

Preparation of Gold-Coated Magnetic Nanorods for Lung Cancer Treatment

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Introduction

- **Purpose:** Evaluate experimental methods used for the preparation of gold coated nanorods.
- Gold-coated magnetic nanorods are a possible means for achieving selective destruction of lung cancer cells through plasmon resonance.
- Particles can be delivered to the lung by inhalation and can be detected by the non-invasive imaging technique of magnetic resonance imaging (MRI).
- Theoretically, temperature of the particle can be raised to 500-1000 ° C by laser irradiation at the proper wavelength, which would cause non-specific thermal death of the cell (photothermal therapy)
- The magnetic core will allow not only detection by MRI but also opens the possibility of aligning the particles to maximize the absorption of light, that is, increase the optical cross sectional area.

Objectives

- Use different experimental methods (reaction conditions) to create iron particles
- Coat magnetite particles with gold
- Determine particle size dynamic light scattering and microscopy

Hypotheses

- The rate of addition of base is inversely related to the resulting particle size
- Changing mole ratio of Fe to OH in KNO₃ oxidation reaction modulates the size and shape of particles

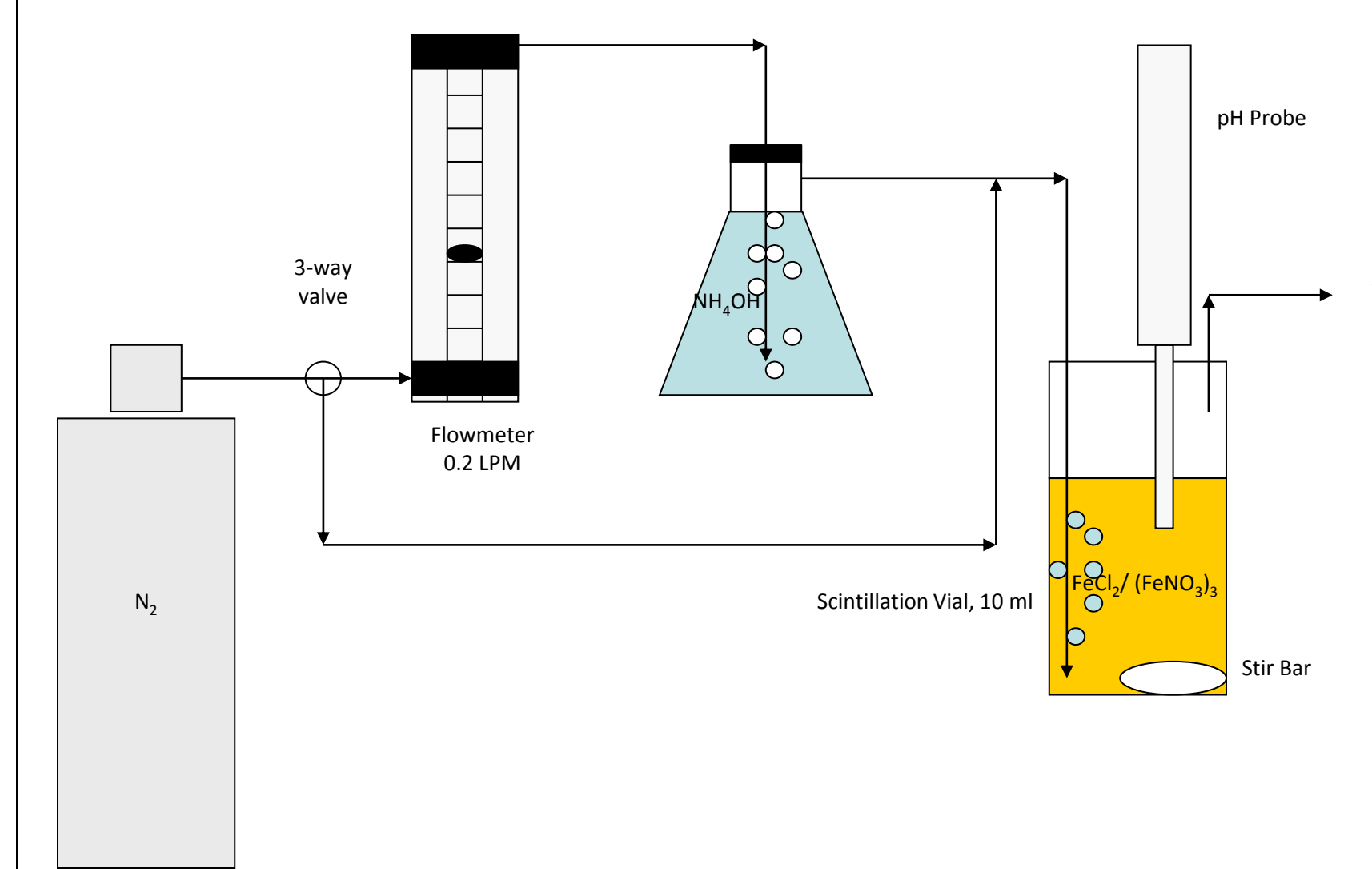


Figure 1: Schematic of reaction set up. Rate of addition of NH₄OH depends on pressure from N₂. More rapid ammonia vapor addition resulted in a more rapid rise in pH and extent of reaction.

pH	1-2	1.5-2.5	1.8-7	3.4-7.1
Color	Yellow/Orange	Reddish brown	Brown	Black

Table 1: Color of reaction mixture at measured values of pH. Target pH level is around 12. The brown color occurred at a pH level of 3.8 and by pH 6, the dispersion was black.

