In Situ Nanoscale Investigation of Catalytic Reactions in the Liquid Phase Using Zirconia-Protected Tip-Enhanced Raman Spectroscopy Probes

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ABSTRACT: Tip-enhanced Raman spectroscopy (TERS) is a promising technique that enables nondestructive and label-free topographical and chemical imaging at the nanoscale. However, its scope for in situ characterization of catalytic reactions in the liquid phase has remained limited due to the lack of durable and chemically inert plasmonically active TERS probes. Herein, we present novel zirconia-protected TERS probes with 3 orders of magnitude increase in lifetime under ambient conditions compared to unprotected silver-coated probes, together with high stability in liquid media. Employing the plasmon-assisted oxidation of p-aminothiophenol as a model reaction, we demonstrate that the highly robust, durable, and chemically inert zirconia-protected TERS probes can be successfully used for nanoscale spatially resolved characterization of a photocatalytic reaction within an aqueous environment. The reported improved lifetime and stability of probes in a liquid environment extend the potential scope of TERS as a nanoanalytical tool not only to heterogeneous catalysis but also to a range of scientific disciplines in which dynamic solid–liquid interfaces play a defining role.

The rational design of novel, chemically functional nanomaterials with tailored properties relies on a deep understanding of structure–behavior relationships. This is especially poignant in the case of heterogeneous catalysis, in which the identification of active sites at spatially nonuniform solid–liquid and solid–gas interfaces under dynamic reaction conditions is key to materials and process optimization. However, conventional analytical techniques, such as Raman, infrared (IR), ultraviolet, and fluorescence spectroscopy, often lack the required sensitivity and spatial resolution to achieve this ambitious goal. In recent years, tip-enhanced Raman spectroscopy (TERS) (Figure 1) has emerged as a potential solution to this challenge, enabling nondestructive topographical and molecular imaging of surfaces at the nanoscale.

In TERS, a metallic scanning probe microscopy (SPM) probe, positioned within the excitation laser spot of a Raman microscope (Figure 1), enhances the incident electric field by several orders of magnitude via a combination of localized surface plasmon resonance (LSPR) and lightening rod effects, confining the field enhancement to a region similar in size to the probe apex. This near-field approach significantly improves the sensitivity of Raman microscopy and pushes its spatial resolution far beyond the diffraction limit. This technique has been successfully used in a wide range of research areas including biology, polymer blends, semiconductors, crystal-line materials, solar energy conversion, nanomaterials research, and single-molecule imaging.

Despite the advantages of high spatial resolution and rich chemical information offered by TERS, only a handful of TERS studies of catalytic reactions have been reported so far. Furthermore, the majority of such studies have been carried out in ambient air or ultrahigh vacuum, with only a few in liquid environments, mostly limited to point spectroscopy measurements. 2D nanoscale chemical imaging of catalytic reactions in liquids represents a key challenge but has not been achieved using TERS, primarily due to the chemical reactivity, short lifetime, and/or instability of TERS probes in liquids. The chemical reactivity of metallic TERS probes is a particularly important consideration when studying catalysis in situ due to the potential for interference in the reaction under investigation. To address this issue, as well as extend the lifetime of metallic TERS probes, the use of an ultrathin coating of dielectric material such as Al2O3 or SiO2 as a protective layer has been reported. However, Al2O3 and SiO2 are not stable over the entire pH range, limiting the conditions in which the TERS probes could be used. Furthermore, both Al2O3 and SiO2
coatings only increased the lifetime of TERS probes for approximately 1 month, and although some increased stability in liquids was observed for the Al2O3-protected probes, no TERS mapping was performed. Recently, a more effective method of protecting TERS probes using atomic layer deposition (ALD) has been reported by Huang et al., where the authors presented three different coatings, SiO2, Al2O3, and TiO2, and successfully showed that the SiO2-coated STM-TERS probes could enhance the Raman signal from a self-assembled monolayer of thiol molecules while suppressing the signal from pyridine molecules present in solution. However, TERS data was presented only for the SiO2-coated probes, and the mechanical stability, long-term stability of plasmonic signal enhancement, and capability of 2D chemical imaging in a liquid environment was not demonstrated. Furthermore, the use of ALD to produce ultrathin dielectric coatings requires expensive experimental apparatus and time-consuming optimization of deposition conditions.

Herein, we present novel atomic force microscopy (AFM)-TERS probes with a multilayer metal coating protected using an ultrathin layer of zirconia (ZrO2) that successfully overcome the key limitations of short lifetime, chemical inertness, and instability in a liquid environment. ZrO2 offers excellent catalytic support properties for several reactions and stability over the entire pH range, making it especially suitable for investigating heterogeneous catalytic reactions under a wide range of conditions. Compared to ALD, the simple wet-chemical method of ZrO2 coating developed in this work is far cheaper and could be easily implemented in any laboratory fume cupboard. Furthermore, using the novel TERS probes, we demonstrate the feasibility of mapping of a catalytic reaction over a heterogeneous metal surface in water with nanoscale spatial resolution, employing the photocatalytic oxidation of p-aminophenol (pATP) to p,p′-dimercaptobenzene (DMAB) as a model reaction (Figure 1).

