

CLINICAL STRATEGIES FOR AVOIDING AND CONTROLLING HEMORRHAGE AND ANEMIA WITHOUT BLOOD TRANSFUSION IN OBSTETRICS AND GYNECOLOGY*

GENERAL PRINCIPLES OF NONBLOOD OB/GYN MANAGEMENT

1. **Formulate an individualized management plan** to facilitate rapid decision making. Exercising clinical judgment, be prepared to modify routine practice (e.g., expeditious control of bleeding, early use of oxytocic drugs, erythropoietin therapy). Planning, prevention, and minimization of blood loss using appropriate multiple interventions are the cornerstones of management without blood transfusion.
2. **Discuss the risks and benefits** of anticipated or potential procedures with the patient.
3. **Ensure the availability of well-trained personnel**, drugs, and equipment for prevention and prompt control of hemorrhage. Considering the available blood management options, refer the patient to another institution if better resources are available elsewhere.
4. **Adopt an organized, multidisciplinary team approach.** Early recognition of complications and immediate involvement of appropriate specialists (e.g., obstetric, gynecologic, vascular, or general surgery; anesthesiology; hematology; intensive care; internal medicine; neonatology/perinatology; nursing; radiology; or urology) experienced in clinical management without transfusion may be essential.
5. **Communicate the plan of care** to all personnel, including the surgical backup team, to avoid treatment delays. Where there are multiple conditions treated by multiple physicians, ongoing interspecialty communication is particularly important.
6. **Maintain frequent, close surveillance for signs and symptoms of postpartum or postoperative bleeding** to facilitate early intervention. Recognition of risk factors before and during labor/delivery may help clinicians to identify women who require appropriate preventive measures and extra vigilance. Nonetheless, complications may occur suddenly and unexpectedly in any patient, even those thought to be at low risk.
7. **Prompt skillful intervention to prevent/control blood loss can be lifesaving.** The clinical urgency of low-level persistent bleeding may not be recognized until compensatory mechanisms fail and blood pressure falls. In general, avoid a “watch and wait” approach to the bleeding patient.
8. **Transfer** a stabilized patient to a major center, if necessary, **before the patient’s condition deteriorates.**

GENERAL THERAPEUTIC PRINCIPLES

1. Thorough history taking, physical examination, and judicious laboratory investigation improve the estimation of risks and facilitate planning and preparation to prevent and control blood loss.
2. Optimize red cell mass preoperatively and during pregnancy. Institute early, aggressive treatment of anemia postoperatively or postpartum.
3. Rapid arrest of blood loss must be a management priority. In the face of severe obstetric hemorrhage, early recourse to definitive surgical or obstetric measures to control bleeding can be lifesaving.
4. In uncontrolled hemorrhage, avoid aggressive fluid resuscitation to restore blood pressure to a normal range until bleeding is controlled.
5. Prevent or treat coagulation disorders promptly.
6. Use appropriate intraoperative blood conservation techniques.
7. Minimize the volume of blood drawn for laboratory analysis during the perinatal or perioperative period.
8. Normovolemic anemia can be well tolerated in hemodynamically stable patients.

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1. CLINICAL EVALUATION/PREOPERATIVE PLANNING^{1,2}

A. Medical History and Physical Examination

1. History of Anemia

2. Hereditary or Acquired Bleeding Disorders³⁻⁵

- a. Family and personal history of abnormal hemostasis or bleeding
 - (1) Easy or spontaneous bruising
 - (2) Frequent nosebleeds or unexplained bleeding gums
 - (3) Bleeding after tooth extractions
 - (4) Surgical history, including postoperative bleeding (e.g., after adenoidectomy)
 - (5) Gynecologic history, including menstrual history, especially menorrhagia
 - (6) Obstetrical history, including complications or previous antepartum or postpartum bleeding⁶

3. Coexisting Disease

- a. Identify disease or conditions that may affect coagulation, hematopoiesis, or clinical management (e.g., renal, hepatic, cardiac, or pulmonary)

4. Medication History

- a. Current medications and identification of prescription or nonprescription drugs that may adversely affect hemostasis (e.g., anticoagulants, platelet aggregation inhibitors, preparations containing ASA or NSAIDs, antibiotics, herbal medications)^{7,8}
- b. Drug allergies

5. Physical Examination

- a. Investigate manifestations of disease known to be associated with hemostatic dysfunction (e.g., purpura, petechiae, ecchymosis, hepatomegaly, splenomegaly)

Note: Clinicians should have a high index of suspicion of a bleeding disorder in women with persistent menorrhagia sufficient to cause iron deficiency or a history of bleeding after hemostatic challenge.⁹

B. Selective Laboratory Assessment¹⁰⁻¹⁴

1. Investigation of Anemia

- a. Complete blood count (including red cell indices, reticulocyte count)
- b. Iron status¹⁵ (e.g., serum ferritin, serum transferrin receptor)

2. Assessment of Bleeding Risk

- a. Bleeding risk may be suggested by the medical history, clinical data, current medications, or degree of hemostatic challenge
 - (1) PT, PTT, template bleeding time
 - (2) More detailed coagulation studies to identify clotting disorders, including specific coagulation factor assays (e.g., von Willebrand's disease)

Notes:

1. Specific laboratory tests may be suggested by the medical history and the degree of anticipated hemostatic challenge, e.g., major surgery, childbirth.
2. Women with mild coagulation abnormalities detected in early labor may be at higher risk of postpartum hemorrhage.¹⁶
3. Conditions that increase risks of complications may warrant prompt referral of the patient to a specialist with experience in management without blood transfusion.

