

# Chemical Science

Volume 16  
Number 3  
21 January 2025  
Pages 983–1470

rsc.li/chemical-science



ISSN 2041-6539



## EDGE ARTICLE

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**15**  
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## EDGE ARTICLE

Cite this: *Chem. Sci.*, 2025, 16, 1115

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# Why do some metal ions spontaneously form nanoparticles in water microdroplets? Disentangling the contributions of the air–water interface and bulk redox chemistry†

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Water microdroplets containing 100  $\mu\text{M}$   $\text{HAuCl}_4$  have been shown to reduce gold ions into gold nanoparticles spontaneously. It has been suggested that this chemical transformation takes place exclusively at the air–water interface of microdroplets, albeit without mechanistic insights. We compared the fate of several metallic salts in water, methanol, ethanol, and acetonitrile in the bulk phase and microdroplet geometry (sprays). Experiments revealed that when  $\text{HAuCl}_4$  (or  $\text{PtCl}_4$ ) is added to bulk water (or methanol or ethanol), metal NPs appear spontaneously. Over time, the nanoparticles grow, evidenced by the bulk solutions' changing colors. If the bulk solution is sprayed pneumatically and microdroplets are collected, the NP size distribution is not significantly enhanced. We find that the reduction of metal ions is accompanied by the oxidation of water (or alcohols); however, these redox reactions are minimal in acetonitrile. This establishes that the spontaneous reduction of metal ions is (i) a bulk phase phenomenon in water and several non-aqueous solutions, (ii) minimally affected by the air–water interface or the microdroplet geometry, and (iii) is not limited to  $\text{Au}^{3+}$  ions and can be explained via the electrochemical series. These results advance our understanding of aquatic chemistry and liquids in general and should be relevant in soil chemistry, biogeochemistry, electrochemistry, and green chemistry.

Received 16th May 2024  
Accepted 13th November 2024

DOI: 10.1039/d4sc03217a

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## Introduction

Biochemistry, chemistry, and chemical engineering textbooks often limit the role of water in chemical reactions to dissolving/hydrating molecules and ions;<sup>1–3</sup> however, this view is rapidly evolving.<sup>4–9</sup> In the last two decades, experimental and computational studies have revealed that the skin of water – the air–water interface – and water–hydrophobe interfaces, in general, have anomalous features such as enhanced speciation of ions<sup>10,11</sup> and electrified interfaces.<sup>12–15</sup> Several studies have

reported on orders of magnitude faster (e.g.,  $10^2$ – $10^6\times$ ) reaction rates in water microdroplets compared with the bulk phase, thereby expanding the function of aqueous interfaces in chemical transformations beyond heat and mass transfer.<sup>5,16–24</sup> This exciting field – microdroplet chemistry – has emerged as a frontier in chemical science with potential implications for atmospheric aerosols,<sup>25</sup> sea sprays,<sup>26</sup> disease transmission,<sup>27,28</sup> life's origins,<sup>16,19,29</sup> the chemistry of clouds, fog, smog, and dew,<sup>30–32</sup> soil processes,<sup>33</sup> green chemistry,<sup>18</sup> hospital disinfection,<sup>34</sup> the food industry,<sup>35</sup> and fizzy beverages;<sup>36</sup> also, one may expect similar trends in microbubbles/foam in fermenter broths,<sup>37</sup> wastewater treatment,<sup>38,39</sup> and electrochemistry.<sup>40,41</sup>

While much excitement exists, several claims for purely interfacial effects being responsible for the rate enhancement have been challenged. This is because the vast majority of reports exploit electrospray ionization mass spectrometry (ESI-MS), which entails rapid solvent evaporation, solute concentration, temperature, electric field gradients, etc., conditions that are far from thermodynamic equilibrium and, therefore, not representative of common systems.<sup>20,23,42,43</sup> For instance, the emergence of superacid chemistry in the microdroplets of mildly acidic water ( $\text{pH} < 4$ )<sup>44,45</sup> and the formation of abiotic sugar phosphates in water microdroplets<sup>16</sup> have been

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† Electronic supplementary information (ESI) available. See DOI: <https://doi.org/10.1039/d4sc03217a>



challenged recently<sup>46–49</sup> (for recent reviews on this subject, see ref. 15, 24, 50 and 51). Controversial reports on microdroplet chemistry are not limited to ESIMS alone. For instance, recently, Zare & co-workers claimed that the air–water interface of microdroplets spontaneously generates H<sub>2</sub>O<sub>2</sub>,<sup>52–55</sup> and the reported amounts varied from ~1 ppm or 30 μM (in 2019 for pneumatically sprayed droplets)<sup>52</sup> to 114 μM (in 2020 for condensed droplets)<sup>54</sup> to 180 μM (in 2021 for microdroplets condensed at 55% or 70% relative humidity).<sup>53</sup> As for the mechanistic insights, computer simulations of Head-Gordon & co-workers<sup>56–58</sup> have suggested the emergence of instantaneous ultrahigh electric fields at the air–water interface that may drive H<sub>2</sub>O<sub>2</sub> formation, while the simulations of Ruiz-López and co-workers<sup>50</sup> and Gong and co-workers<sup>59</sup> suggested that the local electric field was lower at the air–water interface compared to the bulk. Cooks and co-workers have speculated that the air–water interface of microdroplets promotes the formation of water radical cations H<sub>2</sub>O<sup>+</sup>\* and anions H<sub>2</sub>O<sup>-</sup>\*, which constitute the primary redox species.<sup>24</sup> Colussi<sup>60</sup> also proposed a similar mechanism whereby spraying yielded a small fraction of oppositely charged droplets, *i.e.*, comprising excess H<sub>3</sub>O<sup>+</sup> and OH<sup>-</sup>, and these droplets collided to form OH\* and H\* radicals.<sup>60</sup> Our team has challenged these reports<sup>24,27,52–57,60</sup> by pinpointing the artifacts arising due to the ambient ozone contamination,<sup>61</sup> and OH\* radical formation due to ultrasonication;<sup>62</sup> also, we have explained the appearance of trace (~1 μM) H<sub>2</sub>O<sub>2</sub> concentrations in pneumatic sprays based on the reduction of dissolved oxygen gas at water–solid interfaces.<sup>63</sup>

