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Blood Component Therapy: Indications for Transfusion

Allogeneic (Homologous) Blood

- ▶ PRBCs contain the **same amount of Hb** as whole blood, but much of the plasma has been removed.
 - ▶ The Hct value of PRBCs is approximately **60%**
 - ▶ Other than **severe hemorrhage**, most indications for RBCs can be effectively treated with PRBCs, conserving the plasma and the components for other patients
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- ▶ The administration of PRBCs is facilitated by utilizing crystalloid or colloid as a carrier; however, not all crystalloids are suitable. Solutions containing Ca^{2+} may precipitate clotting.

- ▶ **Lactated Ringer** solution is not recommended for use as a diluent or carrier for PRBCs because of the Ca^{2+} although several experimental studies found lactated Ringer solution and normal saline to be equally acceptable.
- ▶ A more important factor may be whether the diluent is **hypotonic** with respect to plasma. In hypotonic solutions, the RBCs will swell and eventually **lyse**

► Solutions that cause hemolysis are :


1. 5% dextrose in water
2. Plasmanate
3. 5% dextrose in 0.2% saline


► Recommended solutions compatible with packed erythrocytes are :


- 1) 5% dextrose in 0.45% saline,
 - 2) 5% dextrose in 0.9% saline,
 - 3) 0.9% saline,
 - 4) Normosol-R with a pH of 7.4.
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- ▶ RBC transfusions are given to **increase O₂-carrying capacity**.
- ▶ Increasing intravascular volume in the absence of significant anemia is not an indication for blood transfusion because volume can be augmented with administration of intravascular fluids that are not derived from human blood (e.g., crystalloids)

- ▶ As such, a sole Hb value should not be the only basis for a transfusion decision.
- ▶ It should be the **overall status** of the patient that prompts transfusion therapy:
 - hemodynamics
 - organ perfusion and oxygen delivery
 - anticipated surgical needs
- ▶ Even so, the Hb value has become the basis for many transfusion strategies. It is the prime criterion for defining restrictive versus liberal transfusion strategies.


- ▶ When a patient is hemorrhaging, the goals should be :
 - ✓ to restore and maintain intravascular volume
 - ✓ cardiac output
 - ✓ organ perfusion to normal levels.
 - ▶ By using crystalloids, colloids, or both to treat hypovolemia, normovolemic dilutional anemia may be created.
 - ▶ Increasing cardiac output enhances O₂ delivery to the tissues only to a limited extent
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- ▶ patients with Hb value **more than 10 g/dL** rarely require perioperative blood transfusions
 - ▶ patients with acute anemia with a Hb value of **less than 7 g/dL** frequently require blood transfusions.
 - ▶ patients with **chronic anemia** (as in renal failure) might tolerate an Hb concentration of less than **6 to 7 g/dL**.
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- ▶ The ultimate determination of the Hb or Hct value at which blood should be given is a clinical judgment based on many factors, such as:
 - cardiovascular status
 - age
 - anticipated additional blood loss
 - arterial oxygenation
 - mixed venous O₂ tension
 - cardiac output
 - intravascular blood volume
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Additional Blood Transfusions

The following key components of information to consider include:

- ▶ 1. Measurement and trend of vital signs
 - ▶ 2. Measurement of blood loss and assessment of anticipated blood loss
 - ▶ 3. Quantitation of intravenous fluids given
 - ▶ 4. Determination of Hb concentration
 - ▶ 5. Surgical concerns.
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factors	class1	class2	class3	class4
Blood loss(ml)	750	750–1500	1500–2000	2000 or more
Blood loss (%blood volume)	15	15–30	30–40	40 or more
Pulse (beat/min)	100	100	120	140 or more
Blood pressure	Normal	normal	decreased	decreased
Pulse pressure (mmHg)	Normal or increased	decreased	decreased	decreased
Capillary refill test	normal	positive	positive	positive
Respiration per minute	14–20	20–30	30–40	35
Urine output (ml/h)	30	20–30	5–10	negligible
CNS:mental status	Slightly anxious	Mildly anxious	anxious confused	confused lethargic


Measurement of Blood Loss

- ▶ Measuring blood loss is obviously important when assessing the need for both the initial and subsequent blood transfusions
- ▶ . standard approach includes a combination of visualization and gravimetric measurements based on weight differences between dry and blood-soaked gauze pads.

Preoperative Anemia


- ▶ low Hb value in women <12 g/dL
- ▶ in men <13 g/dL
- ▶ It is a common comorbidity among patients undergoing major surgery with an incidence up to 40%
- ▶ an independent risk factor for increased perioperative mortality, and postoperative acute kidney injury (AKI)

- ▶ In patients with a moderate to high risk of significant blood loss (defined as >500 mL), the Hb value ideally should be obtained 3 to 8 weeks prior to surgery.
 - ▶ This provides sufficient time for the patient to undergo iron therapy or to correct nutritional deficiencies.
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- ▶ Erythropoiesis-stimulating agents, especially intravenously administered iron therapy, may be beneficial for treatment of preoperative anemia.
 - ▶ The concept of treating anemia preoperatively as a means to decrease the need for intraoperative transfusions is widely accepted.
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- ▶ For example intravascular iron therapy in patients undergoing abdominal surgery significantly
 - ✓ increased **preoperative Hb** levels
 - ✓ reduced the **need for transfusion**
 - ✓ and shortened **hospital length of stay**

- ▶ **Oral therapy**, if given with sufficient time preoperatively and tolerated by the patient, may be just as effective at correcting the anemia as intravenous therapy.

