# Postpartum hemorrhage

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	Class of haemorrhagic shock				
	1	11	III	IV.	
Blood loss (mL)	Up to 750	750-1500	1500-2000	> 2000	
Blood loss (% blood volume)	Up to 15	15-30	30-40	>40	
Pulse rate (per minute)	< 100	100-120	120-140	> 140	
Blood pressure	Normal	Normal	Decreased	Decreased	
Pulse pressure (mm Hg)	Normal or increased	Decreased	Decreased	Decreased	
Respiratory rate (per minute)	14-20	20-30	30-40	> 35	
Urine output (mL/hour)	> 30	20-30	5–15	Negligible	
Central nervous system/ mental status	Slightly anxious	Mildly anxious	Anxious, confused	Confused, lethargic	

Table 2. Clinical Findings in Obstetric Hemorrhage [25] (Open Table in a new window)

Blood Volume Loss	Blood Pressure (systolic)	Symptoms and Signs	Degree of Shock	
500-1000 mL (10-15%)	Normal	Palpitations, tachycardia, dizziness	Compensate	
1000-1500 mL (15-25%)	Slight fall (80- 100 mm Hg)	Weakness, tachycardia, sweating	Mild	
1500-2000 mL (25-35%)	Moderate fall (70-80 mm Hg)	Restlessness, pallor, oliguria		
2000-3000 mL (35-50%)	Marked fall (50- 70 mm Hg)	Collapse, air hunger, anuria	Severe	

- HR & BP  $\rightarrow$  two most commonly used V/S  $\rightarrow$  Dx hemorrhage
- women  $\rightarrow$  hemorrhage may not develop tachycardia or hypotension until significant blood loss (>1,000 mL)
- Signs of a hemorrhage : HR >110 beats/ minute

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BP \le 85/45 \text{ mm Hg}
Spo2 < 95\%
delayed \text{ capillary refi ll}
U/O \downarrow
pallor
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ratio of the heart rate over the systolic BP (HR/Sbp) is called : shock index and may be helpful

A shock index *greater than 1* requires immediate managemen

(lightheadedness, palpitations, confusion, syncope, fatigue, air hunger, and diaphoresis)

## 1- blood loss (vaginal del > 500 cc or C/S > 1000cc & normal V/S)

- angiocath (16 or 18)
- 3.5 lit (maximally) crystalloids
- O2 therapy:

Mask: 6-8 lit/min, cannula: 2-4 lit/min

 $NS \rightarrow reasonable solution in the labor ward setting because: (1)low cost (2) compatibility with most drugs and blood transfusions$ 

(If large amounts (>10 L) of crystalloid are being infused, a change to LRS can be considered)

Dextrose-containing solutions, such as 5% dextrose in water or diluted NS in 5% dextrose in water, have no role in the management of PPH. Remember that the loss of 1 L of blood requires replacement with 4-5 L of crystalloid because most of the infused fluid is not retained in the intravascular space but instead shifts to the interstitial space.

### Continue Blood loss(1000-1500cc) & Normal V/S & Normal Labtest:

- second IV line  $\rightarrow$  16 or 18 (if not available  $\rightarrow$  cv-line)
- invasive monitoring? If necessary
- u/o
- PPH of up to 1500 mL in a healthy pregnant woman  $\rightarrow$  usually be managed  $\rightarrow$  crystalloid infusion alone if the cause of bleeding is arrested
- \* A meta-analysis in the Cochrane Library comparing resuscitation with colloid solutions versus crystalloid <u>favored the</u> <u>use of crystalloids</u> with respect to mortality
- \*The NS groups had a 1% mortality rate, versus an 11% mortality rate in the colloid group
- \* Large volumes of colloid solutions (>1000-1500 mL/d) can  $\rightarrow$  adverse effect on hemostasis

No colloid solution has been demonstrated to be superior to NS

expense and the risk of adverse effects with colloids, crystalloid is recommended

Given these findings  $\rightarrow$  the authors recommend *agains*t the use of colloid solutions in resuscitation outside the setting of an RCT