Extending the Plasmonic Lifetime of TERS Probes. Experimental details are presented in Supporting Information (SI) section S1, including development of the ZrO2 coating procedure (Figures S1–S4). TEM measurements confirmed that the procedure was successful in coating a layer of ZrO2 on Ag-coated TERS probes with a thickness of 1–5 nm (Figure S5). Note that the TEM measurement was conducted after using the probe for continuous TERS measurements in water for ~2 h, which demonstrates the stability of the ZrO2 coating. The ZrO2 coating was also found to be stable in an aqueous environment for a long period of time and over the entire pH range (see Figure S3 and S4).

The lifetime of ZrO2-protected TERS probes was compared with that of the unprotected (Ag-coated) TERS probes using a poly(3,4-ethylenedioxythiophene)-poly(styrenesulfonate) (PEDOT:PSS) thin film as a test sample. Plasmonic enhancement of a TERS probe can be monitored using the ratio of the Raman signal intensity in the near-field (plasmonically enhanced electric field at a TERS probe apex) and far-field (electric field in the entire excitation laser spot), which is known as “contrast” and defined as follows

$$\text{Contrast} = \frac{I_{\text{TERS}}}{I_{\text{FF}}} - 1$$

where \(I_{\text{TERS}}\) and \(I_{\text{FF}}\) are the intensities of a Raman band with the TERS probe in contact with and retracted from the sample, respectively. Time series TERS spectra recorded using unprotected and ZrO2-protected TERS probes stored in ambient air are shown in Figure 2a,b, respectively. Additionally, time series spectra recorded using the unprotected and ZrO2-protected TERS probes are presented in Figures S6 and S7. Contrast was determined using the intensity of the 1454 cm\(^{-1}\) PEDOT:PSS band (\(\nu_{\text{sym,C-S},\text{C=O}}\)) and is plotted in Figure 2c,d for the unprotected and protected probes, respectively. The near-field signal intensity of a fresh unprotected TERS probe is found to be almost 3X higher than the far-field signal, indicating a strong plasmonic enhancement of the electric field at the probe apex. However, the TERS contrast decreases rapidly by 50% of its initial value (half-life) within 4.5 h due to surface oxidation of the Ag coating in air. After a rapid initial decline, the contrast decreases rather gradually by 75% in 10 h, eventually reaching 0 after 170 h of exposure to air, indicating a complete loss of plasmonic enhancement. On the other hand, although the ZrO2-protected TERS probes exhibit a decrease in contrast of almost 50% compared to the unprotected probes (after 2 days of preparation), they show a much higher plasmonic stability; even after 140 days of exposure to the ambient environment, the TERS contrast decreased by only 43%, indicating that the plasmonic sensitivity for TERS measurements is largely preserved. The pinhole-free ZrO2 layer likely blocks contact of the Ag coating with oxygen and moisture present in the surroundings, thereby preventing plasmonic degradation. From extrapolation of the time series plot in Figure 2d, the half-life of the ZrO2-protected TERS probes is estimated to be over 160 days. This corresponds to more than an 850× increase in lifetime compared to unprotected probes and to the best of our knowledge is the longest lifetime of any TERS probe stored under ambient conditions reported to date. Furthermore, unlike STM-based TERS, these novel ZrO2-protected probes could be employed for nanoscale chemical mapping of a wide range of conducting and nonconducting sample surfaces using an AFM-TERS setup.

Nanoscale Mapping of a Photocatalytic Reaction in a Liquid Environment. We next used ZrO2-protected TERS probes to demonstrate spatially resolved nanoscale characterization of a...
catalytic reaction within an aqueous environment. We selected the photocatalytic oxidation of pATP → DMAB as a model reaction due to the ease of monitoring using the distinct Raman bands of the azo group of DMAB, appearing in the 1140–1500 cm⁻¹ spectral region. pATP → DMAB is a well-understood and well-characterized plasmon-assisted reaction wherein hot electrons generated via LSPR of metal nanoparticles cause dissociation of adsorbed oxygen under ambient conditions, facilitating oxidative dimerization. For example, the root-mean-square surface roughness of a spin-coated PEDOT:PSS thin film of pATP molecules on a heterogeneous Ag substrate containing nanostructures varying self-assembled monolayer (SAM) of pATP molecules on a glass substrate was chosen to measure the lifetime of the zirconia-protected TERS tips because of its high chemical stability, low surface roughness, and strong Raman signal. For example, the root-mean-square surface roughness of a spin-coated PEDOT:PSS film has been shown to be 1.2 ± 0.1 nm using tapping mode AFM, confirming a smooth topography. In order to further minimize the effect of surface roughness, the time series contrast plotted in Figure 2c,d was measured from an average of three TERS and far-field measurements conducted at different areas on the PEDOT:PSS thin film.

