

Preparing for an Autologous Stem Cell Transplant

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Disclosures

- The Hospital at the University of Pennsylvania, as our full-time employer, is not financially supporting this presentation
- We have no financial or pharmaceutical contracts to disclose



Agenda

- Understanding Autologous Stem Cell Transplantation (ASCT)
- Indications for ASCT
 - Transplant Eligibility
 - Disease specific criteria
 - Patient specific criteria
 - Pre-testing qualifications
- Preparations for Autologous Stem Cell Mobilization
 - Mobilization regimens
 - Mobilization strategies
- Preparations for Autologous Stem Cell Transplantation
 - High-dose chemotherapy transplant regimens & toxicities
 - Post-transplant follow up care



Understanding Autologous Stem Cell Transplantation (ASCT)

What is ASCT?

- An Autologous Stem Cell Transplant (ASCT) is:
 - the process of removing stem cells from a patient
 - administering a high dose chemotherapy preparative regimen
 - reinfusing the patient's own stem cells back into them self as a rescue therapy
- ASCT is most commonly used in patients with:
 - Multiple Myeloma
 - Relapsed Non Hodgkin's Lymphoma



Disease Indications for Autologous Stem Cell Transplant

Hematologic Malignancies

- Multiple Myeloma
- Amyloidosis
- Lymphoma

Solid Tumors

• Germ Cell Tumor (Chemo-sensitive relapse)

Auto-immune Disease

- Scleroderma
- Multiple Sclerosis



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*Graph from CIBMTR

Patient Selection Criteria For ASCT

In order to determine if a patient is a good candidate for ASCT, the patient's healthcare team will consider the following:

Performance Status

 measure of how well a person is able to carry on ordinary daily activities while living with cancer, and provides an estimate of what treatments a person may tolerate.

Age

Comorbidities

- Obesity
- Diabetes
- Depression
- Organ Function
- Disease Status
- Overall Prognosis

Renal:

- dosing modifications made for renal impairment
- Cardiac:
 - Ejection fraction > 50%
- Lung:
 - Pulmonary Function Test DLCO is <u>></u>50%
- Assessing viral labs:
 - HIV, HSV, HEP C, HEP B antibodies, HEP B antigen, HEP B core, Varicella, Cytomegalovirus

Once we have deemed the patient eligible for auto transplant, we can now proceed onto the next step of the process, which is preparing for autologous stem cell mobilization

DLCO - Diffuse capacity of the lungs for carbon monoxide



Preparation for Autologous Stem Cell Mobilization & Transplant



autologus+transplant+process.png (927×641) (bp.blogspot.com)

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Preparation for Autologous Stem Cell Mobilization

• What is stem cell mobilization?

 Mobilization is a process by which stem cells are released from the bone marrow into the peripheral blood by means of stimulatory hematopoietic growth factors.

What is the goal of stem cell mobilization?

 To move an adequate amount of stem cells from the bone marrow into the peripheral blood in order for the patient to successfully harvest the cells by means of apheresis in order to proceed onto transplant.

Beth Faimain- BMTCN Certification review manual p.22





Medications Used for Mobilization

Medication	Description	Intended Use
Chemotherapy Cytoxan RICE R-DHAC 	Chemotherapy serves as a stimulus for mobilizing hematopoetic progenitor cells to the peripheral blood	Chemotherapy is used in combination with G-CSF to rapidly increase the amount of stem cells available for harvest
G-CSF Zarxio Neupogen Nivestym Granix 	G-CSF (Filgrastim) is a glycoprotein that stimulates production of hematopoietic cells by binding to certain cell surface receptors	G-CSF is commonly used in conjunction with chemotherapy or plerixafor for stem cell mobilization
Plerixafor • Mozobil	Plerixafor is chemokine antagonist used to induce mobilization of stem cells by interfering with the chemokine stromal-derived factor-1 and it's receptor, CXCR4.	When combined with G-CSF, it has proven to increase the number of CD34+ stem cells collected each day

G-CSF: Granulocyte-colony-stimulating factor

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Advantages & Limitations of Mobilization Strategies

Regimen	Advantages	Limitations
G-CSF + Chemo (Chemo mobilization)	 Possible "antitumor" effects High cell yields 	 Toxicities: Chemo related: Nausea, vomiting, headache, neutropenia, electrolyte deficiencies G-CSF related: bone pain, headache, fatigue, muscle aches Requires inpatient hospital stay
G-CSF + Plerixafor	Low toxicityLow risk of failureHigh cell yields	 Cost Additional Plerixafor- specific toxicities which can include severe nausea, vomiting, & diarrhea

