

# NIRS Study Information

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# **Chapter 1**

## **About**

This study is being conducted in collaboration with the PrismaHealth Cancer Institute, Furman University Health Science Department, and University of South Carolina School of Medicine Greenville. The principal investigators are Dr. Jennifer Trilk of USC SoMG and Dr. Randolph Hutchison of Furman.



# Chapter 2

## Literature

### 2.1 Near Infrared Spectroscopy

Continuous wave NIRS devices use the modified Beer-Lambert law to calculate the volume of oxygenated and deoxygenated hemoglobin.

Spatially resolved NIRS devices use multiple transmitters at different distances from a receiver to calculate a Tissue Saturation Index, or TSI.

The NIRS device utilized in this study is the PortaMon by Artinis Medical Systems. The PortaMon could be considered both a continuous wave and spatially resolved device. It uses three continuous-wave transmitters with a single receiver to calculate the TSI.

Each transmitter emits two wavelengths of infrared light. One wavelength is for oxygenated hemoglobin and one wavelength is for deoxygenated hemoglobin, therefore each transmitter measures both oxygenated and deoxygenated hemoglobin.

The transmitters emit light into tissue where the light waves bounce off structures in the tissue and return to the receiver.

The receiver measures the light that returns in Optical Densities. Calculations using several different parameters and assumptions convert OD into usable units.

Subcutaneous adipose tissue has different optical properties than muscle tissue. Most calculations do not account for two separate layers of tissue with different scattering and optical properties. There have been some papers within the last few years examining multi-layered tissues and the effects on NIRS measurements using computer modeling and phantom tissues. Most papers utilizing NIRS in exercise physiology assume a limited or negligible effect of adipose tissue thickness on measurements and report participants' adipose tissue thickness.

Different areas of the body have different types of tissue structure and this have different optical properties. Previous papers have reported the measurements of optical properties at various locations of the body. Consult with the device manufacturer and reports of these optical properties to determine the best settings for your device.

Most studies assume that the tissue structures have constant optical properties. This may not hold true, especially during activities like exercise where blood flow increases and the contents of blood fluctuate. Accounting for varying optical properties has led to the development of time-domain and other measurement techniques. These devices are more expensive than a continuous wave device. Therefore, most NIRS devices assume constant optical properties of the measured tissues.

The differential pathlength factor (dpf) is the measurement of the path that the light waves take through tissue. DPF values have been well researched and measured for multiple body locations and differing adipose tissue thicknesses. This parameter is used in calculating TSI.

## Chapter 3

# Patient Recruitment

Run a report in EPIC with the following parameters:

Select patients in registries

FROM

*Registries:*

Registry ID: CANCER POPULATION REGISTRY

*Level of detail:*

Patients

WHERE

DM TREATMENT PLAN PROVIDER CONTAINS ELDER, JEFFERY WAYNE OR EDENFILED, WILLIAM JEFFERY OR JORGENSON, CARLA WALKER OR STEPHENSON JR, JOE JOHN OR PULS, LARRY EDWIN

AND

CHEMOTHERAPY IN LAST YEAR = NO

AND DIAGNOSIS IN GROUper BREAST OR GYNECOLOGICAL CANCER GROUper

TREATMENT PLAN START DATE GREATER THAN TODAY

- 14

This will return a list of potentially eligible patients who meet the parameters specified in the query above. Take a quick look through the list and eliminate any definitively ineligible patients. Sometimes metastatic disease is listed as a second diagnosis and therefore slips through the exclusion of metastatic disease. Vascular diseases and injuries that would preclude ability to exercise are other

common exclusion criteria. Male patients with breast cancer are not eligible for this study. Take a quick look through the patient's problem list. If the patient does not have any immediate exclusionary problems, pass their name and MRN on to the research coordinators, Sydney Hackwell and Katie Banenas. They will conduct a full chart screening and contact the patient if they are eligible.

After contacting the patient, they will contact you if the patient signs the consent form. After the consent form is signed, we can begin contacting the patient to start scheduling for lab visits. Contact Dr. Julie Martin after obtaining the consent form and she will put all NIRS related blood draws into EPIC. If for some reason we are not able to get the NIRS draws done at the same time as the other pre-chemo blood draws, consult with the patient and schedule this through Dr. Julie Martin.

If a patient is not wearing shorts for a lab visit, there are disposable shorts in the lab for patient use. They are located in a box under the bed in the back corner.

# **Chapter 4**

## **Methods**

### **4.1 Lactate Threshold Protocol**

This protocol is the participant's first visit.

#### **4.1.1 Preparation**

At least 30 minutes prior to when the participant is scheduled to arrive, turn on the metabolic cart.

Print out the following instruments:

- MoCA C.4
- MMSE C.3
- Godin-Leisure Time Exercise Questionnaire C.2
- Physical Activity History C.5
- BFI C.1
- PROMIS Global Health C.6
- Lactate Threshold Data Collection Sheet

Write the patient's study ID number, visit ID, and the date in the upper left corner of each in pen.

Assemble the VO<sub>2</sub> mask except for the blue rubber part as described in B.3.3.

Prepare the PortaMon as described in ??.

Calibrate the metabolic cart as described in B.3.1.

#### 4.1.2 Data Collection

Use the steps described in D.1.1 to measure the participant's subcutaneous adipose tissue thickness above their vastus lateralis.

After the patient arrives, the participant's subcutaneous adipose tissue thickness is measured using an ultrasound. If ATT is less than 2 centimeters, the participant continues through the next steps.

Place the NIRS device over the area measured using ultrasound following the steps described in A.4.4.

Administer the following instruments:

- MoCA C.4
- MMSE C.3
- Godin-Leisure Time Exercise Questionnaire C.2

The MoCA, MMSE, and GLTEQ are exclusion criteria. If the participant does not pass these items, they cannot proceed with the study, therefore they are administered before other instruments. If the participant passes the exclusion instruments, proceed with the rest of the instruments. These instruments are not filled out by the participant- a study staff member must administer and score these instruments.

Have the subject complete the following instruments:

- Physical Activity History C.5
- BFI C.1
- PROMIS Global Health C.6

These instruments are completed by the participant.

Review the completed Godin-Leisure Time Exercise Questionnaire and Physical Activity History instruments. Use your best judgement in collaboration with the other investigators to determine the wattage jumps. See 5.1.2 for further information about deciding wattage jumps.

Measure the subject's face to determine mask size. Record mask size on data collection sheet. Finish assembling mask.

Put the heart rate monitor on the patient.

Have the subject get on the stationary bike and make adjustments to the seat height, seat depth, pedal length, and handlebar placement to the comfort of the participant.

If the PortaMon has been on for at least 10 minutes, proceed.

Place the assembled mask on the participant's face and connect clear tubing to the mask.

Conduct Lactate Threshold Exercise Test as described in 5.1.

## 4.2 On/Off Kinetics Protocol

Use the steps described in ?? and ?? along with the reference image of the previous placement to place the device in the same location as the last visit.

Have the subject complete the following instruments:

- Physical Activity History C.5
- BFI C.1
- PROMIS Global Health C.6

If the PortaMon has been on for at least 10 minutes, proceed.

Place the assembled mask on the participant's face and connect clear tubing to the mask.

Conduct On/Off Kinetics Exercise Test as described in 5.2.



# **Chapter 5**

## **Exercise Testing**

### **5.1 Lactate Threshold Exercise Test**

#### **5.1.1 Specifications**

The lactate threshold test is completed on a cycle ergometer.

Ventilatory measurements are collected breath by breath via metabolic cart. Lactate is measured via finger prick.

Prior to beginning exercise testing, measure resting blood lactate. This should be less than 2.0 mmol/L in order to begin testing. Normal resting blood lactate lies between 0.8-1.2 mmol/L, however stress or nervousness prior to an exercise test can increase resting blood lactate levels. If lactate levels are slightly elevated but still lower than 2.0 mmol/L, then lactate may drop back down as the body uses lactate during warm up stages.

The exercise test is conducted as follows:

Begin with a two minute rest. The subject sits on the stationary bike without pedaling for two minutes. Each subsequent stage is three minutes in duration. The wattage increases by a consistent amount between each stage. During the last 30 seconds of each stage, blood lactate is measured via finger prick. Each lactate measurement is taken in pairs for duplicability.

Lactate will increase as exercise difficulty increases. When the subject's blood lactate exceeds 4.0 mmol/L, have the subject finish the next stage and then end the test. If the subject feels they cannot complete the next stage, end the test immediately. A blood lactate of 4.0 mmol/L should roughly correspond to a moderately difficult exercise intensity but should not be a maximal effort.

### 5.1.2 Selecting Wattage Jumps

Discuss the subject's physical activity levels. If the subject doesn't participate in much cardiorespiratory fitness activities, then start at a very low wattage (10 Watts) and make modest jumps (15 Watts). If the subject participates in long distance running, cycling, or swimming, then the subject may start at a higher wattage than less fit subjects and/or make larger jumps.

When selecting wattage jumps, remember that lactate should increase exponentially as wattage increases linearly. Also, when conducting testing, there is no second chance to get the testing correct. It is better to use a very low starting wattage with modest increases as the initial settings, because if the subject is fitter than expected and their blood lactate levels remain stagnant, the wattage jumps can be increased during testing. Blood lactate will not go down in the time it takes to conduct the testing, and if the difficulty increases too quickly then the blood lactate levels will increase so quickly that it will be difficult to analyze the lactate curve.

## 5.2 On/Off Kinetics Exercise Test

### 5.2.1 Specifications

The on/off kinetics test is completed on a cycle ergometer.

Ventilatory measurements are collected breath by breath via metabolic cart (ParvoMedics OneTrue 2400, ??). Muscle oxygenation measurements are collected via SR-NIRS (Artinis Medical Systems PortaMon, ??)

The exercise test consists of three square-wave transitions from exercise at 110% of wattage measured via lactate threshold testing to rest.

# **Chapter 6**

# **Data Storage**

Follow all appropriate data export instructions.

All physical copies are stored in a locked filing cabinet in the Human Performance Lab.

The study RedCap stores all identifying patient information and digital copies of data.

De-identified NIRS and VO<sub>2</sub> data files can be stored on Prisma Health computers.



# **Chapter 7**

## **Database**

### **7.1 Description of Needs**

Store information about patients, their chemotherapy infusions, their blood-work, visits to the Human Performance Lab for exercise testing, data from exercise testing, and analyzed data from the testing.

GUI for future users who may not be familiar with code or able to learn.

Identifying information will be stored in a RedCap that is secured by Prisma-Health. All information in this database will be deidentified.

Retrieve analyzed testing data for future statistical analysis.

Export data for statistical analysis.

Count patients with breast or gynecological cancer, create calendar for visits, retrieve labs, infusions, or visits associated with one patient. Count type of chemotherapy regimen.

### **7.2 Workload Table**

Action	Query Type	Information	Frequency	Priority
Submit a new patient	write	staging activity, cancer type, age, oncologist, other medical problems, chemotherapy regimen type	2 per month	high
Submit a new infusion	write	date, cycle number, day of cycle, medications administered, dosage	4 per month	low
Submit a new lab draw	write	date, type of lab, values	4 per month	low
Submit a new lactate thresh-old test	write	date, vo2 data collection sheet data, parvo data, nirs data	2 per month	medium
Submit a new on/off kinetics test	write	date, oo data collection sheet data, parvo data, nirs data, associated infusion	4 per month	high
Submit analyzed lactate thresh-old data	update	power output at lactate threshold	2 per month	medium
Submit analyzed on/off kinetics data	write	analysis parameters, final analyzed data	4 per month	high
Count patients by type of cancer	read	cancer type	1 per month	low

Action	Query Type	Information	Frequency	Priority
Count patients by type of chemotherapy	read	chemotherapy regimen type	1 per month	low
View blood draw history for one patient	read	date of lab, values, type of lab	2 per month	low
View chemotherapy infusion history	read	date of infusion, cycle number, day of cycle	2 per month	low
View HPL visit history for one patient	read	date of visit, associated cycle number, weight, att, picture of device placement	4 per month	low



# **Chapter 8**

## **Data Analysis**

Be sure you have exported and stored all data appropriately.

### **8.1 Lactate Threshold**

Take the lactate threshold data collection sheet.

Average the duplicate lactate measurements.

Make a plot of lactate by power (Watts). Find the power at which lactate would have been 4.00. That is the power at which lactate threshold occurred.

Multiply this power by 1.1 to calculate the power at which the patient will conduct subsequent on/off kinetics testing.

### **8.2 On/Off Kinetics**

Read in the exported PortaMon data.

The PortaMon does not record in increments of time, it records the sample number. The PortaMon records data at a rate of 10 hz, so it takes 10 samples a second. The time in seconds from the start of the test is the sample number divided by 10.

The PortaMon was on and collecting data from the time the device was placed until the end of the exercise test. We aren't interested in the data from before the start of the exercise test, so that can be removed.

Next, find the time chunks for each section of the test. An on/off kinetics consists of the following sections, each 2 minutes (120 seconds) in duration: Baseline Rest, Warm Up, Work 1, Rest 1, Work 2, Rest 2, Work 3, and Rest 3.

Plot the exercise test. On the x axis, use time in seconds. Have dual y axes, one with hemoglobin arbitrary units and one with percentages. Plot all signals (O\_2Hb and HHb for all exported transmitters and TSI) over time and check that the transitions between sections line up with the expected times. Adjust slightly as necessary, as the start time recorded for the test may be slightly off due to human error / time taken for the PARVO to begin the test.

Use the plot and notes from the test to examine the data for movement artifacts that need to be removed before analysis. Remove as necessary.

The oxyhemoglobin and deoxyhemoglobin measurements have arbitrary units-they are change in oxy and deoxyhemoglobin. This means that we cannot use the amounts of oxyhemoglobin and deoxyhemoglobin and compare those over time. For example, the average oxyhemoglobin at rest could be measured as 4 for one test and 6 at the next test. This does not mean that at the second test, the amount of oxyhemoglobin in the blood was actually higher than at the first test.

