



## Study in Parkinson Disease of Exercise Phase 3 Clinical Trial: SPARX3

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BIOSPECIMEN COLLECTION & PROCESSING

# Overview

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1. Specimen uniformity and quality
2. Site Equipment
3. Procedures
  - Kit Contents and Ordering
  - Sample Labeling
  - Sample Collection and Processing
  - Shipping Samples
  - Non-Conformance
4. Contact Information

# Specimen Uniformity and Quality

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GENERAL REMINDERS

# Specimen Standardization and Quality

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Most biomarkers are sensitive to *time* and *temperature*

- Standardization of processing across sites is key
- Specimens must be processed within 2 hours of collection
- Reference the *BioSEND Biomarker Specimen Collection, Processing, and Shipment Manual* as needed
- Do not replace or supplement any kit components without first receiving approval from BioSEND/NINDS

Questions? Email [biosend@iu.edu](mailto:biosend@iu.edu)

# Site Consumables and Equipment

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*Sites will need to supply the following items:*

- Gloves
- Alcohol wipes
- Butterfly needles
- Tourniquet
- Gauze pads
- Bandages
- Microcentrifuge tube rack
- Sharps bin and lid
- Crushed ice
- Pipettes and pipette tips
- Centrifuge capable of maintaining 4°C
- -80°C Freezer
- Dry ice

# Procedures

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MAINTAINING SPECIMEN UNIFORMITY AND QUALITY



# Biospecimen Collection Protocol

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Visit & Sample Type	BL	6M	12M	18M	24M
Serum aliquots, 1.5ml	6	6	6	6	6
Plasma aliquots, 1.5 ml	6	6	6	6	6
Buffy Coat	2	2	2	2	2
Whole Blood, 3ml	2	2	2	2	2

# Kit Contents and Ordering

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- All sites will be sent a Supplemental Kit with their first kit shipment
  - Contains extra blood collection tubes and processing supplies
  - May be used to replace items in study visit kits
- Study Visit Kits should be ordered as soon as visits are planned
  - Contains collection, processing, and shipping supplies specific to each visit
  - Includes barcoded labels
  - The supplies/labels in each study visit kit are intended for that visit only



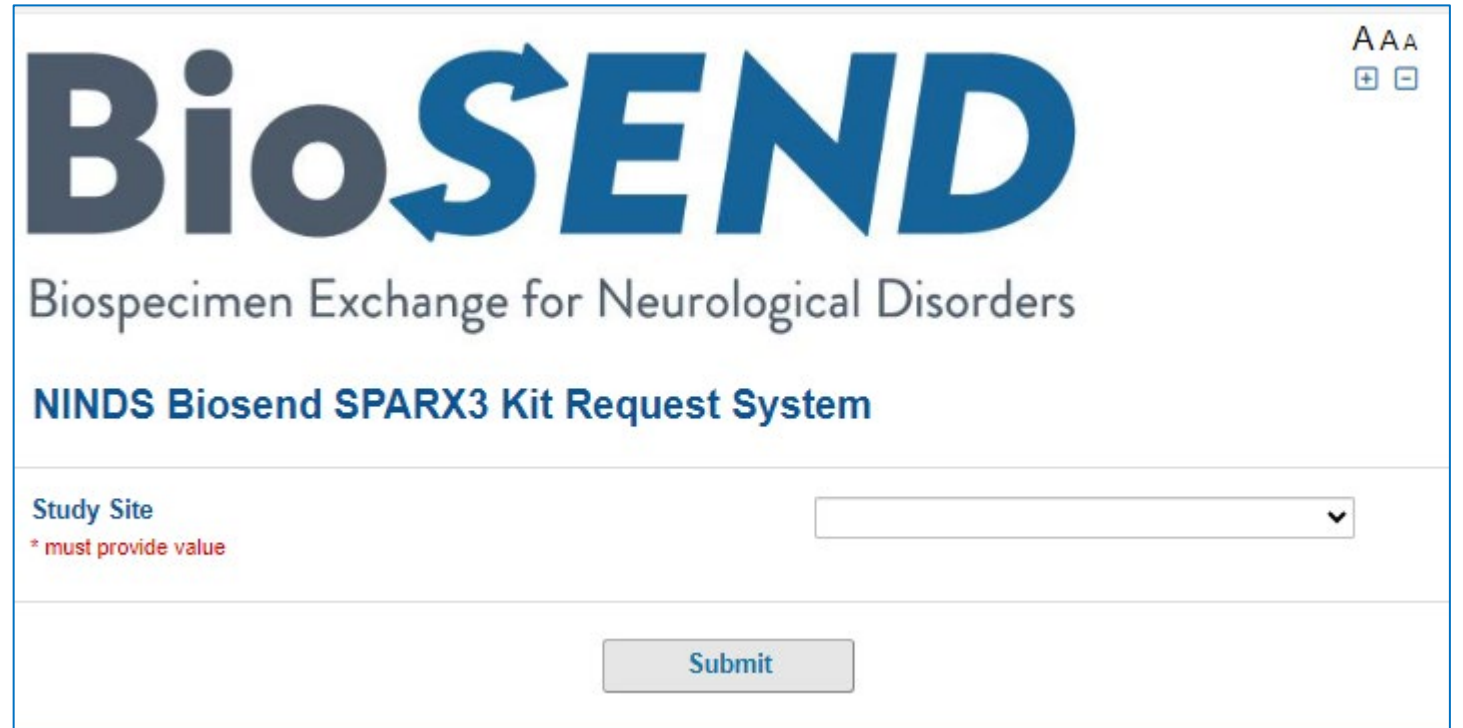
# Kit Contents and Ordering – REDCap Survey

