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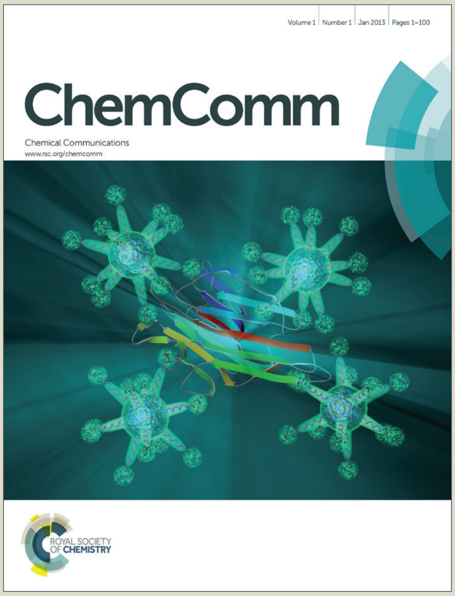

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ARTICLE TYPE

Dual-Responsive Nanoparticles that Aggregate under the Simultaneous Action of Light and CO_2

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Metallic nanoparticles co-functionalised with monolayers of UV- and CO_2-sensitive ligands were prepared and we shown to respond to these two types of stimuli reversibly and in an orthogonal fashion. The composition of the coating could be tailored to yield nanoparticles capable of aggregating exclusively when both UV and CO_2 were applied at the same time, analogously to the behaviour of an AND logic gate.

Stimuli-responsive nanomaterials[1] that self-assemble in a reversible fashion are important for their emerging applications in diverse fields, such as switchable catalysis,[2-3] water purification,[4] detection of analytes,[5] and time-sensitive information storage.[6] Accordingly, a variety of nanoparticles (NPs) have been engineered to aggregate in response to different external stimuli, such as a magnetic field,[7] pH,[8-9] or chemical fuel.[10] For example, dynamic aggregation of metallic NPs was accomplished by reversibly photoswitching the electronic states and geometries of surface-bound chromophores.[11-14] New applications could arise, however, for NPs that respond to two (or more) different external stimuli at once, such as the recently reported dual-responsive NPs that aggregate when either a magnetic field or light is applied.[15-18] Whereas such dual-responsiveness is, in principle, relatively easy to "encode" in the NPs, it is more challenging to design systems capable of aggregating only when two types of external stimuli are at applied at the same time.[19-20] Here, we describe the first system exhibiting such behaviour with light and carbon dioxide as the two "inputs".

Carbon dioxide is a biocompatible, naturally abundant gas that has several advantages over other chemical stimuli: *i*) facile reactivity with various nucleophilic Lewis base functional groups such as amines, NHCs (*N*-heterocyclic carbenes), or guanidines, *ii*) easy removal from the above complexes by mild heating, sonication, or purging with an inert gas, *iii*) no by-product formation through the reversible complex formation, and *iv*) potential applicability to reversible CO_2 capture and/or transformations.[21-24] In addition, nanomaterials that aggregate reversibly in the presence of CO_2 are potentially important as possible anti-cancer therapeutics (taking advantage of the increased production of CO_2 in cancer cells).[25-26] Although many CO_2-sensitive materials,[27] in particular, organic polymers,[28-29] have been reported, the application of amine-functionalised metallic NPs as CO_2-responsive materials remains unknown. Here, we presumed that control of the NP surface polarity by using fine-tuned amounts of photo- and CO_2-sensitive ligands could induce selective and reversible aggregation of NPs with two orthogonal stimuli (Figure 1).

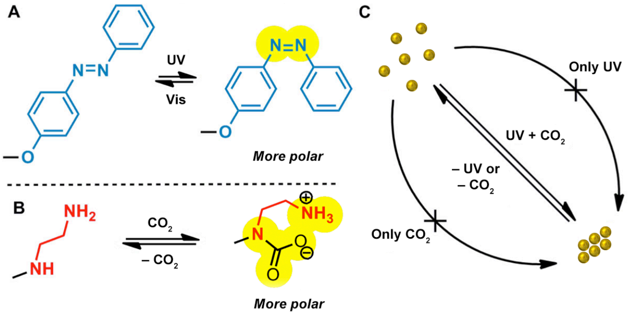


Figure 1. (A) Reversible isomerisation of a photoswitchable ligand. (B) Reversible CO_2 complexation by a diamine ligand. (C) A dual stimuli-responsive AND logic gate.

