



Intraperitoneal glucose sensing – Rapid and accurate

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MOTIVATION

Why intraperitoneal sensing?

- Subcutaneous glucose sensors:
 Slow response, poor robustness towards local tissue effects (mechanical pressure, temperature etc).
- Intravascular glucose sensors:
 Not practically possible outside of the hospital/clinic.
- We need a rapid, accurate and robust glucose measurement for making a safe artificial pancreas.
- Intraperitoneal glucose sensors may react faster than subcutaneous sensors [1,2], while being more practically usable than intravascular sensors.

METHODS

Animal trials

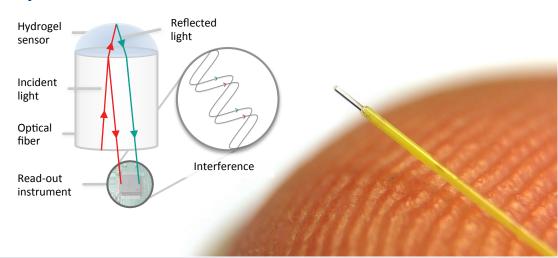
- 3 anaesthetized non-diabetic pigs
- Intravenous infusion of glucose boluses
- Glucose level excursions within the range
 5–22 mmol/L

Sensor types used in trials

- Intraarterial (IA): Optical interferometric phenylboronic acid based sensors, placed in the femoral artery [3,4].
- Intraperitoneal (IP): Using the same sensor as in the IA case. Accessed from below the umbilicus through a common port, directed to 4 different positions (Fig. on the right). 2–4 sensors per animal.
- **Subcutaneous (SC):** Off-the-shelf amperometric enzyme-based (glucose oxidase) sensors. Placed on the belly, above the umbilicus.
- Venous blood was sampled and analyzed on a blood gas analyzer (BGA) for reference and calibration of the other sensors.

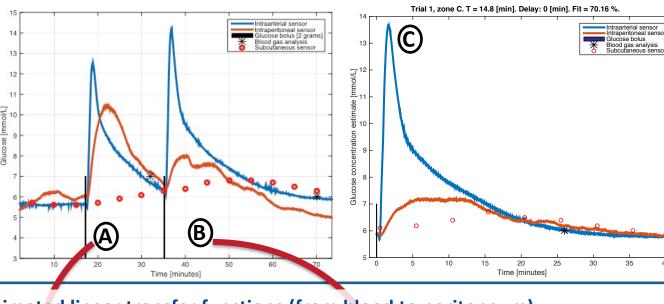
Glucose boluses & BGA IP_{UL} 10 cm 10 cm IP_{LL} sensor IP_{UR} 10 cm 10 cm IP_{LR}

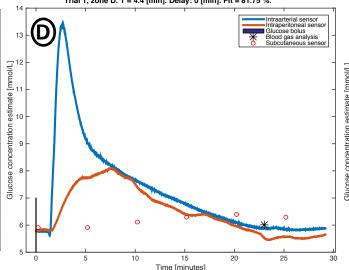
Optical interferometric sensors

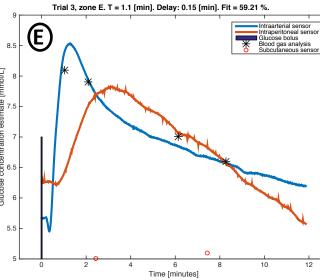


RESULTS

Examples from 6 segments:

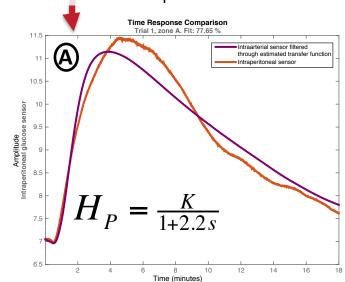


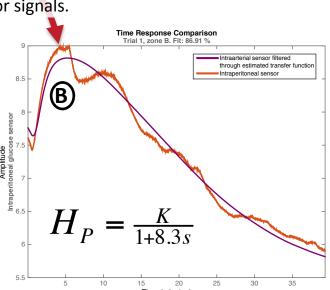


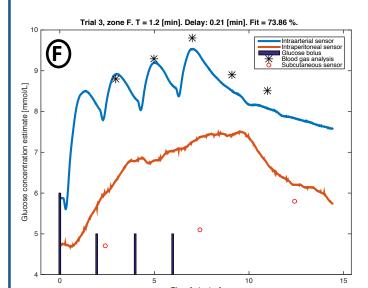


Estimated linear transfer functions (from blood to peritoneum)

Based on relationship between IA and IP sensor signals.







Summary

- 3 trials, 4–7 segments from each trial.
- Time delays: 0–15 seconds (median: 9.0 s, mean: 6.74 s, stdev: 6.76 s).
- Time constants: 1–15 min (median: 4.20 min, mean: 5.03 min, stdev: 4.05 min).

CONCLUSION

- Intraperitoneal glucose sensors can have a substantially faster and more distinctive response than subcutaneous sensors.
- Our results indicate that IP sensors may react even faster than previously shown [2].
- Variable dynamics during the experiments + some sensors had to be relocated -> Need to investigate disturbances & sensor location in more detail.

REFERENCES

- [1] Velho, G., Froguel, P., and Reach, G., "Determination of peritoneal glucose kinetics in rats: implications for the peritoneal implantation of closed-loop insulin delivery systems," *Diabetologia*, vol. 32, no. 6, pp. 331–336, 1989.
- [2] Burnett, D. R., Huyett, L. M., Zisser, H. C., Doyle III, F. J., and Mensh, B. D., "Glucose Sensing in the Peritoneal Space Offers Faster Kinetics than Sensing in the Subcutaneous Space," *Diabetes*, vol. 63, no. 7, pp. 2498–2505, July 2014.
- [3] Skjærvold, N. K., Solligård, E., Hjelme, D. R., and Aadahl, P., "Continuous measurement of blood glucose: validation of a new intravascular sensor," *Anesthesiology*, vol. 114, no. 1, 120–125, 2011.
- [4] Skjærvold, N. K., Østling, D., Hjelme, D. R., Spigset, O., Lyng, O., and Aadahl, P., "Blood Glucose Control Using a Novel Continuous Blood Glucose Monitor and Repetitive Intravenous Insulin Boluses: Exploiting Natural Insulin Pulsatility as a Principle for a Future Artificial Pancreas," *International Journal of Endocrinology*, vol. 2013, Article ID 245152, 2013.