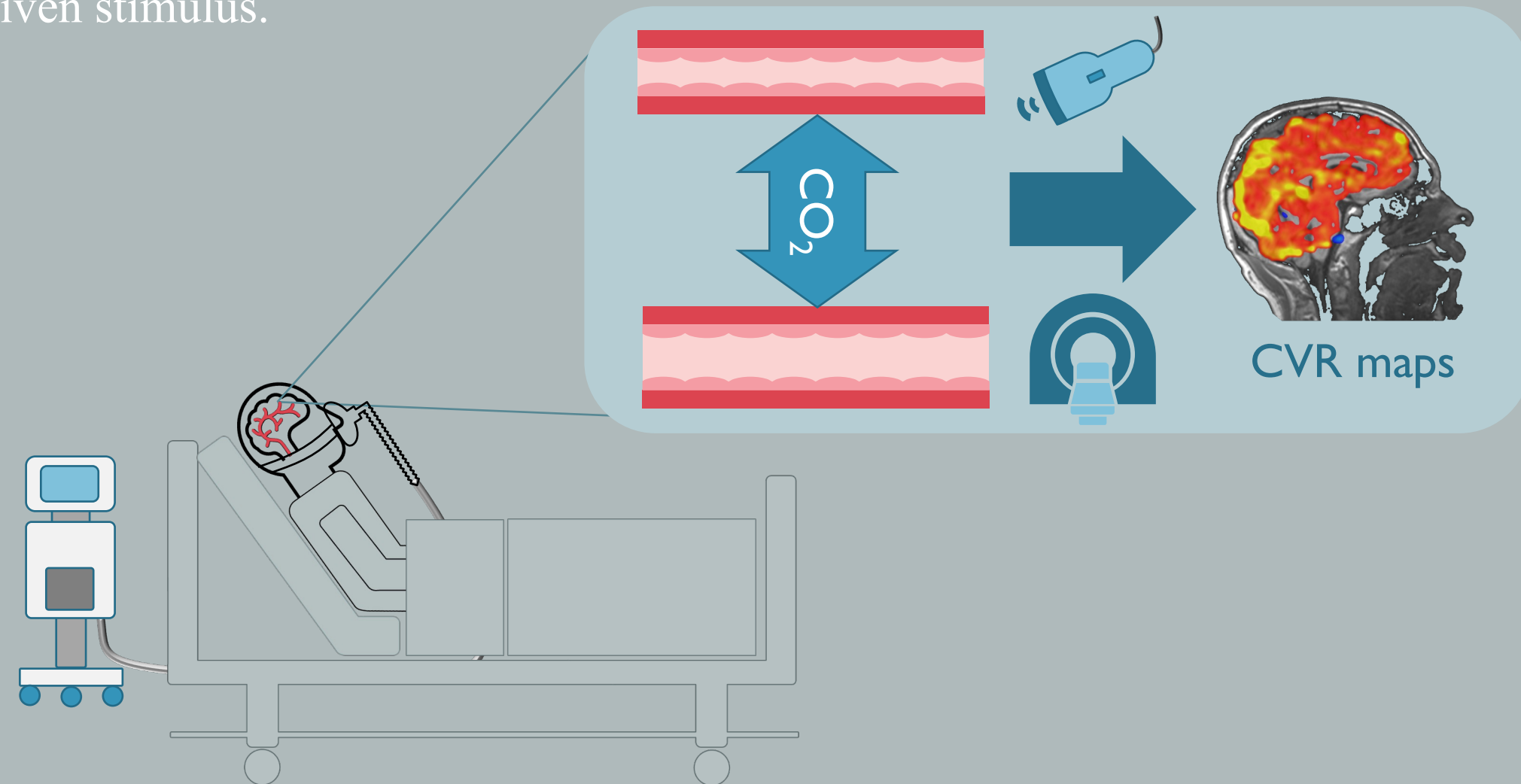


Hypercapnic gas stimulus in ventilators:

