

John R. Corker, MD

Clinical Conundrum:

Using IV estrogen as adjunctive hemostatic therapy in a patient with acute, severe uterine hemorrhage and history of thromboembolic event or risk factors.

Background:

Hormonal management is considered the first line of medical therapy for patients with acute uterine hemorrhage without known or suspected bleeding disorders. However, across indications, it is **generally contraindicated in patients with a history of thromboembolic event or risk factors**. Little data exist regarding the use of IV estrogen in patients with acute severe uterine hemorrhage and pre-existing thromboembolic risk factors.

Findings:

- 1.) In non-pregnant hemodynamically unstable patients, intravenous (IV) or intramuscular (IM) conjugated estrogen 25 mg may be administered, with repeat doses in 4 to 6 hours as needed for hemorrhage control.
- 2.) In one randomized controlled trial of 34 women, IV conjugated equine estrogen was shown to stop bleeding in 72% of participants within 8 hours of administration compared with 38% of participants treated with a placebo.
- 3.) A theoretical risk of thromboembolic complications has been inferred on the basis of previous studies of oral contraceptives and estrogen replacement therapy. However, only 1 case has been reported with use of short-term intravenous CEE therapy. In that case, the patient was treated with intravenous CEE for severe menorrhagia, and she was also taking GnRH agonist treatment combined with add-back therapy consisting of 0.625 mg of CEE plus medroxyprogesterone acetate 10 mg for uterine fibroids.
- 4.) Once hemodynamic stability has been achieved, the patient can be treated with progesterone-only hormonal therapy—10 mg of medroxyprogesterone (Provera) once a day for 14 days—and outpatient follow-up in 2 to 4 weeks.

Conclusions:

IV estrogen therapy can serve as an effective adjunct hemostatic therapy to complement fluid resuscitation in the acute management of a patient with severe uterine bleeding. Although reports of acute complications in patients with history of thromboembolic event or risk factors are rare, **more research is necessary to quantitatively assess the long-term risk of IV estrogen therapy in this population. In all cases, patients should be counseled on potential risks and benefits prior to initiation of therapy.**

- 1.) "Management of Acute Abnormal Uterine Bleeding in Nonpregnant Reproductive-Aged Women." Committee Opinion, Number 557, April 2013. The American College of Obstetrics and Gynecologists (ACOG).
- 2.) DeVore GR, Owens O, Kase N. Use of intravenous Premarin in the treatment of dysfunctional uterine bleeding—a double-blind
- 3.) Zreik T.G., Odunsi K., Cass I., et al: A case of fatal pulmonary thromboembolism associated with the use of intravenous estrogen therapy. *Fertil Steril* 1999; 71: pp. 373-375
- 4.) [Bert Scoccia MD](#), [Habibe Demir](#), [Koray Elter MD](#) and [Antonio Scommegna](#). "Successful Medical Management of Post-hysteroscopic Metroplasty Bleeding with Intravenous Estrogen Therapy: A Report of Two Cases and Review of the Literature." *The Journal of Minimally Invasive Gynecology*, 2009-09-01Z, Volume 16, Issue 5, Pages 639-642, Copyright © 2009 AAGL
- 5.) [John A. Marx](#), [Robert S. Hockberger](#), and [Ron M. Walls](#). "Vaginal Bleeding." *Rosen's Emergency Medicine*, 8th Edition, Chapter 34, 273-277