

Preparation and Characterization of First-Generation Glucose Sensors

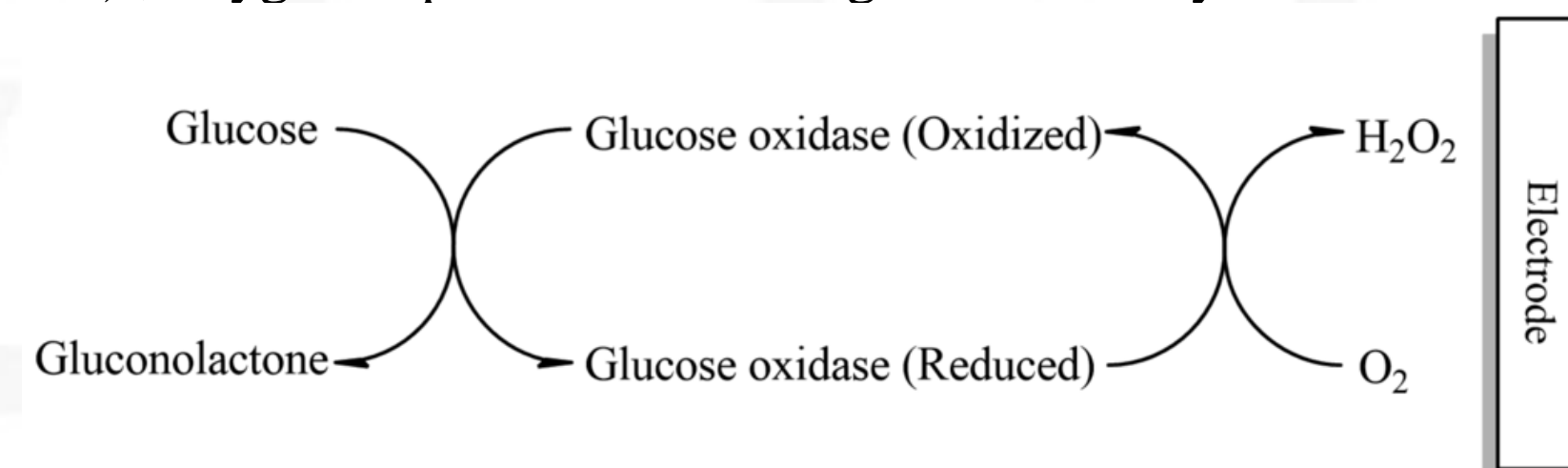
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Introduction¹