First step towards CVR imaging in ventilated patients

1 Cerebrovascular Reactivity

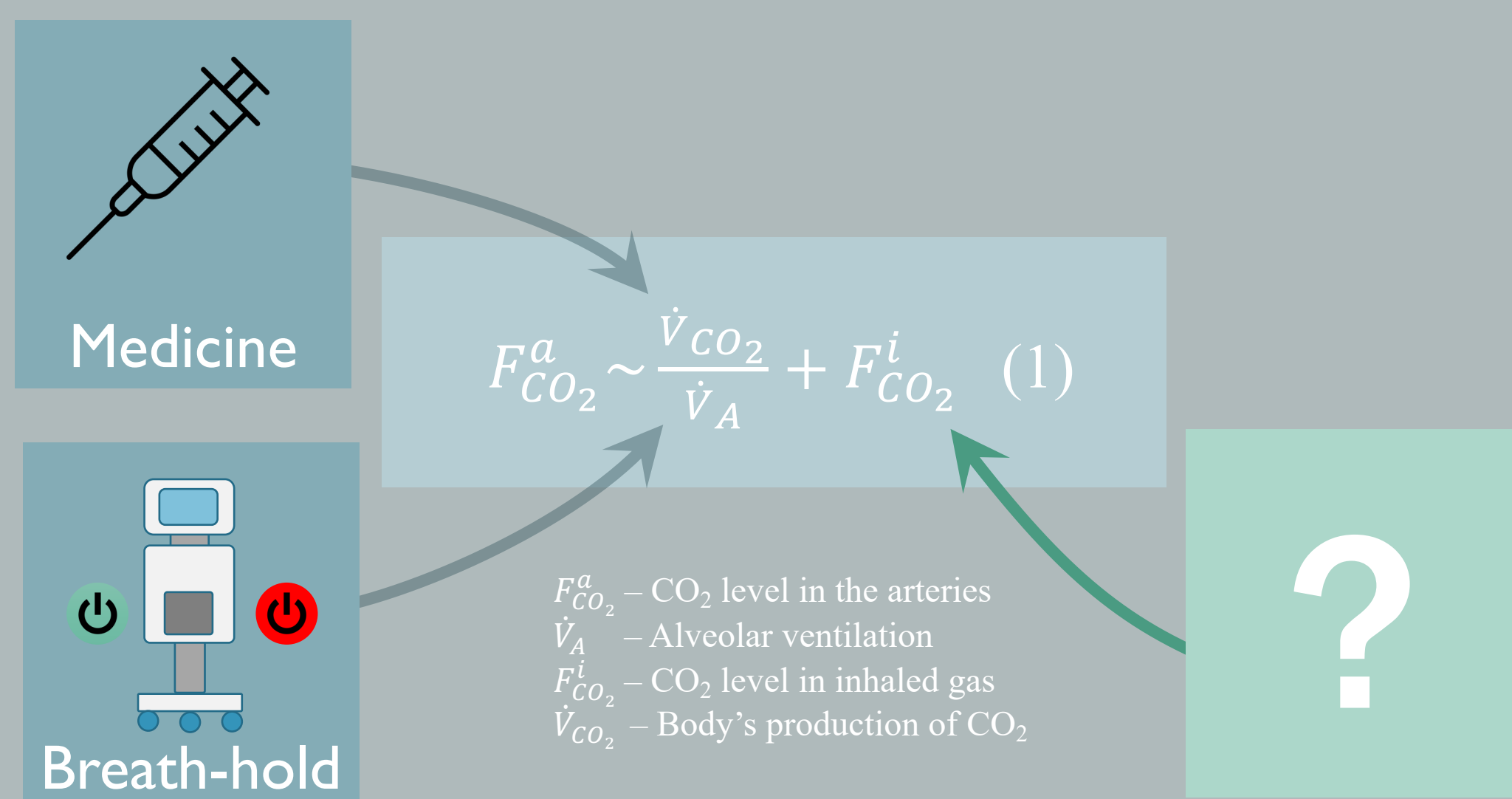
The brain has an advanced system for regulating its blood flow, termed cerebral autoregulation. One way to probe this system is to use a technique called Cerebrovascular Reactivity (CVR) which measures the change in blood to a given stimulus.



- CVR is calculated as: $\frac{\Delta \text{Blood Flow}}{\Delta \text{Stimulus}}$ and can be seen as an indirect measurement of the vascular reserve of the brain.
- Hypo-/hypercapnia is often used as stimulus as the brain has several mechanisms sensing the CO₂ level and alter its blood flow accordingly.
- The changes in blood flow is usually monitored with transcranial doppler or magnetic resonance (MR) imaging to produce CVR maps.
- CVR is today used as a diagnostic tool for multiple patient groups such stroke, brain tumor and small vessel disease.^[1]

2 Research question

The steady state CO₂ level in the blood is controlled by equation (1) below.



To perform hypo-/hypercapnic CVR measurement there exist therefore three main strategies:

- Use medicine, such as acetazolamide, to alter \dot{V}_{CO2} .
- Change the ventilation \dot{V}_A , for example through breath-holding.
- Alter F_{CO2}^i in the inhaled gas.