Administration Guidelines

Regimen	Administration/Dose
Chemotherapy + G-CSF (Chemo mobilization)	High dose G-CSF is usually initiated 24 - 72 hours post mobilization-related chemotherapy (dosing is 10 mcg/kg SC) Injected daily until stem cell collection is complete (Approx.10-14 days of administration)
G-CSF+ Plerixafor	G-CSF is started 4 days prior to stem cell collection Plerixafor is started the night before collection (day 4 of G-CSF) until collection is complete (0.24 mg/kg SC or 0.16 mg/kg SC in renal-impaired patients)



Apheresis Process

Apheresis means "to separate"

Apheresis is the process of which blood of a person is passed through an apparatus that separates out one particular blood component (specifically stem cells) and returns the remainder of the blood back into their body



Apheresis/Collection Process (Cont.)



Apheresis/Collection Process (Cont.)

Apheresis Evaluation Day:

- Takes places after mobilization injections are initiated
- Evaluates basic blood work (CBC & CMP)
- Consents obtained
- Peripheral blood CD34+ sample obtained
- Orientation to the Apheresis Unit
- Home medications reviewed

Apheresis Collection Days:

- Collections approximately 3-5 hours/day
- Blood work assessed daily
- Patient monitored during collection for side effects
- Integrity of apheresis catheter maintained
- Patient receives a call each evening with collection results/further instructions

Importance Of Peripheral Blood CD34

- CD34 is a surface antigen expressed on stem cells
- The peripheral blood (PB) CD34 level measures the amount of circulating stem cells present in the bloodstream
- The PB CD34 level is drawn after:
 - starting mobilization therapies
 - on the day prior to a patient's first collection



*CD34 Cell Level drawn prior to first collection the following day

Mobilized Peripheral Blood: Maximize Your CD34+ Yield | AllCells®



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Importance Of Peripheral Blood CD34



Faramarz Naeim MD, ... Ryan T. Phan PhD, in Atlas of Hematopathology (Second Edition), 2018

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Goal:

- Lymphoma: to eradicate cancer (curative intent)
- Multiple Myeloma: to serve as a therapeutic treatment in the setting of a non-curative disease to provide for significant remission, extending survival.
- High dose therapy/preparative regimen is chosen based on:
 - Disease
 - Prior chemotherapy exposure
 - Chemo-sensitivity

Common Regimens:

- High Dose Melphalan Multiple Myeloma
- BCV, BEAM Lymphoma
- Carboplatin/VP16 Germ Cell

Typical Myeloablative Treatment Plan



🔀 Penn Medicine

Toxicities During ASCT

- Nausea
- Vomiting
- Diarrhea
- Mucositis
- Electrolyte Imbalances
- Pancytopenias (transfusion requirements)
- Alopecia
- Fatigue
- Infection
- Failure to engraft



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Toxicities Associated with High Dose Chemotherapy and Autologous Stem Cell Transplantation in Older Patients with Non-Hodgkin Lymphoma - Biology of Blood and Marrow Transplantation (tctjournal.org)



Discharge/Follow Up Care Post ASCT

Discharge considerations

- Once engraftment occurs ANC >1000 x 2 days → discharged home
- Bi-weekly bloodwork to ensure count recovery
- Weekly follow-up visits
- Home nursing care if warranted
- Prophylactic medications
 - Antivirals- Acyclovir
 - Antifungals- Fluconazole
 - Antibiotics- broad spectrum
 - PCP prophylaxis-Bactrim

• Day 100

- Imaging, restaging, repeat bone marrow biopsy, biochemical response
- Resume dose-reduced maintenance therapy



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Conclusions

- There are several variations to successfully mobilize autologous hematopoietic stem cells
- Type of mobilization does not necessarily influence transplant outcomes
- Both chemotherapy-based and G-CSF/Plerixafor mobilization can show similar rates of success
- Several factors, including cost and toxicity, may influence the decision of which way the oncology team chooses to mobilize a patient's stem cells
- Future studies are still needed for maintaining efficacy, reducing cost and toxicity, and improving transplant outcomes

Thank You For Participating

Please join us at our Adult Nursing Session Q & A



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