

Transfusion Guidelines

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BLOOD CENTER

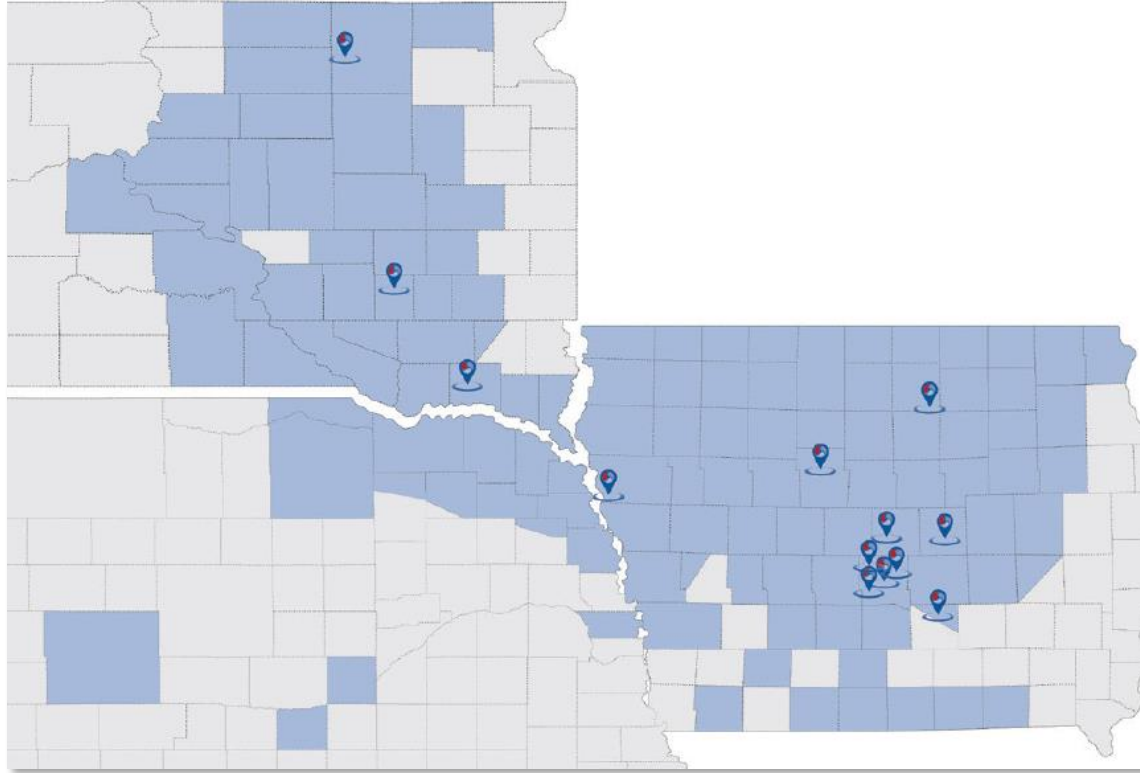
YOUR BLOOD. YOUR HOSPITAL. YOUR NEIGHBORS.

Topics

- Current transfusion guidelines
 - Red blood cells
 - Plasma
 - Platelets
 - Cryoprecipitate
- Blood product modifications
- Platelet refractoriness
- Massive transfusions



What is LifeServe Blood Center?



- Over 136,000 registered donors
 - Over 157,000 blood products per year
- Distribute blood products to 158 hospitals



What is LifeServe Blood Center?



- Freeze and store blood from donors with rare blood types for up to 10 years
- Perform testing for blood banks
 - Crossmatching
 - Antibody ID



Where Do Blood Products Come From?

Two methods to collect from donors

1. Whole blood donations
2. Apheresis



Whole Blood Donation



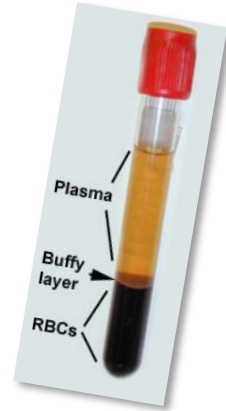
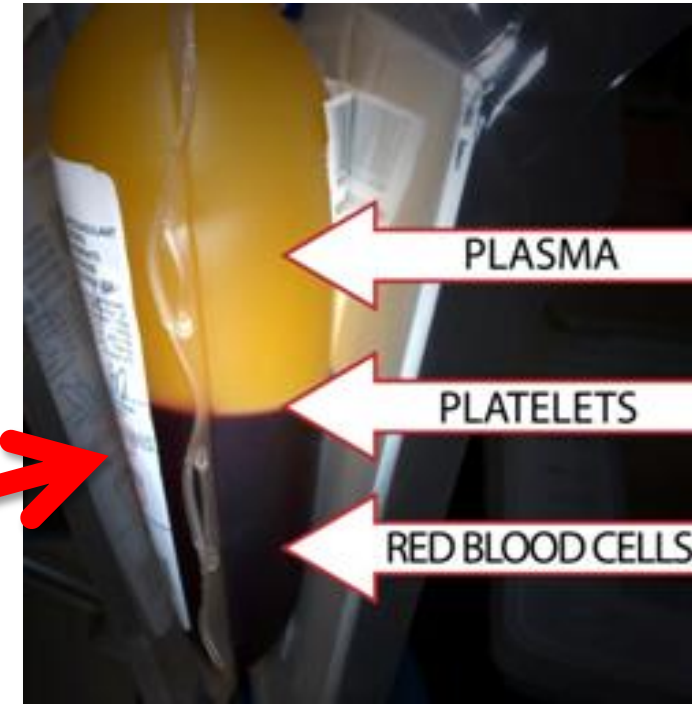
- Most common method