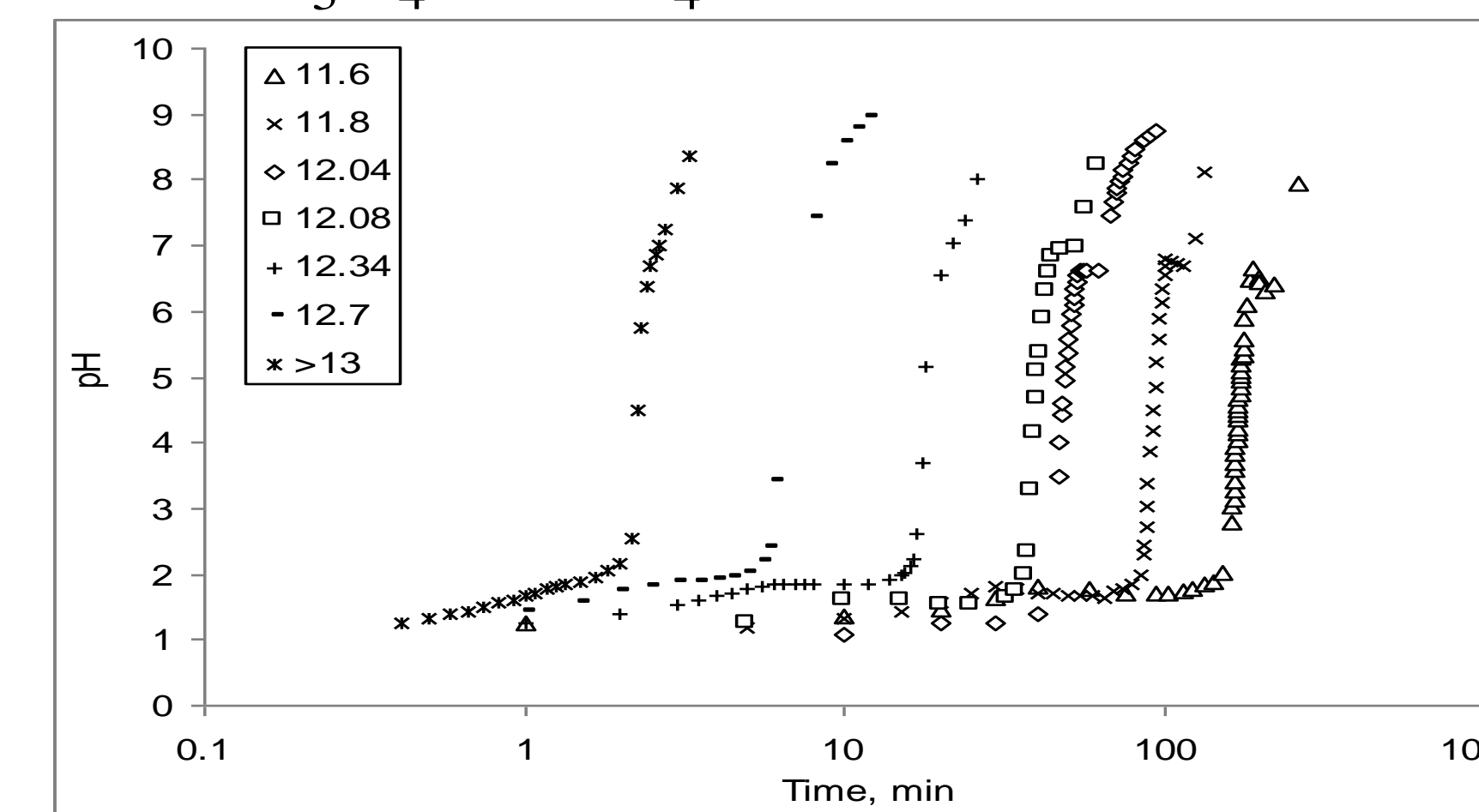