2. OPTIMIZE RED BLOOD CELL MASS AND COAGULATION STATUS

A. Correct Hematinic Deficiencies

1. Iron¹⁷⁻²³ (oral/parenteral)

2. Folic Acid²⁴

3. Vitamin B₁₂^{25,26}

Notes:

1. Consider prophylactic hematinic administration.²⁷⁻³¹
2. Bioavailability of oral iron may be improved with concomitant administration of ascorbic acid.^{32,33} Oral iron absorption may be decreased by concurrent use of milk products, egg yolks, coffee, tea, antacids, calcium supplements, or fiber.³⁴
3. Intravenous iron as a total dose infusion in normal saline may replenish iron stores more quickly and efficiently than oral iron therapy.³⁵⁻⁴⁰
4. Parenteral iron should be considered for patients with low iron stores, intolerance to oral iron, inadequate absorption, or chronic or severe blood loss or for patients who are noncompliant.⁴¹ Consider iron sucrose, sodium ferric gluconate complex (or other parenteral iron products) instead of iron dextran to reduce risk of anaphylaxis. If iron dextran is used, administer a test dose.

B. Recombinant Erythropoietin (r-HuEPO) Therapy

1. Preoperative Optimization of RBC Mass

- a. Consider preoperative use of r-HuEPO to optimize red blood cell (RBC) mass in women scheduled for procedures associated with risk of significant blood loss⁴²⁻⁴⁶

2. Erythropoietin Therapy in Pregnancy

- a. r-HuEPO has been administered to pregnant patients in the third trimester to increase RBC mass without any maternal, fetal, or neonatal adverse effects⁴⁷⁻⁵⁴

Notes:

1. Rate of response to r-HuEPO is dose dependent and varies among patients. Poor response may be overcome by dose escalation.^{55,56}
2. Iron deficiency (or other hematinic deficiency) may diminish or delay response to r-HuEPO therapy.⁵⁷⁻⁵⁹ Virtually all patients should receive iron to support erythropoiesis stimulated by multiple-dose r-HuEPO therapy adequately. Consider r-HuEPO and intravenous iron for severely anemic patients.⁶⁰ (See notes following **2.A.**)
3. Monitor for hypertension and consider initiation of or increase in antihypertensive therapy.

C. Minimize Iatrogenic Blood Loss

1. Perform Only Essential Laboratory Tests^{61,62}
2. Coordinate and Consolidate Blood Tests
3. Minimize Volume of Diagnostic Phlebotomy

D. Treat Coexisting Conditions Associated With Blood Loss

1. Menorrhagia/Dysfunctional Uterine Bleeding

- a. History, physical exam, and systematic consideration of possible causes^{63,64}

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b. Appropriate therapy^{65,66} (See also **3.C.**)

2. Nongynecologic Blood Loss

a. Treat other conditions associated with low-grade bleeding (e.g., hemorrhoids, gastrointestinal lesions) that may become clinically important in some surgical patients

E. Management of Coagulation Status⁶⁷

1. Medication Management

a. Discontinue/substitute medications associated with bleeding complications (e.g., aspirin, NSAIDs)
b. Postpone nonurgent surgery for patients on anticoagulant/antiplatelet medications (See also **7.B.9.**)

2. Management of Congenital/Acquired Coagulation Disorders^{68,69}

3. Thrombocytopenia in Pregnancy⁷⁰

a. Maternal

- (1) Gestational thrombocytopenia
 - (i) Monitor platelet counts; no specific therapy indicated
- (2) Drug-induced
 - (i) Review current medications (e.g., heparin, quinine, quinidine, sulfonamides)
- (3) Idiopathic thrombocytopenic purpura⁷¹ (ITP)
 - (i) Corticosteroids
 - (ii) High-dose intravenous immunoglobulin (IVIG)

(iii) Combination of high-dose steroids (e.g., methylprednisolone) and IVIG

(4) Preeclampsia and the HELLP syndrome⁷² (hemolysis, elevated liver enzymes, low platelet count)

(i) Judicious fluid and volume management
(ii) Corticosteroid therapy⁷³⁻⁷⁵

Notes:

1. Mode of delivery should be based on obstetric indications. Cesarean section delivery has not been shown to reduce the risk of intracranial hemorrhage except in very low birth weight infants.⁷⁶
2. Although limited, data supports the safety of epidural anesthesia in patients with platelet counts greater than 100,000/ μ L.