<http://kits.iu.edu/biosend/sparx3/>

Order kits online through the Kit Request Module for:

- Blood Kits
- Supplemental Kit
- Extra Supplies

Please provide as much notice as possible when ordering kits and/or supplies.



The screenshot shows the BioSEND logo at the top, with the text "Biospecimen Exchange for Neurological Disorders" and "NINDS Biosend SPARX3 Kit Request System" below it. In the top right corner, there are accessibility icons labeled "AAA" and a plus-minus sign. Below the header, there is a "Study Site" label followed by a dropdown menu. A red asterisk and the text "\* must provide value" are positioned below the dropdown. At the bottom center, there is a "Submit" button.

# Kit Contents and Ordering: Confirm Site Info

SPARX3 Kit Request Module

<b>Study Site</b> <small>* must provide value</small>	Indiana Univers	Select your site from the drop-down list
<b>Indiana University School of Medicine</b> Carolyn Dunifon Dept. of Medical & Molecular Genetics 351 West 10th Street, TK-318 Indianapolis, IN 46202-3002 (317) 274-5751 cdunifon@iu.edu		Verify contact information and update if needed
<b>Is the contact name above correct?</b> <small>* must provide value</small>	<input type="radio"/> Yes <input checked="" type="radio"/> No	reset
<b>New Contact Name</b> <small>* must provide value</small>	Claire Wegel	
<b>Is the shipping address above correct?</b> <small>* must provide value</small>	<input type="radio"/> Yes <input type="radio"/> No	reset
<b>Is the e-mail address above correct?</b> <small>* must provide value</small>	<input type="radio"/> Yes <input type="radio"/> No	reset
<b>Is the phone number above correct?</b> <small>* must provide value</small>	<input type="radio"/> Yes <input type="radio"/> No	reset

# Kit Contents and Ordering: Kit Types


## SPARX3 Kit Request Module

- Kits are not specific to a subject or time point. After collection, sites will indicate the subject and time point to which BioSEND should link the samples.

<b>Kit Type</b> <b>**Please allow two weeks for shipment**</b> <i>* must provide value</i>	<input checked="" type="radio"/> <b>Blood Collection Kit</b>
	<input type="radio"/> Supplemental Kit
	<input type="radio"/> Extra Supplies
	Please specify in comments if you need kits before the standard two week shipment time.
<b>Blood Collection Kit Quantity</b> <i>* must provide value</i>	<input type="text"/>
<b>Comments</b>	<input type="text"/>
	<a href="#">Expand</a>

# Kit Contents and Ordering: Kit Breakdown

SPARX3 Kit Request Module

Comments	
Each Blood Collection Kit contains:	Expand
<ul style="list-style-type: none"><li>2 - Purple-top EDTA tube (glass, 10 ml)</li><li>2 - Red-top Serum tube (glass, 10 ml)</li><li>2 - Purple-top EDTA tube (plastic, 3ml)</li><li>6 - Bubble-tube sleeve</li><li>6 - Micronic cryovial (2ml) - purple</li><li>6 - Micronic cryovial (2ml) - red</li><li>2 - Micronic cryovial (2ml) - grey</li><li>2 - Disposable transfer pipet (3ml)</li><li>1- 48-slot micronic cryovial tray</li><li>2 - Biohazard bag with absorbent sheet</li><li>1 - Shipping label packet (Dry Ice, Fragile, UN3373)</li><li>1 - Airway bill envelope</li><li>1 - Dry Ice Shipper</li><li>1- Specimen Label Set-- ST labels</li></ul>	 <p>Kit contents of selected kit will appear at the bottom of the page</p>

# Kit Contents and Ordering: Blood Kit



# Collection Volumes

Total blood volumes

Sample Type	Amount
Whole Blood for Plasma and Buffy Coat	20 ml
Whole Blood for Serum	20 ml
Whole Blood for Banking	6 ml