In a recent feature by the Royal Society of Chemistry's magazine, *Chemistry World*, entitled, "Why is Microdroplet Chemistry Contentious?,"<sup>64</sup> the following excerpt appeared, "Zare's team made its hydrogen peroxide discovery while trying to synthesize gold nanostructures in microdroplet, armed with the knowledge that Michael Faraday had made gold films with lemon juice. They added sodium borohydride – which Zare calls modern lemon juice – to water microdroplets and found that the reaction occurred 100 000 times faster than in bulk liquid. As a control experiment, the team removed the reducing agent – the sodium borohydride – yet still produced gold nanoparticles. That was a shock to us, says Zare." In response, here, we extend our investigation of H<sub>2</sub>O<sub>2</sub> production in water microdroplets to the spontaneous reduction of Au<sup>3+</sup>(aq) to Au nanoparticles (Au NPs) containing chloroauric acid (HAuCl<sub>4</sub>, 100 μM concentration).<sup>65</sup> Zare and colleagues have speculated that this chemical transformation occurs exclusively at the air–water interface. While it has been reported that gold nanoparticles can be formed in bulk water by supplying energy *via* microwaves,<sup>66</sup> how water microdroplets (*i.e.*, the air–water interface) drive this transformation without an external energy source, *e.g.*, electrical voltage or reducing agent, has baffled chemists. Here, we resolve this mystery by addressing the following questions:

- (1) Is the spontaneous reduction of gold in water limited only to water microdroplets, or does it also occur in bulk water?
- (2) If gold ions get spontaneously reduced inside water, does water get oxidized? Is there a peroxide connection?
- (3) What are the effects of the concentration of Au<sup>3+</sup> ions, the water pH, and dissolved oxygen gas on the NP formation?

(4) Do other metallic ions, *e.g.*, Pt<sup>4+</sup>, Al<sup>3+</sup>, Fe<sup>3+</sup>, Gd<sup>3+</sup>, Fe<sup>2+</sup>, Cu<sup>2+</sup>, Mg<sup>2+</sup>, Ni<sup>2+</sup>, Zn<sup>2+</sup>, and Na<sup>+</sup>, also exhibit this behavior?

(5) Does the air–water interface have unique features to form Au NPs, or can other common liquids such as ethanol, methanol, and acetonitrile also drive the spontaneous reduction of metallic ions?

Our experiments reveal that when salts containing Au<sup>3+</sup> (or Pt<sup>4+</sup> or Fe<sup>3+</sup>) ions are added to water they undergo spontaneous reduction in the bulk phase as well as in microdroplets. In fact, in this study, whenever NPs formed for a specific salt–solvent system, they formed both in the bulk phase and in the sprayed microdroplets. There was no scenario wherein the spontaneous reduction of Au<sup>3+</sup> ions occurred exclusively inside microdroplets but not in bulk.

## Results

In this work, we used MilliQ Advantage 10 (18 MΩ cm) deionized water, albeit the results were identical with HPLC grade water. Microdroplets were generated *via* pneumatic nebulization following the protocols of Zare *et al.*<sup>65</sup> Briefly, the spray device comprised two concentric capillaries, and the solution was injected through a 0.1 mm-wide inner stainless-steel capillary at a rate of 25 μL min<sup>-1</sup>. The solution was sheared *via* high-pressure dry N<sub>2</sub> gas flowing through the outer concentric capillary at 100 psi, forming a microdroplet stream (ESI Fig. S1†).

The effects of the air–water interface on NP formation were analyzed by spraying 100 μM aqueous HAuCl<sub>4</sub> solution *via* pneumatic nebulization, which yielded microdroplets of an average size of ~20 μm (ESI Fig. S1†). In a typical experiment, microdroplets were formed and collected in clean glass vials; also, the effects of solvent evaporation during spraying were compensated by adding bulk water to the glass vials to restore the original (bulk) salt concentration. Then microdroplets of the solution were transferred onto transmission electron microscopy (TEM) grids and SiO<sub>2</sub>/Si wafers for TEM and scanning electron microscopy (SEM), respectively. Both techniques revealed NP formation in sprays. Curiously, the transparent mother solution (100 μM aqueous HAuCl<sub>4</sub> stored at 22 °C inside a laboratory) changed its color to red in 2–3 days and blue in 2–3 weeks (Fig. 2a). The TEM analysis of samples drawn from the mother solutions revealed the presence of gold NPs (Fig. 2c and d, and S3†). Dynamic light scattering (DLS) showed the appearance of a bimodal nanoparticle size distribution in the freshly prepared solutions composed of NPs and aggregates (Fig. 2b and S11b†). X-ray photoelectron spectroscopy (XPS) compared the oxidation states of gold in the newly prepared and one-week-old solutions (Fig. 2e and S11(e),† Methods). The oxidation state of gold in the freshly prepared solutions was +3 and +1, which indicated that the reduction was spontaneous, whereas the dominant oxidation state of gold in the 1 week-old samples was 0. These results demonstrate that the reduction of gold ions occurs spontaneously in bulk water and is not exclusive to the microdroplet environment/geometry.

Next, we investigated whether the microdroplet environment, *i.e.*, the air–water interface, considerably accelerated the



rate of reduction and nanoparticle formation compared with the bulk phase. To this end, bulk solutions of the same concentration (100  $\mu\text{M}$   $\text{HAuCl}_4$ ) but different ages, *i.e.*, varying from seconds to minutes to hours, were pneumatically sheared into water microdroplets (Fig. 1). The flying microdroplets were then intercepted and collected (in the liquid state, *i.e.*, before complete evaporation) and analyzed *via* DLS. The observed particle size distributions were compared against those of the mother solution (Fig. S4<sup>†</sup>). Results revealed that the microdroplets afforded no significant enhancement in the particle sizes over the bulk phase, demonstrating that microdroplets do not have any significant effect on the formation or growth of nanoparticles. In summary, the reduction of gold ions in bulk water is spontaneous, and microdroplets are not essential, or the size of microdroplets is not the driving force for the spontaneity of this reduction reaction.

Since the formation of AuNPs from  $\text{Au}^{3+}$  is a reduction reaction, a parallel oxidation reaction must occur simultaneously. Therefore, to reveal the redox reaction's oxidation half-reaction, we performed solution-state NMR spectroscopy to investigate the possible oxidation products in a liquid state. Gas chromatography (GC) was performed to determine the evolution of the formed gases. NMR spectroscopy revealed the spontaneous formation of  $\text{H}_2\text{O}_2(\text{aq})$  in the bulk  $\text{HAuCl}_4$  solution, and GC revealed the gradual evolution of oxygen ( $\text{O}_2$ ) gas in the headspace (Fig. 3a and S5<sup>†</sup>). The formation of  $\text{H}_2\text{O}_2$  was also quantified using a Hydrogen Peroxide Assay Kit (HPAK) (Fig. 3b). The mechanistic insights into this chemistry are presented in the Discussion section.