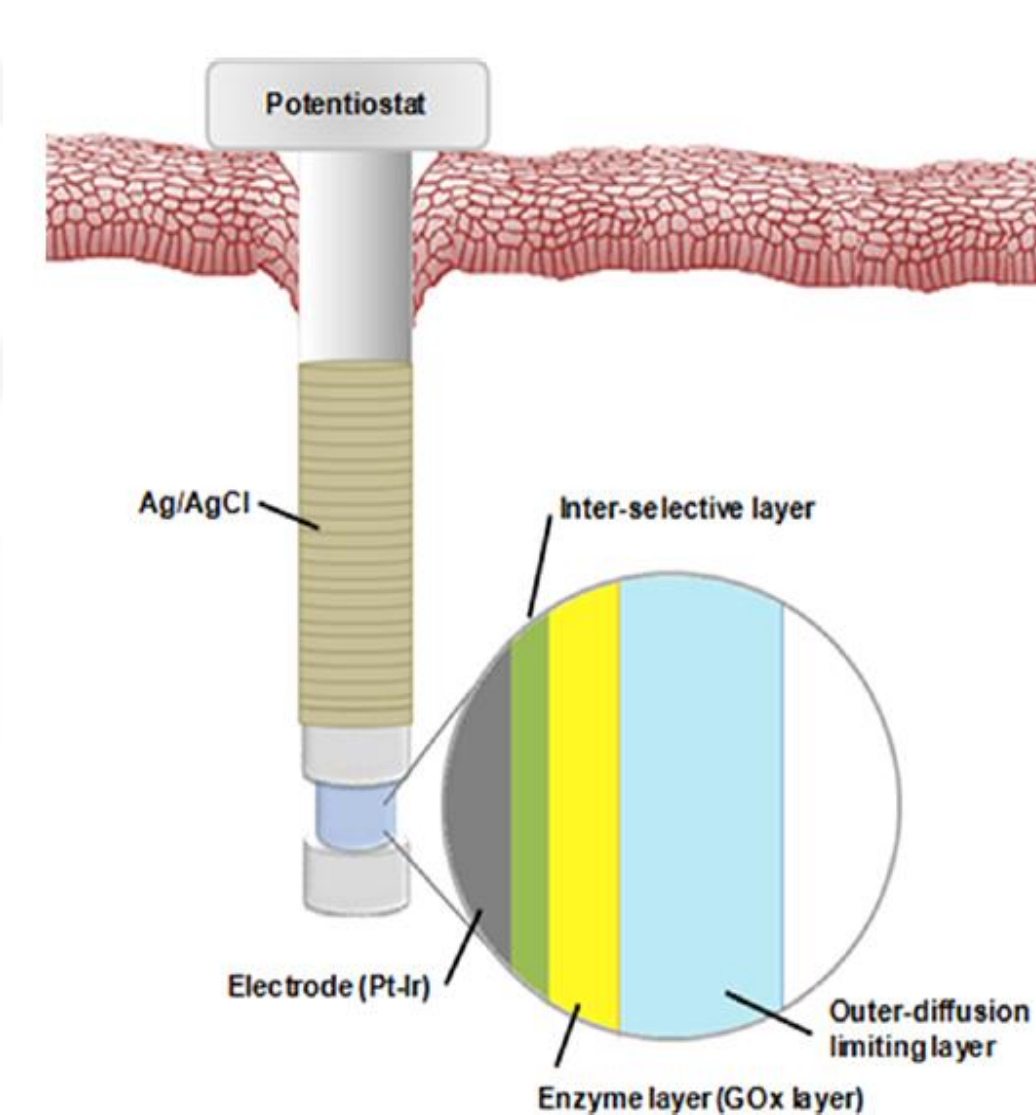
Amperometric enzyme electrodes, based on glucose oxidase (GOx), have played a leading role in the move to simple easy-to-use blood sugar testing and are expected to play a similar role in the move toward continuous glucose monitoring for a worldwide public health problem--diabetes mellitus.

First-generation glucose biosensors rely on the use of the natural oxygen cosubstrate and generation and detection of hydrogen peroxide. The major focuses are glucose linearity, electroactive interferences (like AA: Ascorbic Acid and APAP: Acetaminophen), oxygen dependence and long-term stability.



Keyword: Diabetes, Biosensors, Amperometry

Design²



- Inter-selective layer
 1. PAA: Polyacrylic Acid
 2. CAB: Cellulose Acetate Butyrate
- Enzyme layer
 - i. GOx: Glucose Oxidase Enzyme
 - ii. BSA: Bovine Serum Albumin
 - iii. GA : Glutaraldehyde
 - iv. PVP: Polyvinylpyrrolidone
- Outer limiting layer
 - i. Carbosil: Polyurethane and Silicone
 - ii. PVP: Polyvinylpyrrolidone

Test with +0.6V vs. Ag/AgCl, in 0.1M PBS (Phosphate Buffered Saline)

