pH-based Cancer Detection by Graphene Quantum Dots

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Early and Accurate Detection

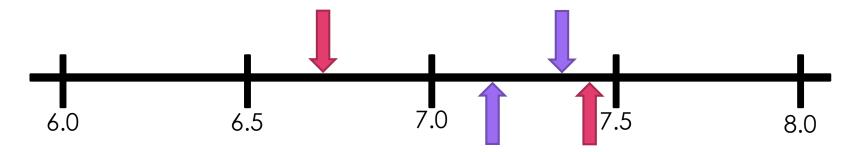
IN 2020:

- OVER 1.8 MILLION
 CANCER DIAGNOSES
 IN THE US
- MORE THAN 600,000 DEATHS

---Early and accurate detection significantly increases survival rates---

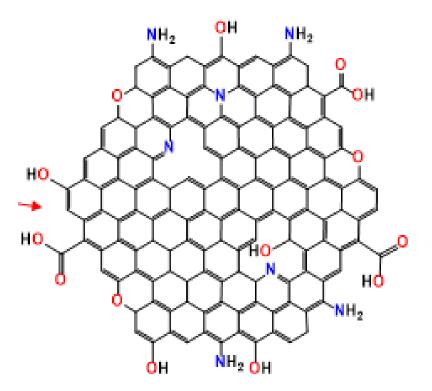
The Acidic Environment

| | Healthy Cells | Cancer Cells |
|---------------------------|---------------|--------------|
| Intracellular Environment | pH 7.2 | pH 7.3-7.6 |
| Extracellular Environment | pH 7.35-7.45 | pH 6.4-7.0 |



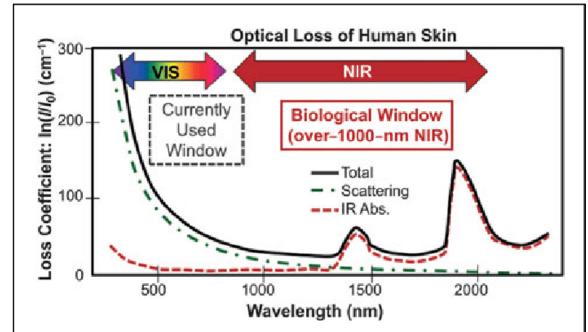
What is a Quantum Dot?

- Man made crystals measuring a few nanometers in size
- Unique optical and electronic properties differing from larger particles
- ► Uses in:
 - ► Transistors
 - Solar Cells
 - Quantum Computing
 - Medical Imaging



Fluorescence Imaging

- Fluorescence can serve as a detection mechanism for nanoparticles in:
 - Drug delivery
 - Cancer sensing
- Fluorescence is advantageous for cancer sensing: allows for non-invasive imagingbased diagnostics
- Near-Infrared fluorescence is advantageous for biological sensing: near-infrared light penetrates several cm of biological tissue: <u>in vivo</u> detection.



Why?

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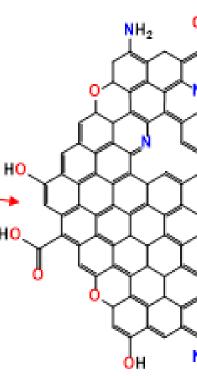
2 ÓH.

Goal

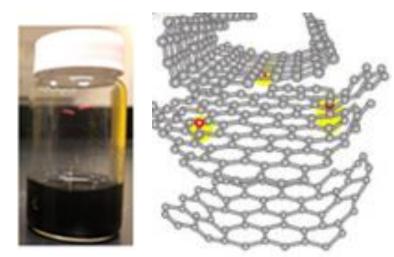
To analyze the fluorescence change with pH in the visible and infrared range of various forms of Graphene Quantum Dots (GQDs) to determine their use in cancer imaging and drug delivery.

Impact

If a relationship can be found between the spectra and pH within the biological range of pH6.00-8.00, these particles be utilized as a highly accurate cancer detection method, in addition to their many other applications for biological imaging



Reduced Graphene Oxide Quantum Dots (RGQDs)





Sodium Hypochlorite



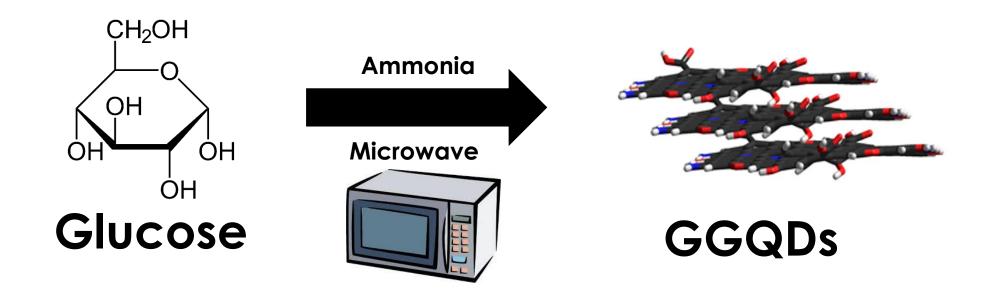


Reduced Graphene Oxide (RGO)