The last alternative is considered to be most reliable and reproducible.^[2]

Problem formulation:

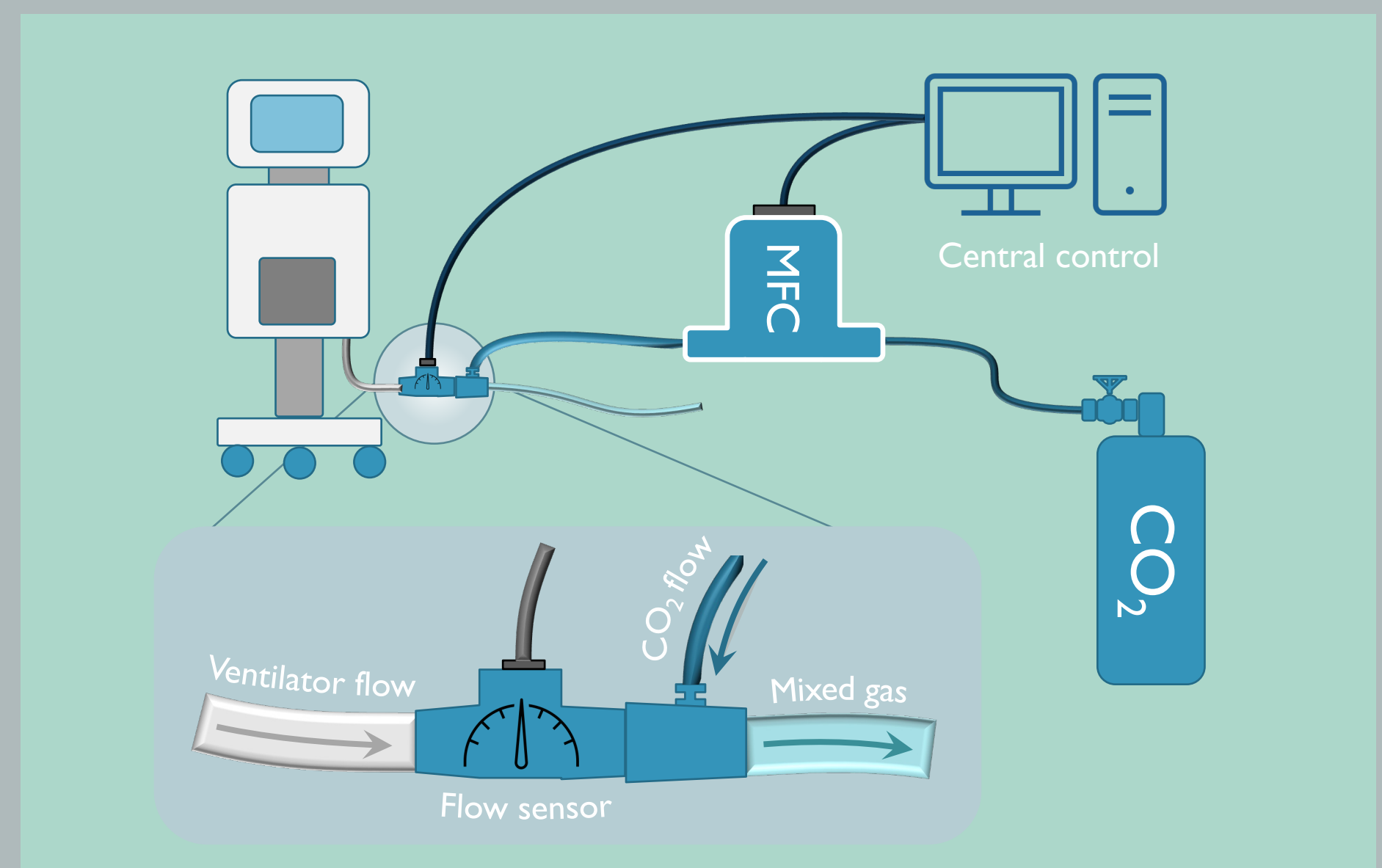
- For ventilated patients there exists today, to our knowledge, no straightforward way to alter F_{CO2}^i .
- Therefore, the only option is to administer drugs or use breath-holding (turning on and off the ventilator).

We therefore set out to construct a system that works together with a ventilator to administer a fix amount of CO₂ in the inhaled gas in order to enable reliable and safe CVR measurements for ventilated patients.

3 Method

We used a solution inspired by INOmax, a system to administer small amounts of Nitric Oxide together with a ventilator.^[3] The idea is relatively simple:

- Measure the flow from the ventilator and control an additional flow of high concentration CO₂ which is added to the ventilator flow.
- By controlling the proportion of the additional flow, a target CO₂-concentration can be achieved.