Results

a) PAA—Hydrophilic—Stable Signal

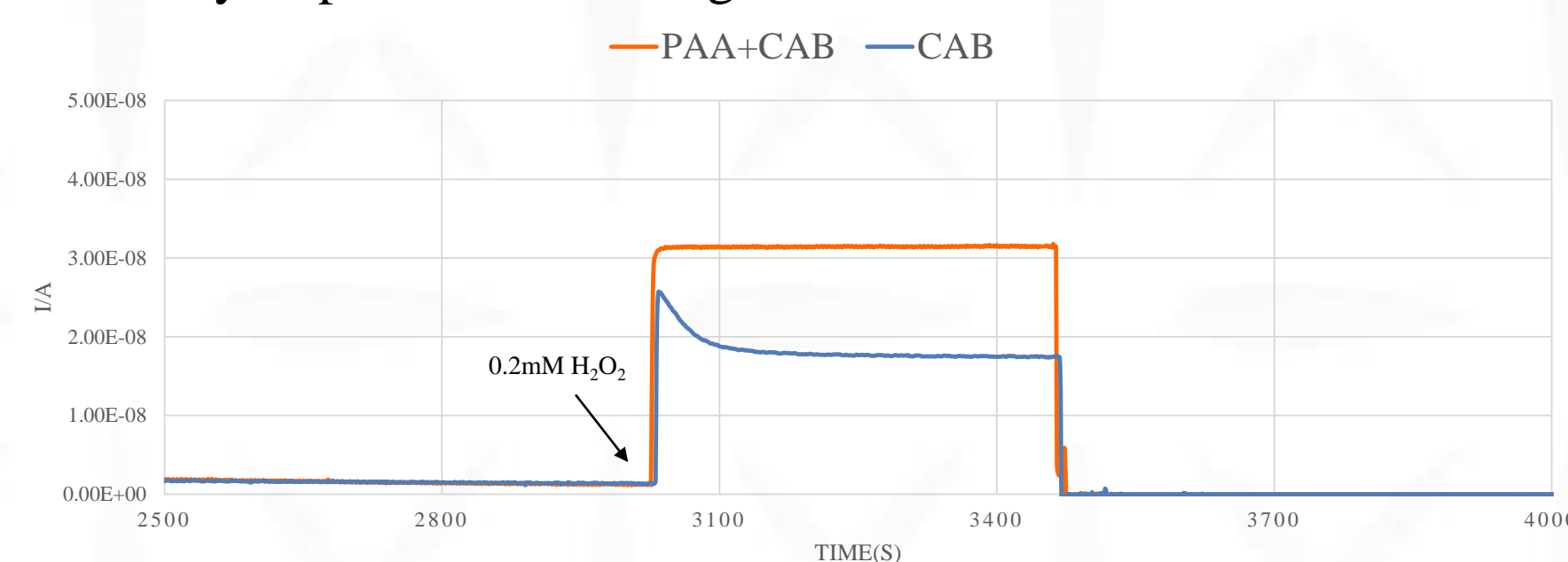


Figure 1. Detect H₂O₂ response from sensors with or without PAA

Conclusions

- PAA could get rid of the sensor signal drift phenomenon effectively
- PAA makes the electrical current more stable and increases sensor sensitivity

