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Objectives:

- Review the diagnostic algorithms of the diagnosis and management of DVT and PE.
- Review appropriate use of Ddimer testing.
- Review the PERC rule
- Discuss the outpatient therapeutic options for DVT and PE, with a focus on the outpatient setting.

Disclosure

• I have no financial or conflict of interest disclosures.

Case

It's Friday afternoon, a 32 yr female presents with leg swelling since 2 days. No fever, no trauma. On physical exam the RT leg is mildly edematous compared to the left (approx 4 cm difference). You notice pain on palpation of the calf and proximal leg with some unilateral pitting edema. She is otherwise healthy, but takes the oral contraceptive pill. She was admitted to the hospital 2 weeks ago for an acute appendicitis.

he best course of action is:

- A. Send her to the ER
- B. Send a d-dimer stat
- C. Give her a prescription for Xarelto and advise her to go the ER on Monday
- D. Reassure her and give a prescription for Naproxen

Case 2

A 44 yr male presents in the evening walk in clinic, complaining of chest pain X 4 days. He denies any cough or fever. He has recently taken on a new exercise routine at the gym. He recently drove to Mt Tremblant for a ski trip. He has no leg swelling, no dyspnea. His grandmother was diagnosed with a PE 3 months and according to Google this is a possibility.

Physical exam shows a HR of 80, Sat of 99% on room air.

The best course of action is:

- A. Send him to the ER to rule out a PE
- B. Reassure him that it is unlikely to be PE
- C. Order bilateral leg dopplers
- D. Refer him for a CT scan to rule out PE as an outpatient

DVT – Why Do I Care

- Progression to PE
- Post thrombotic syndrome
- Phlegmasia cerulea dolens
- Phlegmasia alba dolens





N Engl J Med 2018; 378:658

DVT- Risk factors

- Inherited thrombophilia
- Malignancy
- Trauma
- Pregnancy
- OCP/HRT
- Immobilization
- Heart failure
- Age >65
- Myeloproliferative disorders
- Inflammatory bowel disease

Source: uptodate

DVT – signs and symptoms

- Not very reliable
- Leg edema 97% sensitive, 33% specific
- Pain 86 % sensitive, 19 % specific
- Warmth 72% sensitive, 48 % specific

Source: uptodate

- 1/3 symptomatic DVT have concomitant PE
- 70% confirmed PE have a concomitant DVT

Source: Lancet. 2016 Dec 17;388(10063):3060-3073

DVT- signs and symptoms

- Large calf diameter in a meta analysis doubled likelihood of having DVT
- Calf swelling is measured at 10cm below tibial tuberosity
- Homan's sign unreliable
- Travel should be considered "immobilization" if greater than 4 hours

CMAJ. 2015 Nov 17;187(17):1288-96. doi: 10.1503/cmaj.141614. Epub 2015 Sep 28.



We are SOOO BAD at this



Table 1. Clinical Model for Predicting the Pretest Probability of Deep-Ve	in
Thrombosis.*	