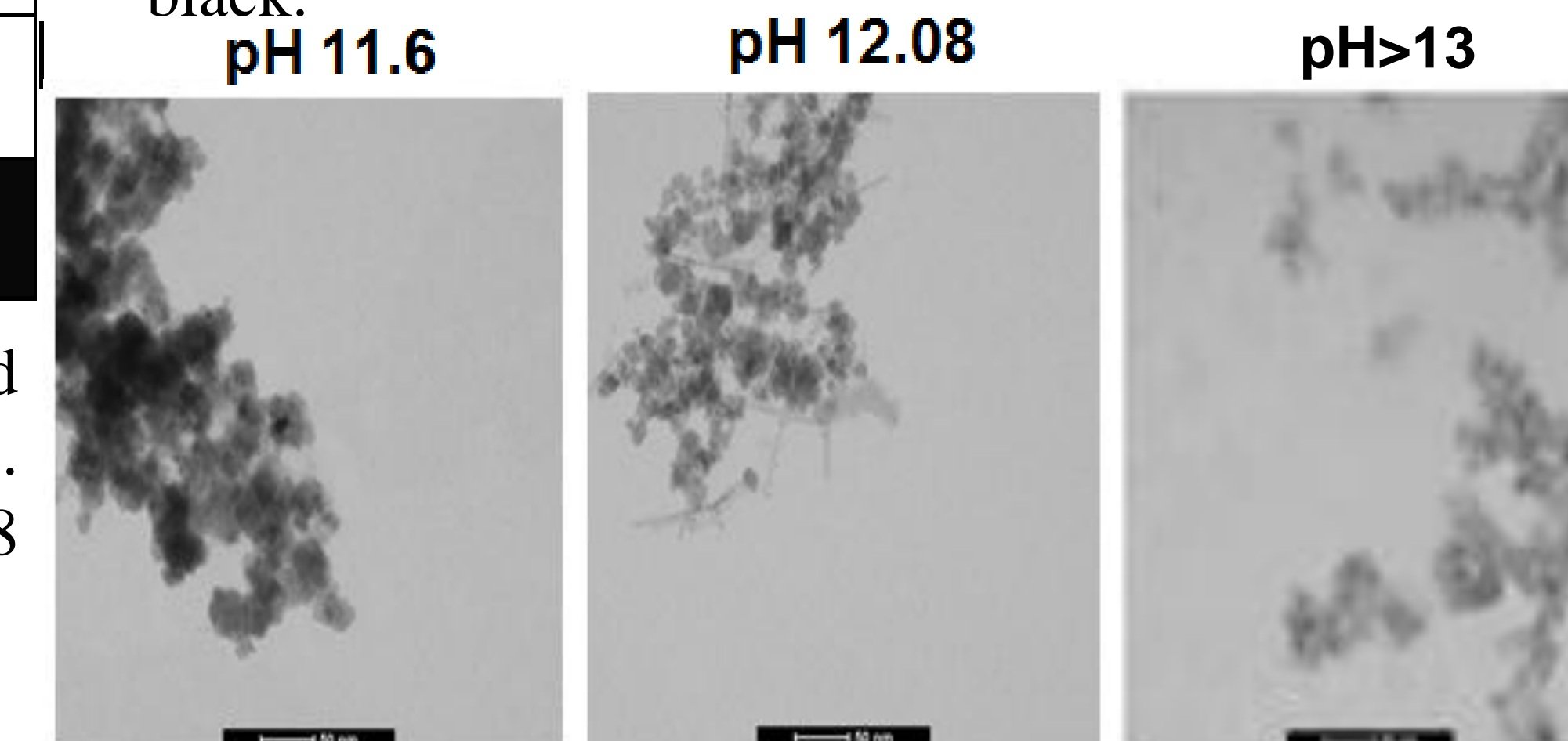


