## Anticoagulation in Pregnancy

## **Background**

Pregnancy is a time of increased risk for venous thromboembolism (VTE), secondary to the physiologic and anatomic changes of pregnancy. The baseline risk for a VTE in pregnancy is about 1 per 1,000 - 1,500 pregnancies. However, if additional risk factors are present, the risk is greater. (Table 1)

## **Medications**

- Heparin
- Lovenox
- Coumadin
- Fondaparinux

## Medical Intervention

- 1. Prophylaxis aim is to prevent development of VTE
  - a. Heparin
    - i. 5,000 u SQ bid, first trimester
    - ii. 7,500 u SQ bid, second trimester
    - iii. 10,000 u SQ bid, third trimester
  - b. Lovenox
    - i. 40 mg SQ once daily
- Prophylaxis is typically started in the first trimester after confirmation of a viable, intrauterine gestation by ultrasound, and is continued for six weeks postpartum.
- Therapeutic treatment of an existing VTE, or prevention of VTE in patients with exceptional risk factors
  - a. Heparin
    - i. 10,000 u or more, every 8 12 hours, until aPTT at desired level (Note: this can be very difficult to achieve and is not preferred for therapeutic treatment)
  - b. Lovenox
    - i. 1 mg/kg every 12 hours, usually rounded off in 10 mg increments
  - c. Coumadin
    - i. Typically reserved for use in pregnancy for patients with mechanical heart valves with exceptionally high-risk of coagulation complication.
  - d. Fondaparinux

i. Typically reserved for patients who have an absolute contraindication to the use of Heparin and Lovenox e.g. allergy, heparin induced thrombocytopenia (HIT).

## **Indications**

- See Table 3 ACOG Practice Bulletin #196. July 2018 (See Appendix A)

## Antepartum Management

- Notify the Anesthesia service if a patient is admitted to L&D and is currently treated with anticoagulation medication.
- Use of compression devices for lower extremities, may be placed prior to cesarean delivery.
- 1. Heparin
  - a. Prophylaxis dosing (10,000 units [or less] bid
- Hold 12 hours prior to admission for delivery
- Obtain CBC, PTT level at time of admission to assess coagulation status
- If Platelet count is normal, and PTT is not elevated, patient may receive regional anesthesia (spinal, epidural).
  - b. Therapeutic dosing (> 20,000 units in 24 hours)
- Hold 24 hours prior to admission for delivery
- Obtain CBC, PTT as noted above.

## 2. Lovenox

- a. Prophylactic dosing (40 mg/d)
- Hold 12 hours prior to admission for delivery
- Obtain CBC for platelet count
  - b. Therapeutic dosing (1 mg every 12 hours)
- Hold 24 hours prior to admission for delivery
- Obtain CBC.

## Intrapartum Management (See Appendix B and C)

1. Vaginal Delivery (See ANMC Obesity in Pregnancy guideline)

-use appropriately sized blood pressure cuff -anticipate higher dose of oxytocin for induction or augmentation of labor -consider placement of prophylactic epidural catheter If BMI <u>></u> 40 kg/M<sup>2</sup> at any point during pregnancy: -Consider early IUPC and FSE

If BMI **<u>></u> 50 kg/M<sup>2</sup>** at any point during pregnancy: -Anesthesia consult on admission

2. Cesarean Delivery (See ANMC Cesarean Thromboprophylaxis Management guideline)

A. Baseline intervention

-Early ambulation

-Intermittent pneumatic compression devices

- Consider consultation with PT to assist in early mobilization of super-obese patients

B. Pharmacologic thromboprophylaxis Begin pharmacologic therapy if:

- previous personal history of deep venous thrombosis or pulmonary embolism
- personal history of an inherited thrombophilia

Treatment will consist of:

BMI < 40 kg/M2

Enoxaparin (Lovenox) 40 mg subcutaneous q 12 hrs, starting 12 hrs post op

BMI  $\geq$  40 and <50 kg/M2 Enoxaparin (Lovenox) 40 mg subcutaneous q 12 hrs, starting 12 hrs post op

BMI <u>></u>50 kg/M2 Enoxaparin (Lovenox) 60 mg subcutaneous q 12 hrs, starting 12 hrs post op

# Postpartum Management

- Early ambulation
- Continue pneumatic compression devices until ambulatory
- Epidural catheter should be removed four hours (or more) prior to restarting anticoagulation medication
- Restart prophylaxis-level anticoagulation about 12 hours after delivery
  - Lovenox 40 mg/d
  - Heparin 5,000 u bid
- Restart therapeutic-level anticoagulation about 24 hours after delivery
  - Lovenox 1 mg/kg every 12 hours
  - o Suggest re-weighing the patient after delivery
  - If therapeutic anticoagulation dosing is essential prior to 24 hours after delivery, suggest using Heparin instead of Lovenox

## **References**

Thromboembolism in pregnancy. ACOG Practice Bulletin No. 196. American College of Obstetricians and Gynecologists. Obstet Gynecol 2018;132:e1—17. (Re-affirmed 2022, Accessed 12/26/24)