RGQDs

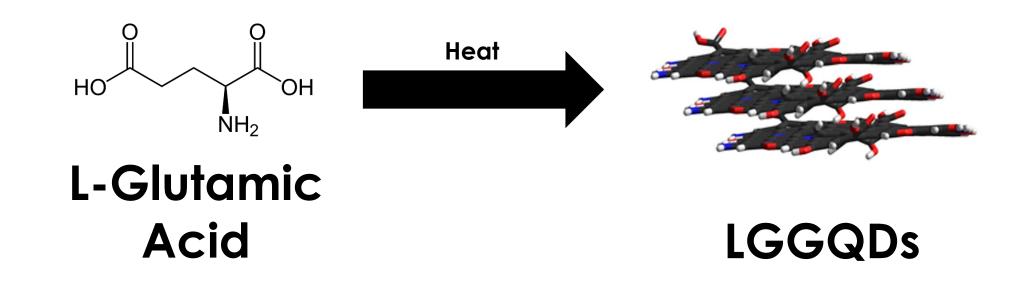
Synthesized by Veronica Lyle

Glucose Graphene Quantum Dots (GGQDs)



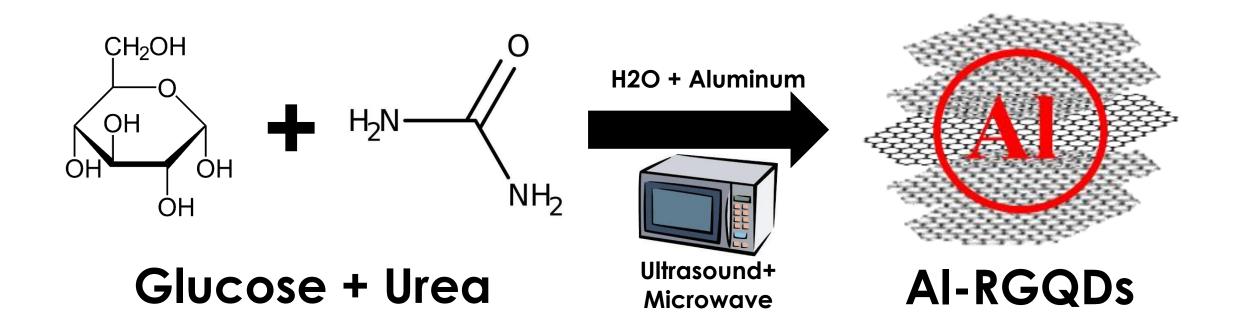
Synthesized by Matt Dadet

L-Glutamic Acid Graphene Quantum Dots (LGGQDs)



Synthesized by Matt Dadet

Aluminum Doped Reduced Graphene Oxide Quantum Dots (Al-RGQDs)



Synthesized by Dr. Roberto Gonzalez Rodriguez

Why These 4?

THEY ALL FLUORESCE IN THE INFRARED RANGE

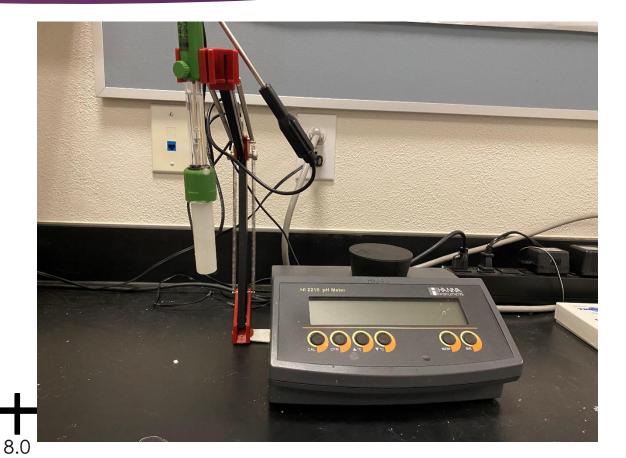
pH Adjustment

- Using the Hanna HI2210 pH Meter, detected initial pH
- Added .075M NaOH or HCl in 5µL increments to adjust pH ∓ 0.25 for each subsequent measurement