Average the O\_2Hb and HHb across the transmitters.

Calculate the steady state at the end of each non-baseline and non-warm up section (6 sections in total- 3 work sections and 3 rest sections) by averaging the last 30 seconds of the section for each signal type (O\_2Hb, HHb, and TSI).

There are 18 steady state values: 3 signal types and 6 sections.

Calculate the delta steady state for each transition. Subtract the rest steady state from the corresponding work steady state. I.e. subtract rest 1 steady state from work 1 steady state.

There are 9 delta steady state values: 3 signal types and 3 transitions.

We use the following equation to describe an asymptotic or monoexponential curve:

$$y = (Asym - B_0) - (Asym - B_0) * e^{-t/\tau}$$

Where *Asym* is the asymptote (or Steady State),  $B_0$  is the value at time 0,  $t$  is the time (in seconds), and  $\tau$  is the time constant.

Calculate the coefficients for the monoexponential curve for each transition. There are 9 monoexponential curve equations: 3 signal types and 3 transitions.

Use the app, shown below, to do all of these steps without having to run any code.

```
knitr::include_app('https://saragracebiddle.shinyapps.io/DataAnalysis/')
```

When you are satisfied with the results obtained from the app, download the report and upload it into the RedCap.

# Appendix A

## PortaMon

### A.1 Specifications

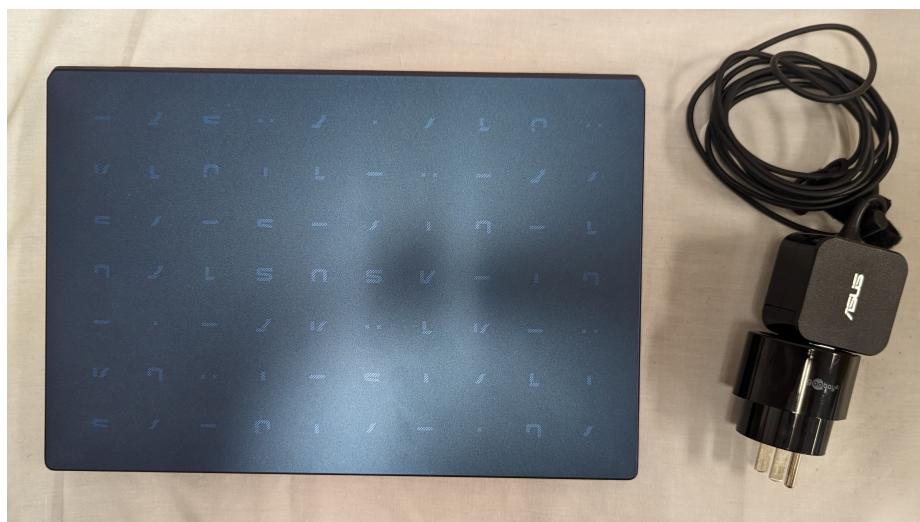
The PortaMon by Artinis Medical Systems is a spatial-resolved near-infrared resonance spectroscopy device. It uses one receiver with three infrared transmitters. Each transmitter is 5 millimeters further from the receiver. This allows the application of mathematical formulas to calculate an absolute tissue hemoglobin saturation, which Artinis Medical Systems refers to as the Tissue Saturation Index, or TSI. In the literature, this is referred to by various acronyms, such as TOI (Tissue Oxygenation Index). A full examination of the mathematics and physics behind the calculation of TSI was published as part of the SPIE Conference on Optical Tomography and Spectroscopy of Tissue III in 1999 (Suzuki et al., 1999).

The yellow case is one unit.



It contains the following:

- ASUS laptop
- laptop charging chord with European to American outlet converter



- yellow Artinis Oxysoft USB - Bluetooth dongle - ROCKEY4ND license key



- PortaMon device



- two batteries - battery charging dock - micro-USB chord for battery charging dock



## A.2 Oysoft

Oxysoft is the software created by Artinis Medical Systems to connect with the PortaMon, collect, and analyze data.

Oxysoft is installed on the ASUS laptops that go with the yellow cases. These laptops do not have any PrismaHealth or USC SoMG software on them. If someone should want to install Oxysoft on a PrismaHealth or USC SoMG computer, Oxysoft will have to be approved by the respective organization's IT team.

Oxysoft has been approved to be on Sara Biddle's PrismaHealth desktop computer by PrismaHealth's IT. No other PrismaHealth or USC SoMG computers have Oxysoft installed.

If you wish to install Oxysoft on a personal computer or have been approved to install Oxysoft on an organization's computer, plug the yellow Artinis Oxysoft USB into the computer and run the installer.

- Oxysoft uses a legacy Bluetooth dongle to connect to the PortaMon. This makes using Oxysoft on a computer with the most up to date Windows difficult. It is easier to collect data on the ASUS laptops, which are default configured to use the Bluetooth dongle, and then export the data for analysis.
- Oxysoft creates files with the extension .oxy4 for every new measurement.

### A.3 Batteries

The PortaMon batteries can hold up to 8 hours of power for the device. Make sure they are fully charged prior to a patient's visit.

Plug the micro-USB into the battery charging dock. Plug this into an outlet. Place the battery into the dock, sticker side down. The three metal connectors on the narrow side of the battery must be touching the three metal connectors on the charging dock. If the battery is correctly placed in the dock, an LED will light up. When the battery is fully charged, the LED will be green.



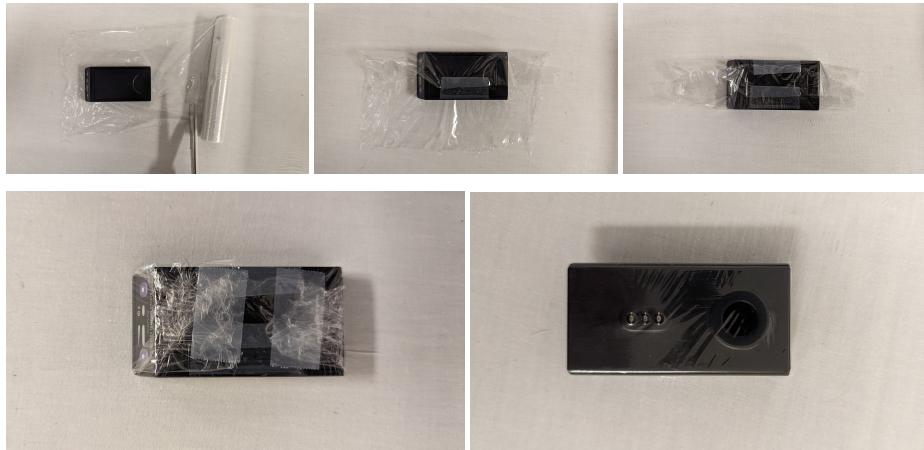
## A.4 Before Data Collection

### A.4.1 Instrument Preparation for Data Collection

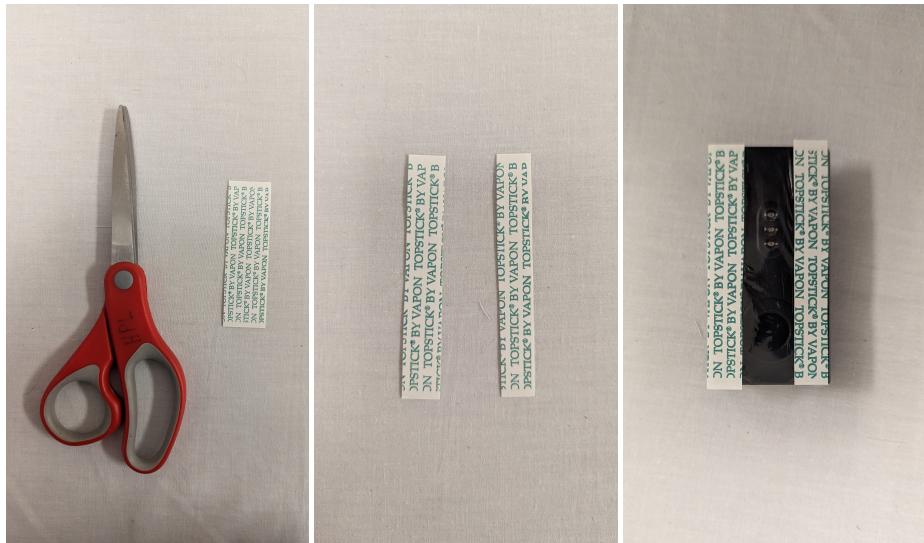
- Place a fully charged battery in the PortaMon, with the three connectors on the battery touching the three connectors in the PortaMon.
- Slide the top of the PortaMon back in until it clicks into place. If the battery is charged and placed correctly, LEDs on the bottom of the PortaMon will light up.



- Wrap the PortaMon in clear saran wrap and tape the saran wrap in place. This prevents sweat and other moisture from getting in to the PortaMon. Since the PortaMon has electronic parts, it cannot be sterilized using liquids and the saran wrap means we do not have to sterilize the device itself. When testing is complete, pull saran wrap and tape off the PortaMon and throw it away.



- Cut the double-sided taupe tape length-wise into two long strips.
- Place one strip to the left of the transmitters and receiver and one strip to the right of the transmitters and receiver. Leave the exterior side of the double-sided tape covered until the placement on the thigh is complete.



#### A.4.2 Bluetooth Troubleshooting

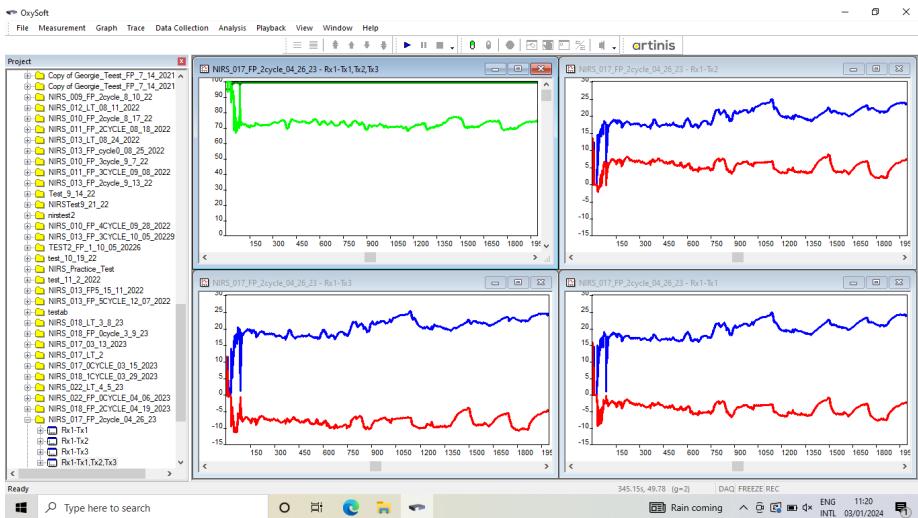
#### A.4.3 Start a Measurement

- Turn on the ASUS laptop.

- Ensure the bluetooth dongle and the Rockey dongle are plugged in to the laptop.
    - The Rockey license key must say ‘ROCKEY4ND’ and NOT ‘ROCKEY4’. The ‘ROCKEY4’ is from an older Artinis device and is not the current license key. You will not be able to use the most up to date Oxysoft software with the incorrect license key.

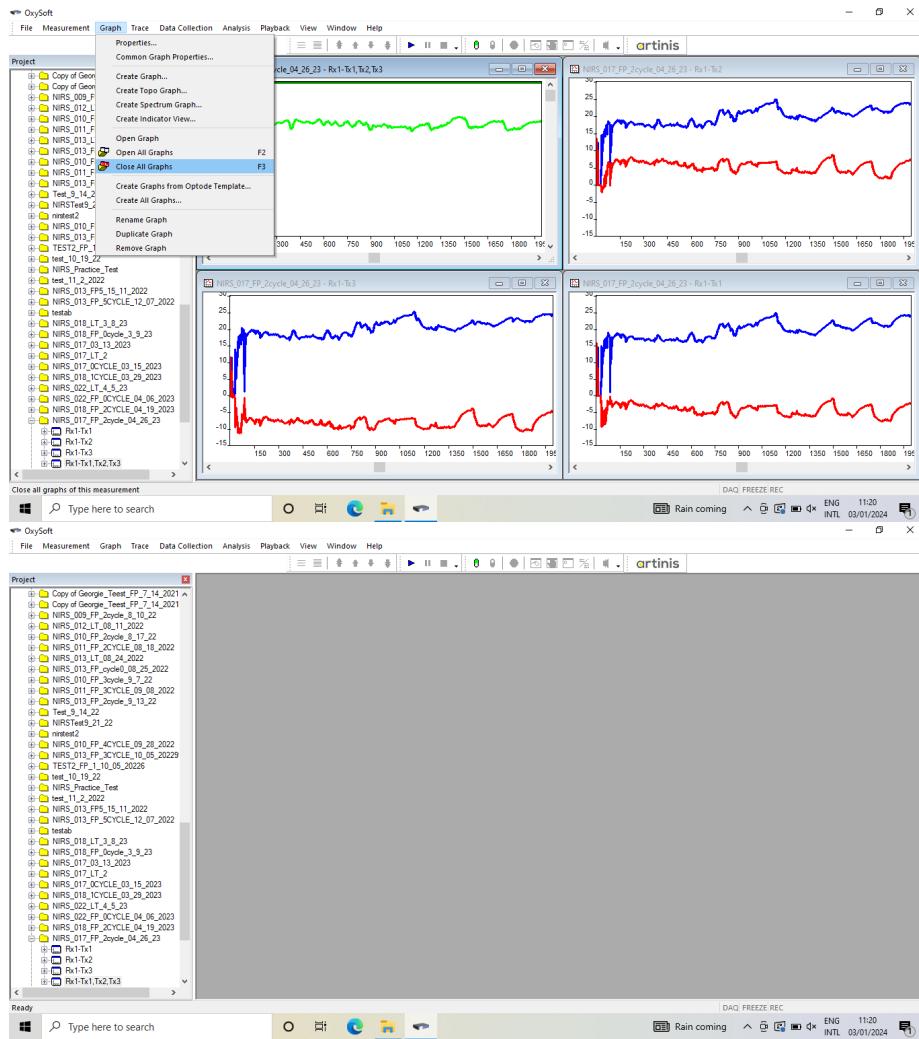


- Open Oxysoft.