b. Perinatal

- (1) Alloimmune thrombocytopenia⁷⁷ (antenatal)
 - (i) Consider maternal administration of IVIG antepartum⁷⁸
 - (ii) Corticosteroid therapy⁷⁹

4. Anticoagulation Management

- a. If an obstetric or surgical patient is at high risk of thromboembolism or if a diagnosis of thromboembolism is established, consider using low-dose heparin to reduce risk of bleeding, close clinical and laboratory monitoring, or mechanical prophylaxis.⁸⁰ Consider use of low molecular weight heparin⁸¹⁻⁸³ (See also **7.B.9.**)
- b. Consider discontinuation of oral contraceptives at least one month before major elective surgery due to risk of thromboembolic complications

3. MANAGEMENT OF MENORRHAGIA

A. Medical Treatment for Acute Menorrhagia⁸⁴⁻⁸⁸

1. Combination of Medications Used Concomitantly

- a. Conjugated estrogens (i.v.)
- b. High-dose oral contraceptives
 - (1) Combined estrogen and progestin
- c. Tranexamic acid

B. Surgical Management of Menorrhagia⁸⁹

1. Emergency Hysteroscopy and Curettage

2. Uterine Tamponade

a. Consider use of Foley balloon catheter

3. Endometrial Balloon Ablation^{90,91}

4. Hysterectomy (See also **5.A.** to **5.H.**)

C. Medical Therapy for Chronic Menorrhagia⁹²

1. Tranexamic Acid⁹³⁻⁹⁵ (alone or with desmopressin^{96,97})

2. Prostaglandin Inhibitors/NSAIDs⁹⁸

Note: NSAIDs may cause gastrointestinal (GI) bleeding.

3. Combined Oral Contraceptives

4. Progesterone-Releasing Intrauterine Device⁹⁹⁻¹⁰¹

Note: Because of their mechanism of action, such devices may not be acceptable to some patients as a contraceptive.

5. Danazol¹⁰²⁻¹⁰⁴

6. GnRH Analogue^{105,106} (preoperative treatment)

4. OBSTETRIC HEMORRHAGE

Be prepared to use a combination of appropriate interventions to prevent or manage hemorrhage. Timely recognition and decisive action (e.g., hysterectomy) is essential for minimizing blood loss. Management of hemorrhagic emergencies requires a coordinated team effort with close cooperation between the obstetrician, anesthesiologist, pediatrician, hematologist, and other specialists.

A. Antepartum Hemorrhage¹⁰⁷

1. General Management¹⁰⁸

a. Clinical assessment (avoiding pelvic examination until a diagnosis of placenta previa is excluded). Evaluate maternal hemodynamic status and fetal health

b. Judicious fluid resuscitation for hypovolemia (See **6.**)

c. Prompt testing to facilitate diagnosis (e.g., ultrasound to confirm placenta previa) and assess fetal status^{109,110}

d. Management should be appropriate to the degree of severity, maternal and fetal health status, and gestational age of the fetus, with a lower threshold for intervention in patients who decline blood transfusion

e. Consider antepartum corticosteroid administration to women at risk of preterm delivery¹¹¹ (for fetal lung maturation)

f. Consider Rh immune globulin prophylaxis for at-risk mothers

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g. If fetal blood monitoring is indicated, consider noninvasive or minimally-invasive fetal blood diagnostic measures to reduce or avoid invasive testing that may result in accidental maternal, fetal, or placental bleeding¹¹²⁻¹¹⁵

h. If preterm delivery or complications are anticipated, consult neonatologist with experience in clinical management without transfusion and pursue appropriate management as indicated (e.g., antibiotic prophylaxis, tocolytic therapy)

2. Ectopic (Extrauterine) Pregnancy¹¹⁶

a. Medical management¹¹⁷⁻¹²⁰

b. Surgical management

(1) Ensure availability of blood salvage equipment¹²¹⁻¹²³

(2) Hysterectomy may be indicated for certain forms of ectopic pregnancy (e.g., interstitial, cervical)

3. Miscarriage (Spontaneous Abortion)¹²⁴

a. Inevitable and incomplete abortion

(1) Prompt evacuation of the uterus

(2) Administration of oxytocic drug (See **4.C.6.**)

(3) Consider prophylactic angiographic embolization¹²⁵

b. Missed abortion

(1) Standard management with appropriate precautions to facilitate rapid control of hemorrhage

4. Placenta Previa

a. Early and aggressive erythropoietin therapy (antepartum/postpartum) (See **2.B.**)

b. Active treatment (rather than expectant) if there is sustained bleeding, if labor begins, or when the patient reaches 37-38 weeks gestation

c. Ensure availability of blood salvage equipment for cesarean section delivery (See **4.E.1.**)

5. Placental Abruption

a. For mild/moderate (grade I/II) placental abruption, standard management, as appropriate

b. For severe (grade III) placental abruption

(1) Supplemental oxygen administration

(2) Judicious fluid resuscitation

(3) Immediate amniotomy to induce or expedite labor followed by carefully monitored oxytocin infusion

(4) Consider early use of aprotinin to treat uterine inertia, expedite delivery, and preempt onset of coagulopathy¹²⁶⁻¹²⁹ (See also **4.E.3.** and **7.B.5.**)

(5) Anticipate postpartum hemorrhage

Note: Cesarean section delivery is seldom indicated (e.g., fetal distress, unripe cervix preventing amniotomy, failed induction).