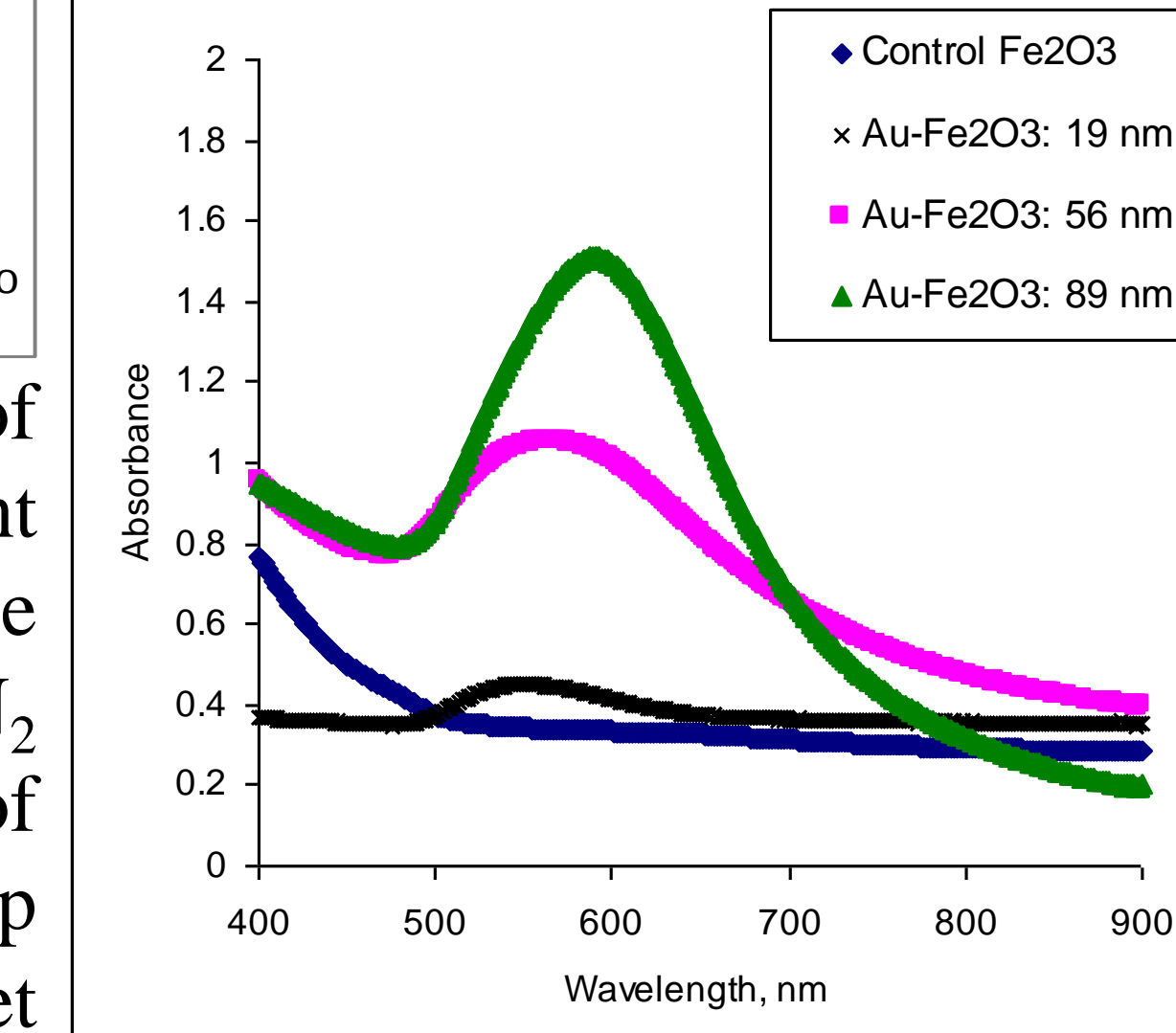
Figure 2: Observed pH plotted as a function of time on a logarithmic scale for seven different reactions. The legend indicates the pH of the ammonium hydroxide solution through which N₂ was passed to control the rate of addition of ammonia vapor to the reaction mixture. The steep increase in pH occurs when the particles turn jet black.