Fig. 3a presents H-NMR data comparing  $\text{H}_2\text{O}_2$  formation in freshly prepared 0.3 M  $\text{HAuCl}_4$  solution in MilliQ water against a freshly prepared 0.3 M  $\text{HAuCl}_4$  solution in 200  $\mu\text{M}$  standard

$\text{H}_2\text{O}_2$  solution. The presence of only one peak in both the solutions, at 9.4 ppm, is observed by following the protocol of Bax and co-workers (detection limit  $\geq 50$  nM).<sup>67</sup> Using the same technique, we found that the concentration of  $\text{HAuCl}_4$  increases in the aqueous solution, so does the spontaneous  $\text{H}_2\text{O}_2(\text{aq})$  formation (Fig. 3b), *i.e.*, they appear to be stoichiometric (see Discussion). We also examined  $\text{H}_2\text{O}_2(\text{aq})$  formation in  $\text{HAuCl}_4$  solutions prepared using water with dissolved oxygen and deoxygenated water. The dissolved oxygen had no measurable effect on  $\text{H}_2\text{O}_2$  formation in these experiments (Fig. S6<sup>†</sup>). Thus, we confirmed that the formation of  $\text{H}_2\text{O}_2(\text{aq})$  in bulk water is the oxidation half-reaction during the spontaneous formation of Au NPs.

Next, we investigated the effects of the solution pH and concentration of  $\text{Au}^{3+}$  ions on the formation of AuNPs and corresponding  $\text{H}_2\text{O}_2$  produced. As the solution turned acidic, the formation of the AuNPs and  $\text{H}_2\text{O}_2$  decreased (Fig. 4a–d). SEM imaging revealed that when a 200  $\mu\text{M}$   $\text{HAuCl}_4$  solution was prepared using deionized water, or 0.1 M HCl, or 1 M HCl, the nanoparticles were well-formed (Fig. 4b), or smaller and fewer (Fig. 4c), or nearly absent (Fig. 4d), respectively. This indicates that gold ions are more stable in lower pH or highly acidic aqueous solutions. Following this observation, we investigated the effects of the gold ion concentration on the formation of Au NPs and  $\text{H}_2\text{O}_2$ . Experimental results revealed that gold ions could undergo complete reduction to form Au NPs in relatively diluted solutions ( $<0.05$  M), whereas in concentrated solutions ( $>0.05$  M),  $\text{Au}^{3+}$  primarily reduced to lower oxidation states (mostly  $\text{Au}^{+1}$ ), while  $\text{H}_2\text{O}_2$  was produced stoichiometrically. XPS pinpointed that as the salt concentration increased, the ratio of hydrated ions  $\text{Au}^{+1}/\text{Au}^{3+}$  decreased (Fig. 4e).

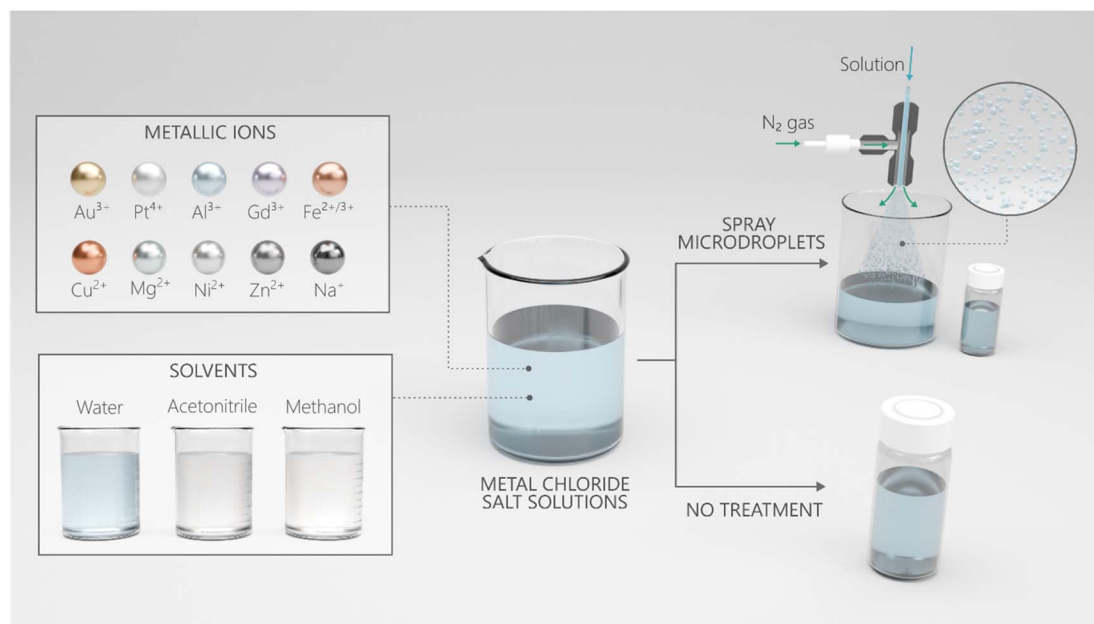


Fig. 1 Experimental design composed of solutions made from a variety of metallic salts and solvents. Nanoparticle formation was probed in bulk solutions and sprayed microdroplets to disentangle the contributions of bulk redox chemistry, microdroplet geometry, and the air–water interface.

To further elucidate whether the mechanism underlying this chemical transformation, *i.e.*, the spontaneous reduction of  $\text{Au}^{3+}$  in water, is unique or in common with other metal ions, aqueous solutions of a few common metal ions such as  $\text{Pt}^{4+}$ ,  $\text{Al}^{3+}$ ,  $\text{Fe}^{3+}$ ,  $\text{Gd}^{3+}$ ,  $\text{Fe}^{2+}$ ,  $\text{Cu}^{2+}$ ,  $\text{Mg}^{2+}$ ,  $\text{Ni}^{2+}$ ,  $\text{Zn}^{2+}$ , and  $\text{Na}^+$  were studied. We found that two metal cations,  $\text{Pt}^{4+}$  and  $\text{Fe}^{3+}$ , also produced  $\text{H}_2\text{O}_2$  in water (Fig. 5c and d), while the remaining metallic ions did not produce  $\text{H}_2\text{O}_2$  in bulk solutions (or microdroplets). Notably, considerably higher amounts of  $\text{H}_2\text{O}_2$  were produced by  $\text{Au}^{3+}$  reduction than those produced *via*  $\text{Pt}^{4+}$  and  $\text{Fe}^{3+}$  reduction.

