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Author Hou, Melody Y

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- 1 Title: Uncomplicated abortion with mifepristone and misoprostol in a hemophilia A
- 2 carrier

3 Author:

- 4 Melody Y. Hou¹
- 5 ¹ Department of Obstetrics and Gynecology, University of California, Davis,
- 6 Sacramento, CA, USA
- 7 Source of Funding: None
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- 10 **Corresponding author:**
- 11 Melody Y. Hou, MD MPH, University of California, Davis
- 12 4860 Y Street, Ste 2500, Sacramento CA 95817
- 13 Phone 916-734-6918; e-mail: myhou@ucdavis.edu
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18 Abstract:

Little evidence exists regarding medical abortion for women with inherited bleeding
disorders. A 21-year old primigravid hemophilia A carrier desired a medical
abortion. After counseling, she chose medical abortion, which occurred without
excess bleeding or surgical intervention.

23

24 Introduction:

25 Medical abortion using mifepristone and misoprostol is a safe and effective 26 method for early first trimester termination of pregnancy.[1] However, medical 27 abortion is associated with heavier and longer bleeding compared to surgical 28 abortion. Women undergoing medical abortion, compared to those having a manual 29 vacuum aspiration, have a mean of 14 days versus 9 days of bleeding, respectively, 30 and have a decrease in hemoglobin of 0.7 mg/dL or less.[2-5] Women with 31 coagulopathies have been excluded from clinical trials; accordingly, mifepristone's 32 product label and protocols from major medical organizations list bleeding disorders as a contraindication for mifepristone.[6] This case report describes a hemophilia A 33 34 carrier, who is a symptomatic hemophilia A heterozygote, undergoing an 35 uncomplicated abortion with mifepristone and misoprostol. 36 37 Case:

A 21 year old gravida 1, para 0 woman presented to our clinic at 46 days gestation requesting an abortion. She is a mildly symptomatic hemophilia A carrier; her occasional symptoms include easy bruising, delayed clotting, and heavy menses. She was diagnosed as a child on routine family surveillance testing due to a family history of hemophilia A. Her menses are light while using combined 43 hormonal contraception. Her surgical history includes an uncomplicated wisdom 44 tooth extraction after using nasal desmopressin acetate, which is a synthetic analogue of vasopressin used in patients with hemophilia A to stop bleeding in 45 46 episodes of spontaneous or trauma-induced injuries. She has two brothers and an 47 uncle with hemophilia A, categorized by factor VIII levels as mild disease, with no 48 history of catastrophic bleeding events. Her last hemophilia panel 7 years prior 49 indicated a factor VIII level of 46% (normal range: 50-150% of normal human 50 plasma, NHP) with all other coagulation factors within normal limits. She recently 51 joined the military and was undergoing basic military training at time of 52 presentation. She reported no bleeding problems or need for desmopressin for 53 several years.

54 The patient had a strong preference for medical abortion for termination, 55 even after extensive counseling regarding her indeterminate bleeding risk. She was 56 aware that need for an emergent surgical intervention was a possibility, and stated 57 that she lived within 10 minutes of an emergency room and that a family member would be staying with her. Her hemoglobin was 12.1 g/dL. She received 58 59 mifepristone 200 mg orally in the office, and she elected to insert misoprostol 800 60 mcg vaginally approximately 6 hours following mifepristone. She was instructed to 61 self-administer desmopressin 150 mcg intranasally 30 minutes prior to misoprostol 62 use.

She returned to clinic one week later for her scheduled follow-up. She used
demopressin and misoprostol as instructed. She reported that she began bleeding
and cramping 2 hours later with the heaviest flow occurring 4 hours after
misoprostol, soaking 2 pads in that hour. The bleeding then quickly ceased. Her
pregnancy symptoms resolved and she reported only spotting at her follow-up visit.

Transvaginal ultrasonography revealed no intrauterine gestational sac and an endometrial stripe measuring 8 mm. The patient had a levonorgestrel 52 mg intrauterine system placed for contraception in the same visit, but did not have her requested postabortion hemoglobin drawn. Six months after her abortion, she reported light menses and high satisfaction with her intrauterine contraceptive.

74 Comment:

Medical abortion is associated with heavier and longer bleeding compared to surgical abortion,[2] but excessive bleeding or need for blood transfusion appear similar.[7] However, clinicians may have concern that the bleeding associated with medical abortion makes medical abortion an unsafe option for women with inherited bleeding disorders.[6]

80 Hemophilia A is a genetic deficiency in clotting factor VIII, and bleeding 81 symptoms inversely correlate with factor VIII levels. Its X-linked recessive 82 inheritance affects males, but female heterozygote carriers may have variable 83 bleeding symptoms. Most hemophilia A carriers have a factor VIII level between 30-84 70% and the majority do not have heavy menstrual bleeding.[8] Very little has been 85 published on hemophilia A carriers' experience with pregnancy termination options, 86 pregnancy complications, or gynecologic issues.[9] Pregnancy complications such 87 as miscarriage, placental abruption, and postpartum hemorrhage have been described in women with inherited bleeding disorders, but primarily among patients 88 89 with von Willebrand disease, a more prevalent coagulopathy caused by a deficiency 90 of von Willebrand factor. Many of these accounts were case reports and series, 91 which do not allow an estimate of rates.[9] Among 43 women affected by inherited 92 bleeding disorders who reported a miscarriage in a survey by the United States

Hemophilia Treatment Centers, 33 (76.7%) self-reported a problem with bleeding
during the miscarriage. However, the report did not describe the women's bleeding
disorders, how their bleeding problems manifested during the miscarriage, or if
their bleeding required medications such as desmopressin, hospitalization, blood
transfusion, or surgical intervention.[10]

98 This patient's medical abortion experience was similar to that described by 99 other women, which suggests that medical abortion may be a reasonable option for 100 some hemophilia A carriers with an undesired pregnancy. Information on identifying 101 whether a woman is an appropriate candidate is limited since women with bleeding 102 disorders have been excluded from medical abortion trials. A clinician's evaluation 103 should include a thorough history of her coagulation disorder and other bleeding 104 risk factors, including any recent traumas that could give an estimate of the 105 patient's likelihood of having a bleeding event with her medical abortion and her 106 previous experience with factor replacement therapies. Consultation with a hematologist may help assess bleeding risk, but if it cannot occur before 107 108 institutional or legal limits for medical abortion, a partial thromboplastin time (PTT) 109 and factor VIII level can provide more information. However, even with a normal 110 factor VIII level of at least 50%, clinicians should still administer desmopressin 111 acetate for bleeding prophylaxis in medical abortion as they would for a surgical 112 procedure. Clinicians should remind patients that emergent curettage remains a 113 possibility and should assess a patient's access to emergency care during the 114 abortion process.

In this case of a symptomatic hemophilia A carrier with access to
desmopressin and emergency care, the patient highly valued having a choice in
abortion options, with a strong preference for medical management. A careful

assessment of her bleeding risk included a confirmation of her hemophilia A carrier status, family history, prior use of and response to desmopressin, and recent history of military training with frequent small injuries without needing desmopressin. Her uncomplicated course supports the feasibility of offering medical abortions to select patients with hemophilia A, and the importance of careful history-taking such as their specific coagulopathy, surgical history, and prior bleeding episodes and treatments, to assess their bleeding risk and candidacy for medical abortion.

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