Janssen Oncology

PHARMACEUTICAL COMPANIES OF Johnson & Johnson

Receipt and Storage Manual

Version 3

April 2022

Requirements for US Commercial CARVYKTI™

Receipt and Storage Manual

This manual provides requirements to support supply chain standards within the CARVYTKI™ commercial manufacturing process related to receipt and storage of the manufactured product.*

The requirements in this manual are intended to be incorporated into each site's operating procedures.

Contents

***Please note that the term receipt as used in this manual is specific to the requirements set forth in this manual and shall NOT apply to any reference or use of the term receipt in any document or agreement outside of this manual that is related to the CARVYTKI™.**

Receipt and Storage Manual

Abbreviations

AWB	Airway Bill
CoC	Chain of Custody
CoI	Chain of Identity
DIN/SEC-DIS	Donation Identification Number/Single European Code
LN2	Liquid Nitrogen
WBC	White Blood Cell
PPE	Personal Protective Equipment
TOR	Temperature Out-of-range

Definition of Terms

CoC/CoI Portal	The Chain of Custody (CoC)/ Chain of Identity (CoI) management portal used by the apheresis site staff for labelling and tracking a patient's order for CARVYTKI™.

Receipt and Handling of CARVYTKI™

Shipment Types

There are two types of shipments related to the receipt and handling of CARVYTKI™:

1. CARVYTKI™ from the manufacturing facility to the site.
2. Empty LN2 shipper back to Janssen's courier for recharge or reuse.

If a situation falls outside of these scenarios, contact your designated Janssen representative for further instructions.

Shipment of CARVYTKI™ From the Manufacturing Facility to the Site

1. **Shipment initiation:** Janssen will initiate pick-up of the product by its designated authorized courier (once it is released from manufacturing quality).
2. **Shipment confirmation:** The courier will then issue an automated shipment confirmation to the site via email.
    - The confirmation will reference the patient number for shipment tracking.
    - The confirmation will also reference the AWB for the shipment.
    - A print-out of the same AWB is provided in the shipper kit pouch inside the shipper.
3. **Shipment notification:** Before delivery, the courier will send one notification to the site.
    - This notification will be received on the same day, within one hour of the delivery, based on the shipment being within a designated range. For example, within a 15-mile (25-kilometer) radius of the site.
    - If there are any delays in shipment, the courier will email or directly call the designated site contact.
4. **Receipt of shipper:** The product will arrive at the site inside the shipper between 9 to 11 a.m. local time, or as designated by the site.
5. **Pick-up of empty shipper:** Janssen will coordinate with the courier to pick-up the shipper container from the Site for return to Janssen the afternoon of receipt at 2 p.m. local time, or as designated by the site.

If the empty shipper will not be ready for pick-up at 2 p.m. local time or the agreed upon time, contact your designated Janssen representative.

Unpacking the Shipment

Before unpacking the shipper, make sure the following items are readily available as they are not provided by Janssen:

- Cryogloves
- Safety glasses
- Wire cutter or heavy-duty scissors

Follow site requirements for PPE when handling any cryopreserved product and equipment

Confirm the following materials are included inside the outer shipper case:

- Reusable LN2 shipper (DV-10 LN2 shipper).
- Labeled consignee kit pouch, which will be needed when the empty shipper is picked-up for return to the courier. The consignee kit pouch is inside the outer shipper case and includes:
  - 1 zip tie for use on the outer shipper case
  - 1 AWB for return shipment
- Cassette rack with CARVYTKI™ inside the LN2 shipper. This will be secured with a red tamper evident seal and wire.

Important—Contact your designated Janssen representative if:

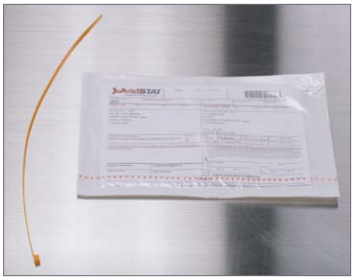
- The shipping container case is not secured or not intact.
- The zip tie is missing from the outer case.
- The patient identifier listed on the AWB does not match the intended patient.
- The LN2 shipper's EVO-IS ID (last 4 digits) listed on the AWB does not match the LN2 shipper.
- Any contents are missing from the consignee kit pouch.

Receiving the LN2 Shipper

Using your own heavy-duty scissors or wire cutters, cut and discard the zip tie securing the zippers. These heavy-duty scissors or wire cutters are **not** included in the shipment.

Unzip and lift open the shipper case lid.

