)}]_{\text{T}}$, but will decrease with $[\text{CEES}]$. On the contrary, if $k_{\text{intra}}^1 < k_{\text{intra}}^2$, then the rate will increase with both $[\text{Au(III)}]_{\text{T}}$ and $[\text{CEES}]$, finally reaching saturation with an observed reaction rate order of <1 . Thus, one of the rate dependences, either that for $[\text{CEES}]$ or that for $[\text{Au(III)}]_{\text{T}}$, can be easily fitted, but not both simultaneously. Importantly, however, the experimental data (Figures 6 and 8) are incompatible with either of these scenarios based on the unimolecular collapse mechanism (eqs 9 and 10), which further rules out this mechanism. In addition, the water-assisted collapse, eq 11, is kinetically the same as eq 10, and incorporation of eq 11 in the above analysis leads to an expression analogous to eq 19 but with k_{intra}^2 replaced by $k_{\text{intra}}^{2/\text{obs}} = \{k_{\text{intra}}^2 (\text{reaction 10}) + k_{\text{intra}}^2 (\text{reaction 11})\}$. Thus, eq 11 is also

incompatible with the data and can also be ruled out as a mechanism.

A third mechanism involving rate-limiting CEES attack at the Au(III) center with immediate fast CEES oxidation and Au(III) reduction is very unlikely. This S_N2-like mechanism does not appear to be operable in any of published stoichiometric Au(III)–thioether systems, nor is it compatible with the relative rates of oxidation of Me₂S, Et₂S, and *n*-Pr₂S by Au(III)(Cl)_{*x*}–(R₂S)_{4–*x*} reported by the Italian group.³⁶ These rates were sensitive to the basicity but not sensitive to the steric bulk of the thioether: Me₂S (slowest) < Et₂S < *n*-Pr₂S (fastest), the opposite trend observed in S_N2-like processes.

A fourth possible mechanism involves bimolecular attack of thioether on a coordinated ligand of Au(III). This is the proposed mechanism for some but not all of the stoichiometric thioether oxidations by Au(III).^{31–33} None of the data on our system rules this out, and it is consistent with the increase in rate when Cl[–] is replaced by Br[–] under otherwise identical conditions. Ligand transfer oxidation processes, including reduction of Au(III) by thioether in our case, are kinetically favored by soft Br[–] (versus harder Cl[–]) ligands.⁷² This kinetic preference for softer ligands is in the direction opposite from the thermodynamics of the reaction. Specifically, the metal centers with Br[–] in place of Cl[–] at parity of other ligands have lower potentials and are weaker oxidants. Additional evidence comes indirectly from the following analysis of our kinetic data.

The overall rate from Scheme 1 and eq 14 is eq 20.

$$\text{rate} = k_1[\mathbf{1}][\text{CEES}] + k_2[\mathbf{2}][\text{CEES}] = k_1[\text{CEES}][\text{Au(III)}]_{\text{T}} + (k_1 - k_2)[\text{CEES}][\mathbf{1}] \quad (20)$$

Combining eqs 17 and 20 gives eqs 21a and 21b for the reaction rate

$$\text{rate} = k_2[\text{CEES}][\text{Au(III)}]_{\text{T}} + (k_1 - k_2)[\text{CEES}][\mathbf{1}] \quad (21a)$$

$$\text{rate} = k_2[\text{CEES}][\text{Au(III)}]_{\text{T}} + ((k_1 - k_2)[\text{CEES}]/4) (\sqrt{K_2[\text{CEES}] + 4[\text{Au(III)}]_{\text{T}}} - \sqrt{K_2[\text{CEES}]})^2 \quad (21b)$$

The best fitting of the experimental data (solid curve in Figure 6) gives $k_1 = 0.12 \pm 0.05 \text{ M}^{-1}\text{s}^{-1}$, $K_2 = 0.04 \pm 0.02$, and $k_2 = 0$. The errors were estimated at the 95% confidence limit. In other words, since $k_2 = 0$, eq 21 simplifies to eq 22.

$$\text{rate} = (k_1[\text{CEES}]/4)(\sqrt{K_2[\text{CEES}] + 4[\text{Au(III)}]_{\text{T}}} - \sqrt{K_2[\text{CEES}]})^2 \quad (22)$$

A low reactivity of complex **2** toward CEES, $k_2 = 0$, is easily demonstrated without curve fitting. At high $[\text{CEES}]$, the majority of Au(III) is present as **2**, which corresponds to $K_2[\text{CEES}] \gg 4[\text{Au(III)}]_{\text{T}}$. Multiplying eq 17 by the unity factor $(K_2[\text{CEES}]/K_2[\text{CEES}])$ affords eq 23.

$$[\mathbf{1}] = (K_2[\text{CEES}]/4)(\sqrt{1 + 4x} - 1)^2 \approx (K_2[\text{CEES}]/4) \times \{(1 + 2x) - 1\}^2 = [\text{Au(III)}]_{\text{T}}^2/K_2[\text{CEES}] \quad (23)$$

where $x = [\text{Au(III)}]_{\text{T}}/K_2[\text{CEES}] \ll 1$.