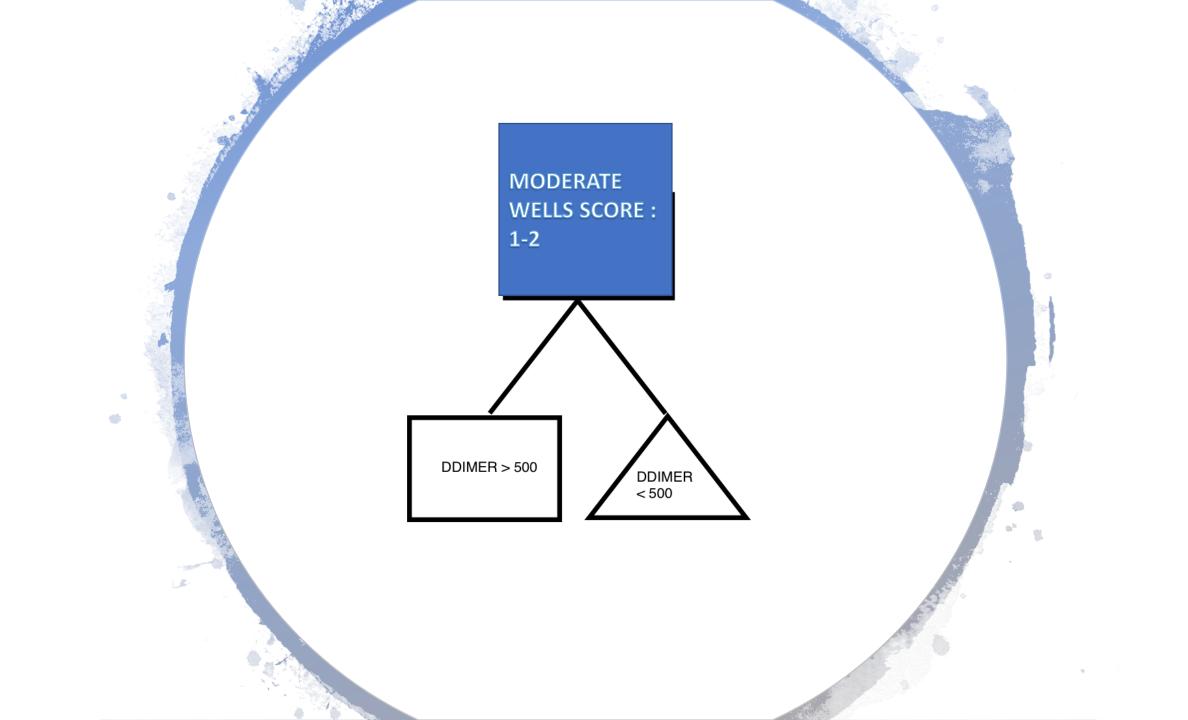
Clinical Characteristic	Score
Active cancer (patient receiving treatment for cancer within the previous 6 mo or currently receiving palliative treatment)	1
Paralysis, paresis, or recent plaster immobilization of the lower extremities	1
Recently bedridden for 3 days or more, or major surgery within the previous 12 wk requiring general or regional anesthesia	1
Localized tenderness along the distribution of the deep venous system	1
Entire leg swollen	1
Calf swelling at least 3 cm larger than that on the asymptomatic side (measured 10 cm below tibial tuberosity)	1
Pitting edema confined to the symptomatic leg	1
Collateral superficial veins (nonvaricose)	1
Previously documented deep-vein thrombosis	1
Alternative diagnosis at least as likely as deep-vein thrombosis	-2

^{*} A score of two or higher indicates that the probability of deep-vein thrombosis is likely; a score of less than two indicates that the probability of deep-vein thrombosis is unlikely. In patients with symptoms in both legs, the more symptomatic leg is used.

N Engl J Med 2003; 349:1227-1235

Nothing is perfect

- Wells may NOT perform well in
 - Hospitalized patients
 - Elderly
 - Recurrent DVT / PE
 - Cancer patients
 - ?primary care setting



D-Dimer

- High sensitivity d-dimer assays outperform non HS assays
- HS D-dimer have a higher NPV
- Should not be used as a stand- alone
- Age adjusted D-dimers
 - Patients > 50
 - Age in years X 10
 - Improves specificity without modifying sensitivity