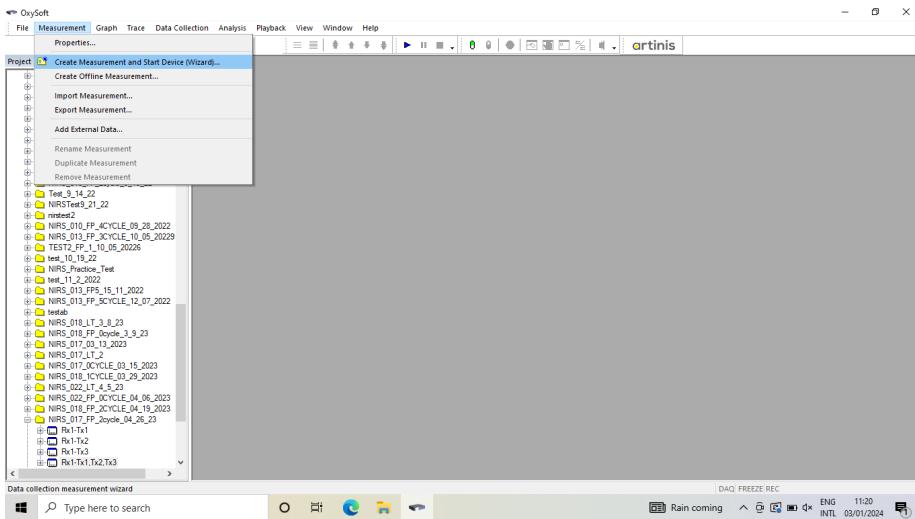


- Close any open graphs in the main panel.

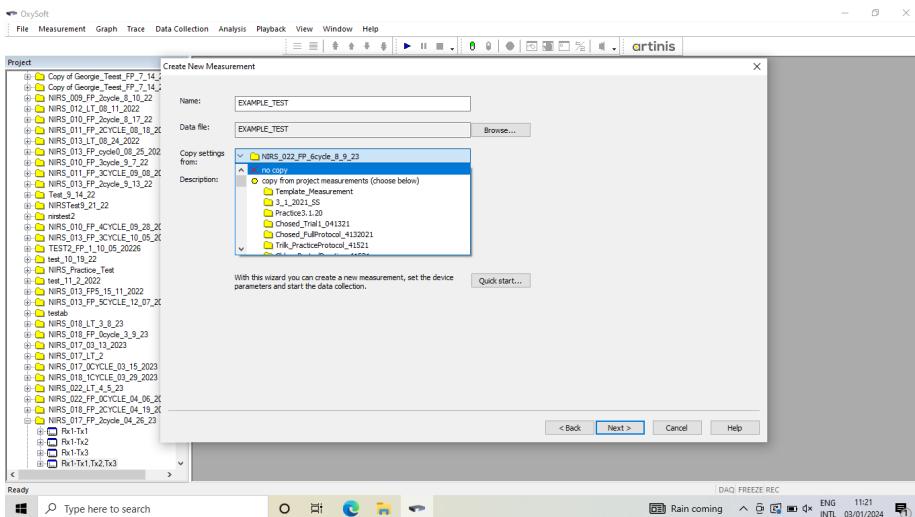
- Select Graphs in the top bar.
- Select Close all graphs. There should be no open graphs in the main panel.



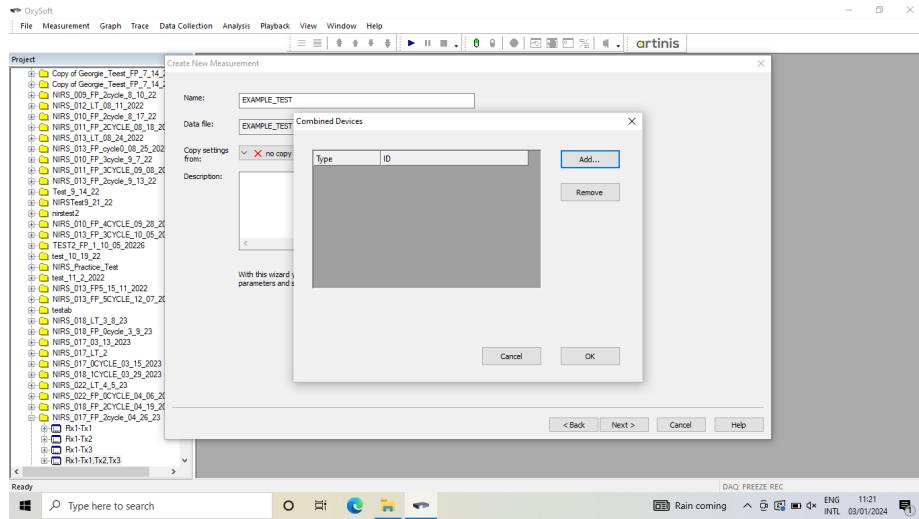
- Select Measurement in the top bar.
- Select Create Measurement and Start Device (Wizard)... from drop down.



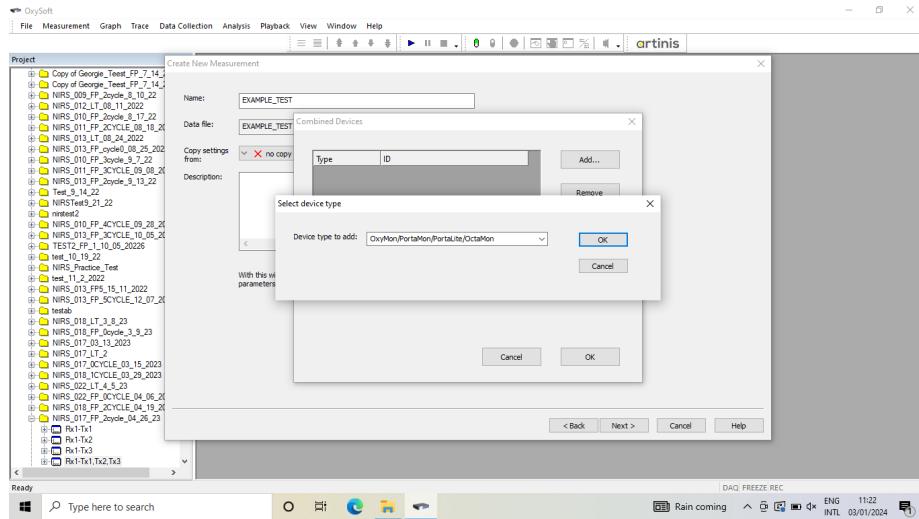
- Name the measurement appropriately.
- Select the **Copy settings from:** drop down.
- Scroll to the top of the drop down and select **no copy**.



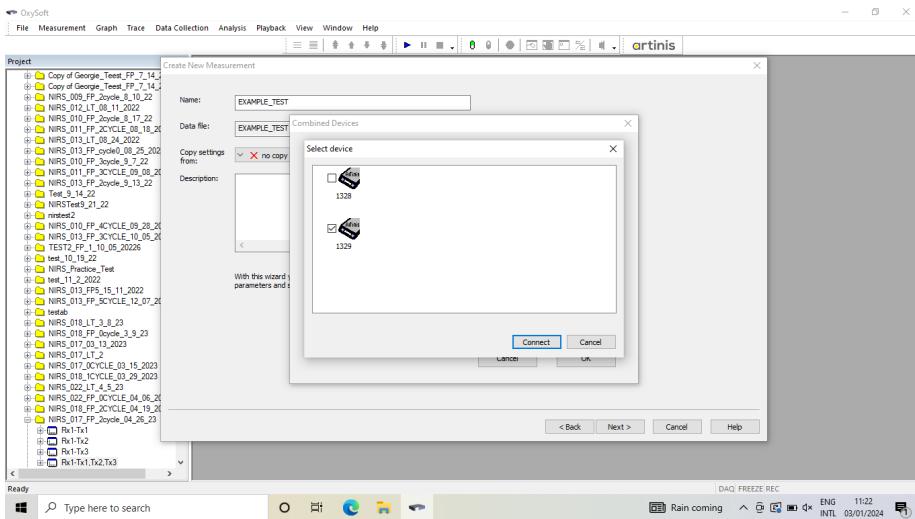
- Click **Next**. Pop up will open to add a bluetooth device.



- Click Add. Pop up will open to select type of device.
- Select OxyMon/PortaMon/PortaLite/OctaMon, which should be auto filled.



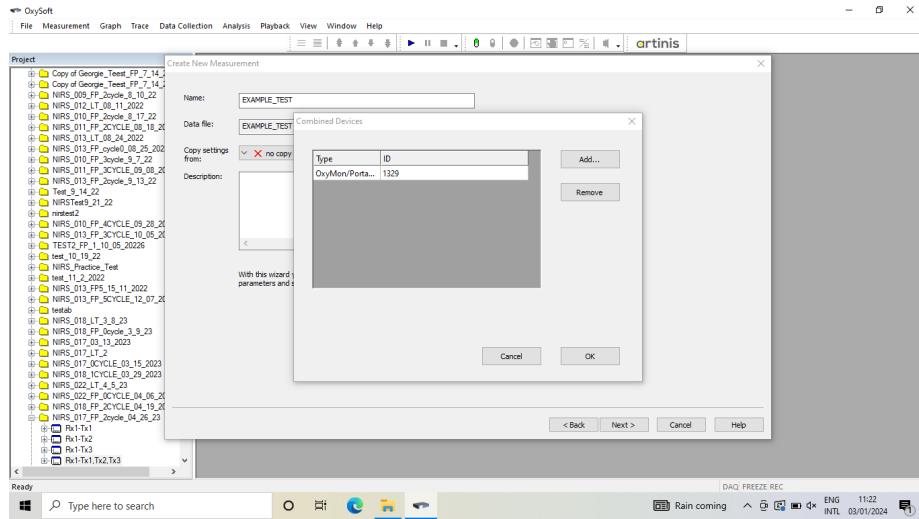
- Click OK.
- Select correct device number, this should match the number on the top of the Portamon.



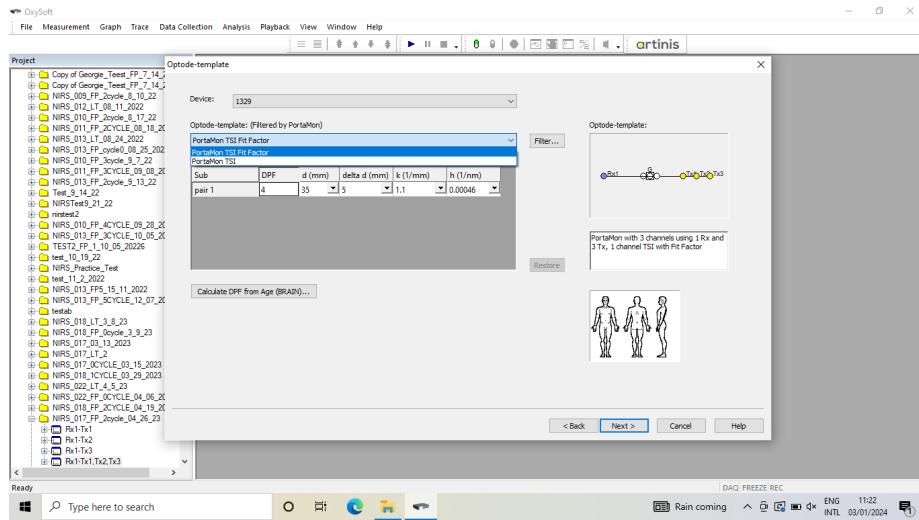
- Click **Connect**.
- Device should pair with the computer. If it does not, try the following steps:

- Check the device is on. The green LED light should be on. If not, turn on by holding down the button on the left until the lights come on (about three seconds).
- Hold the bluetooth button down for about three seconds while you click **connect** on the computer.
- Unpair the device with the computer in the computer's bluetooth settings, then re-pair.
- Restart the computer.
- Try using the other Portamon.

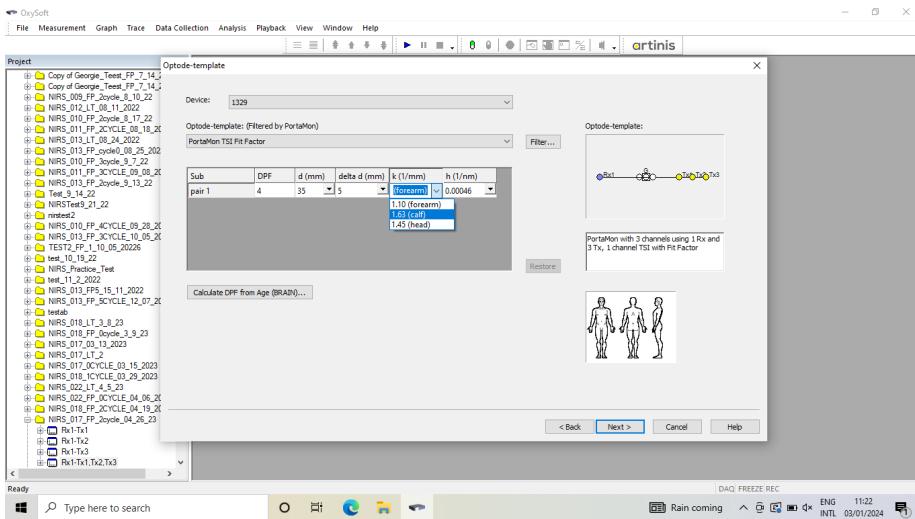
- When successfully connected, a blue LED will turn on on the Portamon and the device type and number will be listed in the **Combined Devices** pop up.