Figures 3: TEM was performed on particles at pH levels ranging from 11.6-13. As the reaction rate increased, a corresponding reduction in size of the particles was observed.

Plasmon Resonance:

- (1) $\text{Fe}_3\text{O}_4 + \text{HNO}_3 \rightarrow \Delta \rightarrow \gamma\text{-Fe}_2\text{O}_3$
- (2) $\gamma\text{-Fe}_2\text{O}_3 + \text{Na}_3\text{Citrate} \rightarrow \gamma\text{-Fe}_2\text{O}_3\text{-Citrate}$
- (3) $\gamma\text{-Fe}_2\text{O}_3\text{-Citrate} + \text{HAuCl}_4 + \text{NH}_2\text{OH} \rightarrow \gamma\text{-Fe}_2\text{O}_3\text{-Au coated}$
- (4) $\gamma\text{-Fe}_2\text{O}_3\text{-Au coated} + \text{HAuCl}_4 + \text{NH}_2\text{OH} \rightarrow \gamma\text{-Fe}_2\text{O}_3\text{-Au increased thickness of Au coat}$



Legend 1: The absorbance increased and underwent a red shift as more gold was added to the particles (■) and (▲). There was also a corresponding increase in the particle size as noted in the figure legend.

Figure 4: Absorbance as a function of wavelength of the control and different levels of coating of the gold (Au) on oxidized magnetite particles with the corresponding size as determined by dynamic light scattering. The control scan consisted of the background scattering of the iron particles (◆ γ-Fe₂O₃, maghemite). The addition of gold chloride (HAuCl₄) and the mild reducing agent, hydroxylamine (NH₂OH), resulted in the formation of the Plasmon resonance band at 550 nm (x).

Summary: A series of reactions were carried out in which the rate of addition of ammonium hydroxide was varied. A series of colloidal dispersions were obtained that contained magnetite. The particle size distribution appeared to vary in that a larger particle size was obtained as the rate of addition of ammonium hydroxide was reduced.