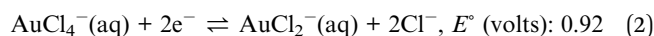
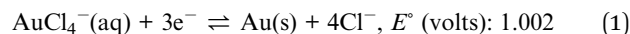
XPS analysis confirmed that  $\text{Pt}^{4+}$  and  $\text{Fe}^{3+}$  were spontaneously reduced in water similar to  $\text{Au}^{3+}$ , yielding  $\text{Pt}^{2+}$  and  $\text{Fe}^{2+}$  (Fig. 5a and b), and the formation of Pt NPs was observed by TEM imaging (Fig. S7†). Notably, the reduction of  $\text{Fe}^{3+}$  to its zero-valent form was not observed. In other words, of all the metal ions studied herein, the pure metallic form of NPs was only observed in  $\text{HAuCl}_4$  and  $\text{PtCl}_4$  solutions.

## Discussion

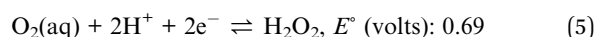
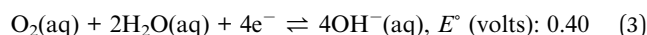
This section consolidates all the findings and presents mechanistic insights. Our experiments have revealed that  $\text{Au}^{3+}$  ions from  $\text{HAuCl}_4$  get spontaneously reduced in bulk water, forming Au NPs and generating  $\text{H}_2\text{O}_2(\text{aq})$ . Spraying bulk solutions (freshly prepared or aged) and collecting the microdroplets reveal no significant enhancement in the size of AuNPs, suggesting that the microdroplet environment's effects, if any, are

minor, such as facilitating solvent evaporation. Based on our experimental results, wherein  $\text{Au}^{3+}$  ions in bulk  $\text{HAuCl}_4$  solutions spontaneously reduce to  $\text{Au}^+$  ions and AuNPs and drive the formation of  $\text{H}_2\text{O}_2(\text{aq})$  (Fig. 2, 3, and S5†) and decrease in the solution pH, we suggest the following half-reactions:<sup>40,68</sup>

Reduction half-reactions:



Possible feasible oxidation half-reactions that may drive the above reduction reactions:



The standard reduction potentials of eqn (1) and (2) are higher than those of eqn (3) and (4); therefore, it is possible that  $\text{OH}^-(\text{aq})$  ions (eqn (3)) and  $\text{H}_2\text{O}$  may be oxidized to form  $\text{H}_2\text{O}_2(\text{aq})$  (eqn (4)), which later decomposes/oxidizes gradually with time and forms  $\text{O}_2(\text{g})$  (eqn (5)) further reducing gold ions.

Out of curiosity, we also probed the behavior of  $\text{HAuCl}_4$  in methanol, ethanol, and acetonitrile, to see if there was something special about water in the nanoparticle formation. We observed that  $\text{Au}^{3+}$  reduced faster in methanol (and ethanol)

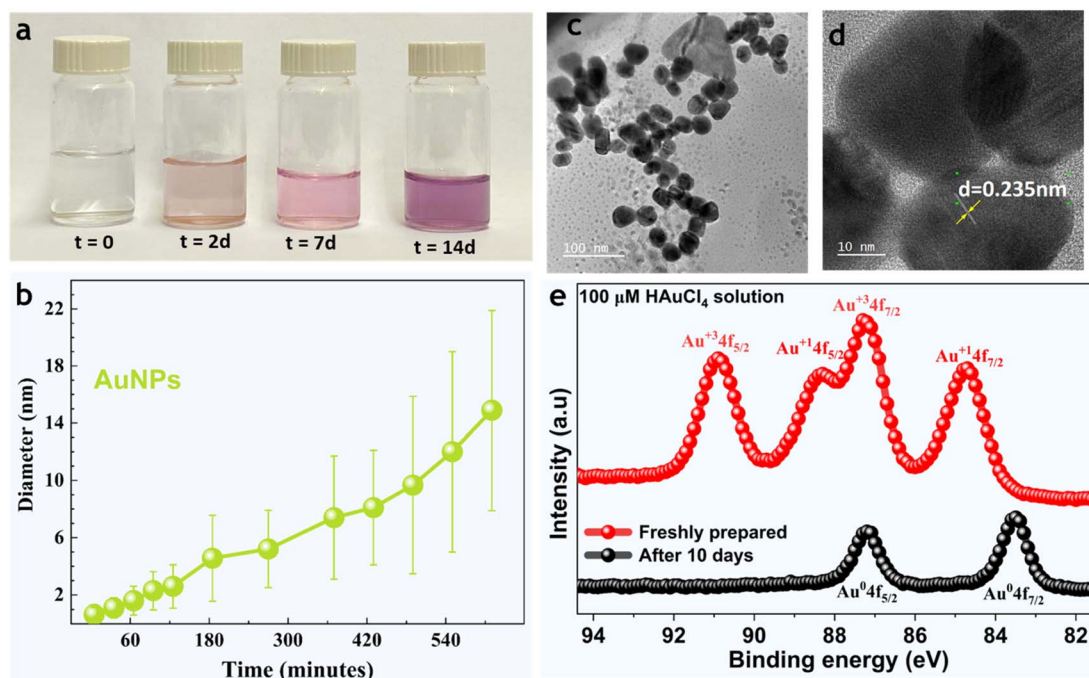


Fig. 2 Spontaneous formation and growth of AuNPs in bulk 100  $\mu\text{M}$   $\text{HAuCl}_4$  in water. (a) Bulk solution changing color due to the nucleation and growth of AuNPs. (b) Formation and growth of particles with time, as measured by DLS. (c) TEM micrograph showing the presence of AuNPs of different sizes. (d) TEM imaging showing an interlayer spacing of  $\sim 0.235$  nm corresponding to the Au(111) atomic plane of metallic gold Au(0). (e) XPS analysis showing the presence of  $\text{Au}^{3+}$  and  $\text{Au}^+$  in the freshly prepared solution and Au(0) in the 10-days-old solution, revealing the complete reduction of gold ions to AuNPs after some time.