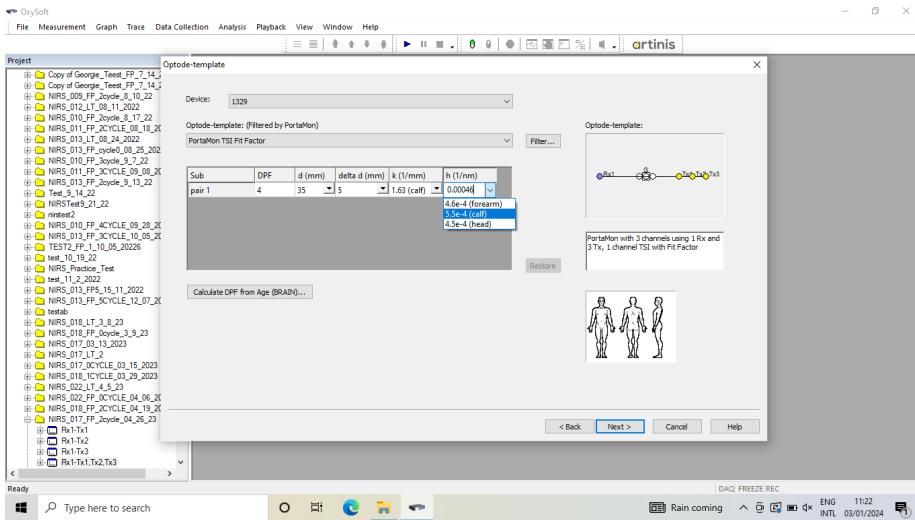
- Click OK.
- The Optode-template window will now be open.
- Under Optode-template: (Filtered by PortaMon) click the drop down.
- Select Portamon TSI Fit Factor.



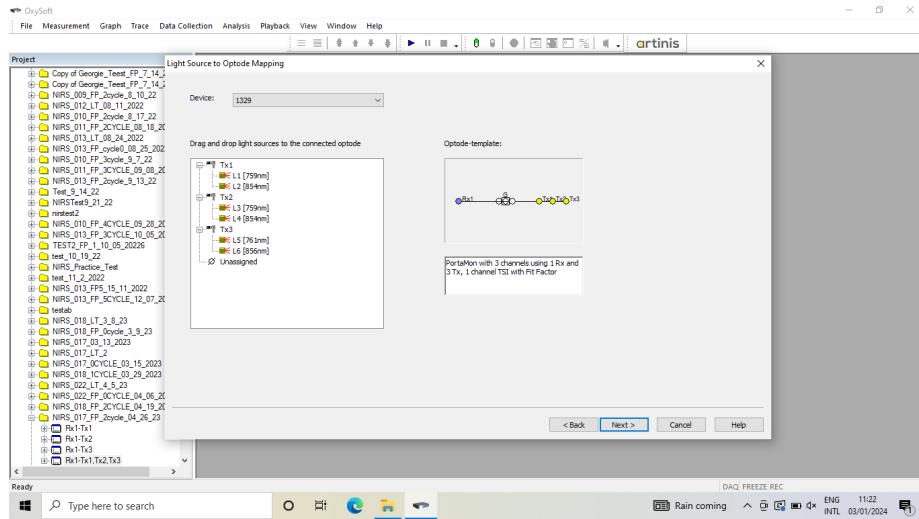
- Under k (1/mm) click the drop down.
- Select 1.63 (calf).



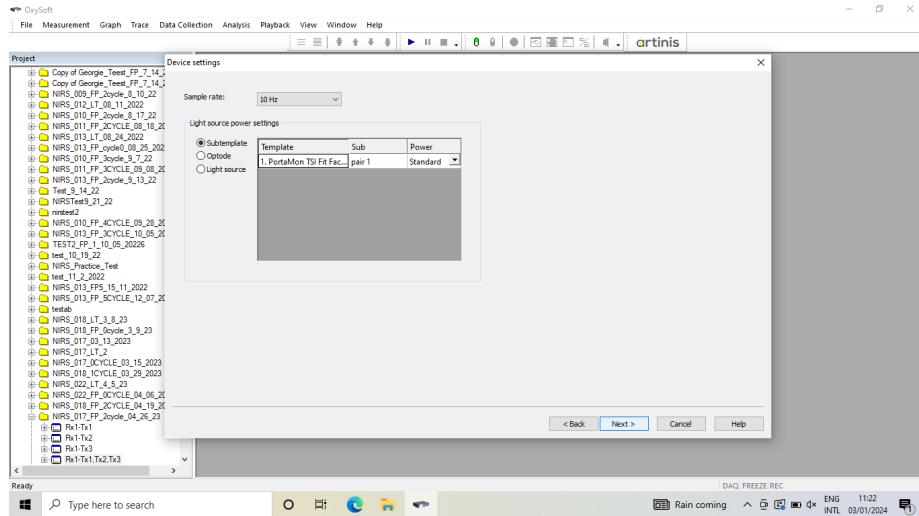
- Under  $h$  ( $1/\text{mm}$ ) click the drop down.
- Select  $5.5\text{e-}4$  (calf).



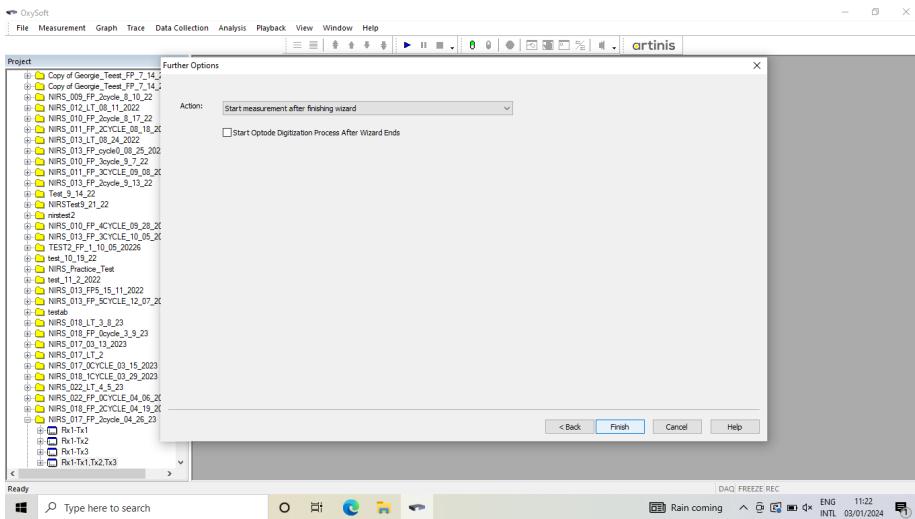
- Click **Next**.
- The Light Source to Optode Mapping window will now be open. Leave all settings as is and click **Next**.



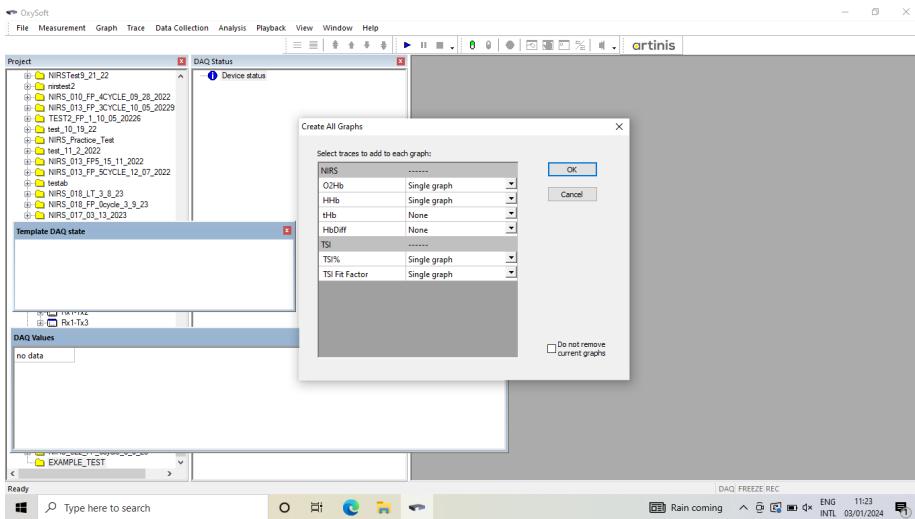
- The Device settings window will now be open. Leave all settings as is and click Next.



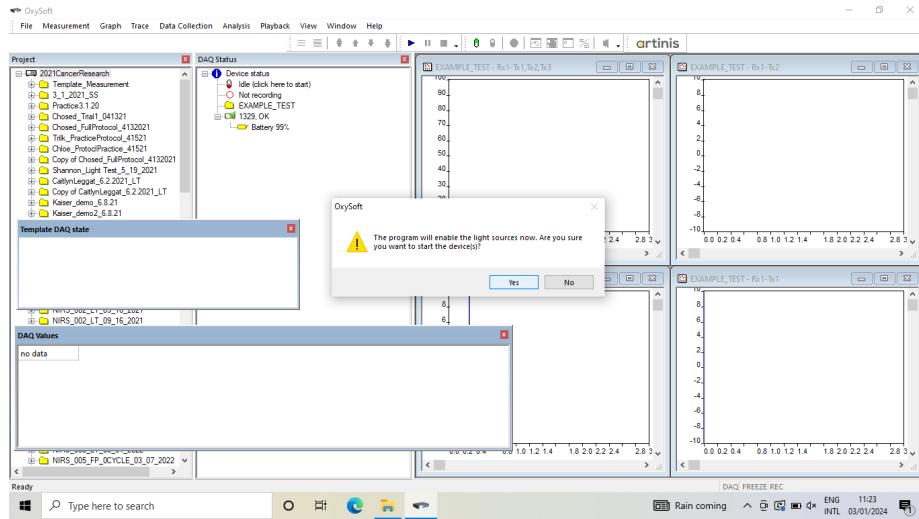
- The Further Options window will now be open. The Action drop down should have Start measurement after finishing wizard selected.



- Click Finish.
- The Create All Graphs window will pop up. Leave settings as is and click OK.



- There will be an Oxysoft pop up. It says ‘The program will enable the light sources now. Are you sure you want to start the device(s)?’ DO NOT click yes yet. Wait until device has been placed correctly. Follow the steps described in A.4.4.



#### A.4.4 Instrument Placement on Vastus Lateralis

The PortaMon can measure up to 2 centimeters depth into tissue. Since we are interested in muscle oxygenation, the muscle belly has to be within 2 centimeters from the surface of the skin. Otherwise, the device is not measuring muscle oxygenation but adipose tissue oxygenation. Therefore, we use an ultrasound to measure the subcutaneous adipose tissue thickness.

Place the device over the location measured by ultrasound.

Connect the NIRS by following the steps under A.4.3.

Click 'yes' to enable light sources and begin data collection. Wait approximately 10 seconds. Check the DAQ (Data Acquisition) values to determine whether the NIRS is receiving good data. If one or more of the DAQ values show as red, then make small adjustments to the placement of the device to attempt to get better DAQ values. If the small adjustments add up to a large difference from the original ultrasound measurement location, re-measure adipose tissue thickness using the ultrasound in the new location.

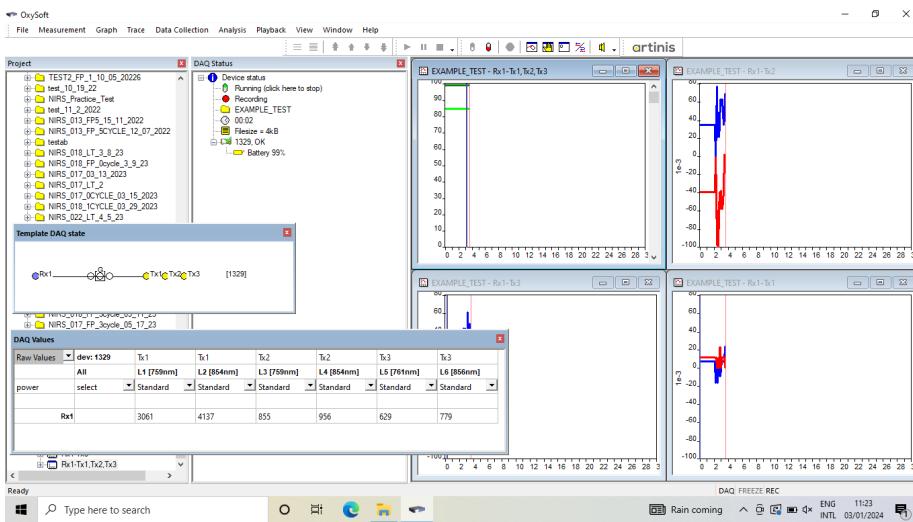
When all three DAQ values remain white when the subject moves around their leg, then the device is in the best location. If one of the DAQ values turns red during movement, that can be okay but only ONE. If two of the DAQ values are red, then TSI cannot be calculated so we do our best to have at least two good signals, three if at all possible.

Rock the device laterally and remove the tape cover on the exposed side. Rock the device back into place and stick the tape to the subject's skin. Repeat on the other side. Double check that the DAQ values remain consistent.