Conclusions

- **Iron Chloride:** The reaction time and thereby particle size was dependent on the pH level of the NH₄OH through which the N₂ was bubbled through.
- **Plasmon Resonance:** Citrate must be added and left to react with the particles for ~10 minutes to allow Au bind to iron
- **Precipitation with base and oxidation with KNO₃:** when excess Fe present (Fe>OH), particles were spherical in shape, and small; when negative excess Fe (Fe<OH), particles were hexagonal and cubical in shape; when Fe≈OH, mixture of larger spherical and hexagonal particles
- When KNO₃ reaction carried out in presence of magnet and Fe>OH, particles grew like beaded nanofibers

References

- Larson, A. Timothy, Bankson James, Jesse Aaron, and Konstantin Sokolov. "Hybrid plasmonic magnetic nanoparticles as molecular specific agents for MRI/optical imaging and photothermal therapy of cancer cells." Nanotechnology 18 (2007).
- Brown, R. Kenneth, Daniel G. Walter, and Michael J. Natan. "Seeding of Colloidal Au Nanoparticle Solutions. 2. Improved Control of Particle Size and Shape." Chem Mater 12 (2000): 306-313.

Acknowledgements

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Precipitation with base and oxidation with KNO₃ (Light Microscopy)

- (1) $\text{Fe}^{\text{II}}\text{SO}_4 \cdot 7\text{H}_2\text{O} + \text{KOH} \rightarrow \text{FeOOH}$
- (2) $\text{FeOOH} + \text{KNO}_3 \rightarrow \text{Fe}_3\text{O}_4$ (ox.)

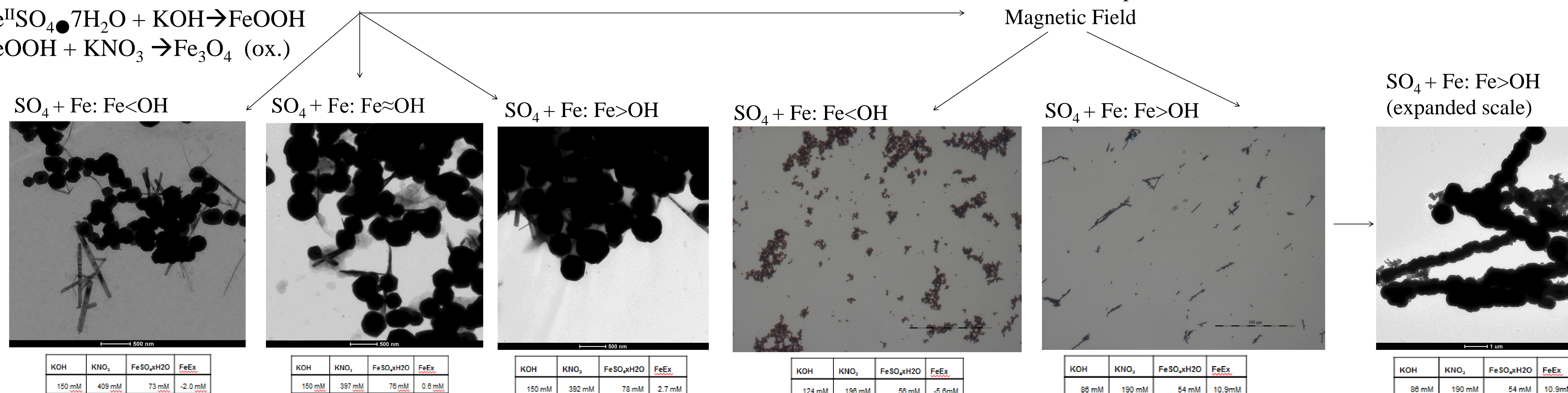


Figure 5: Because there was no expected dependency on the absolute concentration of the Fe^{II}SO₄, a range of 30 to 70mM of Fe^{II}SO₄ was mixed with KOH in first step. In the chemical reaction, 1 mole of Fe reacts with 2 moles of KOH. The outcome of the reaction depends on the ratio of Fe and OH, which was expressed as "excess Fe". When there is a 1:2 Fe/OH ratio, the reaction has zero excess. If there is excess Fe, the particles were on the order of 400nm and spherical; if the ratio is ca 1:2 (no excess), the particles were spherical/hexagonal on the order of 1-2µm; if there is negative excess of Fe (excess OH), small hexagonal particles resulted. If this reaction is carried out in the presence of a permanent magnet, the particles aligned to form very large, beaded nanofibers.