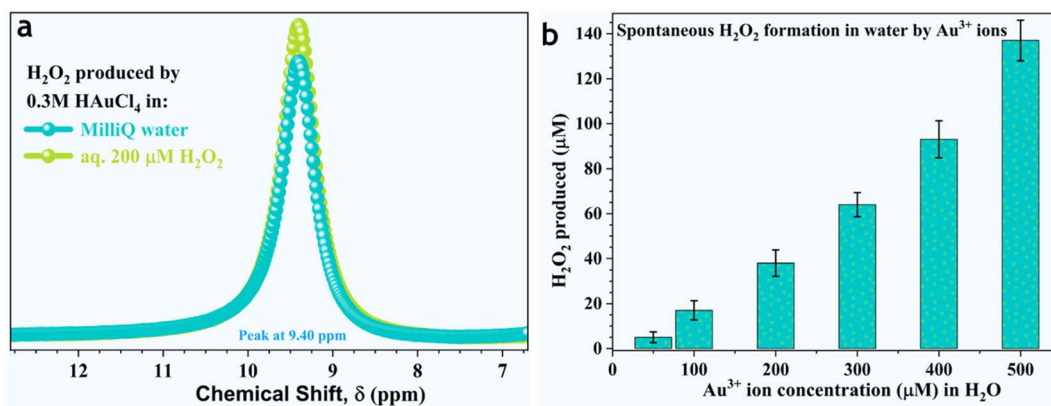


Fig. 3 Production of  $\text{H}_2\text{O}_2$  during the reduction of  $\text{Au}^{3+}$  ions in bulk water. (a) NMR spectroscopy pinpoints the formation of  $\text{H}_2\text{O}_2$ . (b) Quantification of  $\text{H}_2\text{O}_2$  in freshly prepared aqueous solutions as a function of  $\text{HAuCl}_4$  concentration.

than in bulk water, and it was negligible/sluggish in acetonitrile (Fig. 6a and b). The formation of Au NPs in methanol was accompanied by the formation of  $\text{H}_2\text{O}_2$  (Fig. S12<sup>†</sup>), methylal ( $\text{CH}_3\text{OCH}_2\text{OCH}_3$ ), and dimethyl ether ( $\text{CH}_3\text{OCH}_3$ ) (Fig. S8<sup>†</sup>). Note: our investigation of the spontaneous formation of gold nanoparticles in the organic solvents was merely a curiosity, and an in-depth investigation of reaction byproducts and underlying mechanisms falls out of the scope of this manuscript. Lastly, in a complementary experiment, we quantified the amount of the reducing agent (standard hydrogen peroxide, 30%) required to completely reduce a fixed concentration of gold solution ( $0.1 \text{ M HAuCl}_4$ ) in methanol, water, and acetonitrile. The volume percent of  $\text{H}_2\text{O}_2$  required was  $0.4 \pm 0.1$ ,  $1.0 \pm 0.2$ , and  $35 \pm 1$ , respectively.

Now we discuss why some of the metal ions formed NPs in water, while others did not. For instance, we found that  $\text{Pt}^{4+}(\text{aq})$  was reduced to  $\text{Pt}(\text{s})$  NPs and  $\text{Pt}^{2+}(\text{aq})$ , whereas  $\text{Fe}^{3+}(\text{aq})$  was

reduced only to  $\text{Fe}^{2+}(\text{aq})$  ions and  $\text{Fe}^0(\text{s})$  was not observed (Fig. 5 and S7<sup>†</sup>). Our observation about Fe is substantiated by the NIST XPS database<sup>69</sup> and high-resolution XPS spectra (Fig. 5b) in which the Fe 2p peak is broad and asymmetric, suggesting at least two oxidation states of Fe in the solution (Fig. 5b). The bottom line is that the reduction potentials of  $\text{Pt}^{4+}(\text{aq})$  to  $\text{Pt}(\text{s})$  and  $\text{Pt}^{2+}(\text{aq})$  and that of  $\text{Fe}^{3+}(\text{aq})$  to  $\text{Fe}^{2+}(\text{aq})$  were more thermodynamically favored than the water oxidation reactions ((eqn (3) & (4) and ESI Table S2<sup>†</sup>). As the reduction potentials of  $\text{Fe}^{3+}(\text{aq})$  and  $\text{Fe}^{2+}(\text{aq})$  to  $\text{Fe}(\text{s})$ , *i.e.*,  $-0.037$  and  $-0.44$  V, are lower than those listed in eqn (3) and (4), it is not feasible for  $\text{Fe}^{3+}(\text{aq})$  or  $\text{Fe}^{2+}(\text{aq})$  to reduce to  $\text{Fe}(\text{s})$  by water. Similarly, the spontaneous reduction of  $\text{Al}^{3+}$ ,  $\text{Cu}^{2+}$ ,  $\text{Ni}^{2+}$ ,  $\text{Zn}^{2+}$ , and  $\text{Mg}^{2+}$  ions in water was also blocked (ESI Table S2<sup>†</sup>).

Next, we discuss the connection of the present findings with our recent reports unraveling the factors and mechanisms underlying the spontaneous formation of  $\sim 1 \mu\text{M}$   $\text{H}_2\text{O}_2(\text{aq})$  in

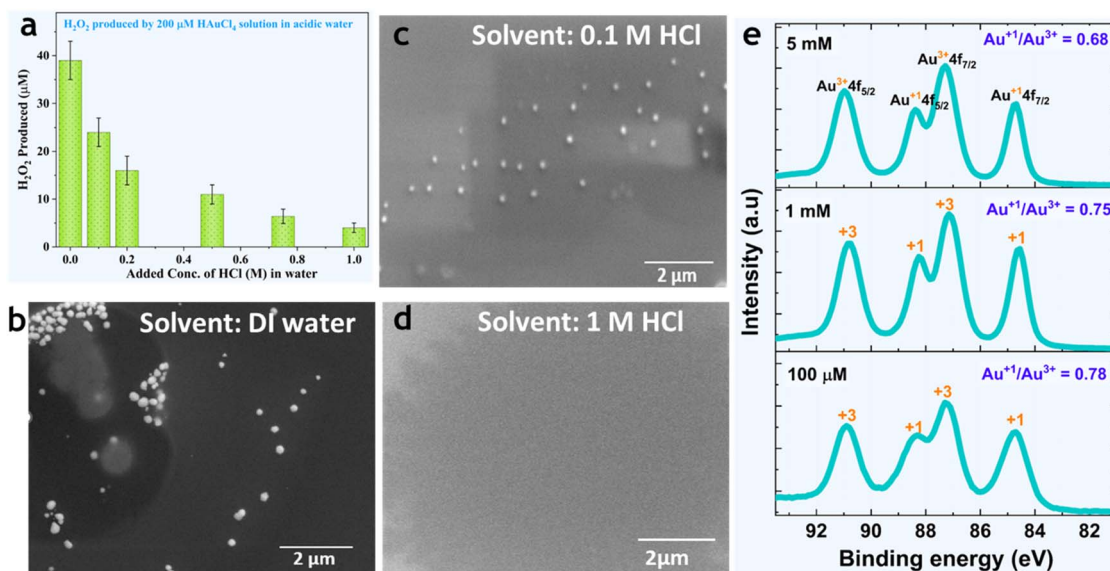


Fig. 4 Effects of the solution pH and concentration on the reduction of gold ions: (a) decrease in the rate of  $\text{H}_2\text{O}_2$  production in bulk solutions as a function of HCl concentration (or pH) for a fixed  $\text{HAuCl}_4$  concentration. Representative scanning electron micrographs (SEM) of gold NPs formed in a  $200 \mu\text{M HAuCl}_4$  solution formed with: (b) deionized water, (c)  $0.1 \text{ M HCl}$ , and (d)  $1 \text{ M HCl}$  solution. (e) XPS reveals the ratio of  $\text{Au}^{1+}/\text{Au}^{3+}$  in freshly prepared  $\text{HAuCl}_4$  solutions of varying concentrations ( $100 \mu\text{M}$ ,  $1 \text{ mM}$ , and  $5 \text{ mM}$ ).