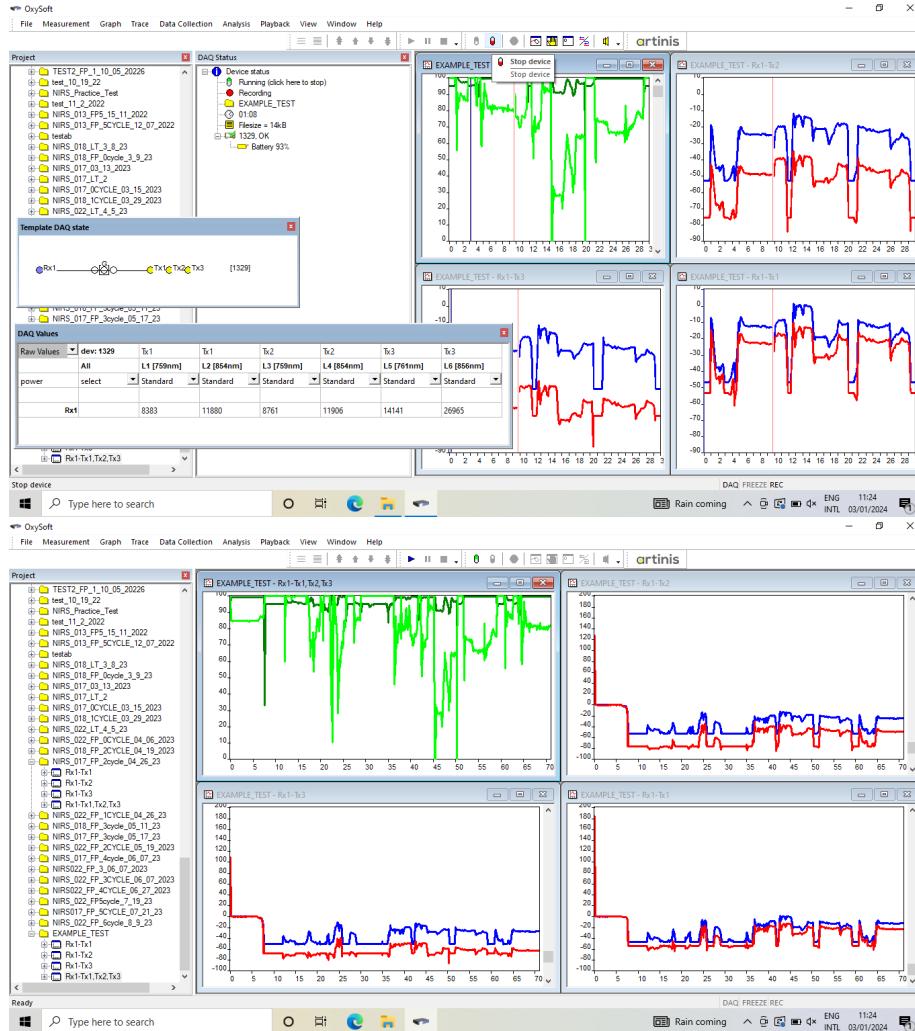
Take a picture of the placement. Use a measuring tape to show the distance from the top of the kneecap to the bottom of the device and use a perpendicular measuring tape to show the distance from the center of the thigh to the right edge of the device. Print the picture and place in the participant's physical file. Upload a .png of the picture into the RedCap.

## A.5 During Data Collection

- When the device is placed correctly, click yes. The new measurement will begin.



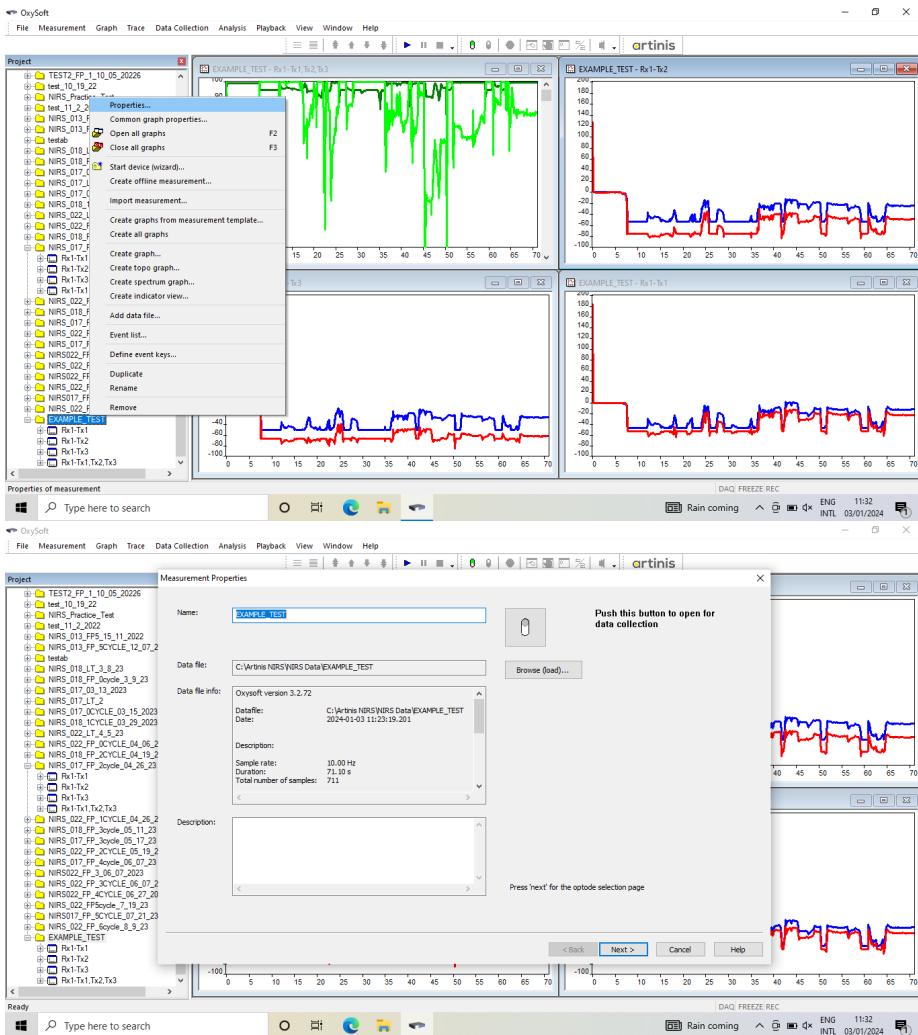
- When you are done with the measurement, click the red pill in the top bar to stop the light sources. Oxysoft will stop the rolling every 30 seconds view and you will be able to see the whole measurement in the panes.



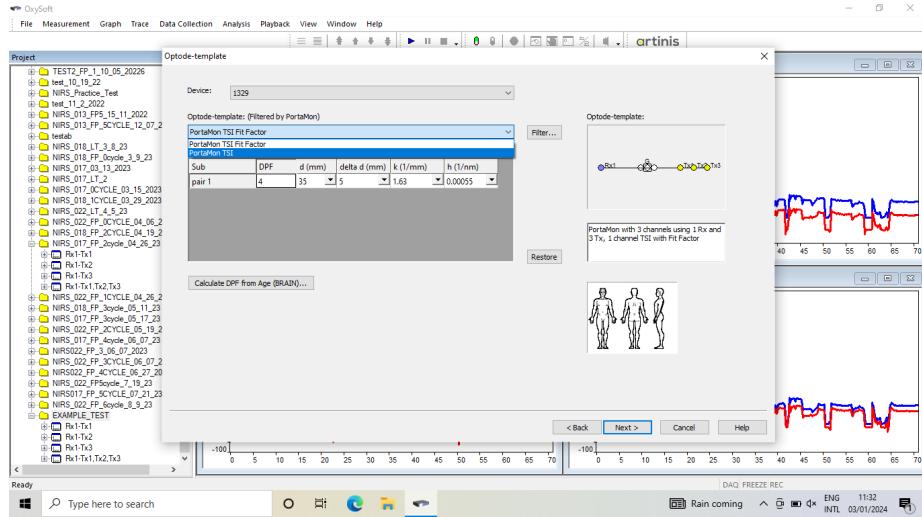
## A.6 After Data Collection

### A.6.1 Change Measurement Properties

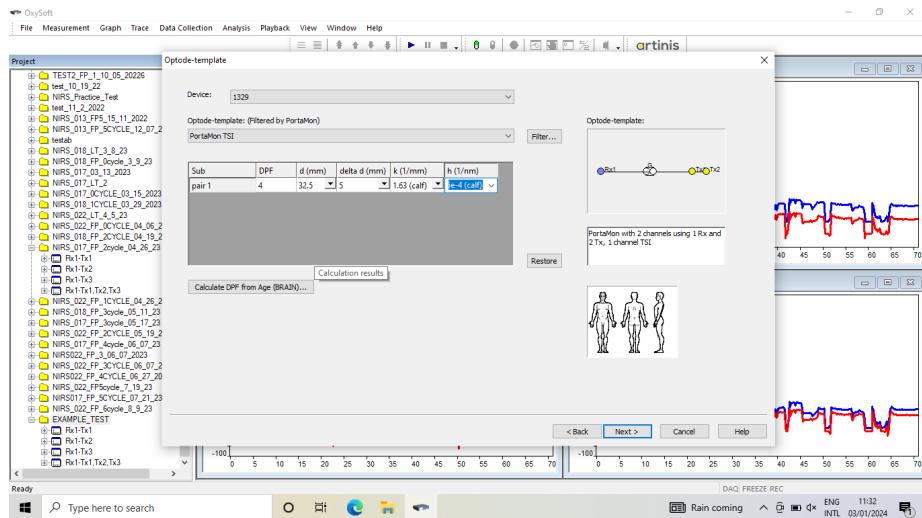
- Right click on the name of the measurement in the Project panel on the left.
- Select Properties.... This will open up a new panel.



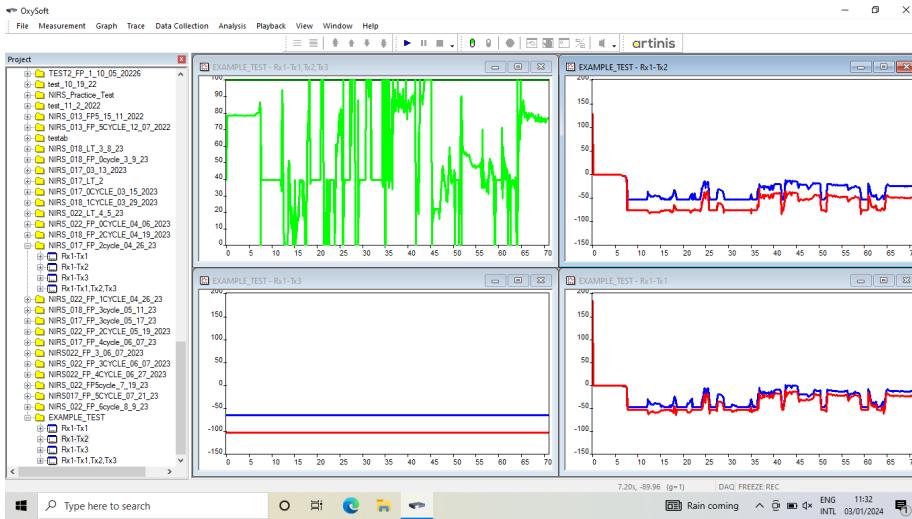
- Leave the measurement name and data file as is.
- Click **Next**.
- Under Optode -template: select PortaMon TSI instead of PortaMon TSI Fit Factor.



- Change the settings as follows:
  - Under  $d$  (mm) type 32.5.
  - Under  $\delta d$  (mm) type 5.
  - Ensure  $k$  (1/mm) is 1.63 (calf). and  $h$  (1/mm) is 5e-4 (calf).



- Click Next.
- When Tx3 has been removed from the TSI calculations, the graph panels will look different. The Rx1-Tx3 panel will have two flat lines, one blue and one red. The TSIFF panel will still have a green line for TSI, and will no longer have a TSIFF line in that panel.



To undo these changes and revert to the original measurement with Tx3 included in the TSI calculations, right click on the file name and open up the **Properties** panel again. Click **Next**.

Next to the **Optode-template** settings table, there is a button on the bottom right that says **Restore**. Click **Restore**. Click **Next**.

The graphs are now restored to their original state.

### A.6.2 Data Export

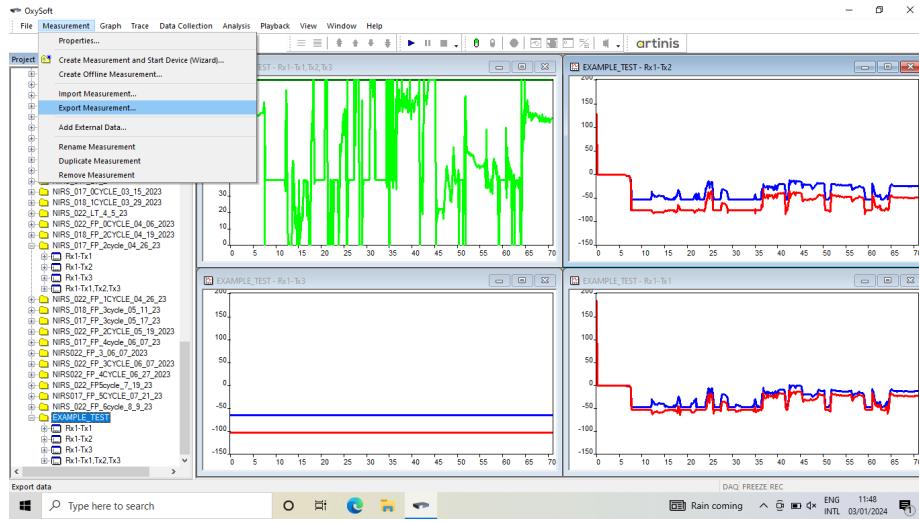
Ensure that only one measurement is open in the Oxysoft graph panels. Do this by closing all graphs, then selecting the name of the measurement you want to open in the file pane on the left.

If you are not sure or cannot tell, under the **Graph** drop down of the header bar, select **Close all graphs**. There should be no visible graphs in the main panel.