# Kit and Supply Ordering

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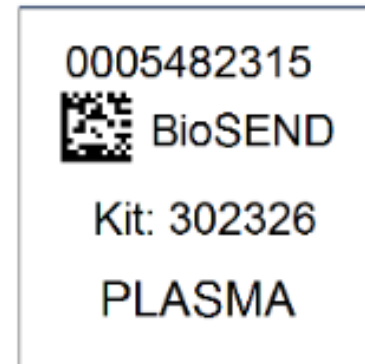
- Click “Submit” to send order to BioSEND; staff will confirm receipt of your order
- Please allow two week turn-around time for kit shipments
- If urgent request needed, please note date needed by in comments and email BioSEND. We cannot guarantee urgent orders, but we will do our best to accommodate.
- BioSEND will send confirmation of shipment and tracking number when supplies ship

# Sample Labeling: Example Labels

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## Labels are provided by Indiana University

- Please check that all samples are properly labelled to ensure correct identification by IU
- If do not have enough labels to complete a visit, please contact IU *immediately*
- Labelling the tubes during processing prevents sample mix-ups



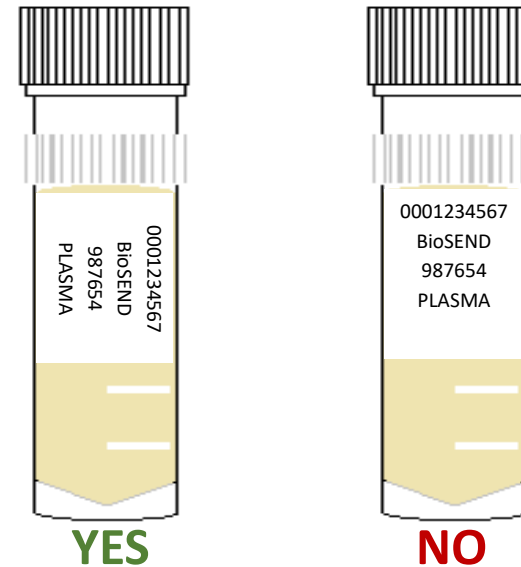


# Sample Labeling: Label Placement

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Please...

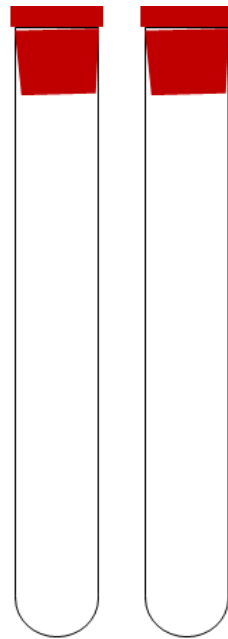
- Label all collection and aliquot tubes before cooling, collecting, processing, or freezing samples
- Label only 1 subject's tubes at a time to avoid mix-ups
- Wrap the label around the tube horizontally - label position is important for all tube types
- Make sure the label is completely adhered by rolling between your fingers