# Continue bood loss(> 1500 cc) or transfusion >2u packed cell or coagulopathy or abnormal lab test or oliguria:

- Newer studies tend to have lower transfusion rates than older studies
- to OPERATING ROOM, TXA
- Ca, inotrope

Cardiovascular collapse(severe hemorrhage, hypovolemic shock amniotic emboli)

# massive transfusion:

- 1-PC > 10 U / 24h
- 2- replacement > 1 blood volume /24 h
- 3- replacement > 50% in 4 h (2h) (adult blood volume = 70 ml/kg)
- $4- \ge 150 \text{ ml /min}$
- goal:
- \* maintain tissue perfusion & oxygenation
- \* stop bleeding to use surgical or other intervention

 Check these parameters early and frequently (e.g. every 30-60 minutes while massive transfusion is ongoing)

Parameters	Values to aim for
Temperature	>35 °C
Acid-base status	pH >7.2, base excess <-6, lactate <4 mmol/L
lonised calcium (Ca)	>1.1 mmol/L
Haemoglobin (Hb)	This should not be used alone as transfusion trigger; and, should be interpreted in context with haemodynamic status, organ & tissue perfusion
Platelet (Plt)	≥ 50 x 10^9 /L (>100 x 10^9 if head injury/ intracranial haemorrhage)
PT/APTT	≤ 1.5x of normal
Fibrinogen	≥ 1.0 g/L

#### First:

- 1- system activity
- 2- CBC, PT, PTT, INR, fibrinogen, ABG
- 3- TXA 1g Qh8
- 4- warming the patients
- 5- preparing 4 *U* packed cell:
- \* compatible type screen or partially cross-match
- \* O –
- \*O + if Rh +

## **Second:**

- $\blacksquare 4u(PC)$ , 4u(FFP), 1u(PltSD) or 10u(RD)
- Lab test every 30 -60 min
- 1g calcium
- if Fibrinogen < 100 mg/dl → cryopersipitate
- Platelet & cryoprecipitate (preferably compatible unless except in emergencies)

#### **Thirth**

Repeat second

If no surgical causes, fibrinogen > 150, Plt > 50000, ABG = Nl but bleeding (+) or unstable

- → recombinant factor 7
- Targets of resuscitation in the setting of massive transfusion include:

Mean arterial pressure (MAP) of 60 to 65 mm Hg

Hemoglobin 7 to 9 g/dL

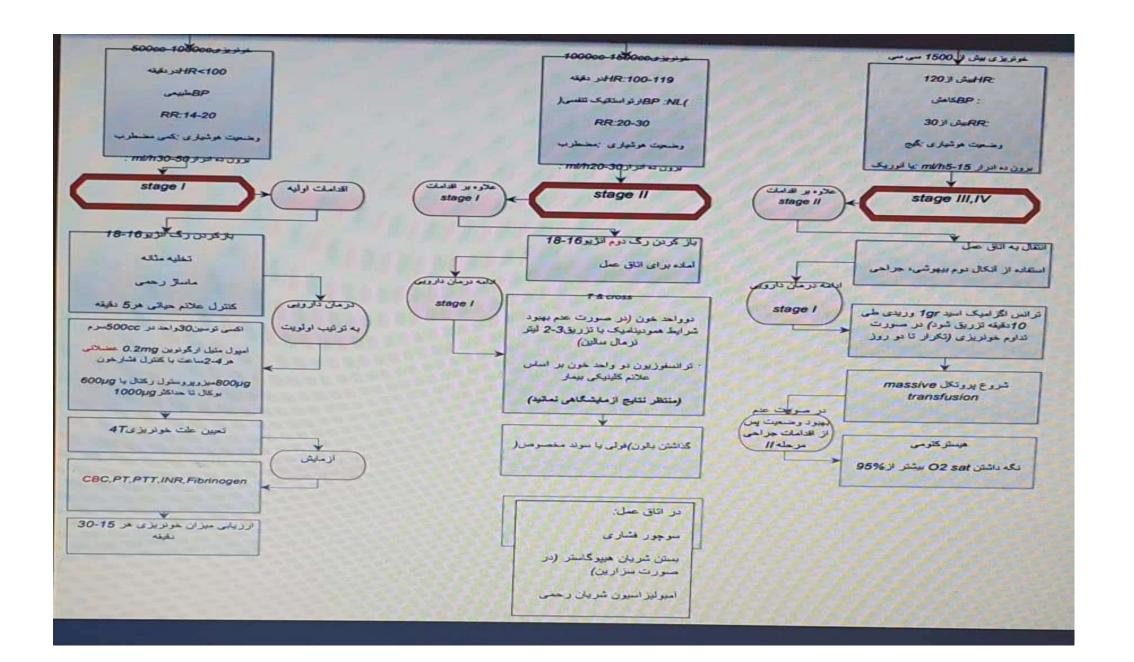
*INR* < 1.5

Fibrinogen > 1.5 to 2 g/L

*Platelets* > 50 000

pH 7.35 to 7.45

Core temperature greater than 35 C



#### Packed cell:

1unit = 250 ml

CPDA -1  $\rightarrow$  shelf life = 35 d ( < 6 °C)

given over 1-2 h but *not longer* than 4 h (child : 2-5 ml/kg/h)

Type – screen & cross-match

1 unit  $\rightarrow$  Hb 1g/dl $\uparrow$ 

CMV-negative or CMV reduced risk (leukocyte reduced) RBCs  $\rightarrow$  *should be used* in pregnant women who are CMV-negative or whose CMV status is unknown.

# Over time:

- Lactic acid K Ca ↑
- 2,3 DPG ATP PH glycolysis ↓
- packed cell :
  - \* young : < 14 -21 d
  - \* old: > 21 d

#### **FFP**

- After thawing(30-37  $\circ$  C) over 20-30 min  $\rightarrow$  use maximally during  $4h \rightarrow$  if not  $\rightarrow$  store 1-6  $\circ$  C  $\rightarrow$  use within 24h(if no use  $\rightarrow$  discarded)
- $\blacksquare 200 300 \text{ ml/h}$
- ABO compatible but need no crossmatch, no (leak clot abnormal color)
- Once thawed  $\rightarrow$  activity of clotting factors, particularly  $\vee$ ,  $\vee$ III  $\rightarrow$  decline gradually  $\rightarrow$ re-administration may  $\rightarrow$  every 6 to 8 hours if there is ongoing bleeding due to the short half-life of factor VII;  $\vee$ II has a half-life of 2 to 6 hours
- 10-20 mL/kg will increase factor levels by 20-30%

#### Platelet:

- pool of 6 whole blood derived (→ referred : random donor) platelets or one apheresis platelet
- \* Random donor : 1 unit  $\rightarrow$  5000 10000  $\uparrow$
- \* single donor(apheresis)  $\rightarrow$  30000 -60000  $\uparrow$
- room T (22  $\circ$  C)  $\rightarrow$  RD : viable  $\rightarrow$  5-7 d
- Thrombocytopenia :
  - \* procedures > 50,000
  - \* CNS & Retina > 100,000
  - \* neuroaxial > 80,000
  - \* LP >40,000

#### Contraindication:

TTP, HUS, HIT

If ABO identical platelets : *not available*  $\rightarrow$  ABO plasma compatible may be use  $\rightarrow$  but will not cause clinically significant problems

ABO incompatible platelets  $\rightarrow$  only minimal risk of hemolysis (unless large doses of ABO incompatible platelets are transfused)

Rh: should be compatible

Pregnancy:

CMV-seronegative or CMV reduced risk (leukocyte reduced) platelets should be used in pregnant women who are CMV-seronegative or whose CMV status is unknown.

# Cryopercipitate:

- prepared from plasma
- fibrinogen, VIII, XIII, von Willebrand factor, fibronectin
- main indication : hyfibrinogenemia

Cryo <u>should not</u> be used for patients with <u>Hemophilia A</u> (Factor VIII deficiency) unless recombinant and/or virally inactivated factor VIII preparations are not available.

It <u>should not</u> be used for patients with <u>von Willebrand disease</u> unless they are proven not to respond to DDAVP.