B. Postpartum Hemorrhage (PPH) Risk Factors¹³⁰⁻¹³⁵

1. Previous Obstetric Hemorrhage

E.g., previous history of postpartum hemorrhage, manual removal of the placenta or retained products

2. Abnormalities of Uterine Contraction

E.g., overdistended uterus (associated with multiple gestation, polyhydramnios, macrosomia), high parity, rapid or prolonged labor

3. Abnormalities of Placentation

E.g., placenta previa, placenta accreta/increta/percreta, placental abruption

4. Retained Products of Conception

E.g., previous uterine surgery, high parity, abnormal placenta observed prenatally on diagnostic imaging, incomplete placenta at delivery

5. Genital Tract Trauma

E.g., previous uterine surgery, especially previous cesarean section, breech presentation, mid-forceps extraction or forceps rotation, lacerations at cesarean section

6. Abnormalities of Coagulation

E.g., congenital or acquired bleeding disorder (such as ITP, HELLP, DIC), therapeutic anticoagulation

7. Other Factors

Emergency or elective cesarean section delivery, preeclampsia or gestational hypertension, nulliparity, obesity

C. Active Management of Third Stage of Labor¹³⁶⁻¹⁴¹

1. Thirty Seconds to Deliver Anterior Shoulder

Note: Unhurried delivery of the infant, allowing for slow uterine retraction, may facilitate smooth placental separation.

2. Immediate Prophylactic Oxytocin Analogue^{142,143} (i.v.)

Note: Early oxytocin administration is associated with lower rates of PPH compared with administration after the third stage.¹⁴⁴

3. Thirty Seconds to Deliver Posterior Shoulder

4. Deliver Body Slowly, Head Down

5. Deliver Placenta by Controlled Cord Traction¹⁴⁵

a. Displace uterus upwards by suprapubic pressure

6. Repeat Oxytocic Drug

a. Oxytocin (or syntometrine)¹⁴⁶⁻¹⁴⁸ (i.v.)

b. Ergot derivative¹⁴⁹⁻¹⁵² (i.m./intramyometrial/i.v.)

c. Prostaglandin analogues (alone or following other oxytocic agent)

(1) Carboprost^{153,154} (intramyometrial injection/i.v.)

(2) Misoprostol¹⁵⁵⁻¹⁶³ (rectal/oral)

Notes:

1. Observe precautions about administration of carboprost to patients with asthma.

2. Observe precautions about administration of ergot derivatives to patients with hypertension.

3. Misoprostol may be more stable than other oxytocic drugs in tropical climates.

7. Examine Placenta for Completeness

a. Inspect placenta for indications of retained tissue or abnormal placentation

8. Monitor Closely Following Delivery¹⁶⁴

a. Maintain close observation, uterine palpation, assessment of bleeding, and sequential vital signs for 2-3 hours or longer following delivery

9. Never Leave a Bleeding Postpartum Patient

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D. Control of Postpartum Hemorrhage¹⁶⁵⁻¹⁷⁰

1. Continuous Uterine Massage

Notes:

1. Fundal massage may be performed by assistants or the woman herself.
2. Elevation of uterus may improve venous drainage.

2. Transabdominal Aortic Compression^{171,172}

3. Bimanual Uterine Compression

4. Stimulate Nipple Manually/Suckle Infant^{173,174}

Note: This may stimulate release of endogenous oxytocin in settings where oxytocic drugs are not available.

5. Repeat/Continue Oxytocic Drug(s)

- a. Oxytocin¹⁷⁵ (or syntometrine) (i.v.)
- b. Ergot derivative (i.v.)
- c. Prostaglandin analogues
 - (1) Carboprost¹⁷⁶⁻¹⁸⁰ (intramyometrial/i.v.)
 - (2) Misoprostol^{181,182} (rectal)

Note: Consider use of prostaglandin analogue in patients unresponsive to oxytocin/ergot (or syntometrine).

6. Judicious Fluid Resuscitation

- a. Nonblood volume expanders, warmed if possible (See 6.)

7. Drain Bladder With Catheter

8. Prompt Extraction of Placenta or Retained Fragments

- a. Manual exploration and removal (See Note 2 below.)