b) CAB plus Nafion—Crosslink—Better Selectivity

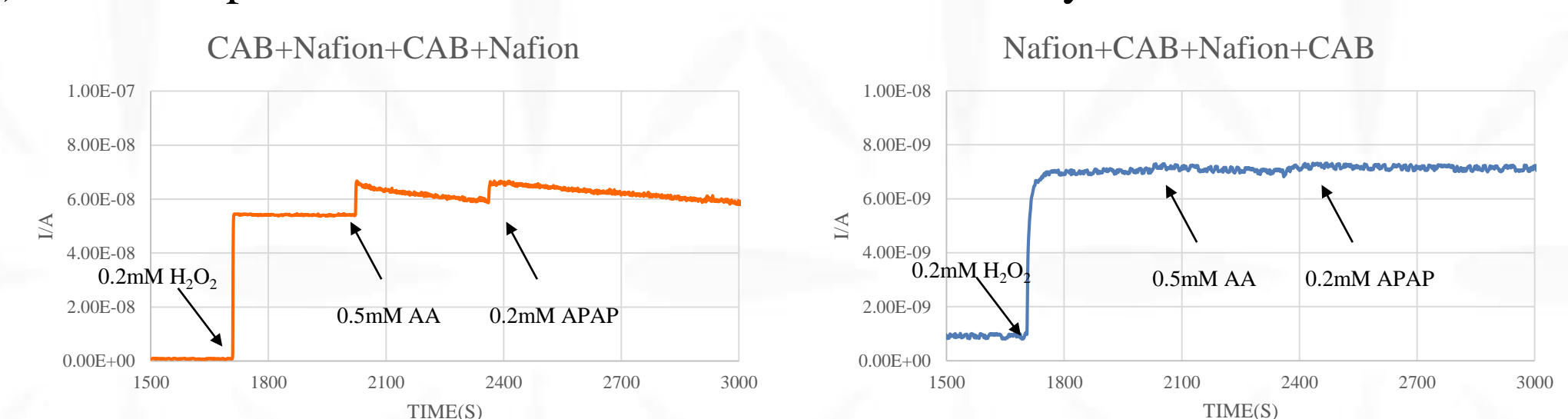


Figure 2. Detect interference response from sensors by coating selective layers in different sequence. Nafion is a sulfonated tetrafluoroethylene based fluoropolymer, which is commonly used polymer to diminish AA interference.

Conclusions

- The combining polymers of CAB and Nafion could improve the sensor selectivity
- The coating sequence of CAB and Nafion contributes to different anti-interference

c) PVP—High O₂ Solubility—Influence O₂ Dependence

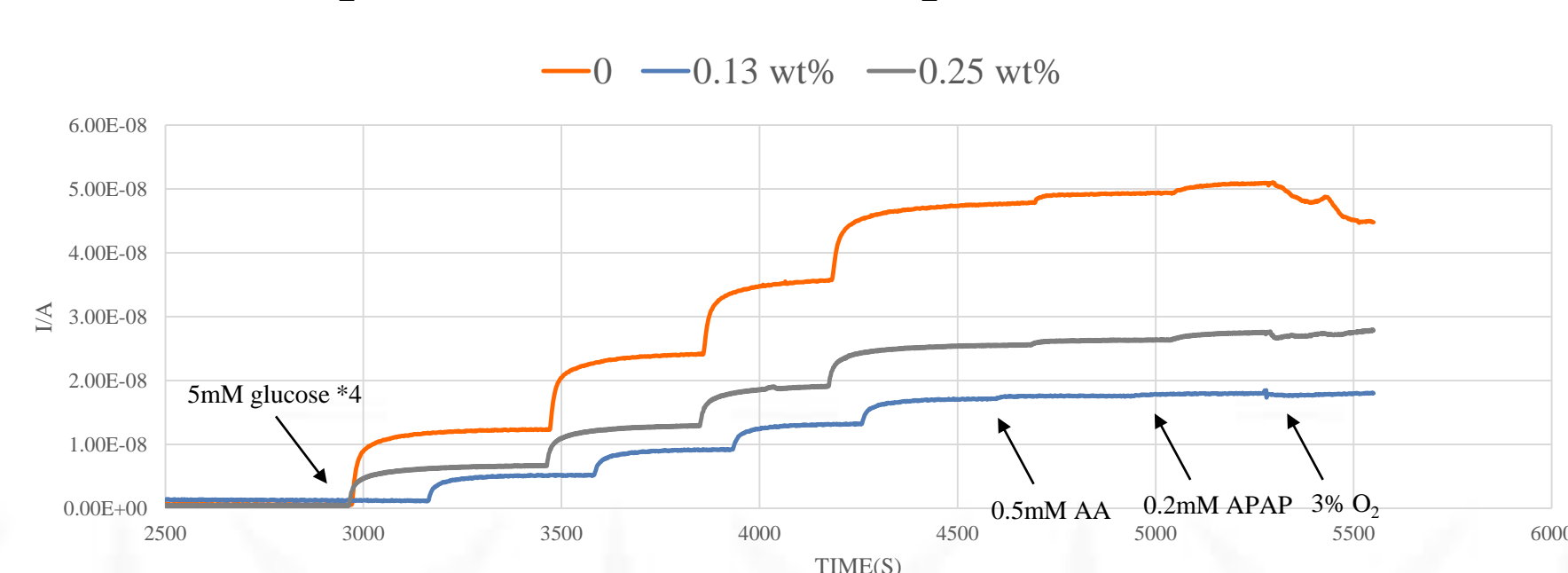


Figure 3. Compare O₂ dependence extent from sensors with different concentration of PVP (in enzyme layer)