Whole Blood Donation

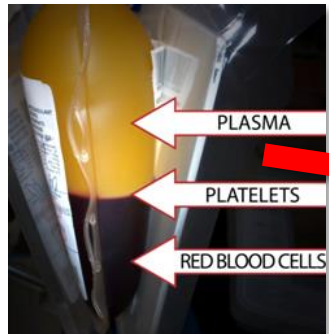
whole blood



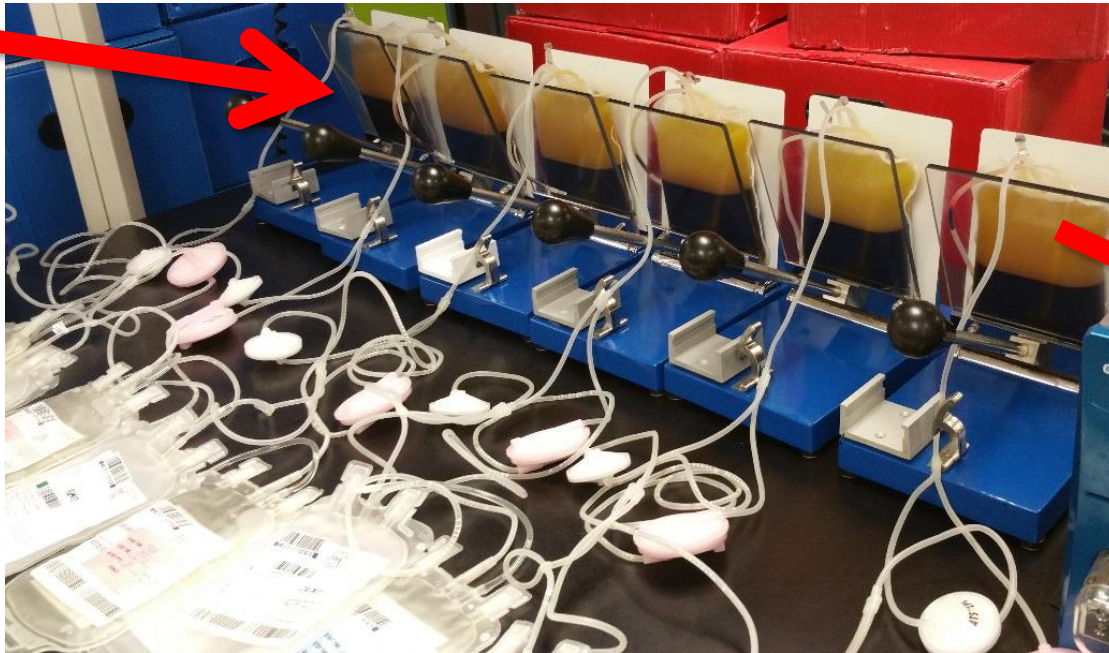
centrifuge



Whole Blood Donation



expresser



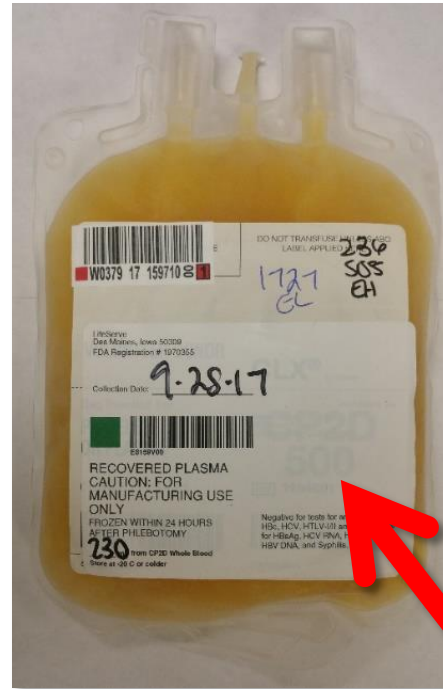
platelet-rich
plasma



RBCs

Whole Blood Donation

plasma



platelets



What remains in the bag are the platelets

Most of plasma is expressed off

Platelets form a pellet at the bottom of the bag

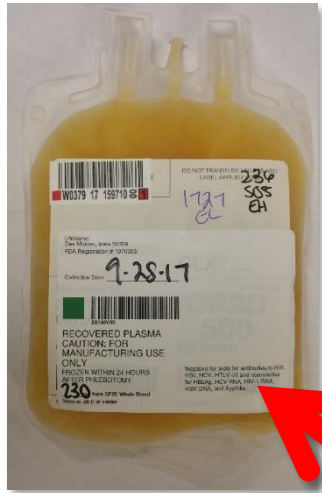
centrifuge

platelet-rich plasma



Apheresis Donation

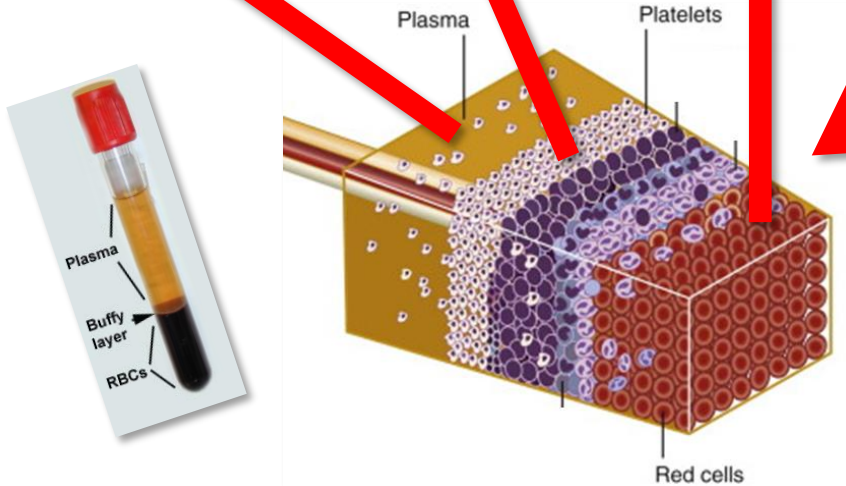
plasma



platelets



RBCs



Red Blood Cells (RBCs)



RBCs

- Most patients can physiologically compensate for a lower hemoglobin, up to a point

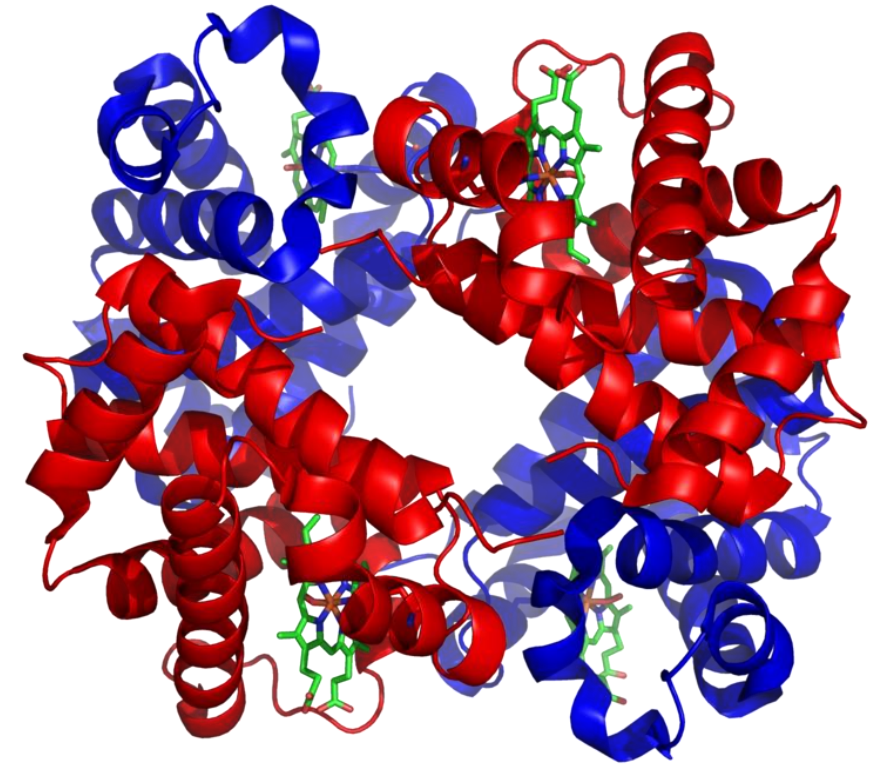
$$\text{oxygen delivery (DO}_2\text{)} = \text{cardiac output (CO)} \times \text{arterial oxygen content (CaO}_2\text{)}$$

$$\text{CO} = \text{heart rate (HR)} \times \text{stroke volume (SV)}$$

- When CaO_2 is low, HR and SV will increase
 - At rest, oxygen delivery is about 4 times the amount needed
- Patients with decreased ability to increase HR and/or SV are at greater risk for ischemic damage

RBCs

- pRBCs (packed red blood cells) and RBCs are the same thing
- Most oxygen delivery occurs via hemoglobin (Hb)
- Low Hb (anemia) results in less oxygen delivery to tissues
 - If oxygen delivery is too low, tissue damage may occur



RBCs – Indications

- Ideal is to transfuse just enough to maximize clinical outcome
 - Decrease risk of side effects
 - Blood is always in short supply → ensures enough for everyone
- Evidence-based transfusion guidelines exist, but all patients are unique, so clinical judgement is required
 - Clinical status
 - Comorbidities
 - Patient's desires



RBCs – Indications

Thresholds for red blood cell transfusion in adults

Condition	Hemoglobin threshold for transfusion
Symptomatic patient (eg, myocardial ischemia, hemodynamic instability)	10 g/dL ^{*[1]}
Hospitalized patient	
Preexisting coronary artery disease	8 g/dL [*]
Acute coronary syndromes, including acute MI	8 to 10 g/dL ^{¶[2]}
ICU (hemodynamically stable)	7 g/dL ^{*[3,4]}
Gastrointestinal bleeding (hemodynamically stable)	7 g/dL ^{*[5,6]}
Orthopedic surgery	8 g/dL ^{*[1]}
Cardiac surgery	7.5 g/dL ^{*[7,8]}
Ambulatory outpatient	
Oncology patient in treatment	7 to 8 g/dL [¶]
Palliative care setting	As needed for symptoms; hospice benefits may vary

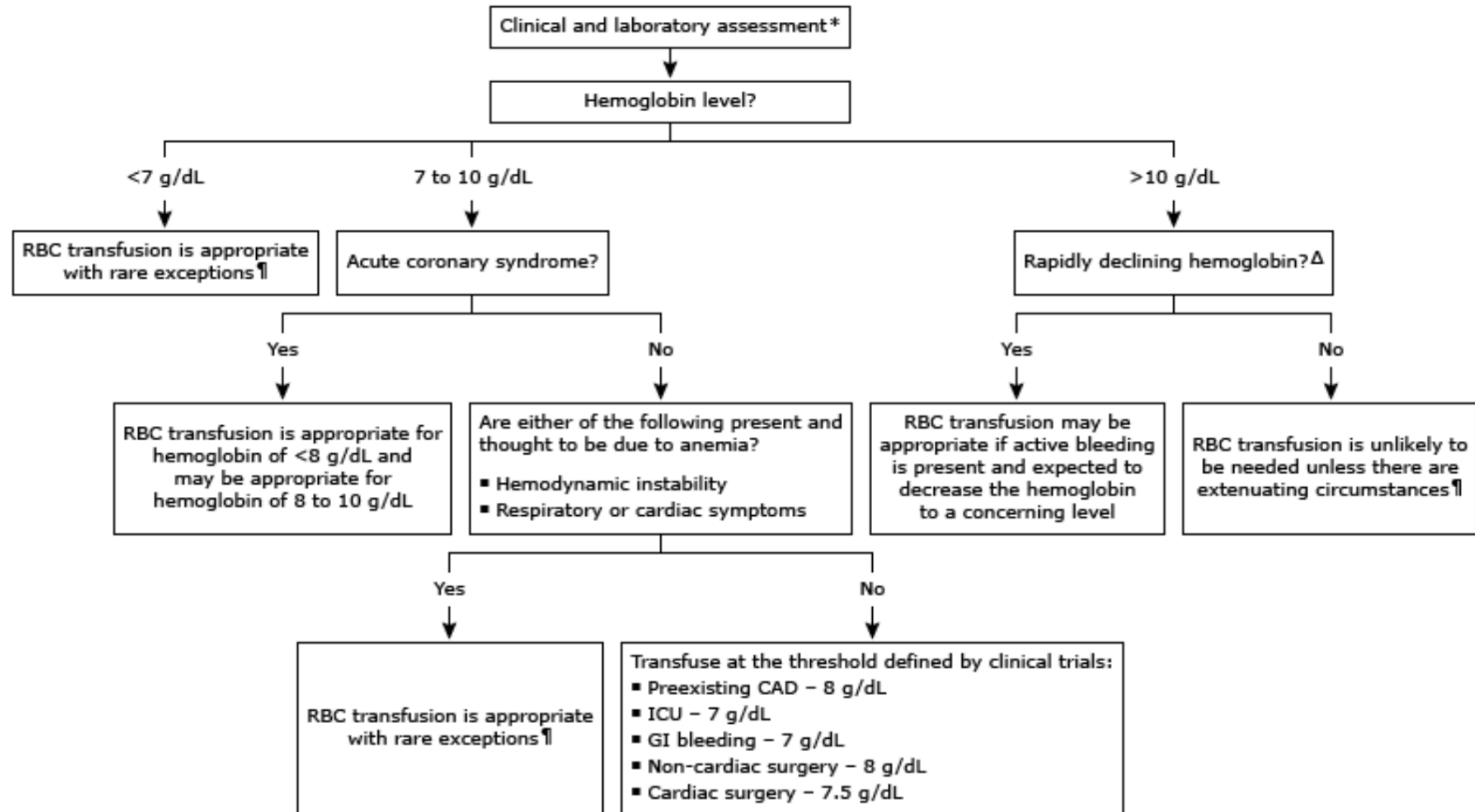
These thresholds are not a substitute for direct assessment of the patient and clinical judgment. Refer to UpToDate topics on red blood cell transfusion and specific clinical settings for further details. Hospitalized patients with heart failure are an especially challenging case because there are no data from large randomized trials, and the improvement in oxygenation from transfusion must be balanced against the risks of worsening heart failure due to the volume of the transfused blood. The authors generally use a threshold of 7 to 8 g/dL in this population, erring on the side of a higher hemoglobin level in those who are expected to be able to better tolerate the volume load. In patients who do not fit into these clinical subgroups, we recommend that transfusion based on the location of care (ICU versus other) or the similarity of their underlying disease to those patient groups where data are available. In most cases, a 7 or 8 g/dL threshold is appropriate.