It is **not usually** given for **Factor XIII deficiency**, as there are virus-inactivated concentrates of this protein available.

Cryo is sometimes useful if **platelet dysfunction** associated with **renal failure** does not respond to dialysis or DDAVP.

Use for **fibrin glue** 

# **Fibrinogen Replacement:**

- 1 unit of cryo per  $5kg \rightarrow will$  increase fibrinogen by about 100 mg/dLNumber of bags =  $0.2 \times weight$  (kg)  $\rightarrow$  provide about 100 mg/dL fibrinogen

  Many institutions use a standard dose of 10 units and then repeat if needed
- lunit of cryo has low volume, ABO compatibility is not required except in neonates & small children unless high volumes of cryo are to be transfused

- Each unit (~10-15mL) provides:
- Fibrinogen 150-250 mg with a half-life of 100-150 hours
- Factor VIII 80-150 U with a half-life of 12 hours
- Von Willebrand factor 100-150 U with a half-life of 24 hours
- Factor XIII (13) 50-75 U with a half-life of 150-300 hours
- Cryo also contains fibronectin; however there are no clear indications for fibronectin replacement

# **Acute hemolytic reactions:**

- **Early signs:** may fever, hypotension, flushing, wheezing, anxiety, and/or red-colored urine
- Late signs: may a generalized bleeding tendency (DIC) and/or hypotension

# **Nonhemolytic febrile reactions:**

**fever** (some times :shaking, chills, hypotension, and vomiting)

# Allergic reaction:

- maculopapular rash and/or urticaria
- Anaphylactic reaction :

Dyspnea, Wheezing ,Anxiety , Hypotension without fever ,Bronchospasm in severe cases

# Transfusion-related acute lung injury (TRALI):

■ rapid onset of *shortness of breath*, hypoxemia, and rales, without signs of acute cardiogenic pulmonary edema and fever during 6 h of transfusion

Or late as 6-72h sfter transfusion

# Circulatory (volume) overload:

■ Shortness of breath, Rales, Orthopnea, Tachycardia, Distended jugular veins, Other evidence of cardiac decompensation

## Acute hemolytic reactions (ABO incompatibility):

Accidental transfusion of RBCs of a different ABO type

# Febrile non- hemolytic reaction:

Cytokines and other normal constituents of leukocytes, platelets, or plasma accumulate in blood components during storage

When transfused  $\rightarrow$  some recipients  $\rightarrow$  which *fever* is the *most common* symptom.

# Allergic reaction

recipient was exposed → foreign substance in the blood product to which the recipient is sensitized.

Studies in the medical literature sugg  $\rightarrow$  causes of allergic reactions  $\rightarrow$  *polymorphic proteins* in the donors' plasma

food (nuts, tomatoes), or medications (penicillin)  $\rightarrow$  donor ingested immediately before collection

# Anaphylactic reaction:

Most cases of anaphylaxis ar  $\rightarrow$  recipients with <u>IgA deficiency</u>  $\rightarrow$  developed anti-IgA

( Not all IgA-deficient persons )

Similar reactions in ahaptoglobinemia → reported

# Transfusion-related acute lung injury (TRALI):

*Neutrophils*  $\rightarrow$  effector cells  $\rightarrow$  adhere to the *pulmonary endothelium*  $\rightarrow$  permeability  $\uparrow$   $\rightarrow$  pulmonary edema

Elements  $\rightarrow$  activation  $\rightarrow$  neutrophils:

transfused human leukocyte or neutrophil antigen (HLA or HNA) antibodies and transfused bioactive substances such as lipids or cytokines

Because pregnancy → common cause of alloimmunization → HLAs and HNAs →

most cases of TRALI have been traced to plasma-containing blood components collected from female blood donors

When the American Red Cross converted to predominantly male-donated plasma, the number of cases of TRALI decreased very significantly from 2006 to 2008

# Circulatory (volume) overload

Increased fluid volume →

susceptible patients→ cardiovascular compromise, elderly patients, and small children→pulmonary edema

A usual **transfusion rate** is 2-2.5 mL/kg per hour  $\rightarrow$  In at-risk patients, blood products can be transfused at a **slower rate**.