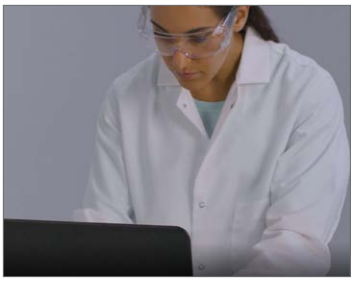
Inside the outer case is another pouch. Verify that it contains the packing insert and the consignee kit pouch.



The consignee kit pouch includes materials for returning the shipper: one zip tie for the outer shipper case and the air waybill to return the empty shipper.

Remove the packing insert and confirm with the CoI record in the CoC/CoI Portal. Scan or enter the packing insert to confirm the product is correctly associated with the CoI record in the CoC/CoI Portal.

You will also be prompted to answer additional questions regarding checks of shipment integrity. Include comments as necessary.



In the CoC/CoI Portal, confirm the EVO-IS ID number (last 4 digits) on the AWB matches the EVO-IS ID number on the LN2 shipper lid.

Confirm the Tamper Seal number on the LN2 shipper lid matches the number on the AWB.

Press and release the light indicator on the DV-Ten lid and wait five seconds.

A steady white light indicates the temperature is within range and you can continue.

If the light indicator shows a flashing white light, a temperature out-of-range, or TOR, has occurred. Indicate this when responding to CoC or CoI receipt questions in the CoC/CoI Portal.

If temperature out of range is noted, complete the TOR process requirements after the product is removed from the shipper and email it to DL-JanssenCAR-T_Vein-to-Vein_US@its.jnj.com

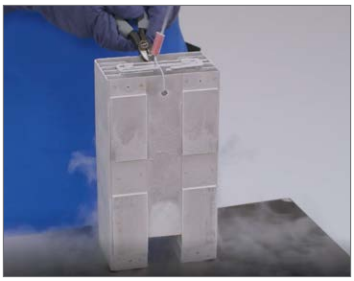
If there was no temperature excursion, continue preparing to remove the product from the shipper.

Unpacking CARVYTKI™

It is recommended to have two personnel work together during the unpacking process. Make sure all necessary PPE is on or in place before unpacking and that you are following your local PPE processes and procedures before unpacking. Additionally, the product must be placed in LN2 storage immediately following unpacking from the shipper and verification of CoC/CoI.

Make sure all necessary PPE is on or in place before unpacking. Also make sure the Drug Product Receipt Checklist for Site form is available for reference and completion.

Prepare these as quickly as possible. It is critical to pack the cassette into the LN2 freezer quickly to avoid thawing of the product.



Use heavy duty scissors or wire cutters, not included in the shipment, cut the tamper evident seal on the lid. Remove the lid and lift the cassette rack out.

Cut and discard the tamper evident seal on the cassette rack and remove the cryogenic containment pouch containing CARVYTKI™ from the rack slots.

Cut open the pouch and remove the cassette.

Before placing the cassette into local LN2 storage, confirm the patient number on the cassette label matches the patient number on the AWB, and the patient number in the Coc/CoI Portal.

Place the cassette with CARVYTKI™ into the LN2 storage and do **not** separate the cassette from the product until preparing for infusion.

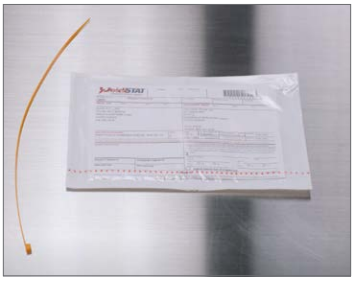
Remove the shipper lid and then place the empty cassette rack back into the shipper. Finally, close the lid.

Complete the Drug Product Receipt Checklist within the CoC/CoI Portal.

Please consider local requirements when the product is received. These may require the quarantine of product until local release steps are complete. This does not apply to the United States.

Returning the LN2 Shipper Container

After completion of the entire receipt process, prepare the empty shipper for shipment:



Gather the unused zip tie and the new AWB from the consignee kit pouch for the shipping container.

Secure the outer case lid with the zip tie and then secure the outer case with buckles.

Place the AWB into the clear packing envelope for courier pick-up. The LN2 shipper is now ready for pick-up.

Janssen will coordinate with the courier to pick up the empty LN2 shipper from the site at the agreed upon pick-up time. Contact your designated Janssen representative immediately if there are any constraints for getting this shipment ready for the agreed upon pick-up time.

Storage of CARVYTKI™

Storing CARVYTKI™

The product must be stored according to the conditions on the label, in a secure location with limited access. During storage, the product must not be separated from the cassette. The packaging is designed to protect the product from breakage and damage and should not be separated.