9. Repair Lacerations

E.g., uterus, cervix, vagina, or perineum

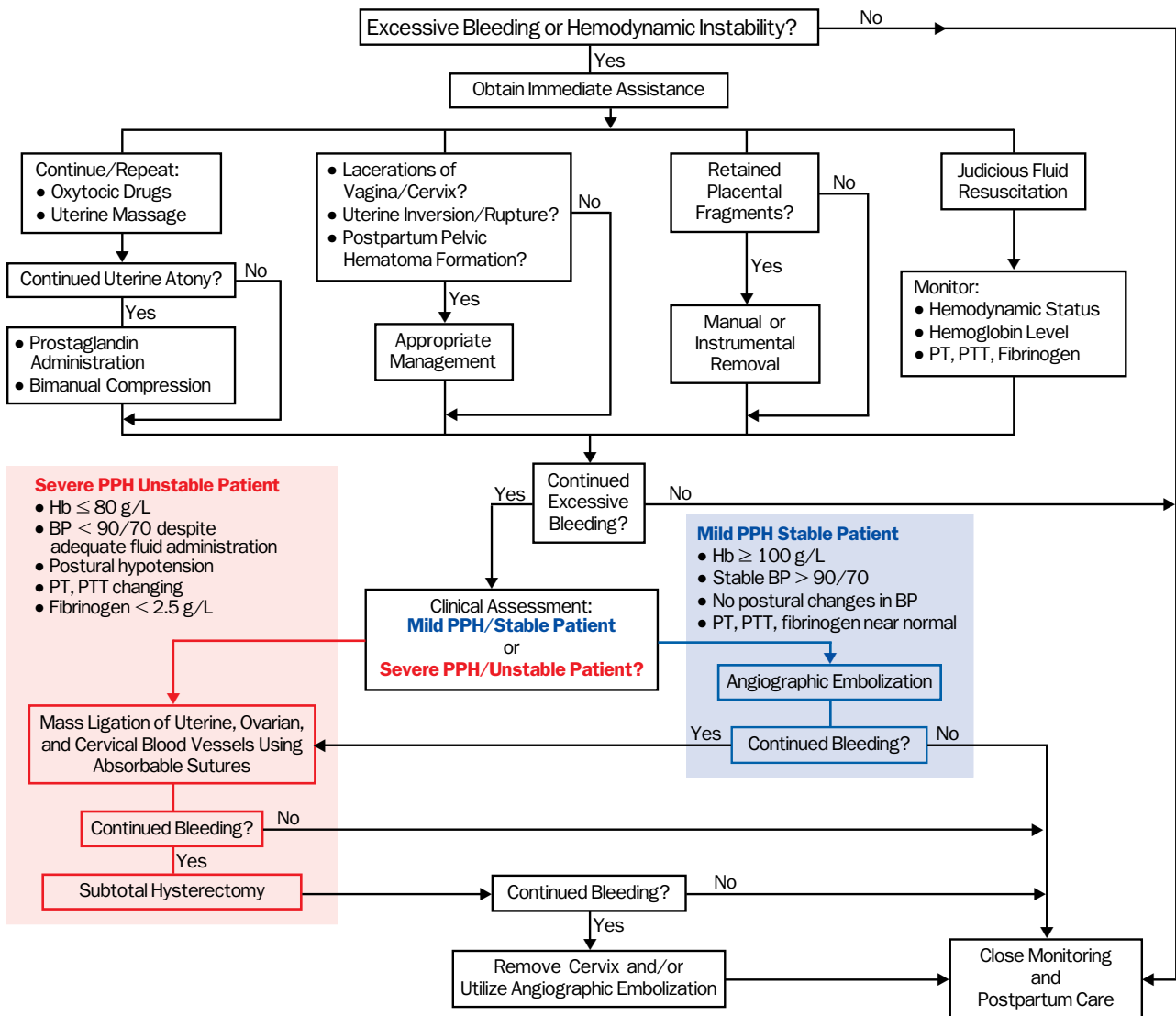
10. Uterine Tamponade

- a. Tamponade should not delay preparation for definitive surgical intervention (See Note 3 below.)

(1) Intrauterine balloon¹⁸³ (Rüsch¹⁸⁴ catheter or Sengstaken-Blakemore^{185,186} tube)

(2) Uterine packing¹⁸⁷⁻¹⁸⁹ (See also 5.1.6.)

ALGORITHM FOR NONBLOOD MANAGEMENT OF POSTPARTUM HEMORRHAGE (PPH)



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11. Expedient Angiographic Embolization

- a. If angiography services can be accessed expeditiously, consider embolization for continuing slow blood loss that is not immediately life threatening in the hemodynamically stable patient¹⁹⁰⁻¹⁹⁴

12. Major Vessel Ligation

- a. Bilateral high/low uterine and ovarian artery ligation¹⁹⁵⁻¹⁹⁷
 - (1) Consider internal iliac artery ligation¹⁹⁸

Note: Internal iliac artery ligation has a lower rate of success than other surgical options for control of PPH. Arterial ligation, particularly hypogastric artery ligation, may preclude embolization.