It is well established that mixtures of ω-functionalised long-chain alkanethiols can form mixed self-assembled monolayers (*m*SAMs) on the surfaces of metallic NPs[30] (here, we focused on gold NPs). We commenced our study with AuNPs functionalised with *m*SAMs comprising an azobenzene-terminated thiol (**AzoSH**) and a "background" ligand, 1-dodecanethiol (**C_{12}SH**) (Figure 2a). As the UV-Vis spectra in Figure 2b show, NPs decorated with critical (>50%) amounts of **AzoSH** aggregate in toluene upon exposure to long-wave (λ = 365 nm) UV light,[31] whereas they exhibited no detectable response to CO_2 (confirmed by dynamical light scattering (DLS) measurements; see ESI†, Figure S2 in Section 5). Furthermore, we observed that the *trans*-to-*cis* photoisomerisation of **AzoSH** did not affect the affinity of these NPs towards CO_2 (Figure S2). Therefore, we concluded that those AuNPs bearing only the **AzoSH** and **C_{12}SH** ligands are inert towards the CO_2 stimulus under these conditions.

The lack of general knowledge on the construction of CO_2-sensitive metal-nanoparticle surfaces encouraged us to investigate the synthesis and application of thiol-terminated amine ligands (Figure 1b).[32] Based on previous literature reports,[33-36] we presumed that ethylenediamine derivatives are capable of CO_2 complexation under low-pressure conditions. To verify the effect of CO_2 binding on the solubility in toluene, we synthesised and studied the behaviour of a model compound, *N*-decylethylenediamine. As detailed in the SI (Section 3), this diamine was readily soluble in toluene; however, rapid precipitation commenced upon bubbling CO_2 through the solution (Fig. S1). The process was reversible: the addition of an inert gas (e.g. N_2) caused the precipitate to redissolve. Encouraged

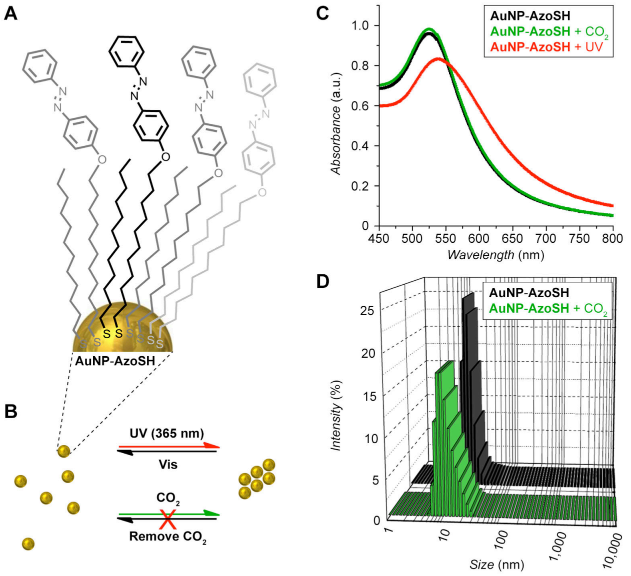
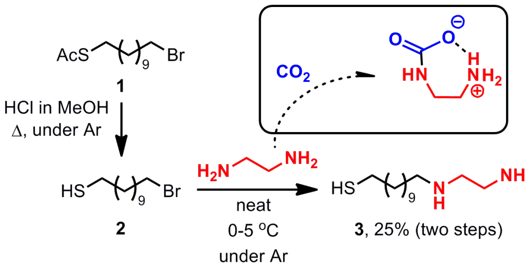


Figure 2. Light-sensitive ligand (**AzoSH**)-decorated gold nanoparticles (AuNP-**AzoSH**) (A) aggregate in response to light why showing to response to CO_2 (B). UV-Vis absorption spectra (C) and DLS (D) of toluene solutions of AuNP-**AzoSH** before and after exposure to UV and bubbling CO_2.

by these results, we then proceeded to synthesise AuNPs decorated with the diamine moieties. Although diamine-functionalised NPs were previously prepared by reacting 11-bromoundecane-1-thiol (**2**)-decorated NPs with 1,2-ethylenediamine, we began our investigation by preparing a well-defined thiol bearing the diamine moiety. As shown in Scheme 1, the synthesis of diamine **3** was straightforward, starting with a known material **1**.[37-39]



Scheme 1. Synthesis of diamine ligand **3**.

