



OLOLRMC Pharmacy Antithrombotic Stewardship



COVID-19 Antithrombotic management guideline

Epidemiology

Caucasians and African Americans have almost double the risk of VTE compared with Hispanics.

Risk factors

Immobility in COVID -19 patients (limited PPE resources for routine ambulation), obesity, advanced age (≥ 50), recent surgery and trauma, sepsis, pregnancy, cancer

Antithrombotic therapy for non-critically ill, hospitalized patients with COVID-19 with no suspected or confirmed VTE¹

All hospitalized patients with COVID-19 should receive at least the standard dose thromboprophylaxis with Enoxaparin (LWMH) or Heparin (UFH)¹

1 For non-critically ill patients hospitalized with COVID-19, prophylactic dose of Enoxaparin (LWMH) or Heparin (UFH) is recommended.

1. Enoxaparin 40 mg SQ daily

OR

2. Enoxaparin 30 mg SQ daily for CrCl < 30 ml/min

Morbid Obesity⁴ – BMI > 40 kg/m²

a. Enoxaparin 40 mg SQ BID

OR

b. Enoxaparin 30 mg SQ BID for CrCl < 30 ml/min

2 In select non-critically ill patients at increased risk of disease progression or thromboembolism and are not high risk for bleeding; consider the use of therapeutic intensity dose of LMWH or UFH

3 For non-critically ill patients hospitalized with COVID-19, intermediate-dose of LMWH or UFH is not recommended

4 For non-critically ill patients hospitalized with COVID-19, add-on treatment with an antiplatelet agent is not recommended

We do not recommend the routine use of post-hospital thromboprophylaxis therapy after discharge following hospitalization for COVID-19.

Antithrombotic therapy for critically ill, hospitalized patients with COVID-19 with no suspected or confirmed VTE¹

<p>1 For critically ill patients hospitalized with COVID-19, prophylactic dose of Enoxaparin (LWMH) or Heparin (UFH) is recommended.</p> <ol style="list-style-type: none"> 1. Enoxaparin 40 mg SQ daily <li style="text-align: center;">OR 2. Enoxaparin 30 mg SQ daily for CrCl < 30 ml/min <p>Morbid Obesity⁴ – BMI > 40 kg/m²</p> <ol style="list-style-type: none"> a. Enoxaparin 40 mg SQ BID <li style="text-align: center;">OR b. Enoxaparin 30 mg SQ BID for CrCl < 30 ml/min
<p>2 In critically ill patients hospitalized with COVID-19, intermediate-dose, or therapeutic dose of LMWH or UFH is not recommended</p>
<p>3 In critically ill patients hospitalized with COVID-19, add-on treatment with an antiplatelet agent to prophylactic dose of LMWH or UFH is not recommended</p>

We do not recommend the routine use of post-hospital thromboprophylaxis therapy after discharge following hospitalization for COVID-19.

Anticoagulation therapy for hospitalized patients with confirmed diagnosis of VTE

Critically Ill with no enteral route	Non-critical ill with enteral route
<ol style="list-style-type: none"> 1. CrCl > 30 ml/min <ol style="list-style-type: none"> a. Enoxaparin 1 mg/kg SQ BID <li style="text-align: center;">OR b. Enoxaparin 1.5 mg/kg SQ daily 2. CrCl < 30 ml/min <ol style="list-style-type: none"> a. Enoxaparin 1 mg /kg SQ daily <p>For Obese patient (weight > 150 kg)</p> <ol style="list-style-type: none"> 1. CrCl > 30 ml/min <ol style="list-style-type: none"> a. Enoxaparin 150 mg SQ BID 2. CrCl < 30 ml/min <ol style="list-style-type: none"> a. Enoxaparin 150 mg SQ daily 	<ol style="list-style-type: none"> <p>Apixaban</p> <ol style="list-style-type: none"> b. Initiate 10 mg PO BID for 7 days c. Followed by 5 mg PO BID d. Hemodialysis: 5 mg PO BID for 7 days; then 2.5 mg PO BID <p>2. Rivaroxaban</p> <ol style="list-style-type: none"> a. Initiate 15 mg PO BID for 21 days b. Followed by 20 mg PO daily c. CrCl < 15 mL/min and HD: avoid use

Monitoring

Anti-factor Xa monitoring on treatment dose will be done automatically by a clinical pharmacist for the following patient populations:

- Obesity (> 150 kg)
- Renal insufficiency (CrCl <30 mL/min).

Using Anti-Factor Xa Monitoring Guideline for Enoxaparin located on formweb

Duration of Therapy

Patients with diagnosis of VTE

1. Treatment dose for minimum of 3 months as recommended for provoked VTE

Patients with no diagnosis of VTE

2. VTE Prophylaxis pharmacotherapy during the course of hospitalization
 - a. Consider benefit vs risk for an extended VTE prophylaxis post discharge.