# Sample Collection and Processing

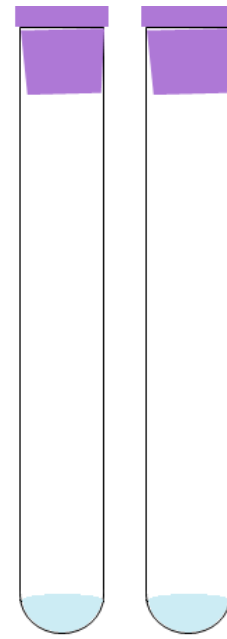
Blood Tube Draw Order

2 x 10ml Serum



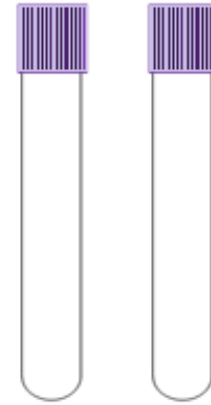
**1**

2 x 10ml EDTA  
for plasma and  
buffy coat



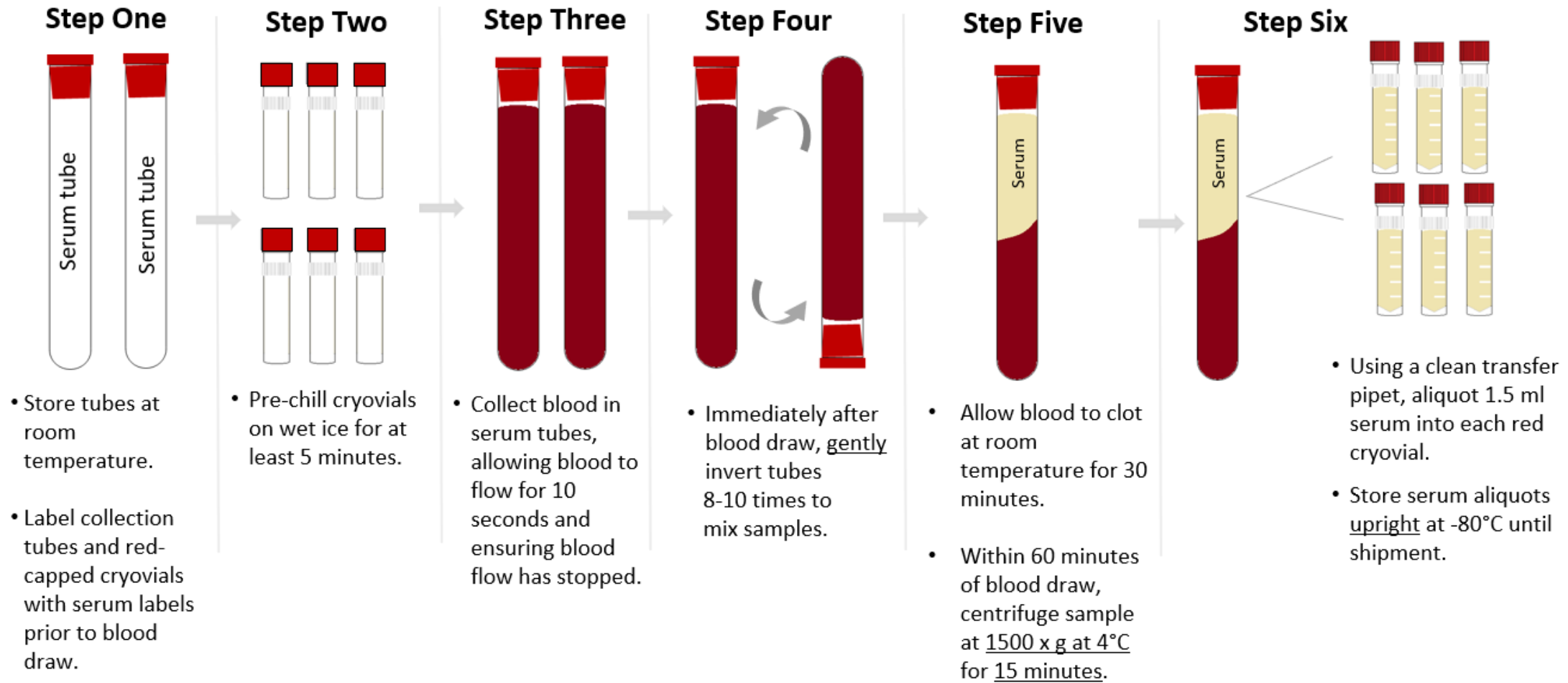
**2**

2 x 3ml EDTA  
for whole  
blood

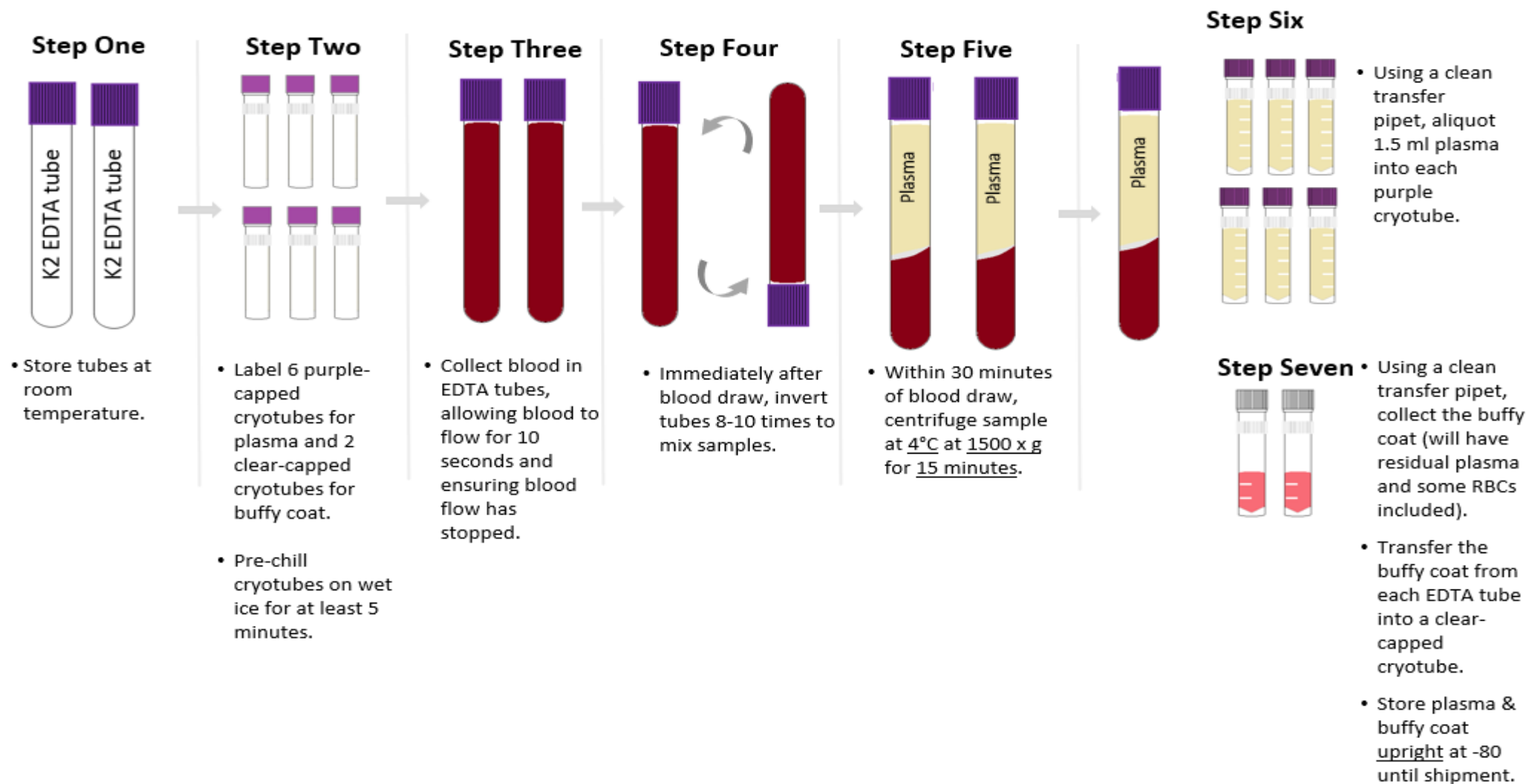


**3**

# Sample Collection and Processing: Serum



# Sample Collection and Processing: Plasma & Buffy Coat



# Sample Collection and Processing: Whole Blood

## Step One



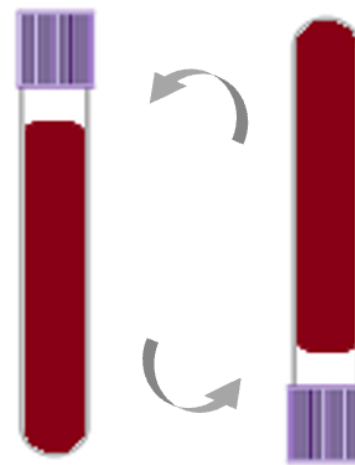
- Store tubes at room temperature.
- Label tubes with preprinted WBLD label prior to blood draw.

## Step Two



- Collect blood into both 3ml EDTA tubes, allowing blood to flow for 10 seconds and ensuring blood flow has stopped.

## Step Three



- Immediately after blood draw, invert tubes 8-10 times to mix samples.

## Step Four



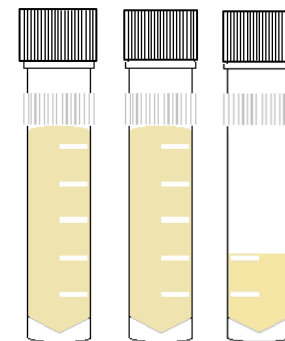
- Transfer to  $-80^{\circ}\text{C}$ . Store upright and keep frozen until shipment to BioSEND.

# Sample Collection and Processing: Aliquots

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Filling biomarker serum and plasma aliquots:

- Fill as many cryovials as possible to 1.5 ml (plasma & serum)
- Over-filled vials may burst in freezer!
- Ship ALL material to IU, even if final vial is less than standard volume