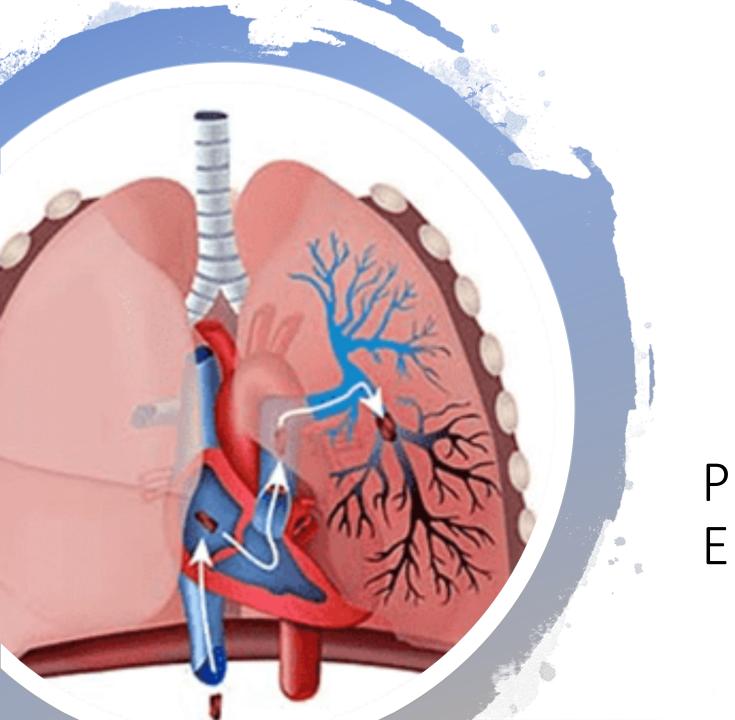
Wells + Ddimer

Clinical Pretest Probability	Sensitivity	Specificity	Negative Predictive Value	Negative Likelihood Ratio
Low	88%	72%	99%	0.18
Moderate	90%	58%	96%	0.19
High	92%	45%	84%	0.16

Reference - JAMA 2006 Jan 11;295(2):199, editorial can be found in JAMA 2006 Jan 11;295(2):213, commentary can be found in ACP J Club 2006 Jul-Aug;145(1):24

Confirmatory testing [+D-Dimer or High risk]

- Duplex ultrasound most widely used
- Controversy between whole leg or proximal veins only
- CT-V and MR-V rarely used
- Contrast venography not used very much at all



PULMONARY EMBOLISM

TAKE A DEEP BREATH...

- PE not as a fatal as initially thought
- Newer data 2011 mortality rate of about 1%, directly related to PE
- 85% of deaths occurred while waiting for diagnostic confirmation, suggesting most patients succumb to their underlying illness

PE risk factors

- Same as those for DVT
- Virchows triad
 - Venous stasis
 - Endothelial injury
 - Hypercoagulable state
- Up to 50% of PE patient haven no apparent risk factors

PE – signs and symptoms

- DYSPNEA!
- Fatigue coinciding with a new dyspnea
- Chest pain
- No chest pain
- Leg swelling
- No leg swelling
- Hemoptysis (although rare)

Syncope and PE

- Probably overstated
- One international study <1% prevalence of PE in ER presentations with syncope
- A Canadian study showed a 1.4% prevalence of PE in a cohort of admitted patients with syncope
- Flipped T waves in anterior and inferior leads more most SPECIFIC finding in PE
- Most common finding -> NSR

Prandoni P, et al. Prevalence of Pulmonary Embolism among Patients Hospitalized for Syncope. The New England Journal of Medicine. 2016;375(16):1524-31.

Approach



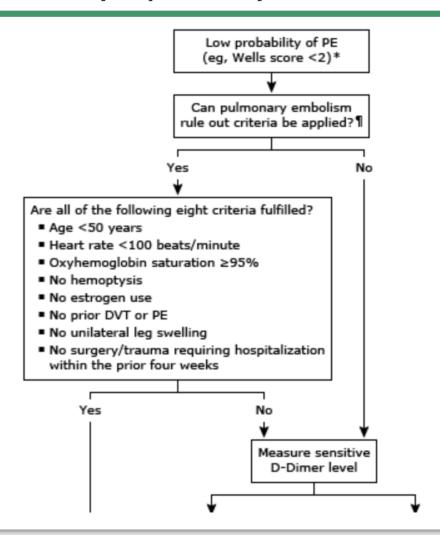
Log in SIGN UP

Wells' Criteria for Pulmonary Embolism 🕸

Objectifies risk of pulmonary embolism.