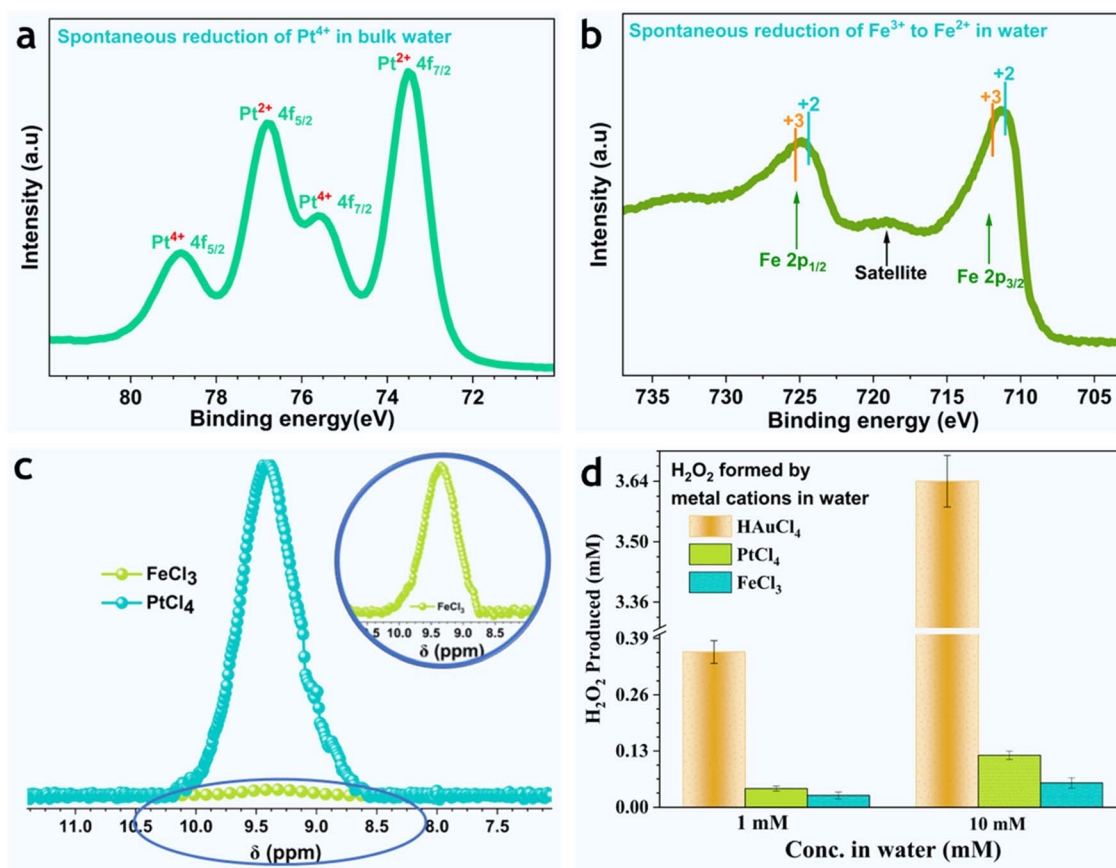


Fig. 5 Spontaneous reduction of metal ions, besides Au<sup>3+</sup>, in water, which yields H<sub>2</sub>O<sub>2</sub>(aq). (a) XPS results of 100 μM PtCl<sub>4</sub> in water showing the reduction of Pt<sup>4+</sup> to Pt<sup>2+</sup>; Pt NPs observed *via* TEM images (Fig. S7†). (b) XPS results of FeCl<sub>3</sub> in water showing the spontaneous reduction of Fe<sup>3+</sup> ions to Fe<sup>2+</sup>. (c) NMR spectra confirming the formation of H<sub>2</sub>O<sub>2</sub>(aq) in aqueous solutions of FeCl<sub>3</sub> and PtCl<sub>4</sub> in water (each at a 100 μM initial concentration). (d) Comparison of H<sub>2</sub>O<sub>2</sub>(aq) formed by different concentrations of Au<sup>3+</sup>, Pt<sup>4+</sup>, and Fe<sup>3+</sup> in water.

water microdroplets.<sup>61–63</sup> We have established that H<sub>2</sub>O<sub>2</sub>(aq) formation is independent of the air–water interface or the microdroplet geometry. Instead, it entails the reduction of dissolved O<sub>2</sub>(aq) at the numerous water–solid interfaces involved in such experiments, such as pipes, tubing, containers, vials, plates, *etc.* Conversely, if dissolved O<sub>2</sub>(aq) is removed from water, then H<sub>2</sub>O<sub>2</sub>(aq) formation is not observed in bulk, films, or microdroplets within a limit of detection of 50 nM. While some reports have corroborated our findings,<sup>70–74</sup> others have disagreed.<sup>75–78</sup> In contrast, the spontaneous NP formation and H<sub>2</sub>O<sub>2</sub>(aq) generation in water (bulk, films, or microdroplets) are driven by the reduction potential of metal ions and their concentration, and can be explained simply *via* the electrochemical series.<sup>40</sup>

Here we list some limitations of this study. First, we did not focus on the shape specificity of the nanoparticles/nanowires as a function of the bulk HAuCl<sub>4</sub> concentration or as a function of solvent evaporation during spraying. Experiments with bulk HAuCl<sub>4</sub> solutions revealed that as the salt concentration exceeded 200 μM, nanowires began to appear – in bulk and in sprayed microdroplets without detectable differences (Fig. S13†). We note that if the solvent evaporation during pneumatic spraying becomes excessive, a regime that we did not study, we anticipate the precipitation of HAuCl<sub>4</sub> salt out of water, *i.e.*, we do not expect

a new interfacial chemistry to arise such that Au<sup>3+</sup> ions specifically start reducing at the air–water interface to form Au NPs. Lastly, we did not take any special precautions to eliminate the existing nanoparticle impurities in the stock salts that may have accelerated the nucleation of nanoparticles in bulk solutions.<sup>79</sup>