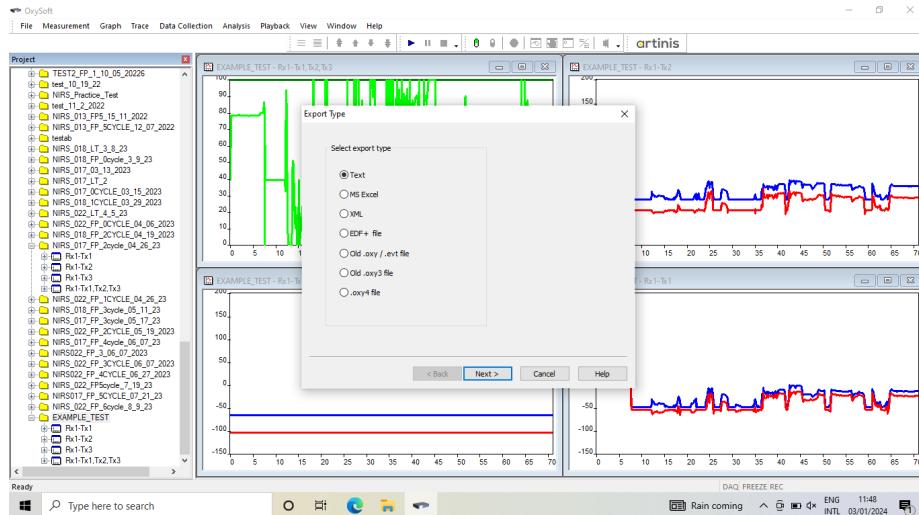
Select the name of the measurement you want to export in the **Project** panel on the left. Click **Open all graphs** under the **Measurement** drop down while the name of the file is highlighted. There should be four panels open, one titled **Tx1-Rx1**, **Tx2-Rx1**, **Tx3-Rx1**, and one **TSIFF**.

If all three transmitters had good DAQ values during data collection and never dropped into red, skip the next steps. If there were issues with only Tx3, follow the steps described in ??.

Select **Export measurement** under the **Measurement** dropdown.

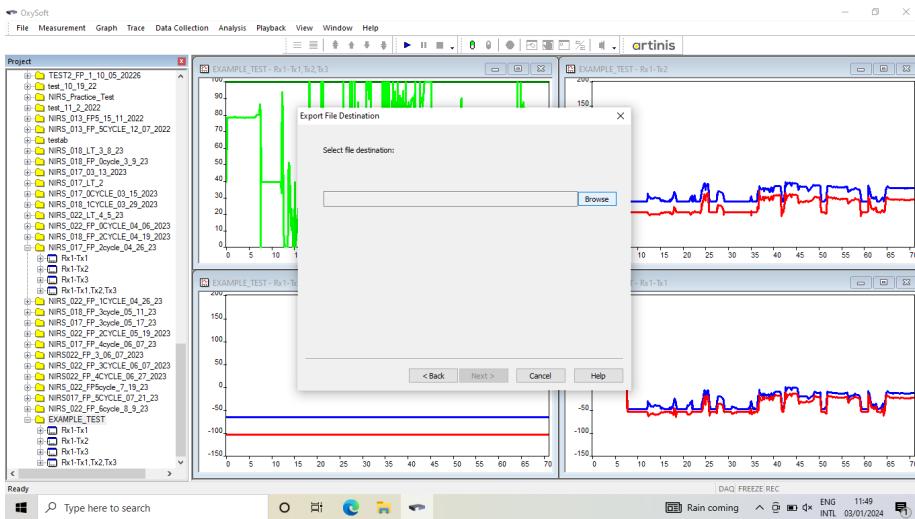


A new panel will open. Select the type of file you wish to export as. The automatic selection is text (.txt), which is the preferred type of export.

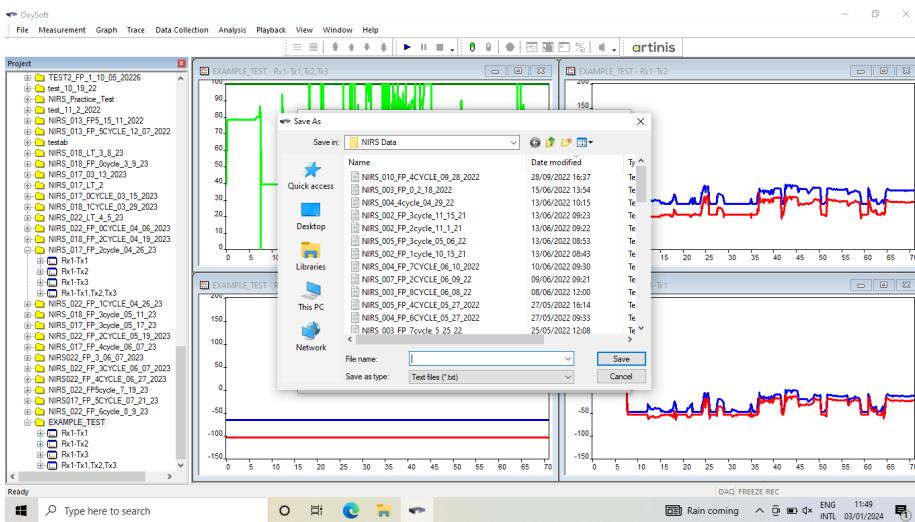


Click Next.

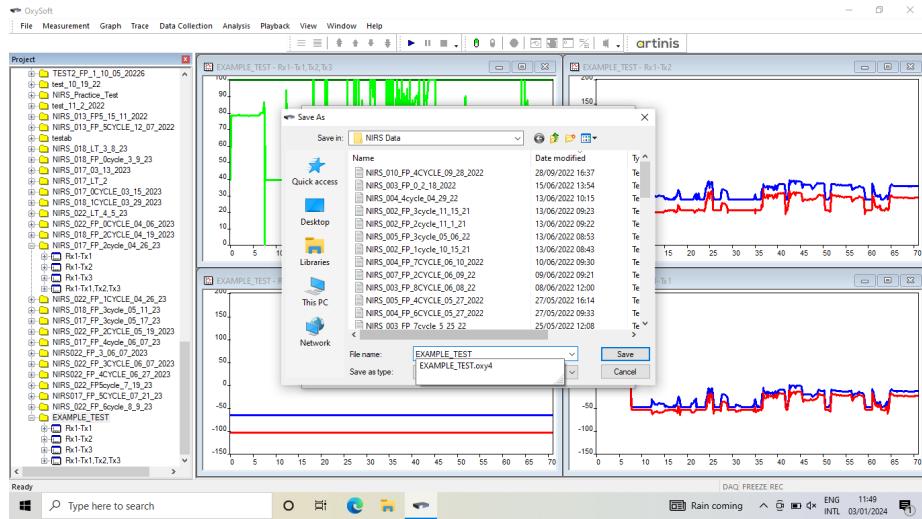
A new popup will prompt you to choose a location for the exported file.



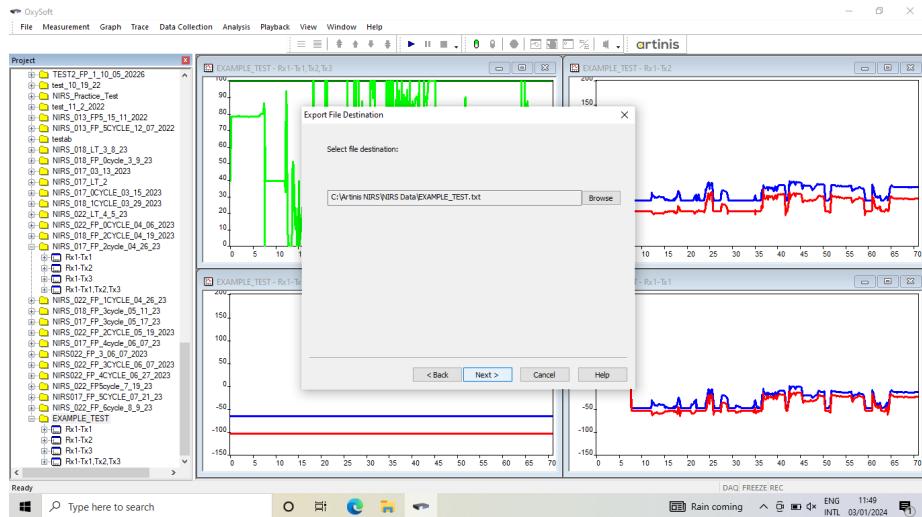
Click Browse.



There is a folder on the desktop for all data for the NIRS study. Select that folder as the destination. In the file name bar, type in the name of the measurement. I prefer for the exported file and the name of the measurement in Oxysoft to match for consistency. When you type in the name of the measurement in oxysoft, it will prompt you to select `nameofthefile.oxy4`. Do not select this. Just type the name of the measurement with no file extension.



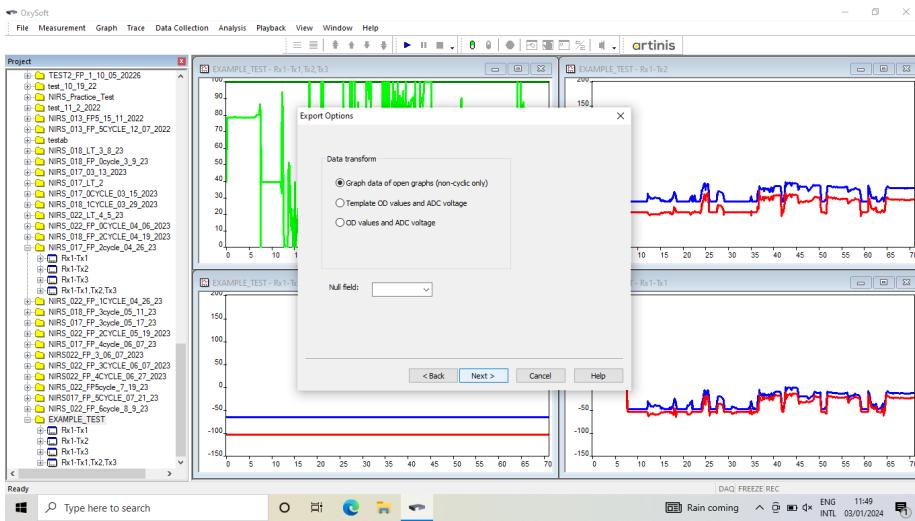
Click Save.



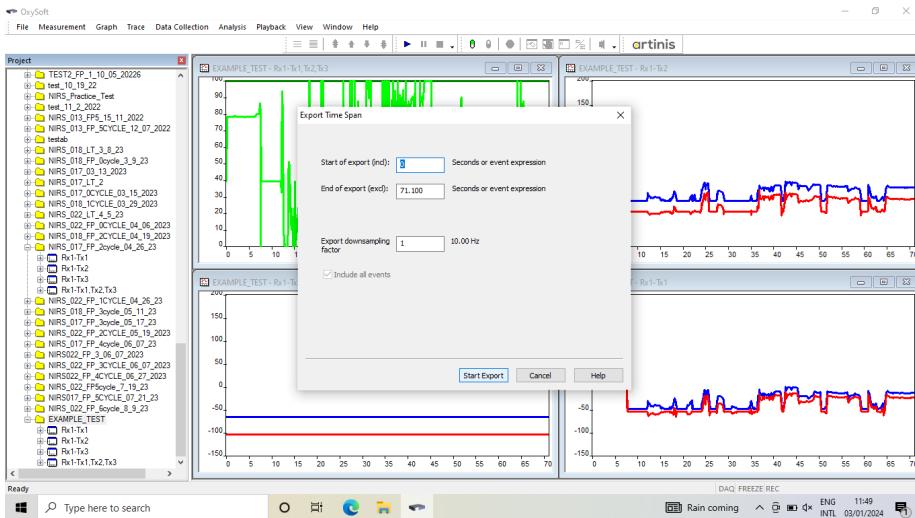
The file path should be to the NIRS data folder and should be named the same as the measurement with a .txt extension.

Click Next.

The Export Options popup will have several selections for the type of data to export. Leave the default selection, Graph data of open graphs. Click Next.



The Export Time Span pop up will have the option to remove data from the beginning or end of the measurement. Leave the time span as it is and export the full measurement. Click Start Export.





## Appendix B

# ParvoMedics TrueOne 2400

### B.1 Specifications

The TrueOne Metabolic Cart by ParvoMedics measures maximal O<sub>2</sub> consumption, indirect calorimetry, and other metabolic conditions.

The computer that has the ParvoMedics software installed is not connected to the internet in any way. It has Windows 7. **Never connect this computer to the internet or update Windows.** This would require attaining approval to install the ParvoMedics software on PrismaHealth computers, which would take several months. The ParvoMedics software is not frequently updated and Windows updates could break the installation. Everything works as it is, and if it ain't broke, don't fix it.

### B.2 Software

### B.3 Before Data Collection

#### B.3.1 Calibration

In order to conduct calibration, the cart must have been powered on for *at least 30 minutes*. Turn on the cart by flipping the switch on the power strip on the back of the cart.

Log in using the password taped to the cart under the keyboard.

The computer on the cart is not connected to the internet. After logging in, you need to manually set the date and time to today's date and time. If you do not manually set the date and time, the data collection software will use

the incorrect date and time and will save the data as such, which will make it difficult to locate later.

Open the ParvoMedics software.

Ensure the filter and other pieces are connected to the gas analyzer as shown below:

*TODO add picture*

#### B.3.1.1 Flowmeter Calibration

Select ‘Flowmeter Calibration’ from the software home page.

Enter room temperature (Celcius), barometric pressure (mmHg), and room humidity (%) from the Davis Vantage Vue on Sara Biddle’s desk.