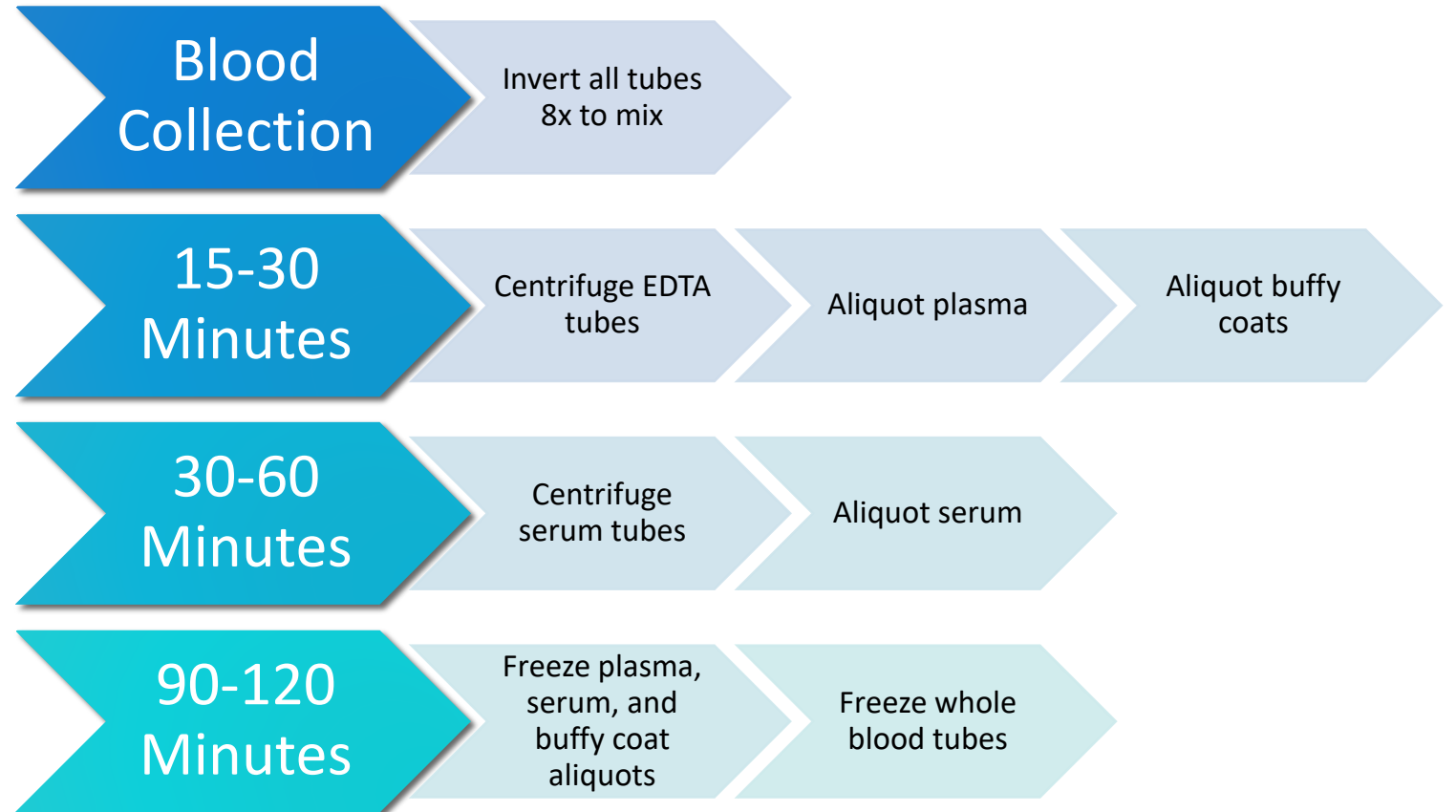
YES



NO

# Sample Collection and Processing: Timeline

Timeline for blood processing



# Sample Collection and Processing: Issue #1

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## Troubleshooting Blood Collection

### Issue #1: Tube with little/no vacuum

- Always check expiration date on the tube before beginning blood draw and discard expired tubes
- Store tubes at “room temperature” – extreme temperature can affect vacuum
- Keep extra vacutainer tubes from supplemental kit nearby during blood draw to replace “bad” tubes
- If this is a frequent occurrence, report tube type and lot number to IU.



# Sample Collection and Processing: Issue #2

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## Troubleshooting Blood Collection

### Issue #2: Hemolyzed serum and/or plasma caused by incorrect collection

Cause: Blood Collection Methods	Corrective Action
Improper venipuncture site	Draw from median cubital, basilic, and cephalic veins from antecubital region of arm
Prolonged tourniquet use	Tourniquet should be released after no more than 1 min, excessive fist clenching should be avoided
Not allowing alcohol to dry on skin before venipuncture	Without touching, allow the venipuncture site to air dry
Use of too large/small bore needle resulting in excess force applied to blood	Avoid using too small/large needle. Needle size dependent on the subject's physical characteristics & amount of blood to be drawn. Most commonly used sizes are 19 – 23.
Pulling/pushing plunger too fast while drawing/transferring blood	Avoid drawing the syringe plunger too forcefully when collecting blood
Ensure all blood collection assemblies are fitted securely, to avoid frothing	

For more information, visit: [http://www.bd.com/vacutainer/pdfs/techtalk/TechTalk\\_Jan2004\\_VS7167.pdf](http://www.bd.com/vacutainer/pdfs/techtalk/TechTalk_Jan2004_VS7167.pdf)

# Sample Collection and Processing: Issue #2 continued

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## Troubleshooting Blood Collection