In summary, our report refutes the claims that water microdroplets or the air–water interface has unique properties that can drive the spontaneous reduction of gold ions. This chemical reduction takes place in the bulk phase, and it is not limited to (i) gold ions, or (ii) water as a solvent, or (iii) microdroplets or the air–water interface. The entire phenomenon can be explained by the standard electrochemical series, *i.e.*, cations with very high reduction potential such as Au<sup>3+</sup> and Pt<sup>4+</sup> are so unstable that they can oxidize the solvent (water, methanol, and ethanol) to form NPs and H<sub>2</sub>O<sub>2</sub>(aq) and other byproducts in the case of alcohols. These findings advance our understanding of aquatic chemistry and warrant caution in investigating microdroplet chemistry and its environmental and practical relevance.

## Methods

### Chemicals

Gold(III) chloride trihydrate (HAuCl<sub>4</sub>·3H<sub>2</sub>O, 520918), platinum(IV) chloride (PtCl<sub>4</sub>, 206113), iron(III) chloride hexahydrate



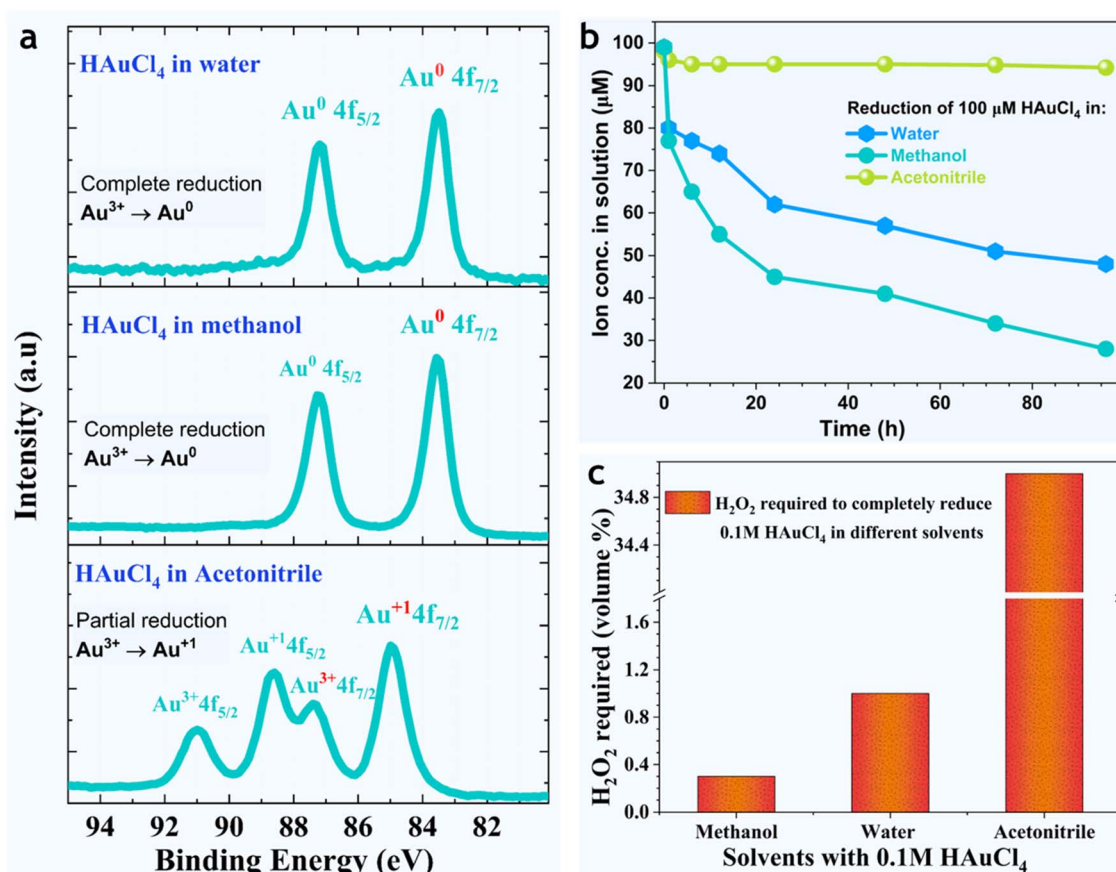


Fig. 6 Reduction of gold ions in different solvents. (a) Reduction of Au<sup>3+</sup> in water, methanol, and acetonitrile. (b) Time-dependence of Au<sup>3+</sup> ion concentration in water, methanol, and acetonitrile (all starting with 100 µM HAuCl<sub>4</sub>) follows the trend: MeOH > H<sub>2</sub>O ≫ CH<sub>3</sub>-CN. (c) As a reducing agent, the amount of concentrated (30% v/v) H<sub>2</sub>O<sub>2</sub> required for the complete reduction of 0.1 M HAuCl<sub>4</sub> in water, methanol, and acetonitrile. Since the amount of H<sub>2</sub>O<sub>2</sub> required follows the trend: MeOH < H<sub>2</sub>O ≪ CH<sub>3</sub>-CN, it demonstrates that gold ions can undergo spontaneous (complete) reduction in water and methanol but not in acetonitrile. Note: the behaviors of methanol and ethanol were similar (Fig. S12†).

(FeCl<sub>3</sub>·6H<sub>2</sub>O, 10025-77-1), iron(II) chloride (FeCl<sub>2</sub>, 37287-0), aluminum(III) chloride (AlCl<sub>3</sub>, 06220), zinc chloride (ZnCl<sub>2</sub>, 211273), magnesium chloride hexahydrate (MgCl<sub>2</sub>·6H<sub>2</sub>O, 13152), sodium chloride (NaCl, 7647145), ammonium chloride (NH<sub>4</sub>Cl, 213330), and copper chloride (CuCl<sub>2</sub>, 7447394) were used. The following liquids were also used: deuterium oxide (D<sub>2</sub>O, 3000007892), methanol-D<sub>4</sub>+0.03%TMS, acetonitrile-D<sub>3</sub> (CAS 2206-26-0), acetonitrile (HPLC grade purchased from Fisher Scientific, batch 1072451), methanol (HPLC LC-MS grade), standard hydrogen peroxide (H<sub>2</sub>O<sub>2</sub>) 30% (Cat. 270733), HPLC grade water (Cat. 2594649), and deionized water (DI) obtained from a MilliQ Advantage 10 set-up (with resistivity 18.2 MΩ cm).