Conclusion

- The PVP in the enzyme layer influences the sensor O₂ dependence, low concentration of PVP results to O₂ decreasing dependence while high concentration of PVP results to O₂ increasing dependence

d) Carbosil-PVP—Diffusion Limit and Unexpected Selectivity

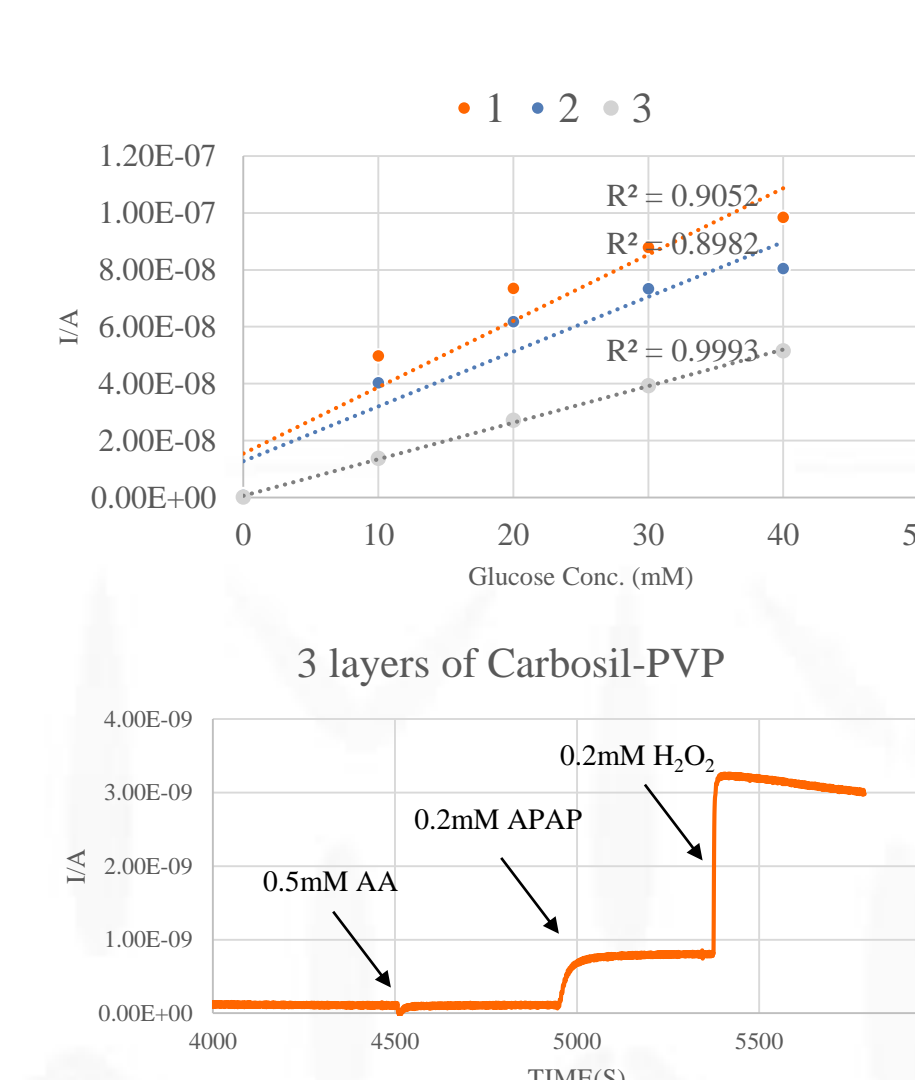


Figure 4. The glucose linearity of regular sensors with 1/2/3 layers of Carbosil-PVP

Conclusion

- Certain numbers of Carbosil-PVP layers could improve the glucose linearity

Figure 5. Detect interference response from sensors with only 3 layers of Carbosil-PVP