(72) Kochi, J. K. In *Oxidation–Reduction Reactions of Free Radicals and Metal Complexes*; Kochi, J. K., Ed.; Wiley: New York, 1973; Vol. 1, pp 591–685.

Equations 29 and 30 afford eq 31, an expression for [1], which is analogous to eq 17 but contains, in addition, DMSO.

$$[1] + (1/4)(\sqrt{K_2[\text{CEES}] + K_2^{\text{DMSO}} + 4[\text{Au(III)}]_{\text{T}} - \sqrt{K_2[\text{CEES}] + K_2^{\text{DMSO}} [\text{DMSO}]}})^2 \quad (31)$$

At high [CEES] and DMSO, the majority of Au(III) is present mainly 2 and 2', which corresponds to $K_2[\text{CEES}] + K_2^{\text{DMSO}}[\text{DMSO}] \gg 4[\text{Au(III)}]_{\text{T}}$. Equation 31 then simplifies to eq 32 (for details, see eq 23), which then gives the rate law in eq 33.

$$[1] \approx [\text{Au(III)}]_{\text{T}}^2 / (K_2[\text{CEES}] + K_2^{\text{DMSO}}[\text{DMSO}]) \quad (32)$$

$$\text{rate} = k_1[1][\text{CEES}] \approx$$

$$k_1[\text{CEES}][\text{Au(III)}]_{\text{T}}^2 / (K_2[\text{CEES}] + K_2^{\text{DMSO}}[\text{DMSO}]) = (k_1/K_2)[\text{Au(III)}]_{\text{T}}^2 / (1 + K_2^{\text{DMSO}}[\text{DMSO}]/K_2[\text{CEES}]) = r_o / (1 + K_2^{\text{DMSO}}[\text{DMSO}]/K_2[\text{CEES}]) \quad (33)$$

where $r_o = (k_1/K_2)[\text{Au(III)}]_{\text{T}}^2$ is the initial rate in the absence of DMSO (for comparison, see eq 25). Thus, under certain experimental conditions, when $K_2[\text{CEES}] + K_2^{\text{DMSO}}[\text{DMSO}] \gg 4[\text{Au(III)}]_{\text{T}}$, the dependence of the reaction rate on [DMSO] is described by a simple hyperbolic function, eq 33. Indeed, the experimental data (Table 1) fitted to eq 33 yield $K_2^{\text{DMSO}} = 12$.

CEESO is likely to inhibit the reaction by the same mechanism as DMSO, eq 28. At [CEES] < 0.4 M, the dependence of reaction rate on [CEES] can be roughly approximated by a linear function $r_o \approx k_{\text{obs}}[\text{CEES}]$ (Figure 6). Taking this and mass balance ($[\text{CEESO}] = [\text{CEES}]_o - [\text{CEES}]$) into account, the rate

law is then eq 34, where $[\text{CEES}]_o$ is the initial CEES concentration. Rearranging eq 34 gives eq 35.

$$\text{rate} = -d[\text{CEES}]/dt = -k_{\text{obs}}[\text{CEES}] / \{1 + K_2^{\text{CEESO}}([\text{CEES}]_o - [\text{CEES}]) / [\text{CEES}]\} \quad (34)$$

$$(d[\text{CEES}]/[\text{CEES}]) \{1 - K_2^{\text{CEESO}} + K_2^{\text{CEESO}}[\text{CEES}]_o / [\text{CEES}]\} = -k_{\text{obs}} dt \quad (35)$$

Since at initial time $[\text{CEES}] = [\text{CEES}]_o$, the integration (a standard integral) gives eq 36:

$$t = (k_{\text{obs}}^{-1}) \{ (1 - K_2^{\text{CEESO}}) \ln([\text{CEES}]_o / [\text{CEES}]) + K_2^{\text{CEESO}}([\text{CEES}]_o / [\text{CEES}] - 1) \} \quad (36)$$

The value $k_{\text{obs}} = 7 \times 10^{-5} \text{ s}^{-1}$ was determined independently from the slope of the plot of initial reaction rate versus $[\text{CEES}]_o$ at $[\text{CEES}] < 0.4 \text{ M}$. The difference between experimental time (after the induction period) and that calculated from eq 36 was minimized by varying K_2^{CEESO} . The solid line in Figure 2 is the result of nonlinear least-squares fitting. The dashed line represents the expected consumption of CEES, assuming that no inhibition by CEESO takes place, and the reaction is first order in [CEES] (an exponential decay with $k_{\text{obs}} = 7 \times 10^{-5} \text{ s}^{-1}$).

The kinetics of CEESO accumulation (solid line plotted through CEESO kinetics experimental points) was not fitted but calculated from eq 37, where the factor of 0.8 accounts for CEESO decomposition in the hot GC injection port.

$$[\text{CEESO}] = 0.8([\text{CEES}]_o - [\text{CEES}]) \quad (37)$$

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