7.0

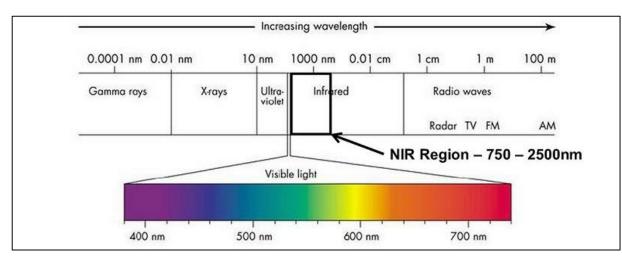
6.5

7.5



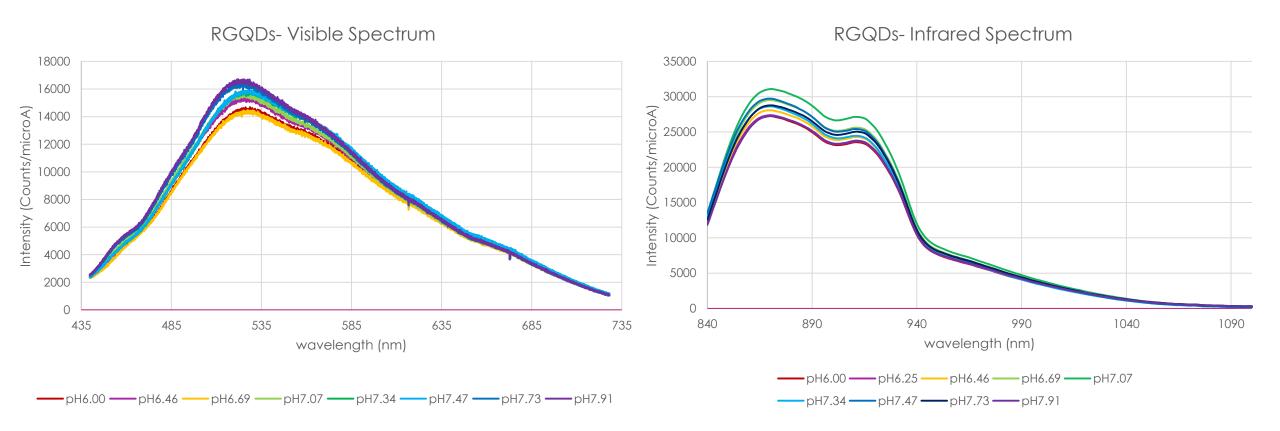
Imaging

- To collect visible spectra, utilized the Horiba spectral array collecting from 440-700nm
- To collect infrared spectra, utilized a combination of the symphony II spectral array, an 808nm laser, and a 400nm laser collecting from 800-1100nm





Reduced Graphene Oxide Quantum Dots (RGQDs)

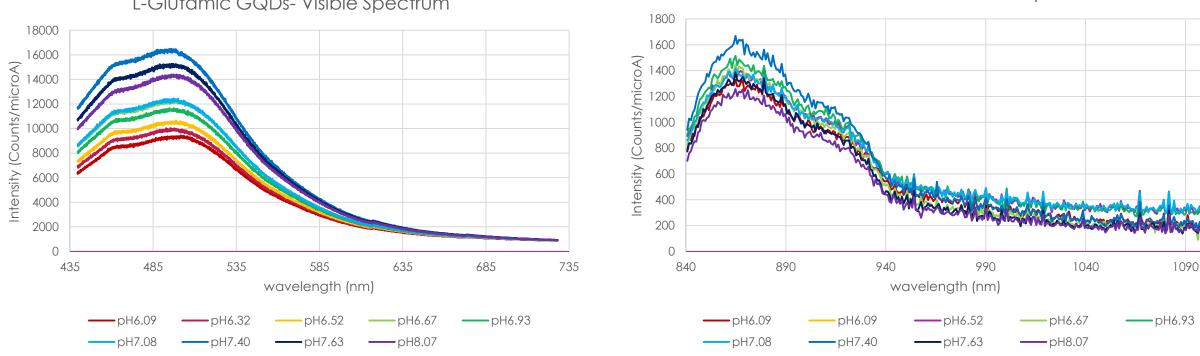


Glucose Graphene Oxide Quantum Dots (GGQDs)

Glucose GQDs- Visible Spectrum

Glucose GQDs-Infrared Spectrum Intensity (Counts/microA) Intensity (Counts/microA) -200 840 wavelength (nm) wavelength (nm) -pH6.28 ---- pH6.44 ---- pH6.75 ---- pH6.95 - pH6.28 ---- pH6.44 ---- pH6.75 ---- pH6.95 pH6.04 pH6.04 pH7.17 — pH7.40 — pH7.80 — pH7.97 pH7.17 — pH7.40 — pH7.80 — pH7.97