MI: myocardial infarction; ICU: intensive care unit.

^{*} Based on results from clinical trial(s). Some experts may use different values. As an example, in individuals with gastrointestinal bleeding, it is often difficult, if not impossible, to estimate what the nadir hemoglobin will be, and some experts recommend a transfusion threshold of 8 g/dL ^[6].

[¶] There are no large clinical trials yet performed in this setting. These recommendations are based on the authors' opinions.

RBCs – Indications



RBCs – Indications

The decision to transfuse always incorporates an assessment by the clinician caring for the patient. Thresholds included here are based on data from clinical trials; refer to UpToDate for details. This algorithm does **not** apply to individuals with hemoglobinopathies (sickle cell disease, transfusion-dependent thalassemia); separate criteria apply to these individuals as discussed in UpToDate. To convert hemoglobin to g/L, multiply by 10 (hemoglobin of 7 g/dL = 70 g/L). Refer to UpToDate topics on indications for transfusion for further details and discussions.

RBC: red blood cell; CAD: coronary artery disease; ICU: intensive care unit; GI: gastrointestinal.

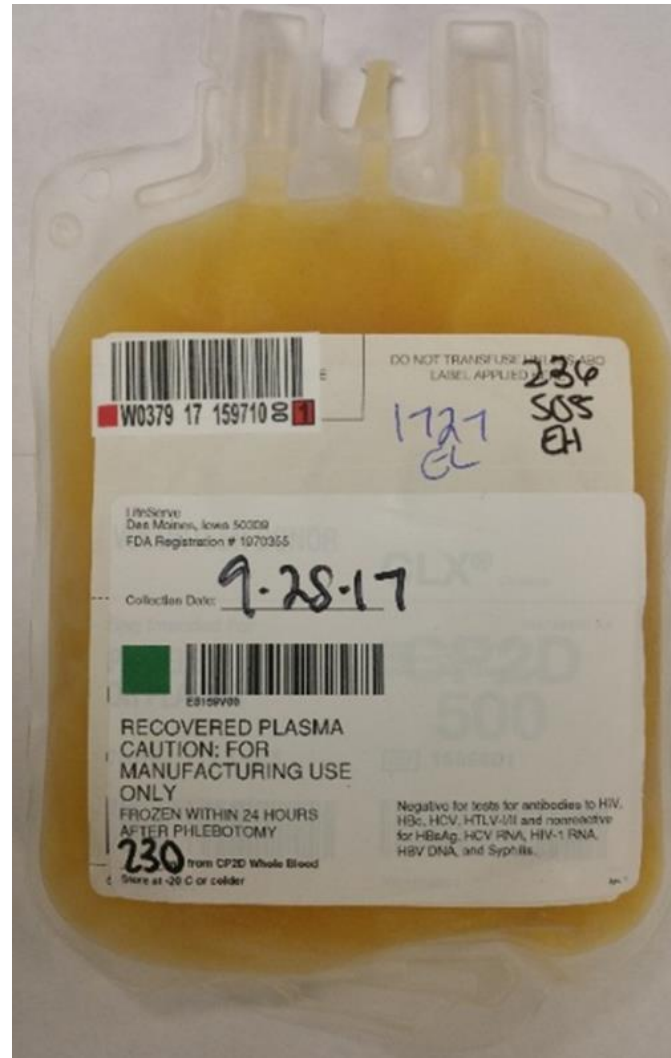
* Assessment includes:

- Symptoms (and whether attributable to anemia)
- Clinical status (vital signs, signs of hemodynamic instability, cardiac and respiratory examination)
- Underlying comorbidities
- Hemoglobin level
- Rate of hemoglobin decline and cause (active bleeding versus ongoing hemolysis versus decreased RBC production)

¶ Rarely, an individual with hemoglobin below an accepted threshold may decline transfusion (Jehovah's Witness, healthy young adult); it is important that they understand the risks and alternatives. Rarely, an individual with a hemoglobin above 10 g/dL may warrant transfusion, such as if there are clear symptoms attributable to anemia and the cause of anemia cannot otherwise be rapidly treated.

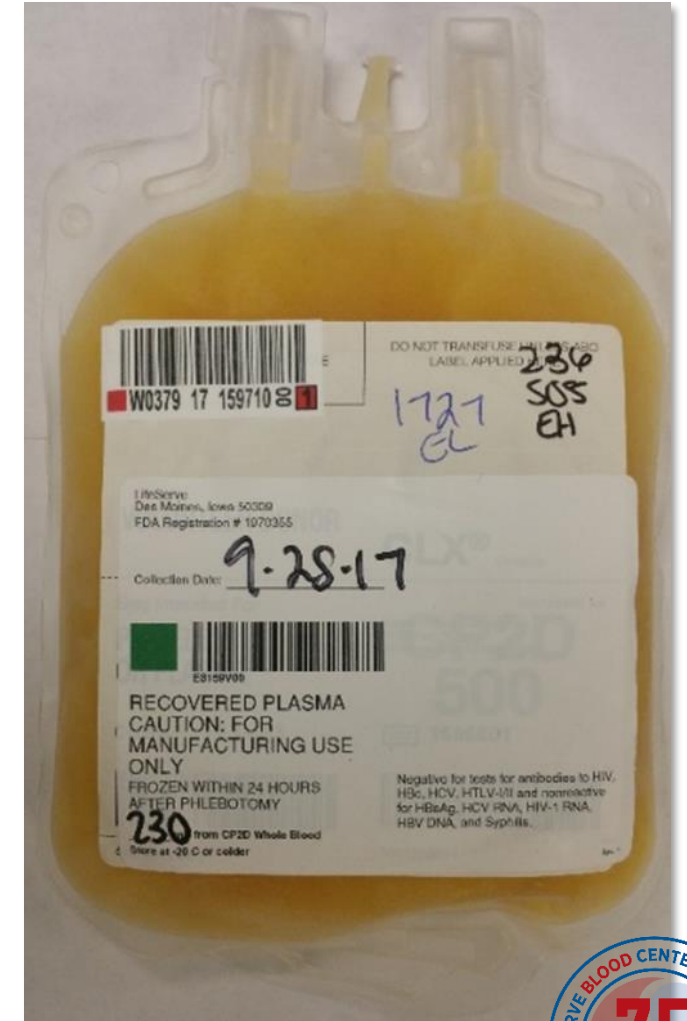
Δ Rapidly declining hemoglobin includes rapid bleeding associated with hemodynamic instability or a fall in hemoglobin of ≥ 2 g/dL per day.