## Lab studies:

acute hemolytic reactions: the workup includes:

Visual inspection of the recipient's plasma and urine

**Retyping** of donor and recipient red blood cells (RBCs)

Direct antiglobulin (Coombs) testing

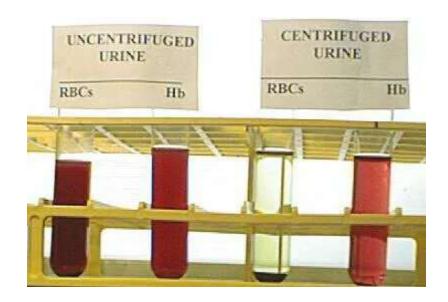
# febrile nonhemolytic reactions:

Every things normal

# Allergic reaction:

Every things normal

Eo may not increase



#### Anaphylactic reaction:

anti-IgA in a pretransfusion sample of the recipient's serum or plasma establishes the diagnosis

## Transfusion-related acute lung injury (TRALI):

plasma levels of brain natriuretic peptide (BNP) may be useful in distinguishing the <u>cardiogenic pulmonary</u> <u>edema</u> present in circulatory overload from the <u>noncardiogenic pulmonary edema</u> present in TRALI

#### **Acute hemolytic reactions:**

- Immediate *DC transfusion* while maintaining IV\_LINEs for emergency management.
- Anticipate hypotension, renal failure, and DIC.
- Prophylactic → reduce the risk of *renal failure* may include *low-dose dopamine* (1-5 mcg/kg/min)

Maintain U/O minimally  $\rightarrow$  75-100 ml/h

- \* vigorous hydration with crystalloid solutions (3000 mL/m<sup>2</sup>/24 h)
- \* If fluid and mannitol  $\rightarrow$  ineffective  $\rightarrow$  *furosemide*
- \* urine alkalization → bicarb

((osmotic diuresis with 20% mannitol (100 mL/m²/bolus, followed by 30 mL/m²/h for 12 h).))

If DIC is documented and bleeding requires treatment, *transfusions of frozen plasma*, *pooled cryoprecipitates* for fibrinogen, and/or *platelet* concentrates may be indicated.

# Febrile, nonhemolytic reactions

fever usually *resolves* in 15-30 minutes without specific treatment

If fever  $\rightarrow$  discomfort  $\rightarrow$  oral *acetaminophen* (325-500 mg)

Avoid aspirin because of its prolonged adverse effect on platelet function

# Allergic reactions

diphenhydramine → effective → pruritus that is associated with hives or a rash

(oral or IV) 25 -100 mg

# Anaphylactic reactions

**SQ** injection of **Epinephrine** (0.3-0.5 mL of a 1:1000) is standard treatment

If patient  $\rightarrow$  sufficiently hypotensive  $\rightarrow$  efficacy of the subcutaneous route?

**Epinephrine** (0.5 mL of a 1:10,000 aqueous solution)  $\rightarrow$  IV

Although **no documented** evidence exists that *IV corticosteroids* are beneficial most clinicians → hydrocortisone or prednisolone if an immediate response to epinephrine does not occur.

# Transfusion-related acute lung injury (TRALI):

- 1- Immediately discontinue → transfusion while preserving venous access IVs
- 2- mild episodes  $\rightarrow$  respond to oxygen administered by nasal catheter or mask If shortness of breath persists after oxygen administration  $\rightarrow$  ransfer to an IC where mechanical ventilation can be employed.
- 3- In the absence of signs of acute volume overload or cardiogenic pulmonary edema diuretics are not indicated.

**No evidence** → corticosteroids or antihistamines are beneficial.

Treat complications with specific supportive measures.

# Circulatory (volume) overload:

- 1- sitting position and administer oxygen to facilitate breathing.
- 2- The *most specific* treatment is *discontinuing the transfusion* and removing the excessive fluid.
- 3- If practical, the unit of blood component being transfused may be lowered to reverse the flow and to decrease intravascular volume by a controlled phlebotomy.
- 3- Less urgent situations may be managed by a *parenteral or oral diuretic (furosemide)*.