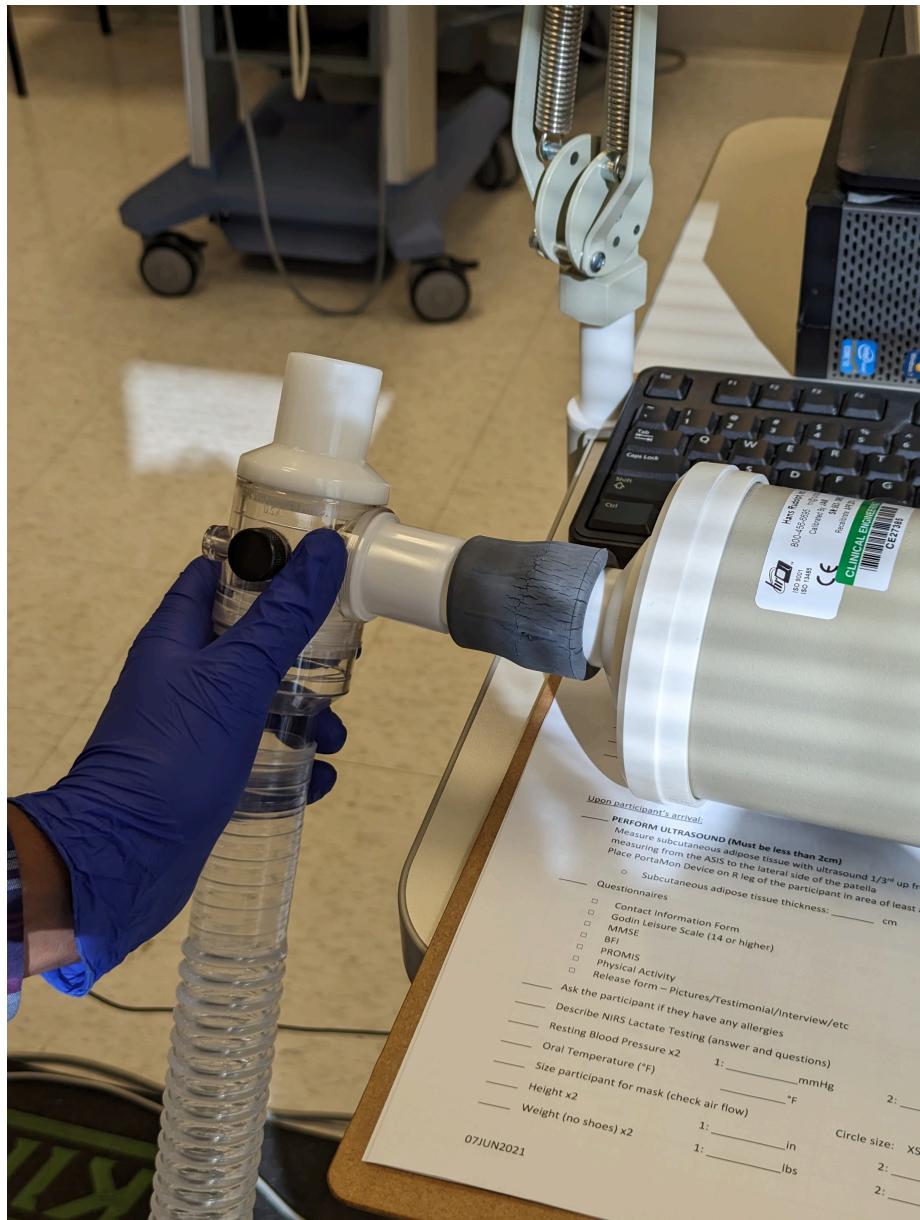
Get a piece of long clear tubing from Parvo equipment cabinet behind the door. The calibration syringe and connection piece are located in the third shelf on the Parvo cart.

Connect one end of the tubing to the clear end of the connection piece.



Connect the other end of the tubing into the gas analyzer module.

Connect the white end of the connection piece to the mouth of the calibration syringe.



Get Frankie Bennett to help with the actual flowmeter calibration.

### B.3.1.2 Gas Calibration

Select ‘Gas Calibration’ from the software home screen.

Enter room temperature (Celcius), barometric pressure (mmHg), and room humidity (%) from the Davis Vantage Vue on Sara Biddle's desk.

Click ‘OK’.

When prompted, find the sample gas (gas container on the bottom left of the cart) and rotate the black handle on top of the gas container counter-clockwise until you cannot turn it anymore. Do not turn the black knob that is taped in place.



Click ‘OK’ and wait for the next prompt on the screen.

Rotate the black handle on top of the gas container clockwise as far as you can to the “OFF” position.



Click 'OK' to see calibration results.

DO NOT SAVE results if the percentage is greater than 1%. This means that this calibration differs from the last calibration by more than we want. If you save the results, since it is a percentage change from the last calibration, the next calibration will have to decrease by greater than 1%.

Example: First calibration attempt results in a percentage of 1.5%. In order to

be within 1% of the attempt prior to the first attempt, the second calibration attempt will need to result in a percentage of -0.5%.

To save us from doing this math in between calibration attempts, just do not save attempts greater than 1% difference.

If results are greater than 1%, try the following steps:

- open the lab door
- turn on the fans to blow air around the room
- wait at least 10 minutes and attempt calibration again

When the gas calibration results in less than 1% change from the previous calibration, save results and exit to the home screen.

If you have tried all of the steps above and the gas calibration still results in greater than 1% change from the previous calibration and you are out of time, save the gas calibration results. After the participant leaves, you can try to redo the gas calibration to get back under the 1% change from the previous calibration. There are several reasons that the calibration will still result in a difference greater than 1%, such as the weather, issues with the air conditioning, and more. The only things we can do to prevent issues with the calibration is limit the number of people in the lab. Have extra people stand out in the hallway or try to schedule fewer people to come in to the lab.

### B.3.2 Mask Sizing

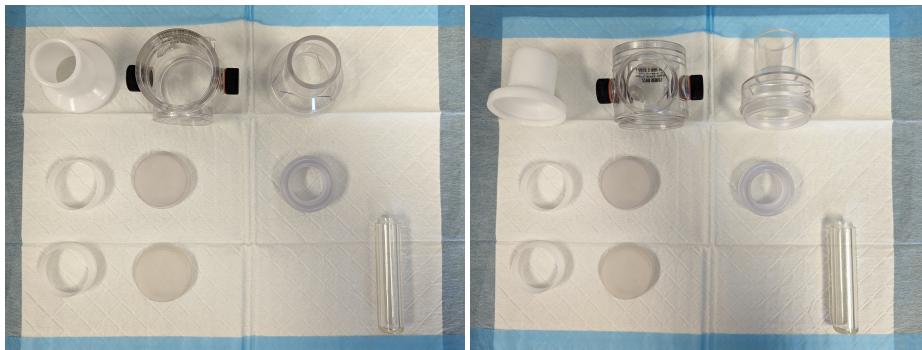
- Find the mask size guide in the caddy that sits on the wheeled cart that sits next to the ultrasound.
- Hold the mask size guide up to the face of the patient. Hold the top of the guide at the bridge of the nose.
- Look at the bottom of the guide and see where the chin sits. Choose the appropriate size. If the chin rests between two sizes, try the smaller size first.

### B.3.3 Mask Assembly

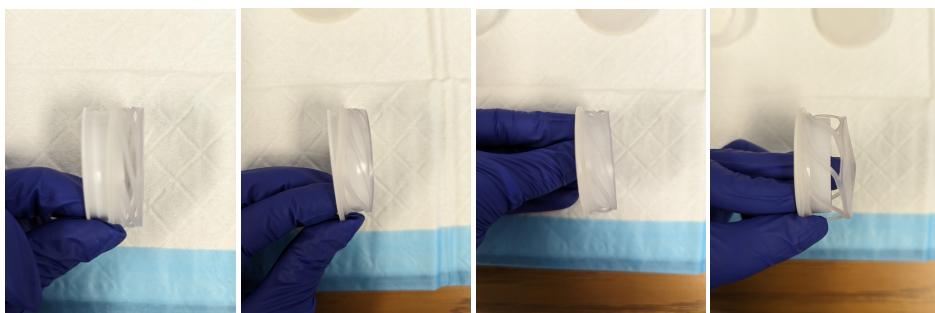
Pink buckets and mask parts are sterile and should NOT be touched without gloves. There are several boxes of latex gloves in different sizes in the lab. Pick whichever size fits you before touching ANY parts.

- Grab a pink bucket and ensure the following components are in the bucket:
  - Clear plastic with two black circles
  - Clear plastic with triangle

- White plastic that narrows
- Semi-transparent circle made of hard plastic
- Two hard plastic white circles
- Two silicone caps

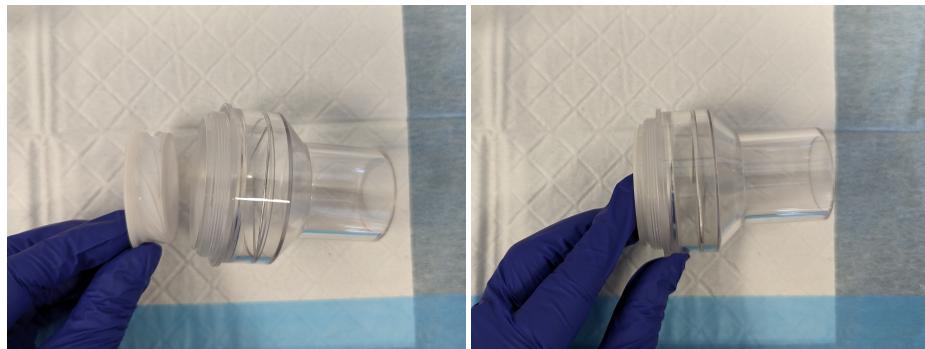


- Take a silicone cap and place on a hard plastic white circle
  - The silicone cap and the hard plastic white circle should face the same direction (the outer rims should be on the same side)
  - Carefully pull the outer rim of the silicone cap over the lip of the hard white plastic circle. Be careful not to tear the silicone, handle these pieces very gently!
  - Rotate the hard white plastic circle to make sure that the outer rim of the silicone cap is pulled over the lip all the way around.
  - You should be able to gently push through the middle of the hard white plastic circle and stretch the top of the silicone cap out from the edge of the hard white plastic circle while the outer rim of the silicone cap stays in place over the lip of the hard white plastic circle.



- Repeat with the other silicone cap and hard plastic white circle. There should be two pairs of silicone cap and hard plastic white circle.
- Take one of the pairs of silicone cap and hard plastic white circle and the clear tapered hard plastic piece.

- Place the silicone cap into the larger end of the clear tapered piece. The outer rim of the silicone cap should sit flush with the rim of the clear tapered piece. The silicone should face the inside of the clear tapered piece.



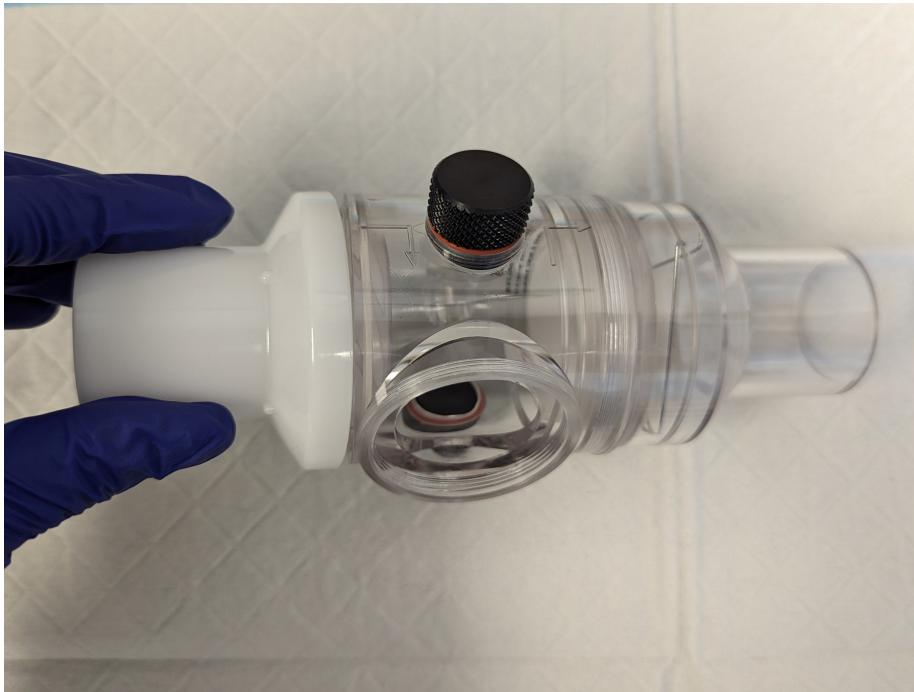
- Take the middle piece of the mask (the clear cylindrical piece with three openings and two black cylinders on it) and rotate it around until you can see the arrows etched into it.
  - The arrows etched into the piece show the direction of air flow through the pieces. The air comes “in” from one opening, into and out of the mask where someone is breathing in and out of one opening, and “out” of one opening into the tubing and finally into the gas analyzer. The silicone pieces allow air to flow in only one direction so that all air the person breathes out is sent into the gas analyzer.
- Find the arrow pointing “out” of the cylinder and rotate it so that the arrow pointing “out” points to your right.
- Take the clear tapered piece with the silicone vent inside it and screw it onto the opening facing the right



- Take the second silicone vent and place it in the left opening of the middle piece. This opening should be directly opposite the opening facing to the right of the middle piece, that now has the clear tapered piece screwed on.
- The silicone vent should have the outer rim flush with the outer rim of the opening, and the silicone should face into the middle piece.



- Take the white tapered plastic piece and screw on to the opening over the silicone vent.



- Take the semi-transparent hard plastic circle and screw onto the remaining opening of the middle piece.



- Select the blue rubber mask of the appropriate size.
  - The size is etched on the “chin” of the mask.
  - See B.3.2 for choosing the correct size.