Plasma



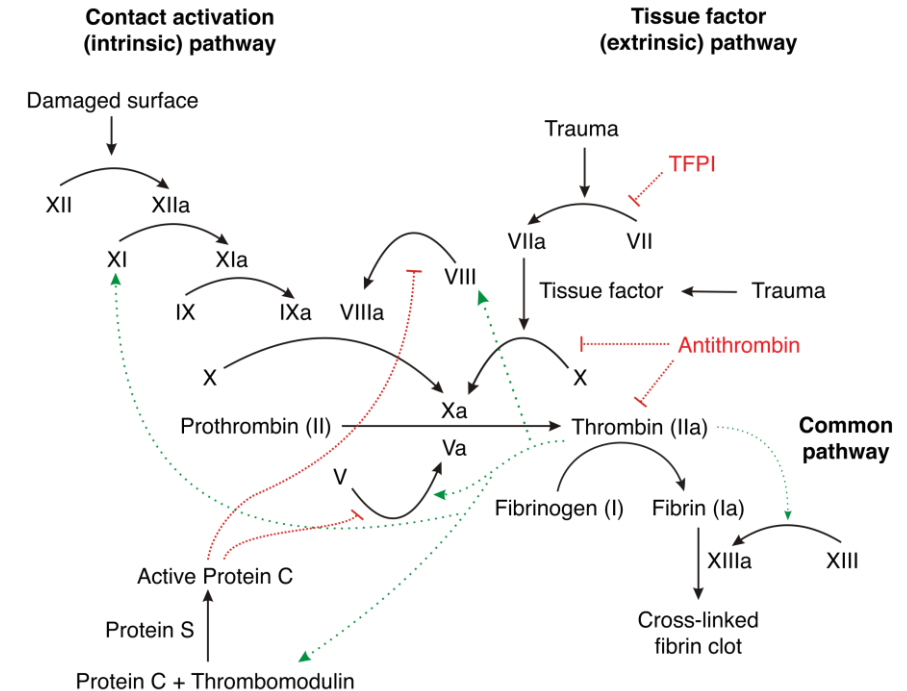
Plasma

- Contains...
 - All soluble plasma proteins and clotting factors
- Often not used appropriately
 - Source of albumin or nutrients
 - Increase blood volume
 - Reverse warfarin when there is no bleeding
 - Correct a minimally elevated INR
 - < 1.85 in a bleeding patient
 - < 2 in a nonbleeding or preop patient



Plasma – Indications

- Patients with multiple coagulation factor deficiencies who are bleeding or expected to bleed (surgery)
 - Liver disease
 - Disseminated intravascular coagulation (DIC)
 - Massive transfusion
 - Warfarin anticoagulated or overdose not corrected by vitamin K and/or prothrombin complex concentrate (PCC)
 - Vitamin K-dependent coagulation factors (II, VII, IX, and X), protein C, and protein S

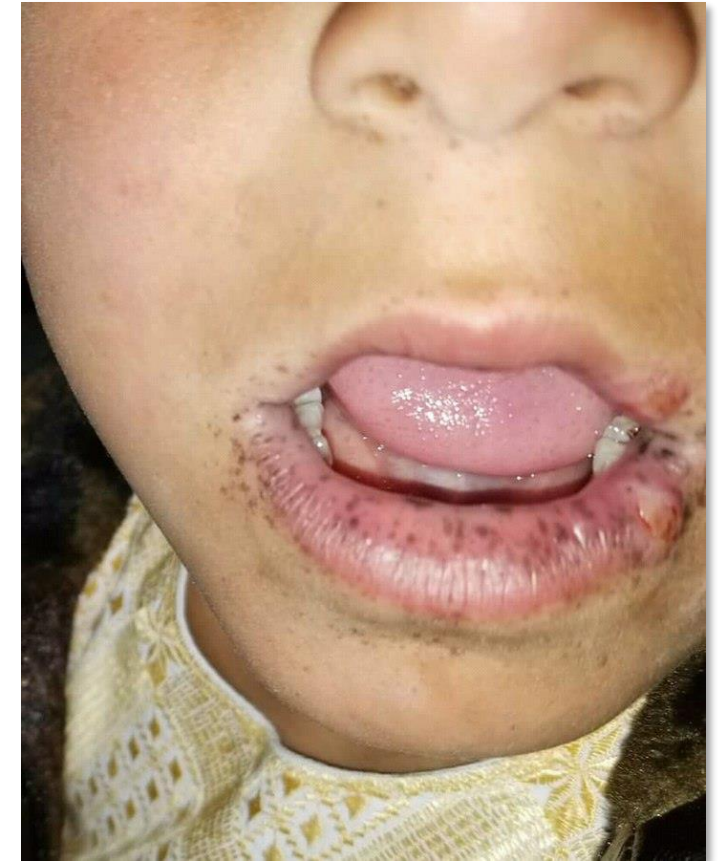


Platelets



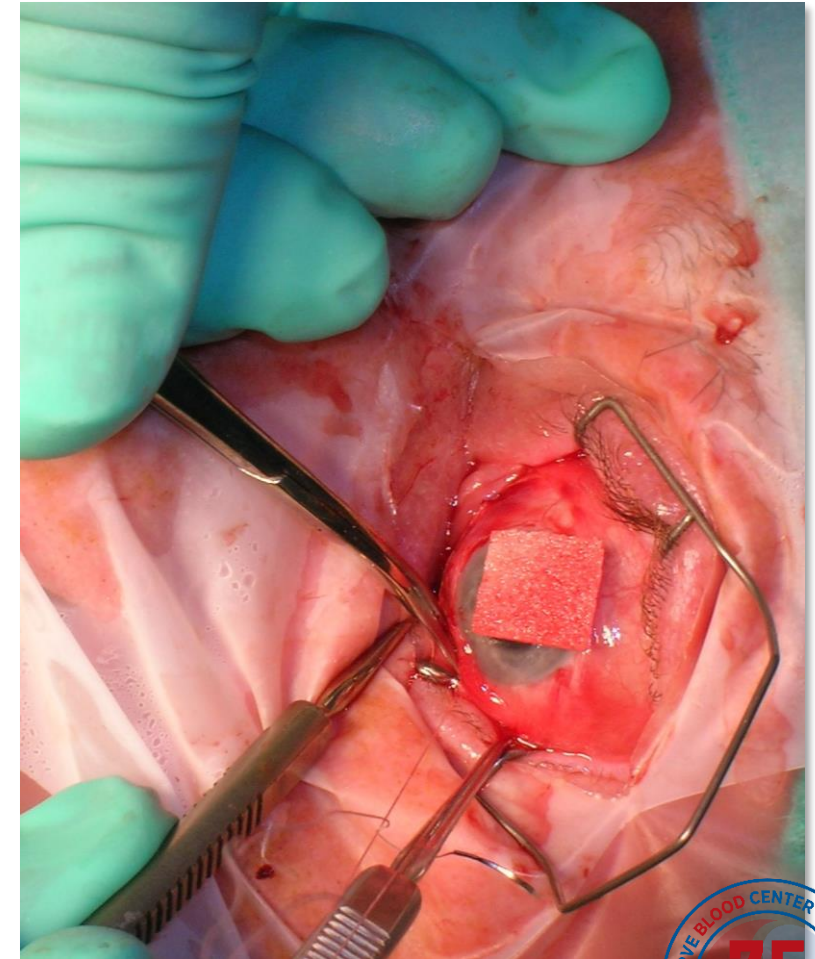
Platelets – When to Transfuse

- Platelet count when previously bled
- Mucosal bleed, epistaxis
- Prevention of spontaneous bleeding
 - Afebrile: $< 10,000/\mu\text{L}$
 - Fever, infection, or inflammation: $< 15,000$ to $20,000/\mu\text{L}$
 - Acute promyelocytic leukemia (APL): $< 30,000$ to $50,000/\mu\text{L}$
- Active bleed
 - CNS: $< 100,000/\mu\text{L}$
 - Elsewhere: $< 50,000/\mu\text{L}$



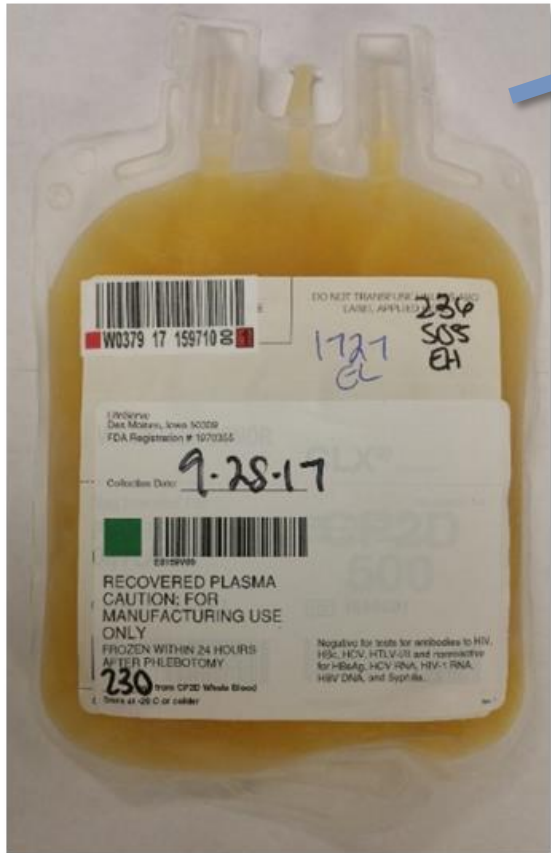
Platelets – Indications

- Preparation for an invasive procedure
 - Neurosurgery or ocular surgery: $< 100,000/\mu\text{L}$
 - Most other major surgery: $< 50,000/\mu\text{L}$
 - Endoscopy: $< 50,000/\mu\text{L}$
 - Bronchoscopy with bronchoalveolar lavage (BAL): $< 20,000$ to $30,000/\mu\text{L}$
 - Central line placement: $< 20,000/\mu\text{L}$
 - Lumbar puncture
 - Without hematologic malignancy: $< 40,000$ to $50,000$
 - With hematologic malignancy: $< 10,000$ to $20,000/\mu\text{L}$
 - Neuraxial analgesia/anesthesia: $< 80,000/\mu\text{L}$
 - Bone marrow aspiration/biopsy: $< 20,000/\mu\text{L}$