### Characterization of NPs

SEM and TEM analyses were performed to characterize the NPs in the bulk solutions. For SEM, a small drop of liquid containing NPs was drop-coated on a silicon wafer substrate, and the solution was dried at room temperature. TEM images were acquired using a Titan ST Image Corrected (Thermo Fisher)

instrument operated at a 300 kV acceleration voltage. EDS spectra and maps were acquired in the STEM mode using a four-quadrant SuperX EDS detector. For sample preparation, a drop of solution was placed on an ultra-thin carbon-coated holey copper grid, blotted with filter paper, and dried under ambient conditions. The presence and growth of NPs were examined *via* DLS. The growth of Au NPs was observed in 100 µM HAuCl<sub>4</sub> bulk solution at varying time scales.

### XPS measurements

For the XPS studies of gold ion reduction in water, 100 µM HAuCl<sub>4</sub> solution was used. The drop coating method was employed to prepare the samples using silicon wafers, and Baer's method was closely followed.<sup>80</sup> A Kratos Axis Supra instrument equipped with a monochromatic Al Kα X-ray source ( $h\nu = 1486.6$  eV) operating at a power of 75 W and under UHV conditions in the range of  $\sim 10^{-9}$  mbar was used to obtain the data. All the spectra were recorded in hybrid mode, using magnetic and electrostatic lenses and an aperture slot of 300 µm × 700 µm. The high-resolution spectra were acquired at



a fixed analyzer pass energy of 20 eV. The samples were mounted in a floating mode to avoid differential charging.

### Quantification of H<sub>2</sub>O<sub>2</sub>

H<sub>2</sub>O<sub>2</sub> concentration in all the diluted salt solutions was quantified using a hydrogen peroxide assay kit (HPAK). This is based on the principle that when hydrogen peroxide maintains contact with the AbIR peroxidase indicator, it causes fluorescence. Its maximum emission and excitation wavelengths are 674 and 647 nm, respectively. The samples were analyzed by mixing 50  $\mu$ L of HPAK reaction mixture with 50  $\mu$ L of samples in a 96-well black/transparent bottom microtiter plate using a SpectraMax M3 microplate reader. For the fluorescence reading, the SoftMax Pro 7 software was used. H<sub>2</sub>O<sub>2</sub> in the samples was quantified using the calibration curve obtained from the standard samples on the same day.

**Peroxide test strips for semi-quantitative analysis.** For the qualitative estimation of H<sub>2</sub>O<sub>2</sub> in the aqueous samples, peroxide test strips (Baker Test Strips purchased from VWR International) with a detection limit of 1 ppm were used. These strips use a colorimetric reagent that changes to blue when in contact with H<sub>2</sub>O<sub>2</sub>.

### NMR spectroscopy for H<sub>2</sub>O<sub>2</sub>(aq) quantification

NMR measurements were performed on a 700 MHz Bruker Avance Neo NMR spectrometer equipped with a 5 mm Z-axis gradient TXO Cryoprobe at 275 K. For water samples, a 6 ms Gaussian 90-degree pulse was applied to excite the proton of hydrogen peroxide, followed by a 50 ms acquisition and a 1 ms recycle delay. For acetonitrile samples, a W5 binomial pulse sequence with gradients or a routine 1D proton pulse was applied for solvent suppression or non-suppression measurements, respectively. Peak quantification and peak position confirmation of H<sub>2</sub>O<sub>2</sub> in salt solutions were performed by comparing the results obtained for standard H<sub>2</sub>O<sub>2</sub> samples.

### GC-MS experiment

The methanol sample was extracted by adding 2 mL each of water and hexane to initiate phase separation. Approximately 1  $\mu$ L of the top hexane layer was analyzed using a single quadrupole GC-MS system (Agilent 7890 GC/5975C MSD) coupled to an EI source with an ionization energy of 70 eV. The ion source and mass analyzer temperatures were kept at 230 °C and 150 °C, respectively, with a solvent delay of 0 min. The mass analyzer was tuned according to the manufacturer's instructions, and the scan was fixed at 15–200 Da. A DB-5MS fused silica capillary column (30 m  $\times$  0.25 mm I.D., 0.25  $\mu$ m film thickness; Agilent J&W Scientific, Folsom, CA) was used for chromatographic separation; the column contained a 5% phenyl and 95% methylpolysiloxane cross-linked stationary phase. Furthermore, helium was the carrier gas at a constant flow rate of 1.0 mL min<sup>-1</sup>. The sample was injected into a 6890A gas chromatograph (Agilent, USA). The oven program was set to 30 °C and held for 10 min; the temperature was ramped at 20 °C min<sup>-1</sup> to 260 °C with a 0 min hold time. The GC inlet temperature was set at 250 °C, and the transfer line temperature to the MS EI source

was kept at 320 °C. The sample was injected using an auto-sampler equipped with a 10  $\mu$ L syringe into a split/splitless inlet with a split ratio 10 : 1.

### Oxygen gas analysis

Headspace analysis of the aqueous HAuCl<sub>4</sub> solution was performed using a gas analyzer (Model 310C, SRI, USA) equipped with a thermal conductivity detector (TCD). About 1 mL of the headspace above the sample was withdrawn using a gas-tight syringe (Hamilton, USA) and injected into a 6-foot molecular sieve 13 $\times$  packed column. The oven temperature of the column was set to 100 °C with argon as the carrier gas.

## Data availability

All data needed to evaluate the conclusions in the paper are present in the paper or the ESI.†

## Author contributions

ME and HM designed the experiments, which ME performed. NK and ME performed LC-MS, GC, and ICP-OES experiments. XG and ME performed NMR spectroscopy experiments. NW and ME performed XPS studies. HM and ME wrote the manuscript.

## Conflicts of interest

The authors declare no competing interests.

## Acknowledgements

HM acknowledges KAUST for funding (Grant No. BAS/1/1070-01-01). The co-authors thank Dr Adair Gallo, Dr Peng Zhang, and Ms Nayara Musskopf for building a robust experimental setup in HM's laboratory; Dr Ryo Mizuta, Scientific Illustrator at KAUST, for preparing illustrations in Fig. 1; Dr Valentina-Elena Musteata, scientist from IAC at KAUST for her assistance with TEM; Mohammed Abdulwahab ElShaer from WDRC at KAUST for his assistance with DLS and Usman Sharif from Laboratory Equipment Maintenance (LEM) team at KAUST for his contribution to maintaining the XPS instrument at the best operating conditions.

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