When to Use 🗸	Pearls/Pitfalls 🗸	Why Use 🗸
Clinical signs and symptoms of DV1	No 0	Yes +3
PE is #1 diagnosis OR equally likely	No 0	Yes +3
Heart rate > 100	No 0	Yes +1.5
Immobilization at least 3 days OR s the previous 4 weeks	surgery in No 0	Yes +1.5
Previous, objectively diagnosed PE	or DVT No 0	Yes +1.5
Hemoptysis	No 0	Yes +1
Malignancy w/ treatment within 6 palliative	months or No 0	Yes +1

Evaluation of the nonpregnant adult with low probability of pulmonary embolism



Source: uptodate

PERC

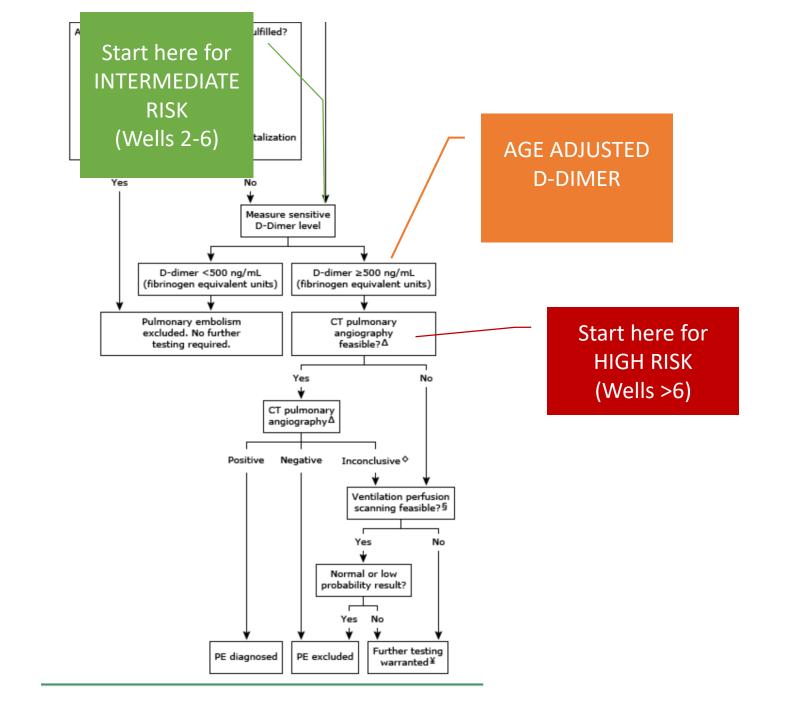
- Low risk patients only with a low prevalence
- Study designed with 1.8% threshold

PERC Rule for Pulmonary Embolism 🗘

Rules out PE if no criteria are present and pre-test probability is ≤15%.

When to Use 🗸	Pearls/Pitfalls 🗸	Why Use 🗸
Age ≥50	No 0	Yes +1
HR ≥100	No 0	Yes +1
SaO ₂ on room air <95%	No 0	Yes +1
Unilateral leg swelling	No 0	Yes +1
Hemoptysis	No 0	Yes +1
Recent surgery or trauma Surgery or trauma ≤4 weeks ago req treatment with general anesthesia	uiring No 0	Yes +1
Prior PE or DVT	No 0	Yes +1
Hormone use Oral contraceptives, hormone replac estrogenic hormones use in males or patients		Yes +1

J Thromb Haemost. 2004 Aug;2(8):1247-55.



CT PE

- The modern "gold standard"
- Over diagnosis
- Almost 100% sensitive for clinically relevant PE
- 5% of high risk patients will develop PE in a few months with a negative CT PE study.

van der Hulle T, van Es N, den Exter PL, et al. Is a normal computed tomography pulmonary angiography safe to rule out acute pulmonary embolism in patients with a likely clinical probability? A patient-level meta-analysis. Thromb Haemost. 2017;117(8):1622-1629.