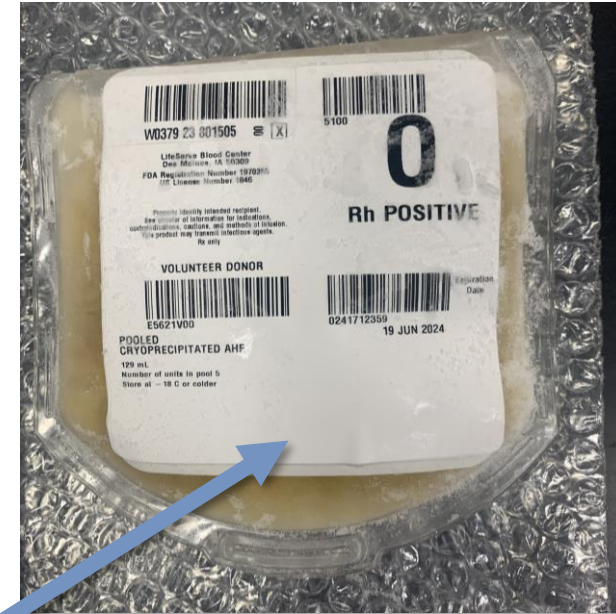
Cryoprecipitate

plasma



thaw between 1 and 6° C

centrifuge

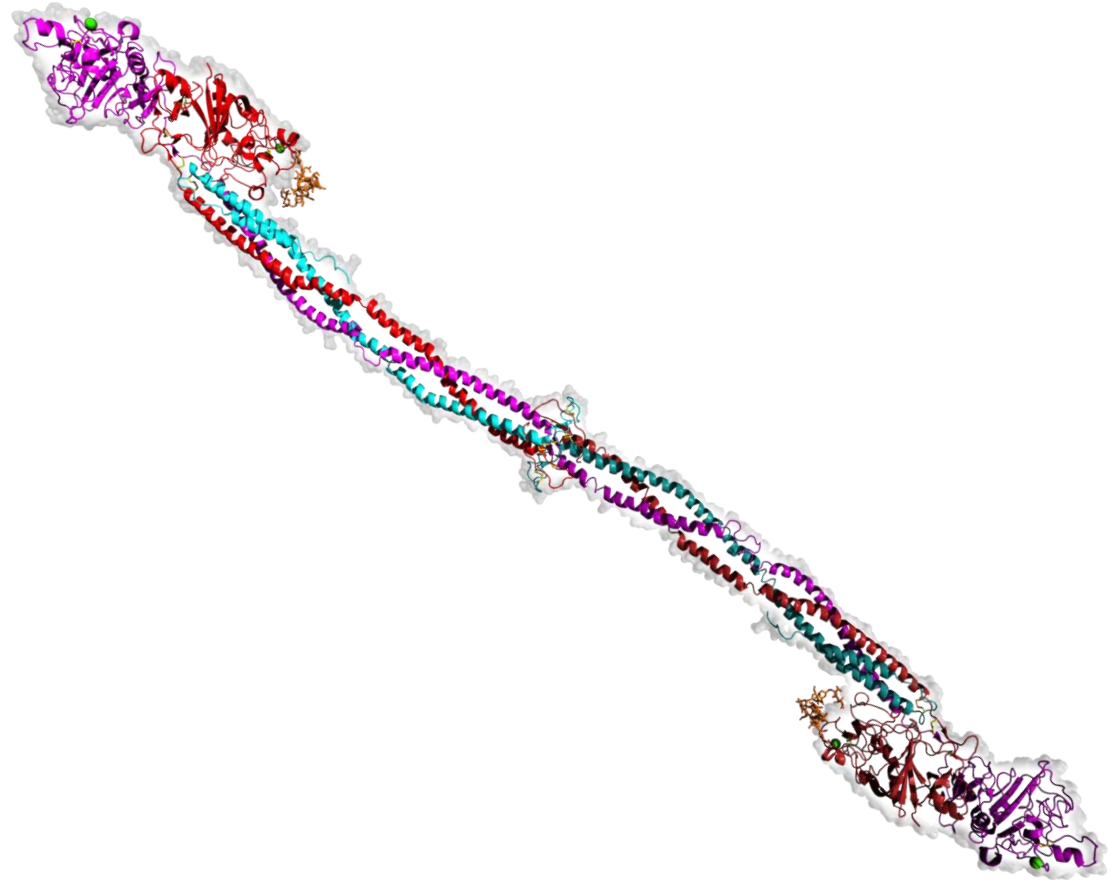


pool four to five units



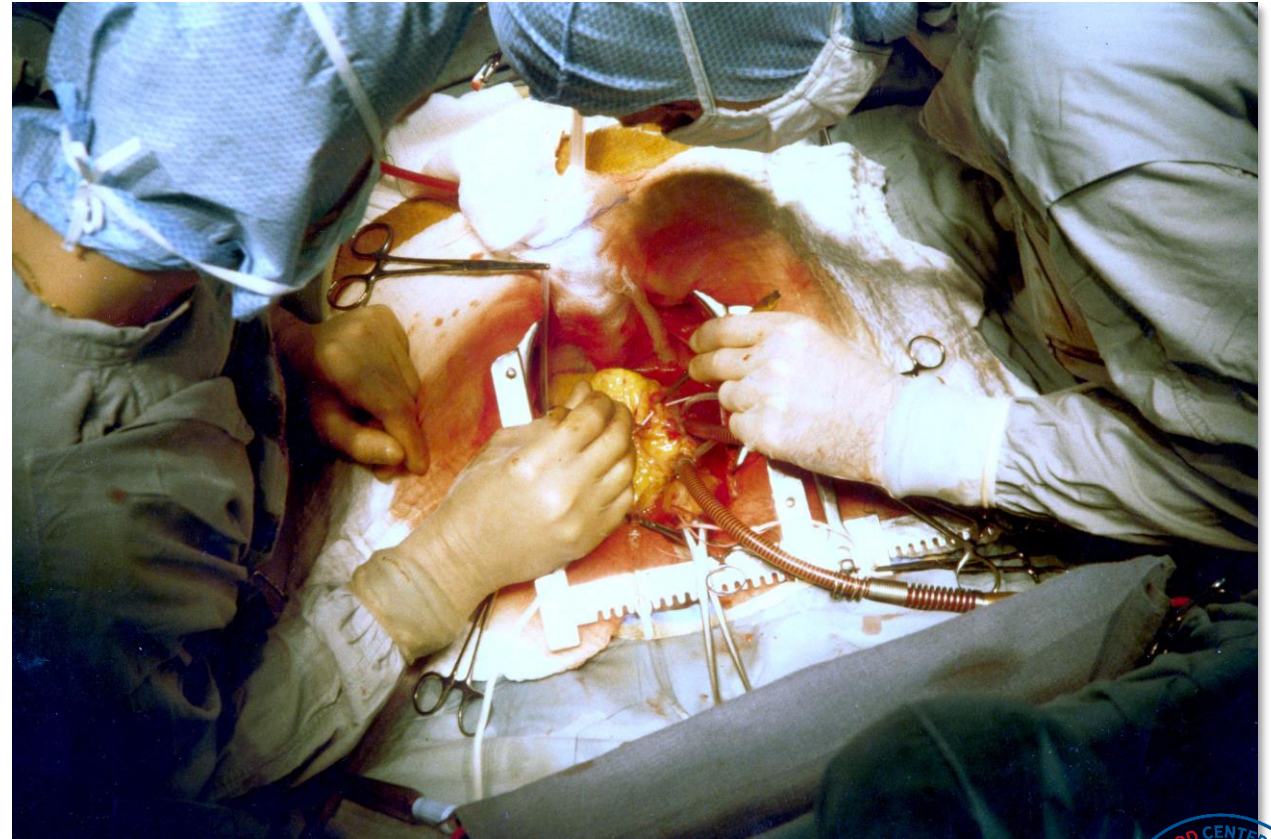
Cryoprecipitate

- Contains
 - Factor VIII
 - Factor XIII
 - von Willebrand factor
 - Fibrinogen
- Increased bleeding risk when fibrinogen is low



Cryoprecipitate – Indications

- Congenital fibrinogen disorder
- Cardiac surgery
- Postpartum hemorrhage
- Trauma
- Disseminated intravascular coagulation (DIC)
- Liver disease
- Kidney disease