Our system: iCO₂ CVR System, was built with three main components:

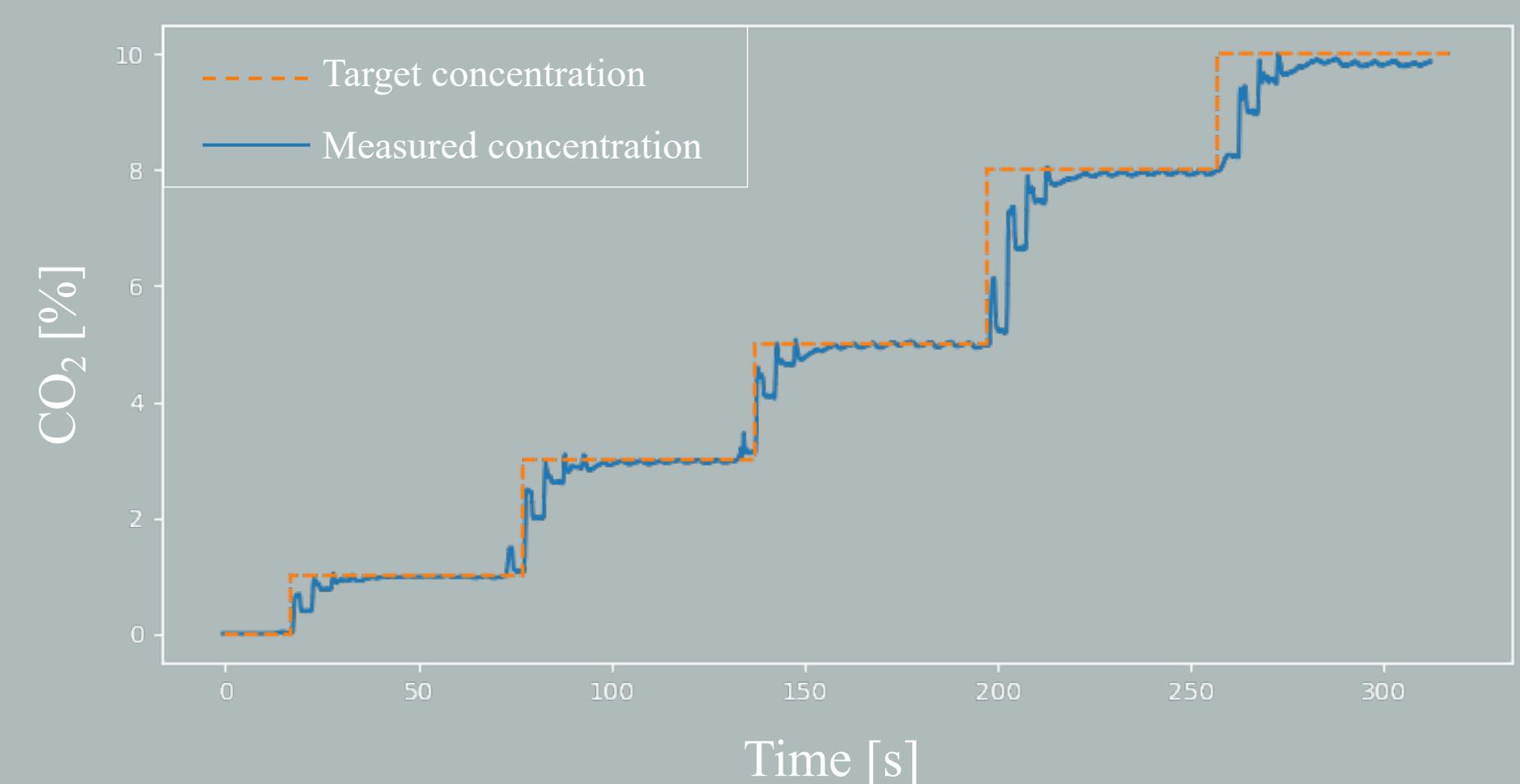
- Flow sensor – Honeywell AWM720P
- Mass Flow Controller (MFC) – Brooks SLA5850
 - Attached to a gas tube containing 10% O₂ + 90% CO₂
- Central control – Computer running a control program written in Python

4 Results and discussion

The iCO₂ CVR System was completed with satisfactory results.

- We were able to achieve target CO₂-concentration ranging from 1–10% with a mean error less than 1% when ventilating a test lung.

Below is an example showing the target CO₂ in orange and the measured CO₂ in blue.



Discussion:

- The timing of the additional CO₂ flow to the ventilator flow turned out to be crucial since the gases mixed very little in the ventilator tube.
- By adding some additional elements to the ventilator circuit, such as a humidifier, mixing of the gases could be increased.
- The length of the CO₂-delivery-tube is a limiting factor since too large compressible volume in the tube will affect the accuracy of the system.

References

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2. Fierstra, J., Sobczyk, O., Battisti-Charbonney, A., Mandell, D. M., Poublanc, J., Crawley, A. P., Mikulis, D. J., Duffin, J., & Fisher, J. A. (2013). Measuring cerebrovascular reactivity: What stimulus to use? *Journal of Physiology*, 591(23), 5809–5821. <https://doi.org/10.1113/jphysiol.2013.259150>
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