We first tested the sensitivity of ZrO2-protected TERS probes for monitoring pATP → DMAB in air and aqueous environments. For measurements in air, we placed the TERS probe in contact with the Ag substrate and carried out Raman mapping the probe apex around. Figure 3a shows the map of the 1437 cm⁻¹ DMAB Raman band intensity, in which a much stronger intensity is observed at the probe apex position. Comparison of average spectra measured at the TERS probe apex in Figure 3a and away from it (Figure S10) showed that the DMAB bands are visible only in the TERS spectra measured at the probe apex, indicating a strong LSPR. Furthermore, the TERS signal measured at the position of maximum intensity in Figure 3a was found to be ~12× stronger than the SERS signal measured at the same position with the probe retracted from the sample, as shown in Figure 3c. To rule out the possibility of the zirconia-protected TERS probe interfering in the reaction at the sample, we conducted similar TERS measurements on a thin film of pATP mixed with PMMA spin-coated on a glass substrate (Figure S11). TERS spectra measured at the TERS probe apex on this sample did not show signal from any DMAB a₁ Raman bands, whereas the a₁ Raman bands of pATP at 1086 and 1591 cm⁻¹ were enhanced by a contrast of 5.3 in the TERS near-field. This demonstrates the chemical inertness of ZrO2-protected TERS probes for investigation of this catalytic reaction.

Next, we examined the sensitivity of the ZrO2-protected TERS probes for monitoring pATP → DMAB in an aqueous environment. In this case, TERS measurements were performed inside of a water droplet placed on the sample surface. Interestingly, compared to the SERS measurements in air, a 210× stronger signal was observed in water, as shown in Figure S12. This is in contrast to previous reports where the far-field Raman signal of molecular SAMs on Au was found to decrease by a factor of >3 in a liquid environment compared to air due to laser focus aberrations. A similar loss of optical coupling in water was also observed in our TERS system during far-field Raman measurements (Figure S13). However, for the pATP SAM on the heterogeneous Ag substrate, we speculate that this effect could arise due to the red shift of the surface plasmon resonance of the SERS substrate in an aqueous environment, which can lead to better matching of the surface plasmon resonance wavelength with the excitation laser, resulting in higher DMAB formation as well as a much stronger SERS signal. However, we cannot rule out other phenomena such as surface molecular diffusion and molecular reorientation, which may be affected by the presence of an aqueous environment. A detailed understanding of this effect is beyond the scope of the present study and warrants a separate investigation, to be pursued as part of our future work. A map of the 1437 cm⁻¹ DMAB band intensity measured around the TERS probe apex is shown in Figure 3b. Once again, a significantly higher DMAB signal intensity is observed at the probe apex, indicating a strong LSPR. See Figure S14 for a comparison of average spectra measured at the probe apex in Figure 3b and away from it. The intensity of the TERS signal in water is found to be ~12× stronger compared to the SERS signal measured at the same location (Figure 3d), clearly showing that the ZrO2-protected TERS probes retain their plasmonic sensitivity in water.

Furthermore, comparison of average spectra measured at the probe apex and away from it in Figure S14 also confirms the chemical inertness rendered by ZrO2 protection. In the Raman...
spectra, the $\nu_{C=S}$ vibrational mode at $\sim 1071$ cm$^{-1}$ is assigned to both pATP and DMAB, whereas the $\nu_{N=N}$ modes at 1390 and 1437 cm$^{-1}$ and the $\beta_{C-H}$ mode at 1142 cm$^{-1}$ are assigned exclusively to DMAB. Therefore, the relative conversion of pATP to DMAB can be assessed using the ratio of the average intensity of DMAB bands ($I_{DMAB}$) to $I_{1071}$, removing the effect of fluctuations in the absolute TERS signal. Calculations of $I_{DMAB}/I_{1071}$ at locations P1–P4 in Figure S14 are presented in Table S1. At all four locations, $I_{DMAB}/I_{1071}$ is found to be $\sim 1.9$, indicating that the conversion of pATP to DMAB is unaffected by the presence of the ZrO$_2$-protected TERS probe, and it can therefore be considered as chemically noninterfering.