This Guideline Review and Revision information

Last Date Revised: 03/2024

Last Date Reviewed: 03/2024

References

1. Schulman S, Donald M, et al: "2023 ISTH update of the 2022 ISTH guidelines for antithrombotic treatment in COVID-19." International Society on Thrombosis and Haemostasis. Journal of thrombosis and Haemostasis (jth). 2024; DOI: <https://doi.org/10.1016/j.jth.2024.02.011>
2. Schulman S, Sholzberg M, et al: "2022 ISTH guidelines for antithrombotic treatment in COVID-19." International Society on Thrombosis and Haemostasis. Journal of thrombosis and Haemostasis (jth). 2022; no.20, pp. 2214-2225. DOI: 10.1111/jth.15808
3. Barnes GD, Burnett A, Arthur A, et al : "Thromboembolic prevention and anticoagulant therapy during the COVID-19 pandemic: updated clinical guidance from the anticoagulation forum." Journal of Thrombosis and Thrombolysis. 2022. <https://doi.org/10.1007/s11239-022-02643-3>
4. Cuker Adam, Tseng E, et al. "American Society of Hematology 2021 guidelines on the use of anticoagulation for thromboprophylaxis in patients with COVID-19." Blood Advances, vol. 5, no. 3, 2021, pp. 872–3291., doi:10.1182/bloodadvances.2020003763.
5. Moores L, Tritschler T, et al. "Prevention, Diagnosis, and Treatment of VTE in Patients with Coronavirus Disease 2019" CHEST 2020; 158(3): 1143-1163. DOI: <https://doi.org/10.1016/j.chest.2020.05.559>

6. Witt, Daniel M., et al. "American Society of Hematology 2018 Guidelines for Management of Venous Thromboembolism: Optimal Management of Anticoagulation Therapy." *Blood Advances*, vol. 2, no. 22, 2018, pp. 3257–3291., doi:10.1182/bloodadvances.2018024893.
7. Kearon, Clive, et al. "Antithrombotic Therapy for VTE Disease: CHEST Guideline and Expert Panel Report." *Chest*, vol. 150, no. 4, 2016, p. 988., doi:10.1016/j.chest.2016.08.1442.
8. Cohen AT, et al. "Rivaroxaban for thromboprophylaxis in acutely ill medical patients". *The New England Journal of Medicine*. 2013. 368(6):513-523
9. Wang, Tzu-Fei, et al. "Efficacy and Safety of High-Dose Thromboprophylaxis in Morbidly Obese Inpatients." *Thrombosis and Haemostasis*, vol. 111, no. 01, 2014, pp. 88–93., doi:10.1160/th13-01-0042.
10. Mavrakanas, Thomas A., et al. "Apixaban Pharmacokinetics at Steady State in Hemodialysis Patients." *Journal of the American Society of Nephrology*, vol. 28, no. 7, 2017, pp. 2241–2248., doi:10.1681/asn.2016090980.
11. Siontis, Konstantinos C., et al. "Outcomes Associated With Apixaban Use in Patients With End-Stage Kidney Disease and Atrial Fibrillation in the United States." *Circulation*, vol. 138, no. 15, Sept. 2018, pp. 1519–1529., doi:10.1161/circulationaha.118.035418.
12. Schafer, Joseph H., et al. "Safety and Efficacy of Apixaban Versus Warfarin in Patients With Advanced Chronic Kidney Disease." *Annals of Pharmacotherapy*, vol. 52, no. 11, May 2018, pp. 1078–1084., doi:10.1177/1060028018781853
13. National Comprehensive Cancer Network. "Cancer-Associated Venous Thromboembolic Disease" NCCN Guidelines Version 1. 2019
14. Mahan, Charles E., and Alex C. Spyropoulos. "ASHP Therapeutic Position Statement on the Role of Pharmacotherapy in Preventing Venous Thromboembolism in Hospitalized Patients." *American Journal of Health-System Pharmacy*, vol. 69, no. 24, 2012, pp. 2174–2190., doi:10.2146/ajhp120236.
15. Nishimura, Rick A., et al. "2017 AHA/ACC Focused Update of the 2014 AHA/ACC Guideline for the Management of Patients With Valvular Heart Disease." *Journal of the American College of Cardiology*, vol. 70, no. 2, 2017, pp. 252–289., doi:10.1016/j.jacc.2017.03.011.
16. Bickford, Annika, et al. "Weight-Based Enoxaparin Dosing for Venous Thromboembolism Prophylaxis in the Obese Trauma Patient." *The American Journal of Surgery*, vol. 206, no. 6, 2013, pp. 847–852., doi:10.1016/j.amjsurg.2013.07.020.
17. Tang, Ning, et al. "Anticoagulant Treatment Is Associated with Decreased Mortality in Severe Coronavirus Disease 2019 Patients with Coagulopathy." *Journal of Thrombosis and Haemostasis*, 2020, doi:10.1111/jth.14817.
18. Nishimura, Rick A., et al. "2014 AHA/ACC Guideline for the Management of Patients With Valvular Heart Disease." *Circulation*, vol. 129, no. 23, Oct. 2014, doi:10.1161/cir.0000000000000031.
19. XARELTO® (rivaroxaban) tablets, for oral use: <http://www.janssenlabels.com/package-insert/product-monograph/prescribing-information/XARELTO-pi.pdf>
20. ELIQUIS® (apixaban) tablets, for oral use: https://packageinserts.bms.com/pi/pi_eliquis.pdf