- b. Single-stage uterine “devascularization”^{199,200}

13. Subtotal/Total Hysterectomy²⁰¹⁻²⁰⁷

14. Pelvic Cavity Pressure Pack^{208,209} or Angiographic Embolization²¹⁰⁻²¹²

- a. Consider for posthysterectomy hemorrhage (See also **5.I.6.**)

Notes:

1. If postpartum hemorrhage is suspected, transfer the patient to the OR and prepare for surgery immediately. **Obtain assistance**, e.g., a second obstetric, gynecologic, or general surgeon, an anesthesiologist, and/or associated staff. **Perform systematic inspection** under anesthesia with adequate lighting, starting from external genitalia, vaginal walls, and cervix, and **suture any lacerations regardless of size, whether bleeding or not.**
2. If retained placental fragments are suspected, consider i.v. nitroglycerin for *transient* uterine relaxation to facilitate manual removal of the placenta.²¹³⁻²¹⁶ Hypovolemia should be corrected with intravenous fluids before administering nitroglycerin. Hemodynamic monitoring, a running i.v. infusion, and immediately available ephedrine are mandatory.²¹⁷
3. **Temporizing or conservative measures to control blood loss** or preserve the uterus **should not delay definitive surgical intervention** to stop bleeding.²¹⁸ The immediate presence of an experienced obstetric surgeon is important to make an early decision to operate before the patient’s condition deteriorates.
4. It is vital to identify and control the source of bleeding expeditiously. The choice of procedure to arrest hemorrhage depends on the availability of facilities and personnel, the extent of hemorrhage, and the condition of the patient.²¹⁹
5. Perform frequent serial clinical monitoring. Assess hematocrit, coagulation profile, blood pressure, pulse, respiratory rate, core temperature, urine output, volume status (e.g., serial central venous pressure measurements, if indicated). In patients with unstable vital signs and little or no external bleeding, suspect concealed hemorrhage in the uterus or a pelvic hematoma.²²⁰
6. Do not delay operative intervention (or reexploration) for postpartum or postoperative hemorrhage that can be controlled surgically.²²¹ Early recourse to hysterectomy may be lifesaving. A subtotal hysterectomy requires less time to perform, thus reducing blood loss.

7. If there is delay before surgical intervention or transport, consider temporary uterine tamponade; use of an antishock garment (MAST) for temporary moderate circulatory support; external aortic compression; direct pressure for lacerations in the perineum, cervix, or vagina; and judicious fluid resuscitation.²²²
8. If gauze packing is used, care should be taken to pack the fundus tightly and systematically to avoid creating space where blood may accumulate.²²³
9. Although intrauterine use of Foley catheters for control of hemorrhage after vaginal delivery has been reported,²²⁴ the volume of the immediate postpartum uterine cavity may be too large for effective tamponade.
10. Consider antibiotic prophylaxis.
11. Consider low molecular weight heparin for thromboembolic prophylaxis.

E. Other Hemorrhagic Emergencies

1. Cesarean Section

Note: Delivery by cesarean section is associated with an increased risk of PPH compared to vaginal birth and should be weighed carefully in women who decline blood transfusion.

- a. Arrange for presence of skilled personnel and appropriate equipment and drugs
- b. Appropriate surgical technique²²⁵⁻²²⁹
 - (1) Misgav Ladach (modified Cohen-Stark)²³⁰⁻²³⁶
 - (2) Pelosi²³⁷⁻²⁴⁰
- c. Intraoperative blood salvage (autotransfusion)²⁴¹⁻²⁴⁴ (See also **4.E.2.g.** and **5.G.**)
 - (1) Employ appropriate cell washing and filtration²⁴⁵⁻²⁴⁷
- d. Acute normovolemic hemodilution²⁴⁸ (ANH)
- e. Aggressive management of PPH (See **4.D.**)
 - (1) Oxytocic drugs²⁴⁹⁻²⁵¹
- f. Maintenance of normothermia²⁵²
- g. Prevention and prompt treatment of infection²⁵³

2. Abnormal Placentation^{254,255}

(Placenta Previa and/or Accreta/Increta/Percreta)

- a. Meticulous planning
 - (1) Optimal management includes detailed planning and collaboration of a multidisciplinary operative team prepared and equipped to prevent and manage hemorrhagic emergencies^{256,257}
- b. Early antenatal detection of abnormalities of placentation
 - (1) Employ ultrasound or magnetic resonance imaging²⁵⁸⁻²⁶⁵
- c. Prophylactic embolotherapy²⁶⁶⁻²⁷⁰ or intra-arterial balloon occlusion²⁷¹⁻²⁷³
- d. Consider erythropoietin therapy (See **2.B.**)
- e. Planned elective delivery and presence of skilled personnel and appropriate equipment and drugs
- f. Acute normovolemic hemodilution²⁷⁴

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g. Intraoperative blood salvage²⁷⁵⁻²⁷⁷
(See also **4.E.1.c.** and **5.G.**)