Inherited thrombophilias in pregnancy. ACOG Practice Bulletin No. 197. American College of Obstetricians and Gynecologists. Obstet Gynecol 2018;132:e18–34. (Re-affirmed 2022, Accessed 12/26/24)

Leffert, Lisa; Butwick, Alexander; Carvalho, Brendan; Arendt, Katherine; Bates, Shannon; members of the SOAP VTE Taskforce The Society for Obstetric Anesthesia and Perinatology Consensus Statement on the Anesthetic Management of Pregnant and Postpartum Women Receiving Thromboprophylaxis or Higher Dose Anticoagulants, Anesthesia & Analgesia: March 2018 - Volume 126 - Issue 3 - p 928-944 doi: 10.1213/ANE.00000000002530. https://journals.lww.com/anesthesia-analgesia/pages/articleviewer.aspx?year=2018&issue=03000&article=00033&type=Fulltext (Accessed 12/26/24)

> Reviewed 12/16/24 njm Revised 10/18/22 njm Approved 3/16/20 dd/njm

	Prevalence in General Population (%)	VTE Risk Per Pregnancy (No History) (%)	VTE Risk Per Pregnancy (Previous VTE) (%)
Factor V Leiden heterozygote	1–15	0.5–3.1	10
Factor V Leiden homozygote	<1	2.2–14.0	17
Prothrombin gene heterozygote	2–5	0.4–2.6	>10
Prothrombin gene homozygote	<1	2–4	>17
Factor V Leiden/ prothrombin double heterozygote	0.01	4–8.2	>20
Antithrombin deficiency	0.02	0.2–11.6	40
Protein C deficiency	0.2-0.4	0.1–1.7	4–17
Protein S deficiency	0.03-0.13	0.3-6.6	0-22

Table 1. Risk of Venous Thromboembolism With Different Inherited Thro

Abbreviation: VTE, venous thromboembolism.

1. Franco RF, Reitsma PH. Genetic risk factors of venous thrombosis. Hum Genet 2001;109:369-84

 Gerhardt A, Scharf RE, Beckmann MW, Struve S, Bender HG, Pillny M, et al. Prothrombin and factor a history of thrombosis during pregnancy and the puerperium. N Engl J Med 2000;342:374–80.

3. Zotz RB, Gerhardt A, Scharf RE. Inherited thrombophilia and gestational venous thromboer Haematol 2003;16:243-59.

4. Haverkate F, Samama M. Familial dysfibrinogenemia and thrombophilia. Report on a study of Fibrinogen. Thromb Haemost 1995;73:151-61.

5. Friederich PW, Sanson BJ, Simioni P, Zanardi S, Huisman MV, Kindt I, et al. Frequency of thromboembolism in anticoagulant factor-deficient women: implications for prophylaxis [publi Intern Med 1997;127:1138]. Ann Intern Med 1996;125:955–60.

6. Vossen CY, Preston FE, Conard J, Fontcuberta J, Makris M, van der Meer FJ, et al. Hereditary a prospective follow-up study. J Thromb Haemost 2004;2:592–6.

7. Paidas MJ, Ku DH, Lee MJ, Manish S, Thurston A, Lockwood CJ, et al. Protein Z, protein S leve thrombophilia and subsequent pregnancy complications. J Thromb Haemost 2005;3:497–501.

8. Dykes AC, Walker ID, McMahon AD, Islam SI, Tait RC. A study of Protein S antigen levels in 3788 of age, sex and hormone use, and estimate for prevalence of deficiency state. Br J Haematol 200

9. Goodwin AJ, Rosendaal FR, Kottke-Marchant K, Bovill EG. A review of the technical, di considerations for protein S assays. Arch Pathol Lab Med 2002;126:1349–66.

10. Bates SM, Greer IA, Middeldorp S, Veenstra DL, Prabulos AM, Vandvik PO. VTE, thrombophilia, pregnancy: Antithrombotic Therapy and Prevention of Thrombosis, 9th ed: American College of Based Clinical Practice Guidelines. Chest 2012;141:e6915–736S.

Inherited thrombophilias in pregnancy. ACOG Practice Bulletin No. 197. American College of Obstetricians and Gynecologists. Obstet Gynecol 2018;132:e18–34. (Re-affirmed 2022)