- Orient the blue mask as if you were going to wear it on your face. The “nose” should be upward with the “chin” downward and the hard clear plastic pieces should face outward. You should only see blue rubber.
- Orient the assembled pieces so that the air flows in the appropriate direction.
  - The “out” arrow direction with the clear tapered end should point towards the station with the gas analyzer, i.e. if the gas analyzer will be on the left then the clear tapered end should point to the left, and if the gas analyzer will be on the right then the clear tapered end

should point to the right. In the HPL, the stationary exercise bike will always have the gas analyzer on the left, therefore have the clear tapered end point to the left.

- The third opening with the semi-transparent circular piece should face the “mouth” of the blue rubber mask.



- Stretch the opening of the blue rubber mask over the semi-transparent circle opening of the assembled pieces.
  - This is difficult and you will have to manhandle the blue rubber mask a bit. It does not stretch super well.
  - Look through the mask and make sure that there is a good seal around the opening. If there is not a good seal, the blue rubber will not make a perfect circle around the opening.



### B.3.4 Heart Rate Monitor

All men leave the room (except family the patient may or may not have brought with them to the lab). Spray water on the part of the monitor that will be in contact with the skin. Have the patient hold the center ‘button’ on the strap to their sternum. Clip the ends of the strap together. Tighten the strap to be snug around the patient’s chest. Push the strap up under the patient’s bra strap and make sure the monitor is in contact with the skin all the way across the chest.

Use the watch to check that the monitor is sending a good signal. Press the middle button to the left of the watch face and wait to see the watch display a heart rate. If the heart rate does not display or shows a number that seems off, try to adjust the monitor to get a better connection. Try to push the monitor up higher on the sternum. This can be difficult on some patients depending on breast tissue, so do your best. Check the signal again.

If the watch is having issues connecting with the heart rate monitor, you can check from the Parvo software. *add steps here*

## B.4 Data Collection

Select ‘Metabolic Testing’.

Enter the following as BOTH the first and last name:

“NIRS\_patientidnumber\_protocoltypeprotocolnumber\_protocoldate”

For example, patient ID 003 participating in the second on/off kinetics on 11/12/2023 would look like: “NIRS\_003\_FP2\_11\_15\_2023”

Enter the participant’s height, weight, and age.

Select the testing protocol: NIRS Lactate Theshold or NIRS Full Protocol.

Enter appropriate testing wattage.

Click ‘OK’ and wait for the software to start the test.

When testing is complete, click ‘end test’.

Save results and print a copy of the report.

Check that the report will be printed with 10-second averages. Click ‘config’ in the bottom left and ensure that the data averaging selected is 10 seconds. Click ‘OK’ and print. The report will be printed on the printer on the bottom shelf of the cart.

## B.5 After Data Collection

### B.5.1 Data Export

The ParvoMedics software only exports metabolic cart data as Excel files. Select the data collection instance you wish to export and save it as an Excel file on the cart. Use a USB drive to transfer the Excel file from the Parvo cart to another computer. Since the Parvo cart is not connected to the internet and never gets Windows or any other software updates, the Excel format is a legacy format. You will have to manually open the Excel file and save it as a modern Excel format.



# **Appendix C**

## **Surveys**

### **C.1 Brief Fatigue Inventory**

The Brief Fatigue Inventory (BFI) is a 9 item survey designed to assess fatigue severity in cancer patients (Mendoza et al., 1999). Each item is measured on a 0-10 numeric scale. The final score is the mean of the 9 measure items and can range from 0-10. A score can be calculated if at least 5 items out of 9 are answered by taking the mean of the completed items.

### **C.2 Godin Leisure-Time Exercise Questionnaire**

The Godin Leisure-Time Exercise Questionnaire (GLTEQ) is a two item instrument developed to asses the leisure-time activity levels in order to classify people into ‘very active’ and ‘sedentary’ categories with the aim of using this instrument to measure activity levels before and after implementation of community health and/or fitness programs in relation to psychosocial variables [@].

### **C.3 Mini-Mental State Examination**

### **C.4 Montreal Cognitive Assessment**

### **C.5 Physical Activity History**

### **C.6 PROMIS Global Health**



## **Appendix D**

# **Instruments**

### **D.1 Ultrasound**

#### **D.1.1 Usage**

Turn on the ultrasound by pressing the power button in the top left of the keyboard.

Locate the vastus lateralis. Find the ASIS (anterior superior iliac spine) by asking the participant to place their hands on their hips and feeling for the bony protrusion at the front of the hip. Measure from the top of the kneecap to the ASIS. Locate the vastus lateralis by moving superior to the kneecap one-third of the distance from the kneecap to the ASIS.

Use the ultrasound gel to cover the wand.

Place the wand over the pre-determined location of the vastus lateralis.

Freeze the screen when you want to measure adipose tissue thickness.

Press the ‘caliper’ button to use calipers on the frozen screen.

Move one end of the caliper to the top of the screen. Move the other end of the caliper to the area of the screen that looks like the boundary between the muscle belly and the edge of the adipose tissue. This will be a silvery line across the screen.

Record the measurement given by the caliper.

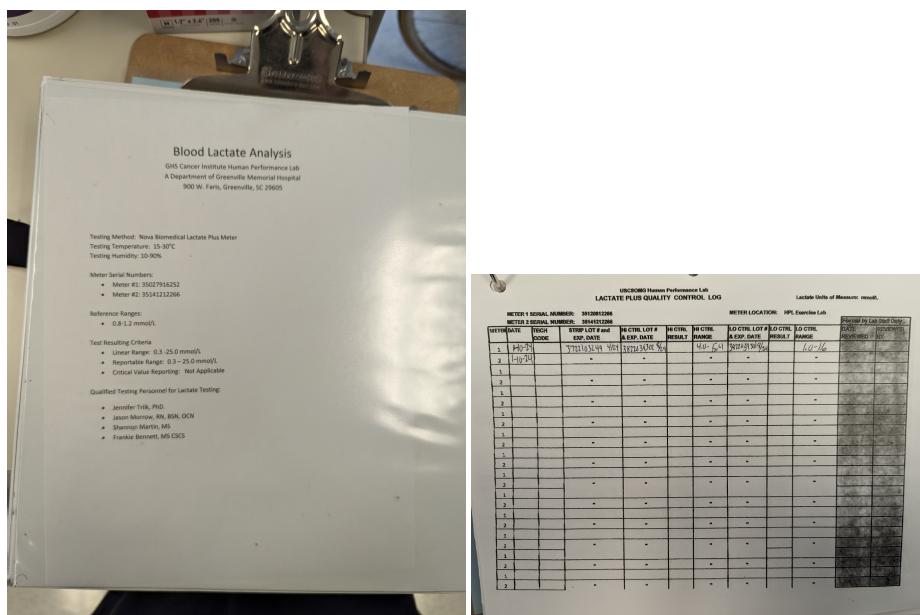
## D.2 Lactate Meter

### D.2.1 Usage

#### D.2.1.1 Quality Control Testing

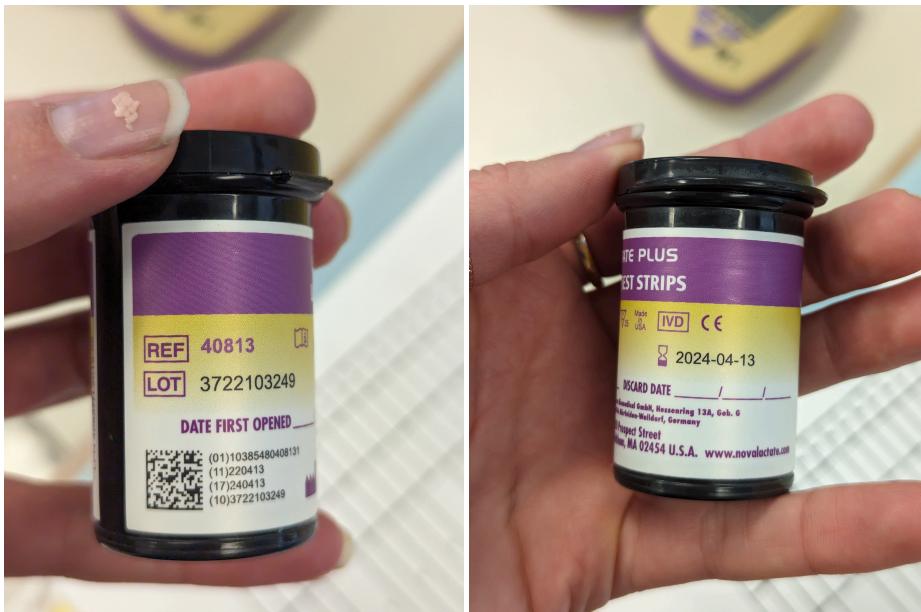
Before using the lactate meters, you must conduct a quality control test.

Get out the ‘Blood Lactate Meter’ binder from Frankie Bennet’s desk. Open the binder to the quality control log.

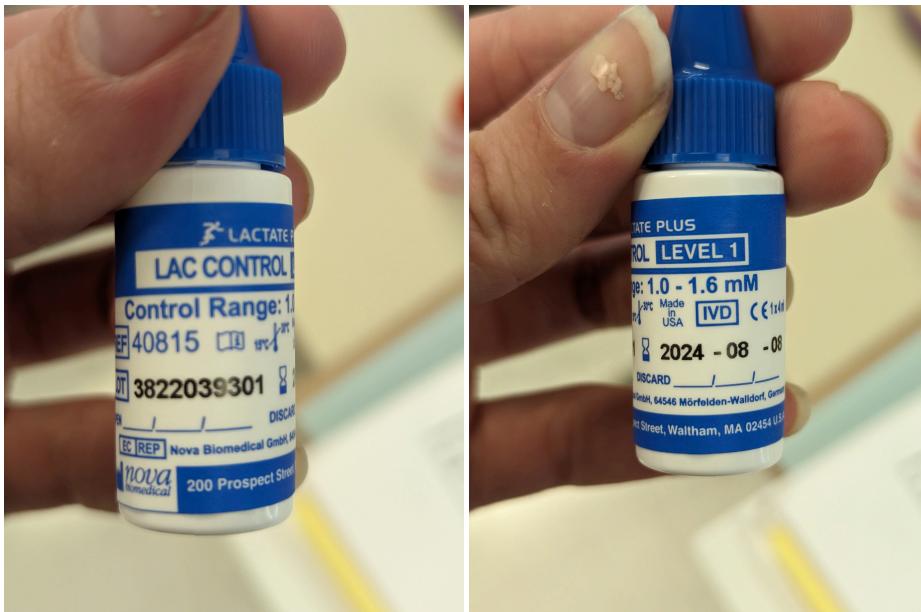


Record the date.

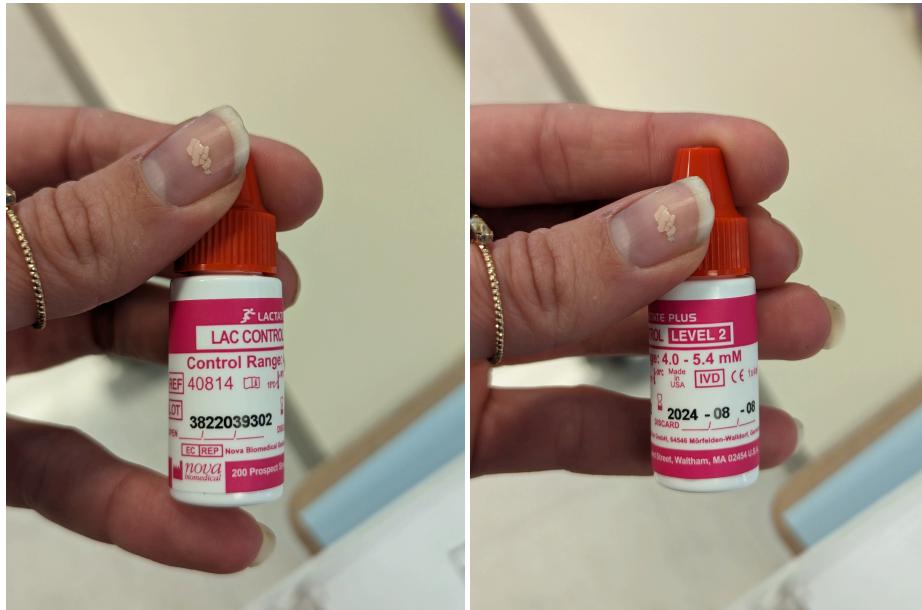
Look on the bottle of lactate strips and find the lot number. Record the lot number in the appropriate space.



Look on the low control bottle (blue dropper). Record the expected range for this bottle. Record the lot number and expiration date.



Look on the high control bottle (red dropper). Record the expected range for this bottle. Record the lot number and expiration date.



Turn on the lactate meter by pressing the center triangle-shaped button. Use the button to click through the screens until you reach the control screen, which has a star in the upper left corner and says 'CRL-1'.

Put a lactate strip in the lactate meter. The white end should stick out from the meter and the metal end should go in the meter.



Use the red control dropper to place a drop on the lactate strip. Wait for the lactate meter to give results (about 15 seconds). Record the results in the binder. Discard the used lactate strip in the trash and do another test with the red control with a new lactate strip.

If you get an error, you probably did not use enough liquid. Make sure the drops are of a size to completely soak the tip of the lactate strip.

Repeat with the blue control dropper.



Make sure results are recorded correctly in the log.

USC SOMG Human Performance Lab LACTATE PLUS QUALITY CONTROL LOG										Lactate Units of Measure: mmol/L		
METER	DATE	TECH CODE	STRIP LOT # and EXP. DATE	HIGH CTRL. LOT # & EXP. DATE	HIGH CTRL. RESULT	HIGH CTRL. RANGE	LOW CTRL. LOT # & EXP. DATE	LOW CTRL. RESULT	LOW CTRL. RANGE	REVIEWED by Lab Staff	DATE REVIEWED	REV'D BY
1	10-24		3722103249 4/24	3872034302 8/24	4.9	4.0-5.1	3872039301 9/24	1.3	1.0-1.6			
2	10-24		-	-	4.3	-	-	1.2	-			
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