3. Disseminated Intravascular Coagulation (DIC)^{278,279}

- a. Anticipation/preparation
 - (1) Obstetric causes of DIC include placental abruption, intrauterine fetal death, amniotic fluid embolism, gram-negative infections, and eclampsia
 - (2) Prolonged hypovolemia or hypothermia may trigger or exacerbate DIC

b. Urgent medical/hematological consultation

c. Identify and promptly treat the underlying process triggering the coagulopathy (e.g., early evacuation of the uterus for placental abruption) (See **4.A.5.b.**)

d. Consider use of recombinant activated factor VII (r-FVIIa)²⁸⁰ or clotting factor concentrates

e. Consider use of cryoprecipitate

5. PERIOPERATIVE BLOOD CONSERVATION²⁸¹⁻²⁸⁵

A. Preoperative Planning

1. Tolerance of Anemia

- a. Moderate levels of normovolemic anemia are well tolerated in hemodynamically stable patients²⁸⁶⁻²⁹⁰
- b. 10/30 rule for minimum hemoglobin/hematocrit level has no scientific basis^{291,292}

2. Optimization of Preoperative RBC Mass

- a. If the expected surgical blood loss may render the patient significantly anemic, consider preoperative optimization of RBC mass as part of an overall blood conservation plan (See **1.** and **2.**)

B. Operative Approaches to Minimize Blood Loss

1. Meticulous Hemostasis and Surgical Technique²⁹³⁻²⁹⁵

- a. Minimize duration of surgery/use enlarged surgical team
- b. Review and rehearse procedures for managing massive hemorrhage
- c. Ensure availability of equipment to manage contingencies
- d. Consider combining techniques for hemostasis

2. Hemostatic Surgical Instruments

- a. Electrosurgery/electrocautery²⁹⁶⁻²⁹⁸
- b. Laser
- c. Microwave hemostatic instruments²⁹⁹⁻³⁰¹
- d. Ultrasonic scalpel³⁰²⁻³⁰⁵

3. Mechanical Occlusion of Blood Vessels

- a. Prophylactic use of ligation, vascular clips, clamps^{306,307}

C. Angiographic Embolization³⁰⁸⁻³¹⁶

- a. Consider prophylactic embolization for patients with high bleeding risk³¹⁷ (See also **4.E.2.c.** and **5.I.7.**)

D. Minimally Invasive Approaches^{318,319}

1. Endometrial Resection/Ablation³²⁰⁻³²²

- a. Electrosurgical ball³²³⁻³²⁵
- b. Laser^{326,327}
- c. Thermal balloon³²⁸⁻³³¹

E. Spinal/Epidural/General Anesthesia³³²⁻³³⁶

Note: Regardless of the choice of anesthesia (general, regional), the anesthetic technique must be well planned and executed so as to minimize blood loss.

F. Intraoperative Hemodilution³³⁷⁻³⁴⁰

G. Blood Cell Salvage³⁴¹⁻³⁴⁷

(See also **4.E.1.c.** and **4.E.2.g.**)

1. Oncologic Surgery^{348,349}

- a. For oncologic surgery with potential for high blood loss, consider use of blood salvage with leukocyte depletion filters^{350,351} or intraoperative blood irradiation³⁵²

H. Controlled Hypotensive Anesthesia³⁵³

I. Management of Surgical Hemorrhage/Shock³⁵⁴

1. Immediate Control of Hemorrhage³⁵⁵

- a. Apply direct pressure. Call for assistance

2. Elevate Legs/Apply Blood Pressure Cuffs

3. Administer Oxygen

4. Consider Permissive Hypotension

- a. Until bleeding is controlled, consider use of permissive hypotension (moderate fluid therapy sufficient to maintain minimally acceptable perfusion) to avoid normalization of blood pressure, which may accelerate hemorrhage^{356,357} (See also **6.C.** Note 2.)
- b. Assess volume status by evaluating clinical signs, including mean arterial pressure, heart rate, and urine output, and by appropriate monitoring (e.g., central venous line, pulmonary artery catheter)

5. Prompt Laparoscopy/Surgery/Vascular Ligation

6. Pelvic Packing³⁵⁸⁻³⁶⁰ (See also **4.D.14.**)

Note: Packing or balloon catheterization may be used as a temporary measure while awaiting prompt surgical intervention to arrest bleeding or to tamponade persistent postoperative hemorrhage. Avoid premature packing for repairable major vascular injury.

7. Angiographic Arterial Embolization³⁶¹⁻³⁶³

Notes:

- 1. Insert two large-bore intravenous catheters prior to major surgery and warm i.v. fluids to avoid hypothermia and coagulopathy.³⁶⁴
- 2. Laparoscopic surgery should be undertaken by an experienced operator with a rehearsed plan of action for immediate control of hemorrhage.
- 3. During the early postoperative hours, allow a slow, gradual return to normal blood pressure after bleeding is controlled.³⁶⁵
- 4. Consider postoperative serial monitoring of vital signs, urine output, and hemoglobin/hematocrit to facilitate early clinical recognition of ongoing bleeding.
- 5. Angiographic embolization should be undertaken by an experienced interventional radiologist when bleeding is not immediately life threatening.