Appendix A

# Table 3. Recommended Pharmacologic Thromboprophylaxis in Pregnancy and the Postpartum Period

Clinical Scenario	Antepartum Management	Postpartum Management
No history of VTE, no thrombophilia	Surveillance <sup>*</sup> without anticoagulation therapy	Surveillance without anticoagulation therapy or postpartum prophylactic anticoagulation therapy if the patient has multiple risk factors. <sup>†</sup>
VTE diagnosed during pregnancy	Adjusted-dose LMWH/UFH	Adjusted-dose LMWH/UFH for a minimum of 6 weeks postpartum. Longer duration of therapy may be indicated depending on the timing of VTE during pregnancy, prior VTE history, or presence of a thrombophilia. Oral anticoagulants may be considered postpartum based upon planned duration of therapy, lactation, and patient preference.
Single provoked VTE (precipitated by a specific event such as surgery, trauma, or immobility) unrelated to estrogen or pregnancy due to a transient (resolved) risk factor, no thrombophilia	Surveillance <sup>*</sup> without anticoagulation therapy	Surveillance without anticoagulation therapy or postpartum prophylactic anticoagulation therapy if the patient has additional risk factors. <sup>4</sup>
History of single unprovoked VTE (no identified precipitating factor present; includes prior VTE in pregnancy or associated with hormonal contraception), not on long-term anticoagulation	Prophylactic, intermediate- dose, or adjusted-dose LMWH/UFH	Prophylactic, intermediate-dose, or adjusted- dose LMWH/UFH regimen for 6 weeks postpartum
Low-risk thrombophilia <sup>‡</sup> without previous VTE	Surveillance <sup>*</sup> without anticoagulation therapy	Surveillance without anticoagulation therapy or postpartum prophylactic anticoagulation therapy if the patient has additional risks factors <sup>4</sup>
Low-risk thrombophilia <sup>‡</sup> with a family history (first-degree relative) of VTE	Surveillance <sup>*</sup> without anticoagulation therapy or prophylactic LMWH/UFH	Postpartum prophylactic anticoagulation therapy or intermediate-dose LMWH/UFH
Low-risk thrombophilia <sup>‡</sup> with a single previous episode of VTE—Not receiving long-term anti- coagulation therapy	Prophylactic or intermediate-dose LMWH/ UFH	Postpartum prophylactic anticoagulation therapy or intermediate-dose LMWH/UFH
High-risk thrombophilia <sup>§</sup> without previous VTE	Prophylactic or intermediate-dose LMWH/ UFH	Postpartum prophylactic anticoagulation therapy or intermediate-dose LMWH/UFH
High-risk thrombophilia <sup>§</sup> with a single previous episode of VTE or an affected first-degree relative—Not receiving long-term anti- coagulation therapy	Prophylactic, intermediate- dose, or adjusted-dose LMWH/UFH	Postpartum prophylactic anticoagulation therapy, or intermediate or adjusted-dose LMWH/UFH for 6 weeks (therapy level should be equal to the selected antepartum treatment)
Two or more episodes of VTE—Not receiving long-term anticoagulation therapy (regardless of thrombophilia)	Intermediate-dose or adjusted-dose LMWH/UFH	Postpartum anticoagulation therapy with intermediate-dose or adjusted-dose LMWH/UFH for 6 weeks (therapy level should be equal to the selected antepartum treatment)
Two or more episodes of VTE—Receiving long- term anticoagulation therapy (regardless of thrombophilia)	Adjusted-dose LMWH or UFH	Resumption of long-term anticoagulation therapy. Oral anticoagulants may be considered postpartum based upon planned duration of therapy, lactation, and patient preference.

Abbreviations: LMWH, low-molecular-weight heparin; UFH, unfractionated heparin; VTE, venous thromboembolism.

\*VTE risk assessment should be performed prepregnancy or early in pregnancy and repeated if complications develop, particularly those necessitating hospitalization or prolonged immobility.

<sup>†</sup>First-degree relative with a history of a thrombotic episode, or other major thrombotic risk factors (eg, obesity, prolonged immobility, cesarean delivery).

<sup>‡</sup>Low-risk thrombophilia: Factor V Leiden heterozygote; prothrombin G20210A mutation heterozygote; protein C or protein S deficiency, antiphospholipid antibody.

<sup>§</sup>High-risk thrombophilias include Factor V Leiden homozygosity, prothrombin gene G20210A mutation homozygosity, heterozygosity for factor V Leiden and prothrombin G20210A mutation, or antithrombin deficiency.

# Appendix B

# Figure 3.:

Decision aid for urgent or emergent neuraxial procedures in the obstetric patient receiving UFH.



Leffert, Lisa; Butwick, Alexander; Carvalho, Brendan; Arendt, Katherine; Bates, Shannon; members of the SOAP VTE Taskforce The Society for Obstetric Anesthesia and Perinatology Consensus Statement on the Anesthetic Management of Pregnant and Postpartum Women Receiving Thromboprophylaxis or Higher Dose Anticoagulants, Anesthesia & Analgesia: March 2018 - Volume 126 - Issue 3 - p 928-944 doi: 10.1213/ANE.00000000002530

# Appendix C

## Figure 4.:

Decision aid for urgent or emergent neuraxial procedures in the obstetric patient receiving LMWH.



Leffert, Lisa; Butwick, Alexander; Carvalho, Brendan; Arendt, Katherine; Bates, Shannon; members of the SOAP VTE Taskforce The Society for Obstetric Anesthesia and Perinatology Consensus Statement on the Anesthetic Management of Pregnant and Postpartum Women Receiving Thromboprophylaxis or Higher Dose Anticoagulants, Anesthesia & Analgesia: March 2018 - Volume 126 - Issue 3 - p 928-944 doi: 10.1213/ANE.00000000002530