CARVYTKI™ must be kept frozen at less than or equal to -120°C, vapor phase of LN2.

What are storage temperature monitoring requirements?

1. Temperature conditions during on-site storage of CARVYTKI™ must be monitored and recorded (e.g., temperature logs or data, charts or graphs from temperature monitoring equipment or devices).
2. The use of a temperature monitoring device is required for the storage of all products. All documentation and data of the storage temperature must be retained in accordance with your local records retention policy, and applicable legal and regulatory requirements.
3. Temperature monitoring equipment must be calibrated in accordance with the manufacturer's recommendations.
    - When the temperature monitoring device is equipped with audible and visible alarms to attract immediate attention in the event of a TOR occurrence, the proper functioning of these alarms must also be periodically tested and documented.
4. The output of the temperature monitoring device must be verified and recorded on a temperature log or temperature alarm log daily, during site working days.
    - It is not necessary to check the temperature monitoring equipment on weekends and holidays. If a TOR event occurs during the weekend or holiday, Janssen should be notified immediately.
5. Temperature logs must identify or link to the temperature monitoring device and, if applicable, the LN2 storage system.
    - Links can be through the serial number or other unique identifier.

6. CARVYTKI™ must be kept frozen at less than or equal to -120°C, vapor phase of LN2 Temperature monitoring device with audible and/or visible alarms:
    - The audible and/or visible alarm will alert the cell processing laboratory staff of a TOR situation and the procedure for reporting a TOR must be followed.
    - Even when the system has not alarmed, the temperature records generated by the system should be periodically reviewed to verify all temperature measurements were within range.

Temperature Out-Of- Range (TOR) Events

What To Do

Immediately upon discovery of a TOR reading(s), via the white light on the LN2 shipper lid, take the following steps:

Quarantine the affected product and affiliated supplies. It is important not to use supplies that have experienced a temperature excursion (See *Quarantining* sub-section).

1. Remove the product from the cryoshipper and place the product in the cryofreezer following the instructions provided in the *Packaging and Shipment of White Blood Cell Apheresis Collection Manual.*
2. Remove the product from the LN2 Shipper and place the product in the LN2 freezer.
3. Immediately contact your designated Janssen representative of the TOR occurrence and discuss the potential impact. Provide them with the last 4 digits of the EVO-IS # and patient number.
4. Complete the Temperature Out-Of-Range (TOR) Report Form Immediately. Instructions are provided on the form for completion and submission to Janssen.
5. Janssen will complete the bottom of the TOR Report Form and indicate whether the product is acceptable for use or not.

   For an LN2 shipper TOR event, there is no need to attach the temperature reports to the TOR Report Form. The Janssen Vein-to-Vein Team will extract the data from the EVO-IS shipper readout.
6. For Storage TORs, which may occur while the product is in storage on site, contact Janssen at 1-800-526-7736 if in the United States. For all other regions, please contact your local Janssen representative.

Quarantining

If a concern is found by the site upon receipt and it is necessary to quarantine the product (e.g., damaged, TOR, product quality complaint), please follow the steps below:

1. Immediately contact your designated Janssen representative.
2. Physically separate the product being quarantined. Follow local quarantine procedures for maintaining product segregation.
3. While in quarantine, the product must be stored and handled according to the manufacturing requirements (e.g., LN2 freezer) and procedures.

   This will prevent further deterioration or damage to the product while the viability of the product is being assessed.
4. Once quarantined, the product must remain separated until further notification is received from Janssen.
5. **If the product was quarantined due to a TOR,** the completed TOR Form will indicate if the product is acceptable for use or not.
6. **If the product has been quarantined due to product quality complaint,** provide the details and a photograph of the product and/or packaging to the appropriate Janssen representative.
7. After an assessment, Janssen will review outcome and next steps.

   If indicated that the product is acceptable for use, follow your local procedures to moving product into available inventory.

   If indicated that the product is not acceptable:

   - Identify the product as 'ready to destroy'
   - Contact your Janssen representative to discuss the Credit and Replacement Policy.

Damaged or Lost Product

If the product is damaged while stored at the site, follow the instructions for Quarantining. Contact your designated Janssen representative to review instructions for next steps.

Loss of product is considered a critical situation by regulatory authorities. If the product is lost while stored at the site, as soon as loss is confirmed immediately:

- Contact your designated Janssen representative within 24 hours.
- Conduct an investigation as per your local procedures.
- In the United States, Contact Janssen at 1-800-526-7736. All other regions, please contact your local Janssen representative.

Receipt and Storage Manual

Version 3

For questions related to the information in this manual, contact your designated Janssen representative.