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6. NONBLOOD VOLUME EXPANDERS

A. Crystalloids

1. Ringer's Lactate
2. Normal Saline
3. Hypertonic Saline

B. Colloids

1. Pentastarch^{366,367}/Hetastarch^{368,369}
2. Gelatin
3. Dextran (See Note 7.)

C. Oxygen-Carrying Red Cell Substitutes

(when available for clinical use)

1. Perfluorocarbon Solutions
2. Hemoglobin-Based Oxygen Carriers

Notes:

1. In active bleeding or oozing, take prompt measures to arrest hemorrhage.
2. In the actively bleeding patient, excessive fluid resuscitation may accelerate hemorrhage by dilution of coagulation factors or disruption of clots. In uncontrolled hemorrhage, adequate perfusion can be achieved with moderate underresuscitation and permissive mild hypotension (in patients without severe head injuries).³⁷⁰

3. Avoid circulatory overload, especially in severely anemic patients. Closely monitor fluid balance and vital signs.
4. Maintain a high index of suspicion of bleeding when a patient shows evidence of hypovolemia despite reasonable hydration. Poor response to fluid therapy may indicate continuing hemorrhage.
5. Pentastarch and low molecular weight starches or starches in balanced electrolyte solutions may be more effective in optimizing macro- and microcirculation than crystalloids and other colloids. High molecular weight hydroxyethyl starches may increase the risk of bleeding in surgical patients who have either congenital or acquired coagulation abnormalities.^{371,372}
6. Administration of large volumes of saline-based fluids has been associated with coagulation abnormalities, decreased urine output, and a hyperchloremic metabolic acidosis that may be clinically relevant.^{373,374}
7. Dextrans should be avoided in obstetrics due to their anticoagulant effects and risk of anaphylactic reactions. In surgical patients, the anticoagulant effect of dextrans may be partially offset by desmopressin.³⁷⁵

7. PHARMACOLOGICAL ENHANCEMENT OF HEMOSTASIS

A. Topical Hemostatic Agents

1. Tissue Adhesives/Fibrin Glue³⁷⁶⁻³⁷⁸
2. Collagen Hemostat (e.g., Avitene®, Instat®)
3. Oxidized Cellulose (e.g., Oxycel®, Surgicel®)
4. Gelatin Foam/Sponges (e.g., Gelfoam®, Surgifoam®)
5. Vasopressin³⁷⁹⁻³⁸¹
 - a. Vasopressin infiltration or soaked uterine packing^{382,383} (See also 5.I.6.)
6. Thrombin³⁸⁴

B. Systemic Hemostatic Agents

1. Vitamin K^{385,386}

Note: Consider prophylactic treatment of patients with vitamin K before invasive procedures.

2. Tranexamic Acid³⁸⁷⁻³⁸⁹
3. Epsilon-Aminocaproic Acid³⁹⁰
4. Desmopressin³⁹¹⁻³⁹⁴
5. Aprotinin^{395,396} (Use test dose.)

Notes:

1. Aprotinin or desmopressin may be used for control of bleeding due to drug-induced platelet dysfunction (e.g., due to ASA, NSAIDs, beta-lactam antibiotics, and antithrombotics).^{397,398}
 2. Use of aprotinin has been reported for the management of obstetric bleeding due to fibrinolysis associated with severe placental abruption.^{399,400}
 6. Conjugated Estrogens^{401,402}
 7. Recombinant Activated Factor VII (r-FVIIa)⁴⁰³⁻⁴⁰⁵ (e.g., eptacog alfa [activated], NovoSeven®, NiaStase®)
 8. Coagulation Factor Replacement Therapy⁴⁰⁶
 - a. Consider recombinant preparations of factors VIIa, VIII, IX^{407,408}
 9. Prothrombin Complex Concentrate⁴⁰⁹
- Note: Prothrombin complex concentrate and i.v. vitamin K have been used for urgent reversal of anticoagulation in surgical patients.^{410,411} Consider protamine for reversal of heparin anticoagulation.
10. Cryoprecipitate

8. MANAGEMENT OF PROFOUND ANEMIA⁴¹²⁻⁴¹⁶

A. Stop Any Bleeding

1. Permissive Mild Hypotension
 - a. Avoid aggressive normalization of blood pressure in the bleeding patient
2. Surgical Control of Bleeding
 - a. Do not defer operative intervention if active bleeding can be controlled surgically, even if the patient is anemic
3. Maintain/Restore Normothermia

B. Restrict Diagnostic Phlebotomy⁴¹⁷

C. Maximize Oxygen Delivery

1. Optimize Circulating Volume
2. Supplemental Oxygen/Hyperbaric Oxygen Therapy^{418,419}

D. Minimize Oxygen Consumption

1. Analgesia and Sedation
2. Mechanical Ventilation

E. Stimulate Red Cell Production⁴²⁰⁻⁴²⁶ (See 2.A. and 2.B.)

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Database Abbreviations:

PMID: Medline®/PubMed® Unique Identifier (Index Medicus)

EMBASE: Embase® Accession Number (Excerpta Medica)

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