Finally, we performed nanoscale chemical imaging of this reaction on a Ag substrate in water. Figure 4a shows an AFM topography image of the surface, and Figure 4b shows a TERS map of the pATP $\rightarrow$ DMAB conversion, obtained using the $I_{DMAB}/I_{1071}$ ratio in an area of 500 $\times$ 500 nm$^2$. The conversion exhibits a high degree of spatial heterogeneity across the Ag substrate employed, and a histogram analysis of the $I_{DMAB}/I_{1071}$ ratio across this map is consistent with a broad range of catalytic activities (Figure 4c). Furthermore, the TERS map in Figure 4b exhibits a number of highly localized regions of notably high pATP $\rightarrow$ DMAB conversion. Some of these "reaction hotspots" are labeled in Figure 4b (marked as 1–3), and corresponding spectra are shown in Figure 4d. For comparison, example spectra from regions of low conversion are also shown in Figure 4d and labeled as 4–6 in Figure 4b. The locations of the most active spots can be seen in the map presented in Figure S15a, in which the contrast has been adjusted to improve visualization, and additional example spectra are shown in Figure S15b. We propose that these extremely confined regions of high conversion reflect the plasmon-assisted nature of the pATP $\rightarrow$ DMAB reaction, in which LSPR at or in between individual Ag nanoparticles can confine light to a $\sim 1$ nm$^3$ volume. We note that a high conversion of pATP to DMAB at a particular position on the Ag substrate could be caused by two factors: (1) strong LSPR coupling between the ZrO$_2$-protected TERS probe apex and the Ag substrate or (2) LSPR within the SERS hotspots present on the Ag substrate. In the current study, it is not possible to distinguish between these two contributing factors as deconvoluting the influence of the TERS probe LSPR on the surface reaction remains a challenging problem beyond the scope of this work. Line profile analysis of the high activity regions (Figure S16 and Table S2) indicates that they are typically confined to one or two pixels (100–200 nm$^2$), consistent with nanoscale spatial resolution. The stability of the ZrO$_2$-protected TERS probe during TERS mapping was also assessed by comparing the first and last spectra recorded in the TERS map (see Figure S17), in which a similar signal intensity was observed.

In summary, in this work, we have addressed the key challenges of plasmonic degradation, chemical inertness, and...
instability within a liquid environment of Ag-coated TERS probes by developing a novel method of protecting them with an ultrathin dielectric layer of zirconia. This successfully extends the lifetime of the probes over 850×, from a few hours to >4.5 months, while rendering them chemically inert for the investigation of catalytic reactions in liquids at the nanoscale. Finally, using these robust and durable probes, we have demonstrated mapping of a model catalytic reaction on a heterogeneous substrate within a liquid environment with nanoscale spatial resolution. This work opens up the possibility of using TERS as a nanoanalytical tool to map molecular heterogeneities and interfacial dynamics in situ in diverse areas of scientific research such as heterogeneous catalyst systems, biology, and electrochemistry, in which nanoscale features at solid–liquid interfaces play a primary role in governing chemical behavior. Furthermore, this clear improvement in the probe lifetime and structural and chemical stability paves the way for using AFM-TERS for nondestructive, label-free, and nanoscale chemical characterization on a wide range of samples and environments.

**ASSOCIATED CONTENT**

**Supporting Information**
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**Figure 4.** (a) AFM topography image of a heterogeneous Ag substrate functionalized with pATP. (b) TERS map of the I_{DMAB}/I_{1071} intensity ratio for the area marked in (a). Integration time: 1 s. Laser power: 117 μW. Pixel size: 10 nm. (c) Histogram showing the % frequency of the I_{DMAB}/I_{1071} ratio in the pATP → DMAB TERS map in (b). (d) TERS spectra from the locations marked as 1–6 in (b) showing different degrees of conversion across the TERS map. Spectra have been normalized to the intensity of the 1071 cm⁻¹ band for comparison.

Experimental details of the TERS system, sample preparation, and preparation of TERS probes including ZrO₂ coating; time series TERS measurements of unprotected Ag-coated TERS probes in air; time series TERS measurements of ZrO₂-protected Ag-coated TERS probes in air; AFM topography of the Ag substrate; comparison of the Raman spectrum of bulk pATP and SERS spectrum of pATP adsorbed on a heterogeneous Ag substrate; analysis of pATP → DMAB at the apex of the Ag-coated TERS probe in air; chemical inertness test of the ZrO₂-protected TERS probe; comparison of pATP → DMAB SERS spectra measured in air and water environments; comparison of far-field Raman spectra measured from a polystyrene thin film on glass in air and water; analysis of pATP → DMAB at the apex of the Ag-coated TERS probe in water; further analysis of the pATP → DMAB TERS map; and comparison of the first and last spectra measured in the TERS map (PDF).
Author Contributions
N.K. and C.S.W. contributed equally to this work.

Notes
The authors declare no competing financial interest.

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