Outcomes following a negative computed tomography pulmonary angiography according to pulmonary embolism prevalence: a meta-analysis of the management outcome studies. J Thromb Haemost. 2018 Jun;16(6):1107-1120.

Subsegmental PE

- Likely not clinically significant
- Consensus recommendations to treat based on individual risk
- Observational study in 2015 of 2213 patients
 - No difference in rate of recurrent PEs between treatment vs non treatment
 - 5% of anti coagulated patients had life threatening bleeds



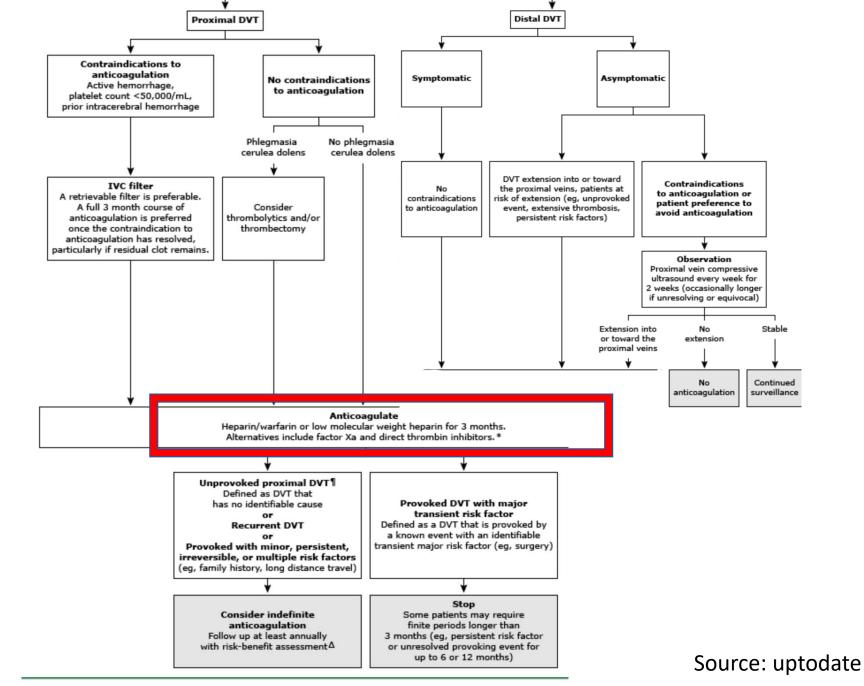




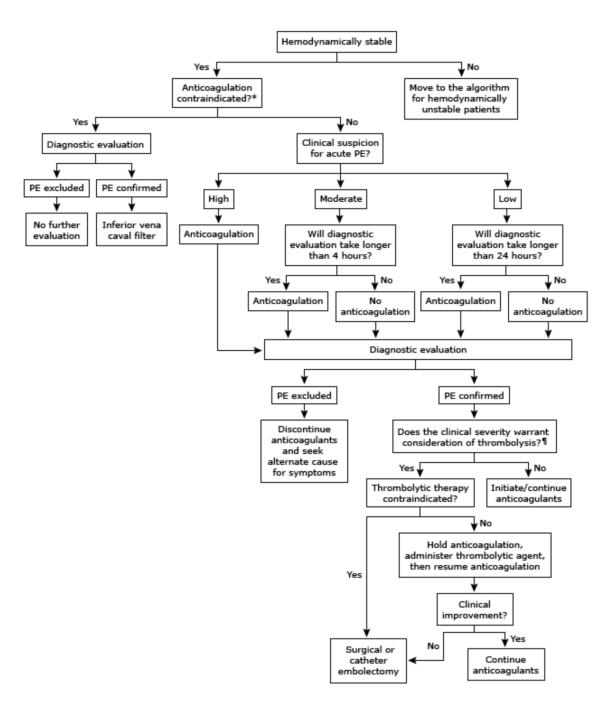


TREATMENT

DVT



This algorithm only applies to patients with a **first** episode of DVT.