Blood Product Modifications

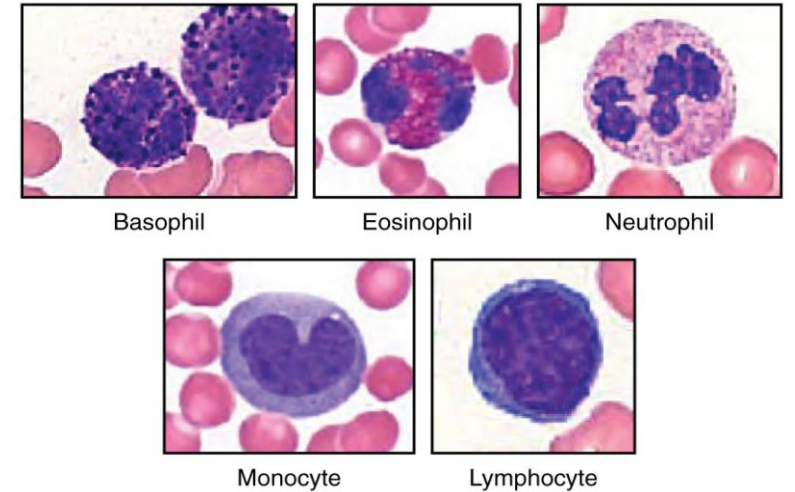


Modifications

- Leukoreduction
- Irradiation
- Washed
- Volume reduction

Leukoreduction

- Each whole blood or RBC unit contains 2.0 to 5.0×10^9 leukocytes
- Leukocytes in blood products can cause adverse effects
 - Fever
 - Human leukocyte antigen (HLA) antibody formation
 - Transmission of intracellular viruses, e.g. cytomegalovirus (CMV)
- Leukoreduction removes 99.9 to 99.99% of leukocytes
- All blood products are leukoreduced
 - Bedside leukoreduction no longer occurs



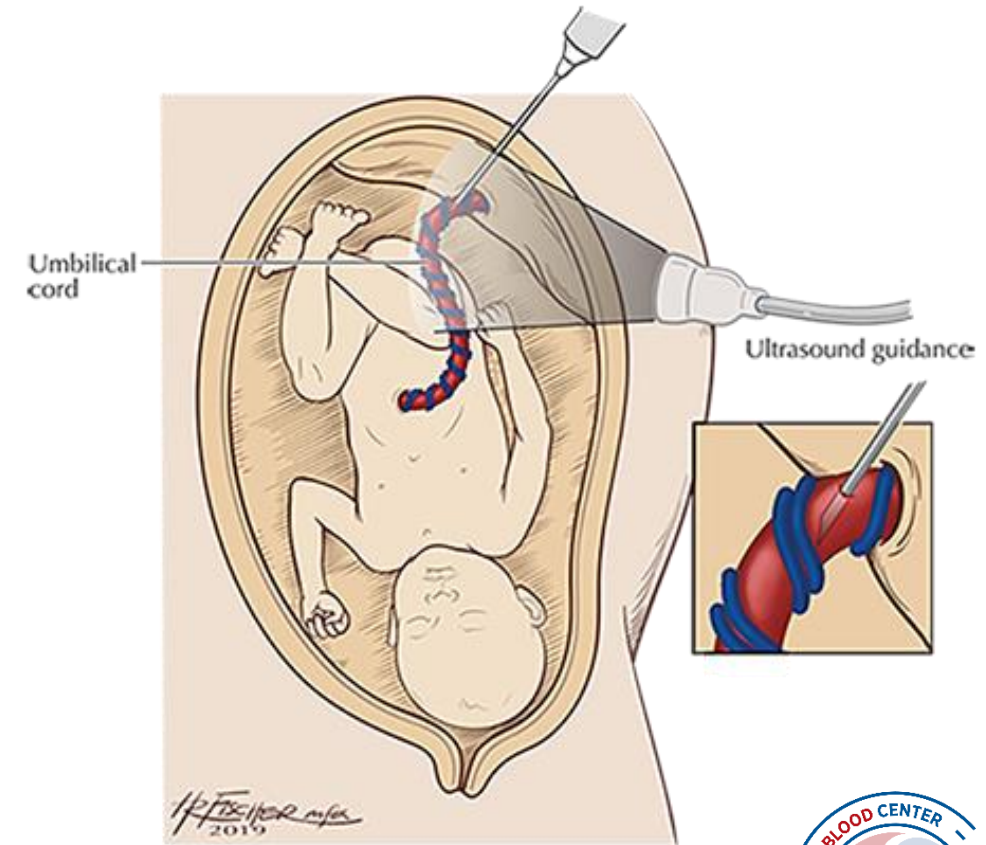
Irradiation

- Transfused leukocytes can attack a patient's cells
 - Transfusion-associated graft-vs-host disease (TA-GVHD)
- Irradiation sources
 - X-rays
 - Gamma rays (being phased out)
 - Cesium-137
 - Cobalt-60
- Disadvantages
 - Increases potassium leakage from RBCs
 - Problematic in certain populations
 - Reduces shelf life to 28 days
 - Adds cost



Irradiation – Indications

- Patients who cannot mount an attack against transfused leukocytes
 - Intrauterine or neonatal exchange transfusion
 - Premature neonate
 - Congenital cell-mediated immunodeficiency
 - Patients on certain immunosuppressive therapies
 - Purine analogs
 - Antithymocyte globulin
 - Monoclonal antibodies
 - Stem cell transplant
 - Hodgkin lymphoma
 - At risk for partial HLA match with blood donor
 - Directed donations
 - HLA-matched products
 - Genetically homogeneous population



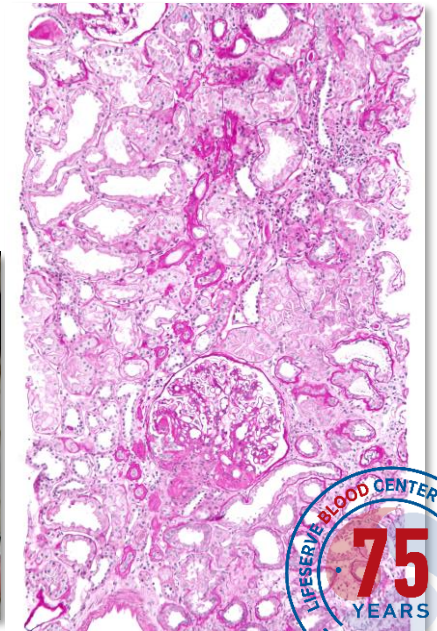
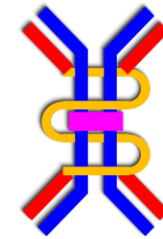
Washed

- RBCs are stored in an “additive solution” that keeps them healthy, but a small amount of the donor’s plasma will remain
 - Platelets are stored in plasma
- The residual plasma can be problematic for some patients, so the unit can be “washed” with saline
 - Adds cost
 - Reduces the shelf life to four hours



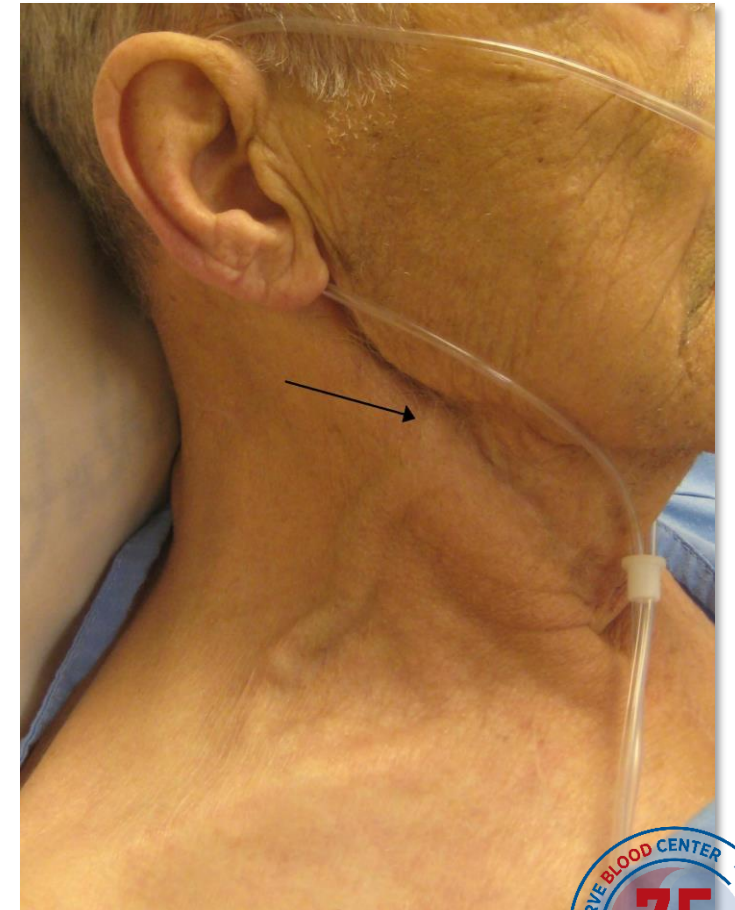
Washed – Indications

- Plasma contains...
 - Substances that may, rarely, cause an allergic reaction
 - Indication: severe or recurrent allergic reactions
 - IgA
 - If a patient does not make their own IgA, they may see a donor's IgA as "foreign" and make an antibody against it
 - May result in a severe anaphylactic reaction
 - Indication: IgA-deficiency
 - Potassium
 - Over time, RBCs in a unit will release potassium → increases the potassium in the unit
 - This can be problematic for patients at risk for hyperkalemia
 - Indications
 - Severe renal impairment
 - Newborns and infants