With the diamine-functionalised thiolate ligand **3** in hand, we prepared a family of dual stimuli-responsive AuNPs by varying the ratio of the two responsive ligands (**AzoSH** and diamine **3**) and the background ligand 1-dodecanethiol (**C_{12}SH**), as shown in Figure 3. Although **3** could potentially bind to AuNPs via both the mercapto and the amine functionalities, the latter scenario is unlikely given the ca. two-orders-of-magnitude higher affinity of Au to thiols as compared to amines.[12] In addition, binding to Au via the amine functionalities would cause NP crosslinking and irreversible aggregation, which we did not observe.[40]

Owing to the high polarity and low toluene solubility of the diamine ligand **3**, the ligand exchange reaction was conducted under diluted (ca. 10 times) conditions, compared with typical ligand exchange reaction conditions.[41] Moreover, **3** was always used after drying under vacuum at 40 °C to ensure that it was not complexed with CO_2. The functionalised AuNPs were precipitated with methanol (1:1 v/v) in the presence of didodecyldimethylammonium bromide (0.5 mg per 0.716 mg of Au), washed extensively with methanol to remove any unbound molecules, dried under vacuum and kept in an inert atmosphere prior to use. A good indication of the presence of **3** on the functionalised AuNPs is their good solubility in polar solvents (e.g., ethanol, methanol and even water), which confirms a successful ligand-exchange reaction (for a comparison, AuNP-**AzoSH** are only soluble in non-polar solvents such as toluene, chloroform, or dichloromethane). The functionalisation procedure did not affect the size or size distribution of the NPs, whose monodispersity was confirmed by transmission electron microscopy (Figure 5a).

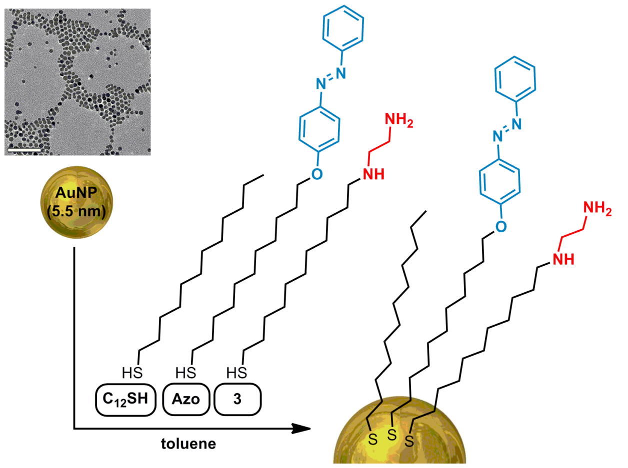


Figure 3. Preparation of dual-responsive AuNPs functionalised with an *m*SAM comprising **C_{12}SH**, **AzoSH** and diamine **3** (the scale bar in the TEM image is 50 nm).

Fine-tuning of the ratio of the three ligands was critical for obtaining NPs exhibiting the behaviour shown in Fig. 1c (in this context, it is worth mentioning that the previously adapted[32, 34] on nanoparticle reaction between ethylenediamine and **2**-functionalised

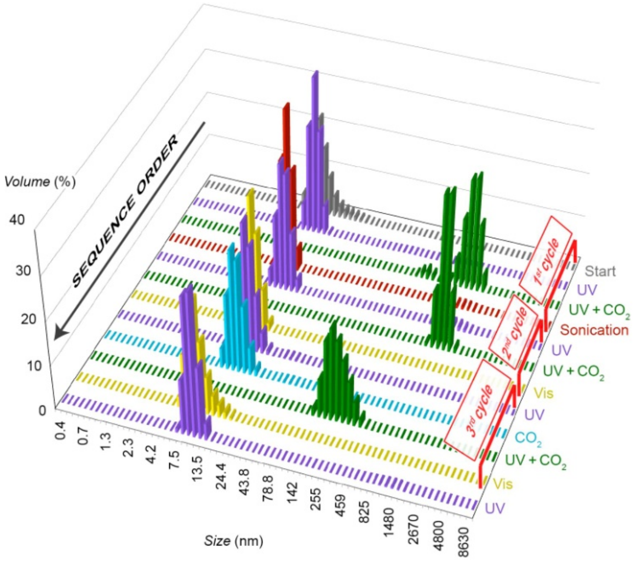


Figure 4. Reversible, dual stimuli-responsive clustering of AuNPs studied by dynamic light scattering.