- ▶ Erythropoiesis-stimulating agents (ESAs), such as **darbepoetin alfa**, act by stimulating red cell progenitor cells in the bone marrow and inducing erythropoiesis
 - ▶ They are frequently prescribed for patients with anemia who have **end-stage renal disease** or who are undergoing **chemotherapy** treatment to increase their Hb levels and reduce the incidence of transfusion
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Liberal Versus Restrictive Transfusion Strategy

- ▶ It is based on the Hb value when a transfusion decision is made.
- ▶ A **restrictive** policy is the administration of blood transfusion when the Hb value is **7 to 8 g/dL or less**.


- ▶ In contrast, **a liberal policy** is the administration of blood transfusion when the Hb value is **9 to 10** g/dL or greater.
 - ▶ The most recent randomized controlled studies continue to show **no benefit** to a liberal strategy compared with a restrictive strategy
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- ▶ One conclusion is that if no clinical advantages are associated with the liberal transfusion policy, perhaps the restrictive approach should be used.
- ▶ Certainly, **fewer transfusion reactions** would be expected with the restrictive approach

- ▶ In the presence of incomplete data, the ASA's 2015 updated practice guidelines offer these recommendations:

- ▶ 1. Transfusion is rarely indicated when the Hb concentration is more than 10 g/dL and is almost always indicated when it is less than 6 g/dL, especially when the anemia is acute.

- ▶ 2. A restrictive transfusion strategy (Hb <8 g/dL) should be employed to reduce the patient's transfusion requirements and decrease the potential harmful effects of transfusions

- ▶ 3. Multimodal protocols and algorithms should be employed to reduce intraoperative blood loss and transfusion requirements.
 - ▶ These pathways include point-of-care testing to direct care
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- ▶ 4. The use of a single Hb trigger for all patients and other approaches that fail to consider all important physiologic and surgical factors affecting oxygenation is not recommended


► 5. When appropriate, intraoperative and postoperative blood recovery, acute normovolemic hemodilution (ANH), and measures to decrease blood loss (i.e., deliberate hypotension and pharmacologic drugs) may be beneficial

Platelet Concentrates


- ▶ Platelet concentrates are obtained either as **pooled concentrates** from 4 to 6 whole-blood donations or as **apheresis concentrates** obtained from one donor.
- ▶ If platelets are stored at room temperature, they can be used **up to 7 days** after collection with constant and gentle agitation.


- ▶ **Bacterial contamination**, mainly from platelet concentrates, is the third leading cause of transfusion-related deaths although the incident rate has steadily declined over the last 15 years.


- ▶ Because the use of multidonor platelet products stored for 5 days results in an incidence of sepsis five times higher than use of those stored for 4 days, shorter storage times are being emphasized.

- ▶ At present, platelet concentrates are routinely tested for bacteria and are **the only blood product stored at room temperature.**
 - ▶ For any patient who develops **a fever within 6 hours** after receiving platelets, sepsis from platelets should be considered.
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The most recent guidelines published in 2015 by the ASA Task Force on Perioperative Blood Management provide the following recommendations regarding management for platelet transfusions:



- ▶ 1. Monitor **platelet count**, except in situations of massive transfusion.
 - ▶ 2. Monitor **platelet function**, if available
 - ▶ 3. Consider use of **desmopressin** in patients with excessive bleeding or suspected platelet dysfunction.
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- ▶ 4. Platelet transfusion may be indicated despite an adequate platelet count if there is known or suspected platelet dysfunction:
 - ▶ cardiopulmonary bypass
 - ▶ bleeding
 - ▶ recent use of antiplatelet therapy
 - ▶ congenital platelet dysfunction
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- ▶ 5. **Prophylactic** platelet transfusion is rarely indicated in surgical or obstetric patients when the platelet count is **greater than $100 \times 10^9/L$** and is usually indicated when the platelet count is **less than $50 \times 10^9/L$** .
- ▶ The determination of whether patients with intermediate platelet counts (**$50- 100 \times 10^9/L$**) require therapy should be based on the patient's risk for bleeding.

- ▶ Many institutions have strict thresholds targeted to the patient's condition that outline the minimum platelet count needed for the categories of :
 - ▶ (1) prophylaxis
 - ▶ (2) periprocedural (based on type of procedure)
 - ▶ (3) active bleeding
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- ▶ In the first category, a required platelet count may be $10 \times 10^9/L$ in patients receiving chemotherapy.

In the second category, patients undergoing bone marrow biopsy or lumbar puncture should have platelet counts between 20 and $30 \times 10^9/L$.