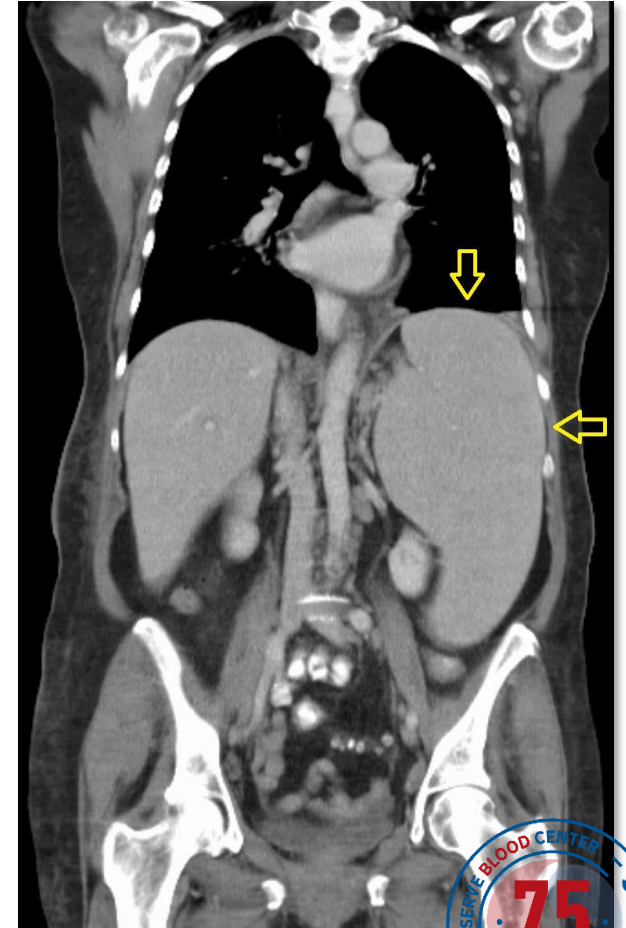
Volume Reduction

- Transfusions increase a patient's blood volume, and are often overlooked as a source of hypervolemia
- Most of the solution in an RBC unit can be centrifuged out immediately prior to transfusion
- Indication: patients at risk for transfusion-associated circulatory overload (TACO)
 - Small stature
 - Low body weight
 - Very young or old
 - Cardiac dysfunction
 - Renal dysfunction
 - High number of units already transfused (trauma)
 - High rate of transfusion (trauma)



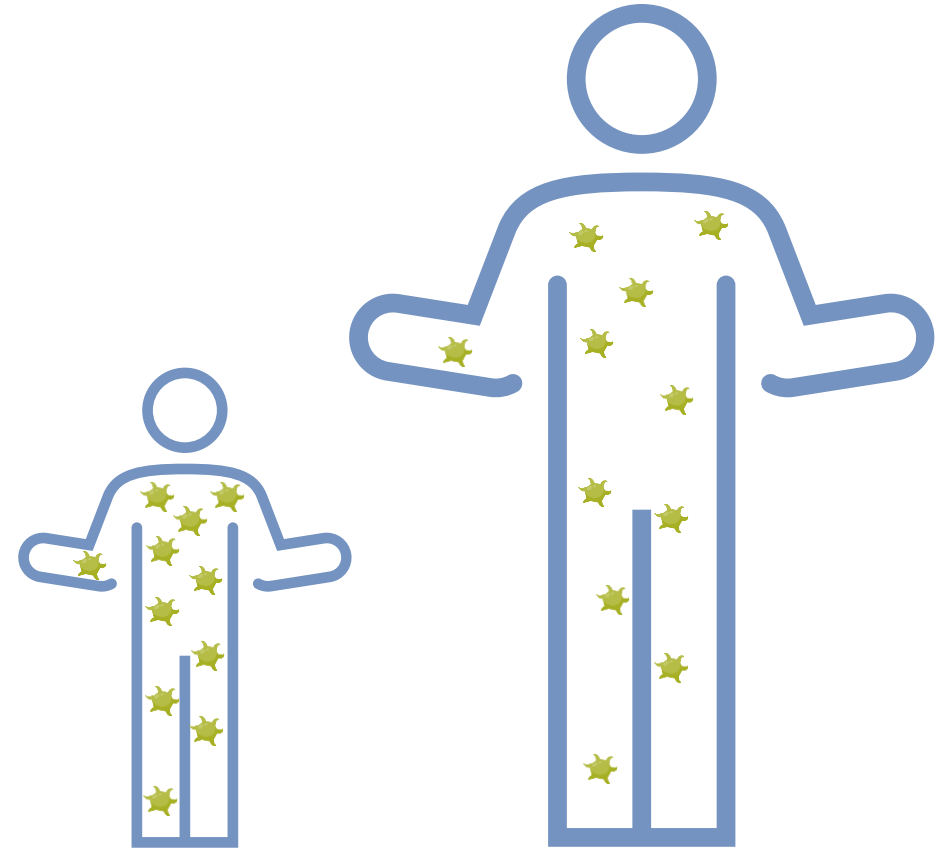
Platelet Refractoriness

- Definition
 - Lower than expected platelet count increase after two platelet transfusions
- Causes
 - **Non-immune**
 - Active bleeding
 - Fever, infection, sepsis
 - Splenomegaly
 - DIC
 - Medications
 - Stem cell transplant
 - **Immune**
 - Antibodies against human leukocyte antigens (HLA) on the platelet
 - Antibodies against human platelet antigens (HPA) on the platelet



Platelet Refractoriness – Evaluation

- Post-transfusion platelet count
 - Check platelet count after the transfusion is completed
 1. 10 minutes to one hour, and
 2. After 24 hours
 - Platelet count is measured as the number of platelets per μL
 - Larger patients have a higher blood volume, so the same number of platelets would be more diluted than a smaller patient
 - This results in a lower platelet count



Platelet Refractoriness – CCI

- To accommodate for differences in patient volume, calculate a corrected count increment (CCI)

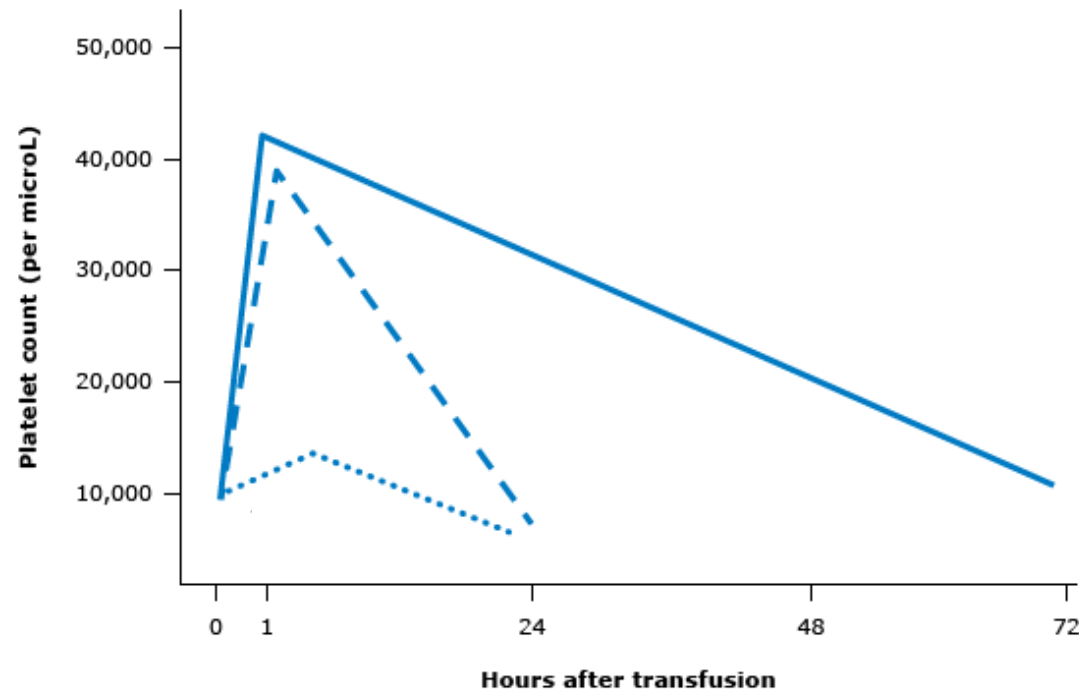
$$\frac{(\text{pretransfusion platelet count} - \text{posttransfusion platelet count}) / \text{body surface area}}{\text{\# of platelets transfused}}$$

- The hospital blood bank can give you the number of platelets that were in the unit
 - Otherwise use 3.0×10^{11}