Conclusion

- Carbosil-PVP layers have special ability of preventing AA

Performance

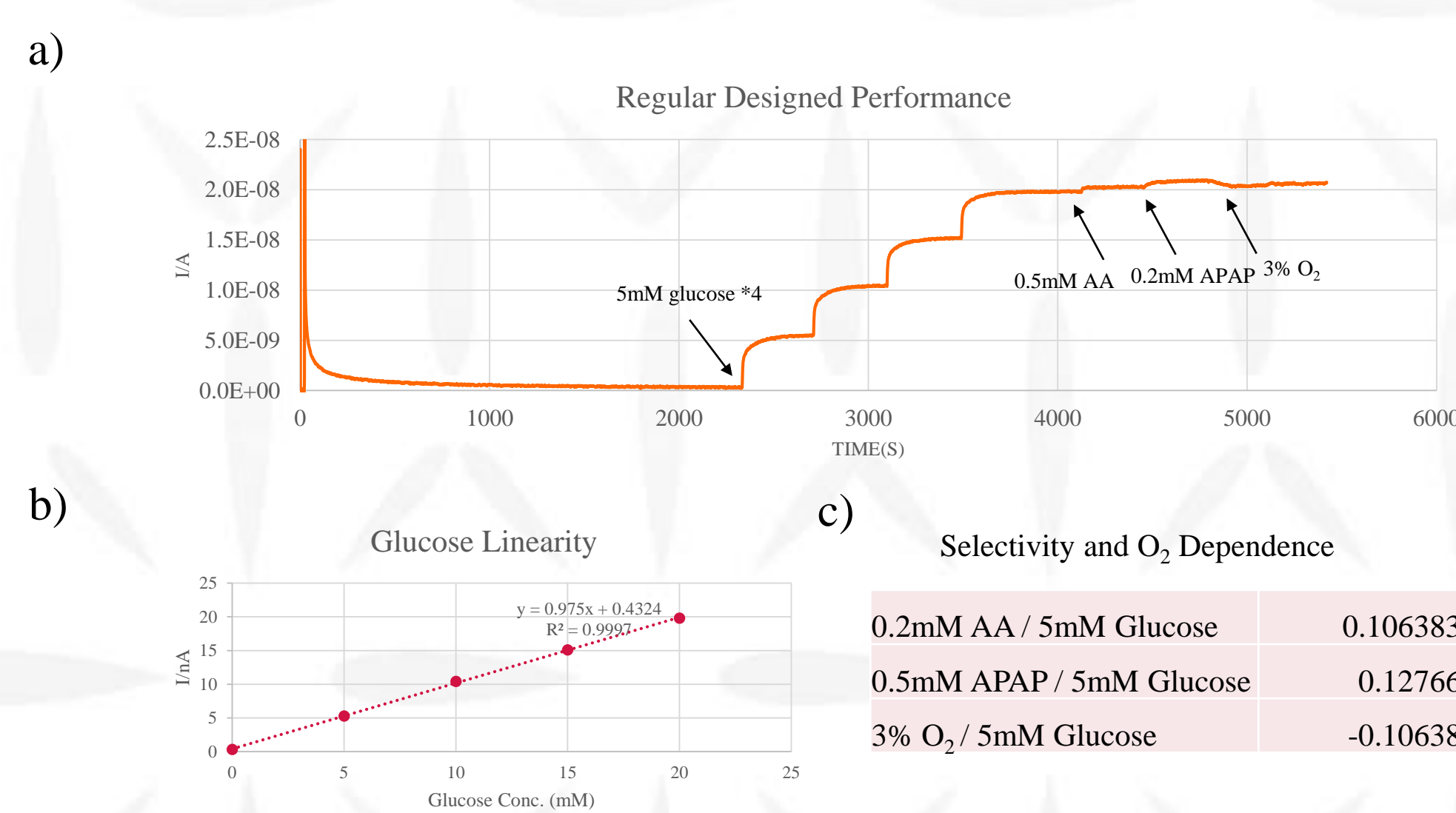


Figure 6. The performance of regular well-designed sensors

Others

Future works

- Optimize the fabrication and coating method
- Characterize the polymers by Atomic Force Microscopy (AFM)
- More sensors for blood and *in vivo* test
- Combine with NO release and insulin cannula

References

- [1] Wang, Joseph. "Electrochemical Glucose Biosensors." *Chemical Reviews* 108.2 (2008): 814-825.
- [2] Nichols, Scott P., et al. "Biocompatible Materials for Continuous Glucose Monitoring devices." *Chemical Reviews* 113.4 (2013): 2528-2549.

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