For neurosurgery, a platelet count of $100 \times 10^9/L$ may be targeted.

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- ▶ Patients with **severe thrombocytopenia** ($<20 \times 10^9/L$) and clinical signs of **bleeding** usually require platelet transfusion.


However, patients may have **very low platelet** counts (much lower than $20 \times 10^9/L$) and **not have clinical bleeding**.

These patients probably do **not need** platelet transfusions

- ▶ When possible, **ABO-compatible** platelets should be used.
- ▶ The platelet membrane has immunoglobulins, and any additional deposit of recipient antibodies is difficult to detect

- ▶ ABO-incompatible platelets produce very adequate hemostasis.
- ▶ The effectiveness of platelet transfusions is difficult to monitor

- ▶ Under ideal circumstances, **one platelet concentrate** usually produces an increase of approximately **$7 \text{ to } 10 \times 10^9/\text{L}$** at 1 hour after transfusion in the 70-kg adult.
- ▶ **Ten units** of platelet concentrates are required to increase the platelet count by **$100 \times 10^9/\text{L}$**

- ▶ However, many factors may lead to **decreased survival** and decreased recovery of transfused platelets including
 - ❖ splenomegaly
 - ❖ previous sensitization
 - ❖ fever
 - ❖ sepsis
 - ❖ active bleeding,
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Fresh Frozen Plasma

- ▶ FFP is **the most frequently** used plasma product.
- ▶ It is processed shortly after donation, generally frozen within **8 hours or 24 hours** (PF24)

- ▶ It contains **all the plasma proteins**, particularly **factors V and VIII**, which gradually decline during the storage of blood.
- ▶ **PF24** is comparable to FFP, except for a slight reduction in **factor V** and approximately 25% decrease in **factor VIII**.

- ▶ Thawed plasma is stored at 1 °C to 6 °C for up to 5 days

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- ▶ The use of FFP carries with it the same inherent risks that are observed with the use of any blood product, such as sensitization to foreign proteins.

- ▶ Although FFP is a reliable solution for intravascular volume replacement in cases of acute blood loss, alternative therapies are equally satisfactory and considerably safer.

- ▶ The risks of FFP administration include
 - TRALI
 - TACO
 - allergic or anaphylactic reactions.

- ▶ In 2015 the ASA Task Force recommended the following guidelines regarding the administration of FFP:
- ▶ 1. **Prior to the administration of FFP, coagulation studies** should be obtained when feasible.

- ▶ 2. For the correction of coagulopathy when the international normalized ratio (INR) is greater than 2, in the absence of heparin

- ▶ 3. For the correction of coagulopathy due to coagulation deficiencies in patients transfused with **more than one blood volume (approximately 70 mL/kg)** when coagulation studies cannot be easily or quickly obtained

- ▶ 4. Replacement of known **coagulation factor deficiencies** with associated bleeding, disseminated intravascular coagulation (DIC), or **both**, when specific components are not available.

- ▶ 5. Reversal of **warfarin anticoagulation** when severe bleeding is present and prothrombin complex concentrations are not available.


Cryoprecipitate


- ▶ Cryoprecipitate is prepared when FFP is thawed, and the precipitate is reconstituted.

The product contains :

- factor VIII
- vWF
- fibrinogen
- factor XIII
- fibronectin

- ▶ Cryoprecipitate is frequently administered as **ABO compatible**; however, this probably **is not very important** because the concentration of antibodies in cryoprecipitate is extremely low


- ▶ Cryoprecipitate may contain **RBC fragments**, and cryoprecipitate prepared from Rh-positive donors can possibly **sensitize Rhnegative** recipients to the Rh antigen.
 - ▶ Cryoprecipitate should be administered through a **filter** and as **rapidly** as possible.
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- ▶ The rate of administration should be at least **200 mL/h**, and the infusion should be completed within **6 hours** of thawing.
 - ▶ transfusion of cryoprecipitate is **rarely** indicated when the fibrinogen levels are greater than **150 mg/dL** in nonobstetric patients.
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- ▶ The following indications were provided regarding the administration of cryoprecipitate:

1. When testing of fibrinogen activity reveals evidence for **fibrinolysis**

- ▶ **2. When **fibrinogen** concentrations are less than **80 to 100** mg/dL in patients experiencing excessive bleeding**

- ▶ 3. **Obstetrical** patients who are experiencing excessive bleeding despite a measured fibrinogen concentration **greater than 150 mg/dL**
 - ▶ 4. In patients undergoing **massive transfusion** when the timely assessment of fibrinogen concentrations cannot be determined
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- ▶ 5. In patients with **congenital fibrinogen deficiencies** and when possible, in consultation with the patient's hematologist
 - ▶ 6. In **bleeding** patients with **von Willebrand** disease types 1 and 2A who fail to respond to desmopressin or vWF/FVIII concentrates (or if not available)
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- ▶ 7. In **bleeding** patients with von **Willebrand** **disease types 2B, 2M, 2N, and 3** who fail to respond to vWF/FVIII concentrates (or if concentrates are not available)