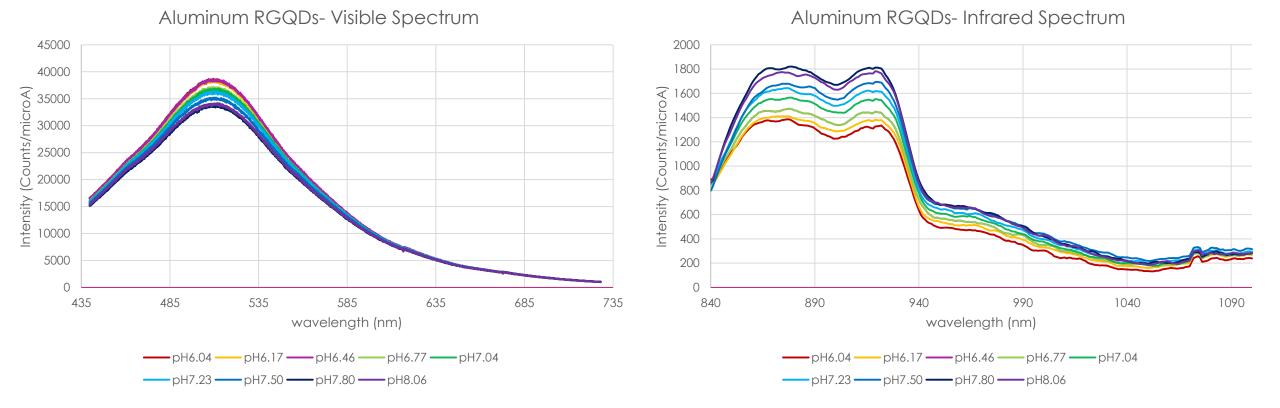
L-Glutamic Acid Graphene Oxide Quantum Dots (LGGQDs)



L-Glutamic Acid GQDs- Infrared Spectrum

L-Glutamic GQDs- Visible Spectrum

Aluminum Doped Reduced Graphene Oxide Quantum Dots (Al-RGQDs)

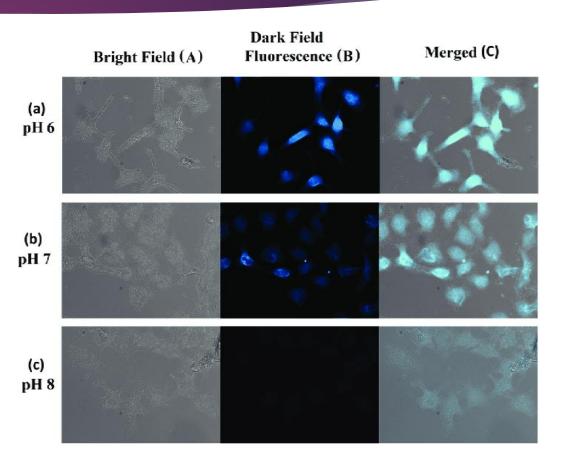


Conclusions

| No Detectable | Visible | Infrared and Visible |
|----------------|----------------|----------------------|
| pH Sensitivity | pH Sensitivity | pH Sensitivity |
| RGQDs | GGQDs, LGGQDs | AI-RGQDs |

Next Steps

- Cytotoxicity studies for GGQDs and LGGQDs
- Live Cell Imaging Studies (example right)
 - ▶ Time Lapsed
 - Cancerous and Noncancerous Cell Lines
 - Collect data regarding:
 - ► Internalization
 - Excretion
 - ► Fluorescence



References

- "Cancer Facts & Figures 2020." American Cancer Society, www.cancer.org/research/cancer-facts-statistics/all-cancer-factsfigures/cancer-facts-figures-2020.html#:~:text=Estimated%20numbers%20of%20new%20cancer,deaths%20in %20the%20United%20States.).
- Hasan, Md Tanvir, et al. "Near-Infrared Emitting Graphene Quantum Dots Synthesized from Reduced Graphene Oxide for in Vitro/in Vivo/Ex Vivo Bioimaging Applications." 2D Materials, 2021, doi:10.1088/2053-1583/abe4e3.
- Piasentin, Nicola, et al. "The Control of Acidity in Tumor Cells: a Biophysical Model." 2020, doi:10.1101/2020.03.22.002113.
- Trafton, Anne. "How Tumors Behave on Acid." MIT News | Massachusetts Institute of Technology, news.mit.edu/2019/how-tumors-behave-acid-0320#:~:text=Scientists%20usually%20attribute%20a%20tumor's,have%20an%20a dequate%20blood%20supply.