AuNPs would hardly be applicable). We found that when the molar percentage of **3** on the NPs exceeded 25%, the particles were insoluble in toluene due to their high surface polarity. These aggregated NPs could not be redissolved using visible light, sonication, or by purging with N_2 (to remove any possibly complexed CO_2). Therefore, the amount of the diamine **3** was fixed at 25% throughout the optimisation of the *m*SAM composition. Having screened the ratio of the remaining two ligands (**AzoSH** and **C_{12}SH**) from 1:2 to 2:1, we found that NPs containing 33-35 mol% of **AzoSH** within the *m*SAM exhibited the behaviour shown in Fig. 1c. According to DLS, these NPs had hydrodynamic diameters of 7-10 nm in toluene, and did not aggregate with either only UV irradiation or CO_2 treatment (bubbling into the solution) (red and blue bars, respectively, in Figure 3). Only when both stimuli were applied at once (4 min of UV irradiation during CO_2 bubbling), however, did the NPs rapidly assemble into aggregates of 800-1000 nm in size, as shown by DLS (green bars in Figure 3). The presence of aggregated NPs was also evident from TEM and spectroscopic analysis (Figure 5b, c). Interestingly, these aggregates were stable in a closed vial placed in the dark for more than 30 minutes.

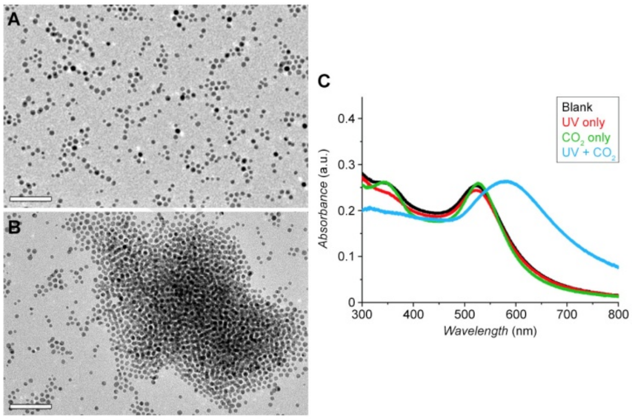


Figure 5. (A) TEM images of AuNPs before aggregation. (B) The same NPs after applying two stimuli (CO_2 and UV) at once. (C) UV spectra of AuNPs with only one stimulus (red and green) and both stimuli at once (cyan). The scale bars in TEM images are 50 nm.

Further demonstration of the AND gate-like behaviour was provided by removing one of the two stimuli. For example, when a solution of aggregated NPs was sonicated, facile disaggregation was observed even under constant UV irradiation (yellow bars in Fig. 4; ca. 10 nm). The resulting NPs did not respond to only CO_2 or UV, but they could again be quickly aggregated by applying UV and CO_2 simultaneously (second-cycle green bars in Figure 4 and the TEM image in Figure 5b). Interestingly, our NPs also assembled as a result of CO_2 bubbling (4 min), followed by exposure to UV light, or via the opposite sequence, whereby UV-irradiated NPs were treated with CO_2 in the dark: these results indicate that both *i*) *cis*-azobenzene and *ii*) the CO_2-diamine complex is relatively stable, *i*) in the dark, and *ii*) unless other stimulants (such as sonication, heating or inert gas bubbling) are applied.

Thus assembled AuNPs readily deaggregated under ambient light (5-6 minutes) or upon exposure to visible light (fluorescent bulb, <30 seconds), and consequently, additional assembly-disassembly cycles could be performed. Overall, we concluded that this study represents the first example of CO_2- and light-responsive nanoparticles, whose behaviour constitutes an AND logic gate. Noteworthy, NPs having higher surface concentrations of **AzoSH** did respond to UV light by assembling into aggregates, which could aggregate *further* upon the delivery of CO_2 (SI, Figure S6C). In other words, only upon pre-exposure to UV light were these specific NPs responsive to CO_2.

Conclusions

We prepared gold nanoparticles co-functionalised with azobenzene- and diamine-based ligands, which rendered the particles responsive to UV irradiation and CO_2, respectively. Exposure to UV and CO_2 each led to a stepwise increase in the nanoparticle surface polarity, triggering aggregation in a non-polar medium (toluene).[42] The composition of the mixed monolayer of ligands could be tuned in order to afford NPs capable of aggregating only when both types of stimuli are applied at once. Both of these stimuli could easily be turned off / removed without the generation of any by-products / chemical waste, after which the assembly-disassembly cycles could be repeated. The aggregates only exist when both light and CO_2 are delivered simultaneously. An important future direction concerning these and other man-made dynamically aggregating materials is to study / engineer their emergent behaviours under non-equilibrium conditions.

Notes and references

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† Electronic Supplementary Information (ESI) available: Synthesis and charaterisation of ligand **3**; synthesis of functionalisation of gold nanoparticles; control experiments. See DOI: 10.1039/b000000x/

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