PE Algorithm

Source: uptodate

Choices of out patient anti coagulation

- LMWH
- DOAC (oral Xa inhibitors) Rivaroxaban or apixiban
- Warfarin can be used but not acutely
- (IVC filter)

Generalities about anti coagulation choices

- LMWH is preferred for cancer patients, liver disease, pregnancy
- Avoid DOACs with renal disease
- UFH if admitted, high risk of bleeding

 Bottom line: pick the one that you feel most comfortable with initially and above all DO NO HARM

HAS-BLED score

- Developped to assess bleeding risk in AFIB with anti coagulation
- Estimates 1 yr risk of major bleeding
- Can likely be extrapolated to PE population
- Helpful in decision making for pre testing treatment vs sub segmental PE empiric treatment

https://www.mdcalc.com/has-bled-score-major-bleeding-risk#next-steps

HAS-BLED Score for Major Bleeding Risk 🕸

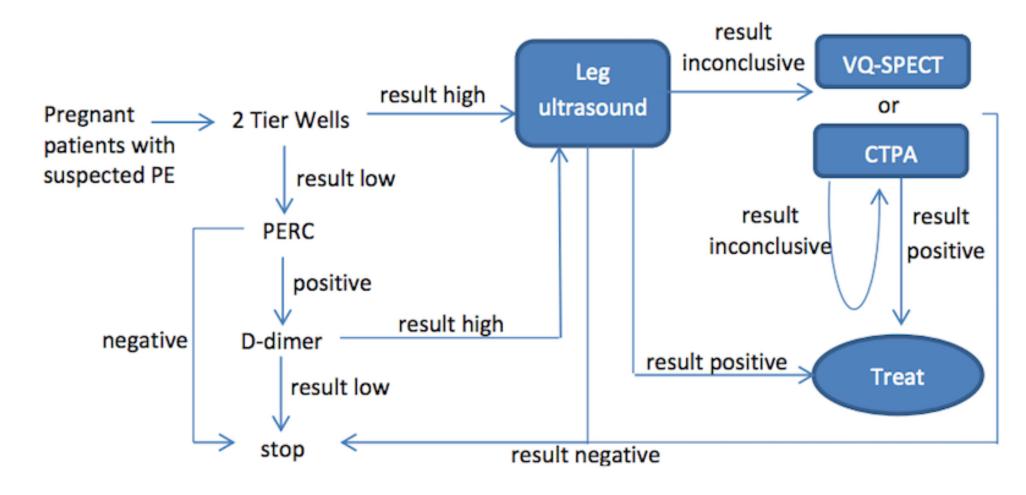
Estimates risk of major bleeding for patients on anticoagulation to assess risk-benefit in atrial fibrillation care.

When to Use ✓	Pearls/Pitfalls 🗸	Why Use 🗸
Hypertension Uncontrolled, >160 mmHg systolic	No 0	Yes +1
Renal disease Dialysis, transplant, Cr >2.26 mg/dL or >20 µmol/L	No 0	Yes +1
Liver disease Cirrhosis or bilirubin >2x normal with AST/ >3x normal	ALT/AP No 0	Yes +1
Stroke history	No 0	Yes +1
Prior major bleeding or predisposition t bleeding	No 0	Yes +1
Labile INR Unstable/high INRs, time in therapeutic ra <60%	nnge No 0	Yes +1
Age >65	No 0	Yes +1
Medication usage predisposing to bleed Aspirin, clopidogrel, NSAIDs	ding No 0	Yes +1
Alcohol use	No. 0	Yes +1

Pregnancy

- Pregnant patients excluded from decision rules including Wells
- American thoracic society discourages use of D-dimer
- CADTH developed guideline for PE work up in Pregnancy

CADTH suggested guideline for PE in Pregnancy



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