### Issue #2: Hemolyzed serum and/or plasma caused by incorrect processing

Cause: Sample Processing Methods	Corrective Actions
Vigorous mixing/shaking	Gently invert blood collection tube when mixing additive with specimen, follow guidelines in Biologics Manual regarding number of times to invert each type of tube
Not allowing serum to clot for recommended time	Serum tubes without clot activator should be allowed to clot for 60 min in a vertical position
Exposure to excessive heat or cold	Keep samples at ambient temperature until processing
Prolonged contact of serum/plasma with cells	Do not store uncentrifuged samples beyond recommended time

For more information, visit: [http://www.bd.com/vacutainer/pdfs/techtalk/TechTalk\\_Jan2004\\_VS7167.pdf](http://www.bd.com/vacutainer/pdfs/techtalk/TechTalk_Jan2004_VS7167.pdf)

# Sample Collection and Processing Form

Direct link:

<https://redcap.link/SPARX3SampleForm>

First page captures basic subject and visit information

# BioSEND

## Biospecimen Exchange for Neurological Disorders

Please complete the Specimen Collection and Processing Form, below.

Returning?

AAA



Page 1 of 2

### BioSEND SPARX3 Study

Study Site

Email address of staff member completing this form

*Note: A copy of the completed sample form and the shipping manifest will be sent to this address.*

GUID:

Sex (used for DNA quality control)

Visit

IU Kit Number

6 characters remaining

Next Page >>

Save & Return Later

# Sample Collection and Processing Form

Direct link:

<https://redcap.link/SPARX3SampleForm>


Second page captures processing information

## Blood Collection and Processing

Date of venipuncture blood collection

  Today M-D-Y

Time of venipuncture blood collection

  Now H:M  
Use 24 Hour clock

Date participant last ate

  Today M-D-Y

Time participant last ate

  Now H:M


SERUM (red-top tubes, 10 mL)

Was blood collected and processed for SERUM?

 Yes  
 No

reset

Time of SERUM tube centrifugation

  Now H:M  
Use 24 Hour clock

Duration of SERUM tube centrifugation

  
minutes

Rate of SERUM tube centrifugation

  
x g

Temperature of SERUM tube centrifugation

  
degrees Celsius

Total volume of SERUM collected

  
ml

# Sample Collection and Processing Form

Direct link:

<https://redcap.link/SPARX3SampleForm>

PDF form of responses will be emailed to you. **Print a copy of the Frozen Shipping Manifest and include with shipment.**

## SPARX3 Frozen Shipping Manifest

Please verify/update the information below. When you click the "Submit" button below, a PDF copy of the Frozen Shipping Manifest will be emailed to you for Subject [subj\_id].

Please print a copy of that document and include it in the Kit #[kit\_num] shipping container.

Study Site:

- Boston University (Charles River Campus)
- Cleveland Clinic
- Columbia University Medical Center
- Emory University
- Iowa State University
- Kent State University
- Louisiana State University
- Mayo Clinic, Rochester
- New York University Langone Health
- Northwestern University
- Ohio Health
- Oregon Health & Science University
- Rush University Medical Center
- University of Alabama at Birmingham
- University of Alberta
- University of California, San Francisco
- University of Cincinnati
- University of Colorado, Denver
- University of Florida
- University of Michigan
- University of Minnesota
- University of Pennsylvania
- University of Pittsburgh
- University of Southern California
- University of Texas Medical Branch
- University of Texas San Antonio
- University of Utah
- University of Virginia
- Washington University St. Louis
- Wildred Laurier University of Canada

GUID:

\_\_\_\_\_

Visit:

- BL
- 6M
- 12M
- 18M
- 24M

IU Kit Number:

\_\_\_\_\_

Date of blood collection:

\_\_\_\_\_

# Sample Collection and Processing Form

Submission of the Shipping Manifest portion of the form serves as shipment notification to BioSEND

- Must be completed prior to shipment
- If samples are shipped and not received, BioSEND will follow-up with courier. It is recommended that sites also track shipment to ensure safe delivery

# Shipping Frozen Samples: Tips

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## Packing and Shipping Frozen Samples

- SPARX3 samples all ship frozen on dry ice
- Frozen samples should be shipped **only** Monday through Wednesday
- Always fill carton to **top** with dry ice
- Do not pack shipment until the day of pickup



Class 9 Dry Ice Label should not be covered with other stickers and must be completed, or UPS will reject/return your package!

# Shipping Samples

Packing and Shipping Frozen Samples

Shipper's Declaration not Required.

Dry Ice amount must be in kilograms.

Note: 2 lbs. = 1 kg.