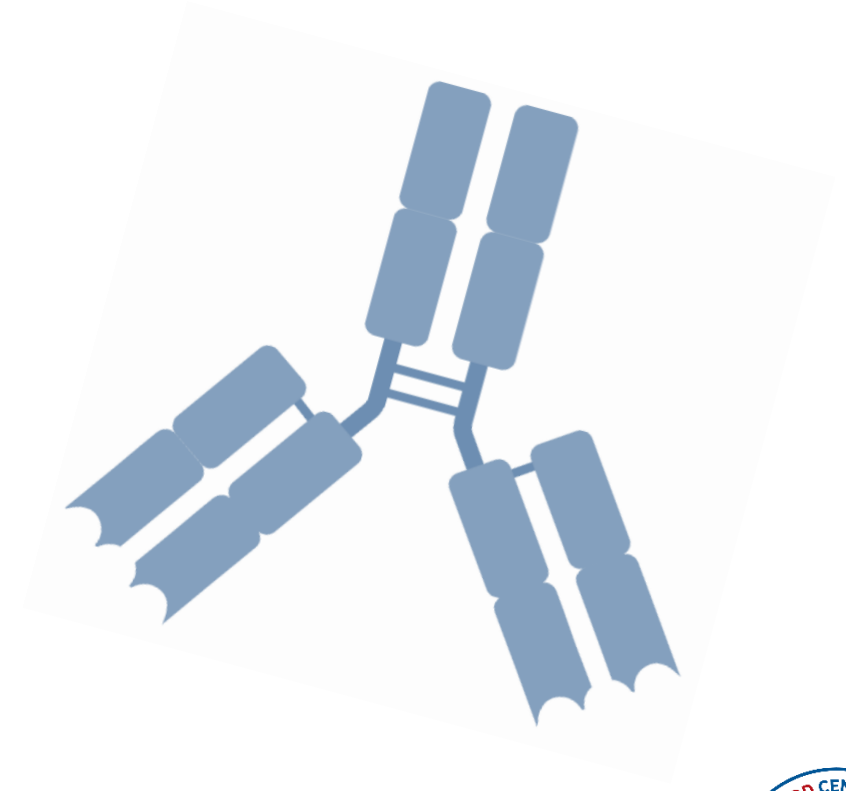
Platelet Refractoriness – CCI

- Calculate a CCI for two subsequent platelet transfusions at least 24 hours apart
 - If CCI $\geq 10,000$ after each transfusion \rightarrow likely non-immune
 - If CCI $< 10,000$ after either transfusion \rightarrow likely immune



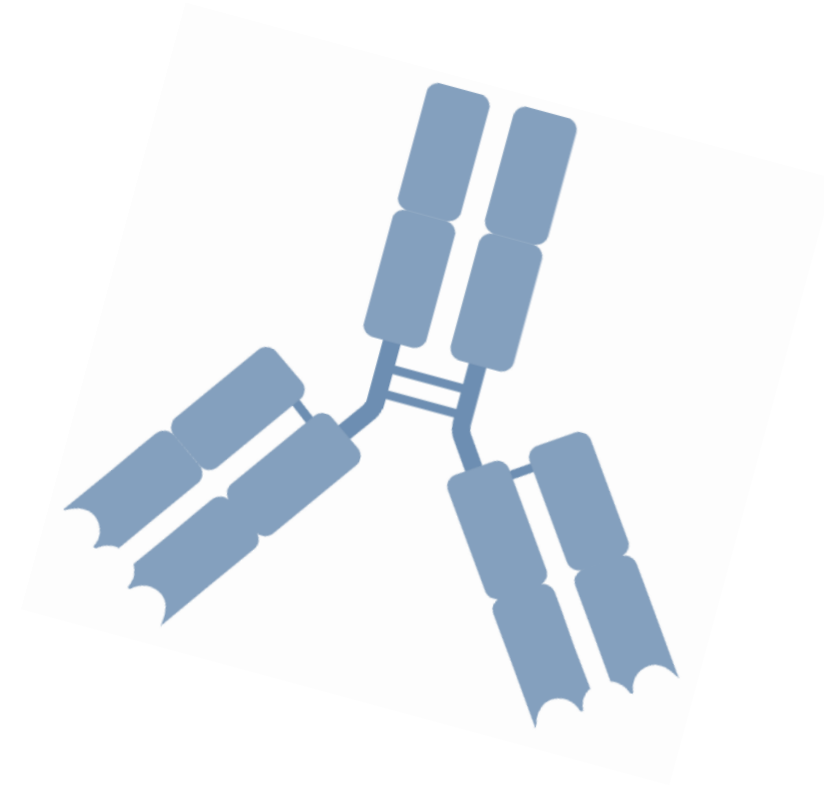
Platelet Refractoriness – Management

- Non-immune
 - Treat the underlying cause
- Immune
 - Random platelet unit
 - If the patient does not destroy the transfused platelets, the person who donated the platelet can be asked to donate more
 - Disadvantage: the patient is already forming HLA antibodies → higher risk of creating antibodies against the donor's HLAs



Platelet Refractoriness – Management

- Immune (cont.)
 - **Crossmatched platelets**
 - Mix the patients serum with a sample from a platelet unit
 - If the platelets in the sample are not destroyed, the unit is compatible
 - Disadvantage: not always accurate
 - **HLA-matched platelets**
 - Determine what HLA antibodies the patient has
 - Find a donor without the concomitant antigens
 - Disadvantages
 - Time to receive platelets
 - Cost



Massive Transfusions



Massive Transfusions

- A treatment for massive hemorrhage
 - **Definition**
 - Tends to be arbitrarily defined and dependent on the institution
 - Blood loss is a continuum
 - Clinical judgement is required
 - Examples
 - Ten or more RBC units transfused in 24h
 - Three or more RBC units transfused in one hour
 - 4 or more blood components (RBCs, plasma, platelets) transfused in 30 minutes



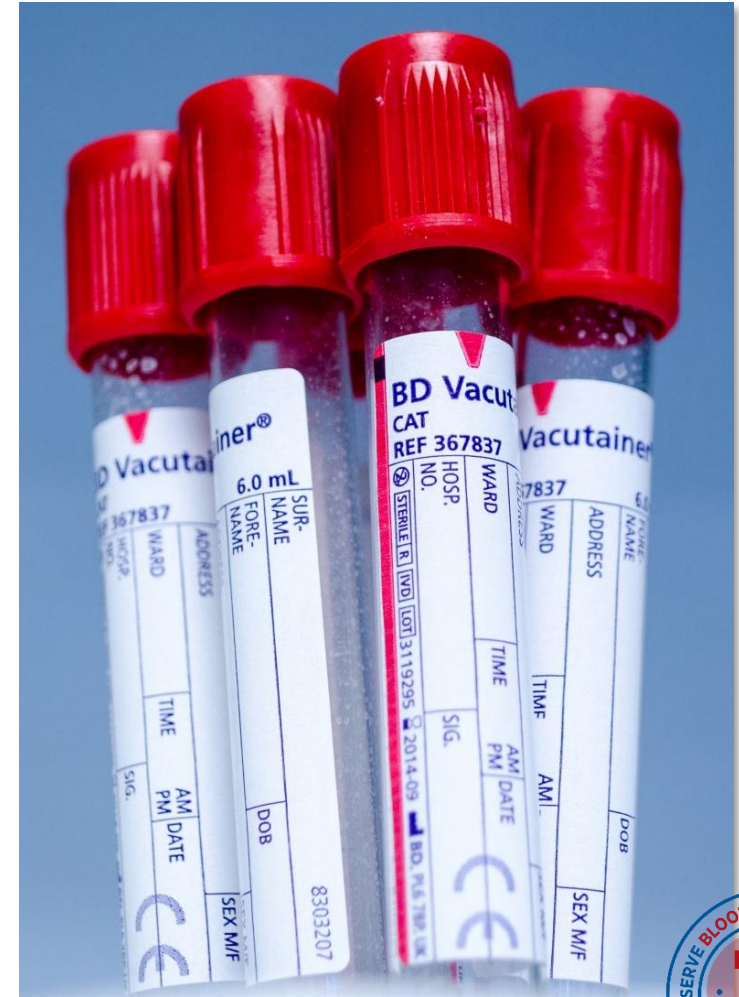
Massive Transfusions

- Common indications
 - Trauma
 - Cardiac surgery
 - Obstetric bleeding
 - Liver disease
- Massive transfusion protocol (MTP)
 - Instead of ordering individual blood components, the blood bank will send a cooler of blood components to where the massive transfusion is occurring
 - Dependent on the facility
 - Typically, the ratio is 2:1:1 (RBCs to plasma to platelets)
 - 10 RBC units, 5 plasma units, 5 platelet units
 - Cryoprecipitate is often included



Massive Transfusions

- Lab testing to monitor progress
 - CBC
 - PT
 - PTT
 - Fibrinogen
 - Thromboelastography (TEG)
 - pH
 - Blood gases
 - Electrolytes
 - Glucose
 - Lactate



Massive Transfusions

- Complications
 - Coagulopathy
 - Acidosis
 - Hypothermia
 - Hypocalcemia
 - Hyperkalemia

Questions?



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Outreach



- Physician available 24/7
 - Practitioners with transfusion-related questions/issues
 - Blood bank-related questions/issues
 - **(515) 309-4840**
- Keep up to date on transfusion-related topics and educate the medical community

Thank you!



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References

- Carson J. Indications and hemoglobin thresholds for red blood cell transfusion in the adult. UpToDate. Accessed September 26, 2023. <https://www.uptodate.com/contents/indications-and-hemoglobin-thresholds-for-red-blood-cell-transfusion-in-the-adult>
- Kaufman R. Refractoriness to platelet transfusion. UpToDate. Accessed September 27, 2023. <https://www.uptodate.com/contents/refractoriness-to-platelet-transfusion>
- Tobian A. Cryoprecipitate and fibrinogen concentrate. UpToDate. Accessed September 26, 2023. <https://www.uptodate.com/contents/cryoprecipitate-and-fibrinogen-concentrate>
- Uhl L. Clinical use of plasma components. UpToDate. Accessed September 26, 2023. <https://www.uptodate.com/contents/clinical-use-of-plasma-components>
- Yuan S. Platelet transfusion: Indications, ordering, and associated risks. UpToDate. Accessed September 26, 2023. <https://www.uptodate.com/contents/platelet-transfusion-indications-ordering-and-associated-risks>