Airwaybills / airbills must have the following:

- Dry Ice; 9; UN 1845
- $\frac{\text{Number}}{\text{(Number pkgs)}} \times \frac{\text{wt}}{\text{(wt)}} \text{ Kg}$

Net weight of dry ice in kg

Dry Ice  kg.

Your name & address

Shipper's Name and Address

UN 1845

Consignee Name and Address

IU information and address

19

06426 1/01 RRD



# Shipping Samples: Frozen

Do not ship more than 4 biohazard bags in a single shipper (equivalent to two subject-visits).

- Allows room for dry ice to keep samples frozen in transit
- Minimizes loss in the rare but unfortunate event of courier issues



# Shipping Samples – UPS: <https://kits.iu.edu/UPS>

The screenshot shows the shipping interface for Indiana University School of Medicine. The header includes the IU logo and the text "INDIANA UNIVERSITY SCHOOL OF MEDICINE". Below the header, the text "and Affiliated Biorepositories" is displayed. The interface is divided into two main sections: "Ship From" and "Shipment Information".

**Ship From Section:**

- Search for address: A search bar with a magnifying glass icon and a "Clear" button.
- Code: Input field.
- Company: Icahn School of Medicine - Mt. Sinai
- Contact: Kenny Persaud
- Address 1: 1425 Madison (Icahn Building)
- Address 2: Icahn L4 39
- Address 3: Input field.
- City: New York
- State/Province: NY
- Postal Code: 10029
- Country/Territory: United States

**Shipment Information Section:**

- Study Group: SSBC (dropdown menu)
- Weight: 20 (input field) LB (dropdown menu)
- Dry Ice Weight: 10 (input field) LB (dropdown menu)
- Description of Return: Biologic Specimens
- Pickup Request: Button

**Annotations:**

- "Search for address": Points to the search bar.
- "Choose Study": Points to the Study Group dropdown.
- "Enter weight": Points to the Weight and Dry Ice Weight input fields.
- "Schedule Pickup": Points to the Pickup Request button.
- "Click 'Ship'": Points to the Ship button at the bottom right.

**Buttons:**

- Clear
- Pickup Request
- Reset
- Ship

# Shipping Samples via UPS

IU UPS ShipExec Shipping Portal

- Print out UPS air waybill
- Ensure all elements (barcode, return address, etc.) printed clearly
- Fold and insert UPS air waybill into clear plastic sleeve on package

JOHN SMITH  
INDIANA UNIVERSITY  
410 WEST 10TH STREET  
INDIANAPOLIS IN 46202

2 LBS

1 OF 1

RS

SHIP TO:

SCHOOL OF MEDICINE  
317-278-2694  
INDIANA UNIVERSITY  
TK 217  
351 W 10TH ST  
INDIANAPOLIS IN 46202



IN 461 9-01



UPS NEXT DAY AIR

1

TRACKING #: 1Z 976 R8W 84 3985 8595



BILLING: P/P  
DESC: Biological Specimens  
RETURN SERVICE

Reference No. 1: 4087277

XOL 20.03.09 NV45 83.0A 12/2019



# Shipping Samples: Closures

Date	Holiday
January 1	New Year's Day
3 <sup>rd</sup> Monday in January	Martin Luther King, Jr Day
4 <sup>th</sup> Monday in May	Memorial Day
June 19	Juneteenth (observed)
July 4	Independence Day (observed)
1 <sup>st</sup> Monday in September	Labor Day
4 <sup>th</sup> Thursday in November	Thanksgiving
4 <sup>th</sup> Friday in November	Friday after Thanksgiving
December 25	Christmas

Please also consider weather when shipping. UPS will post service updates on their webpage. Reach out to BioSEND if you are unsure if it is safe to ship.

# Non-Conformance

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Non-conformance to standard procedures may reduce the utility of the biospecimens:

- Not processing serum/plasma within 2 hours of collection allows for breakdown of certain proteins and small molecules
- Over/under centrifuging changes plasma, serum composition



# Non-Conformance Reporting con't

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Most common non-conformance issues:

- Shipment notification not sent
- Samples shipped for weekend/holiday delivery
- Sample form incomplete/inaccurate
- Low volume
- Unlabeled or mislabeled tube(s)
- Sample hemolysis



# Non-Conformance and Inventory Reporting

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Most common non-conformance issues:

- BioSEND will notify sites directly of any issues upon receipt
- BioSEND will email sites a monthly inventory report of all samples received from that site to date
- ***If you are experiencing issues, please reach out to us for help! It is much easier to prevent an issue before sample collection & shipment than trying to fix it after the fact***

# Contacts

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## Indiana University

General Questions/Shipment Notifications:

[biosend@iu.edu](mailto:biosend@iu.edu)

317-278-6158

Request kits:

<http://kits.iu.edu/biosend/sparx3/>