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## **Bleeding Following Pregnancy Loss Prior to Six Weeks Gestation**

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## Abstract

**BACKGROUND**— Pregnancy loss prior to six weeks gestation is common, but little has been reported about the associated bleeding. We compared women's bleeding following a pregnancy loss before six weeks gestation with their typical menstruation.

**METHODS**— Women provided daily urine samples while trying to become pregnant and recorded the number of pads and tampons used each day. Thirty-six women had complete bleed data for a loss prior to six weeks gestation and one or more non-pregnant cycles.

**RESULTS**— Mean bleed length following a pregnancy loss was 0.4 days longer than the woman's average menstrual bleed, due primarily to more days of light bleeding. Although there was no overall increase in the total number of pads plus tampons used, women with losses bled less than their typical menses following pregnancies of very short duration and more than usual for the pregnancies lasting the longest.

**CONCLUSIONS**— Overall, the bleeding associated with pregnancy loss before six weeks gestation is similar to menstrual bleeding and unlikely to be recognized as pregnancy loss. The intriguing finding that pregnancies of very short duration were associated with less bleeding than the woman's typical menses might reflect endometrial factors associated with loss.

## Keywords

pregnancy loss; menses; bleed length; spontaneous abortion

## Introduction

Studies of women seeking medical care for miscarriage have reported that the bleeding that accompanies miscarriage is heavier and more prolonged than typical menses (Haines *et al.* 1994;Nielsen and Hahlin 1995;Chung *et al.* 1998;Nielsen *et al.* 1999;Bagratee *et al.* 2004). However, most of these clinically reported miscarriages occurred after six weeks gestation; the bleeding that accompanies earlier pregnancy loss has not been carefully described. Such early losses are common: approximately a quarter of all pregnancies end prior to six weeks gestation, often before the pregnancy has received any medical attention or even become apparent to the woman (Wilcox *et al.* 1990;Zinaman *et al.* 1996;Wang *et al.* 2003). Using data from a prospective study of women who were attempting pregnancy, we describe in detail the bleeding that accompanies pregnancy loss prior to six weeks gestation, and we compare those patterns of bleeding with the women's ordinary menstrual bleeding.

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## Materials and methods

Study participants were members of the North Carolina Early Pregnancy Study, a prospective cohort study conducted from 1982 to 1985 involving 221 women who were attempting pregnancy (Wilcox *et al.* 1988). These women had no known fertility problems or major chronic disease. Participation began when women discontinued contraception and continued until 8 weeks past their last menstrual period if they became pregnant, or for 6 months if they did not. During this time, participants collected daily first-morning urine specimens and kept daily diary records of their menstrual bleeding (number of pads and tampons). The study was approved by the Institutional Review Board of the National Institute of Environmental Health Sciences, and participating women provided informed consent.

Ovulation was identified using measurements of estrogen and progesterone metabolites in urine (Baird *et al.* 1991). Pregnancy was detected using a highly sensitive and specific polyclonal radioimmunoassay for urinary human chorionic gonadotrophin (hCG) (Armstrong *et al.* 1984). This assay measured intact and beta hCG with <1% cross-reaction with human luteinizing hormone (McChesney *et al.* 2005). The lowest hCG concentration detectable by the immunoradiometric assay was 0.01 ng per milliliter. Three consecutive days of urinary hCG greater than 0.025 ng/mL was the criteria for pregnancy. This criterion ensured high specificity and was determined from a parallel study of women who had either undergone a tubal ligation. Pregnancy loss was detected by a subsequent fall in urinary hCG. We restricted our study to pregnancies that ended less than 6 weeks after the last menstrual period, with the day of loss considered to be the first day of vaginal bleeding associated with the fall in hCG (Wilcox *et al.* 1988). Over-the-counter pregnancy test kits were not available at the time of the study, and only one woman who lost a pregnancy prior to six weeks gestation reported that it was clinically detected.

Following World Health Organization guidelines (World Health Organization Task Force on Adolescent Reproductive Health 1986), we defined length of a bleed period as the number of days from the start of bleeding up to and including the day before a bleed-free interval of at least 2 days. Menstrual discharge consists of tissue fluid (20–40% of the total discharge) and fragments of the endometrium in addition to blood (50–80% of the total discharge); we refer to this discharge simply as bleeding, as is customary (Oats and Abraham 2005).

We used the number of pads and tampons per day as a measure of bleeding quantity, focusing on comparisons within women. Women can differ from one another in their use of pads and tampons for reasons related more to personal habits than to volume of menstrual discharge. Because this variability between women is controlled in a within-woman analysis, it is more informative to conduct within-woman analyses of bleeding (comparing an early loss with the same woman's regular menses) than to make comparisons across women. In support of this, studies comparing pad and tampon counts with quantitative measurement of blood loss across women have reported considerably higher correlation coefficients when the studies included multiple cycles per woman (0.61 (Higham and Shaw 1999) and 0.74 (Higham *et al.* 1990)) than when they did not (0.14 (Fraser *et al.* 1981) and 0.30 (Warner *et al.* 2004)).

Women may also differ in their patterns of use of pads versus tampons. However, women in this study were fairly consistent from one period to the next in their use of pads versus tampons. The mean percentage of tampons used per period  $[100 \times tampons/(pads + tampons)]$  was 62% and the mean within-woman standard deviation in the percentage of tampons used per period was 9%.

Forty-four women had 48 pregnancy losses prior to six weeks gestation during their study participation. One of these women had a single loss for which she reported no associated bleeding; because her reporting was incomplete in other respects, we believed her report of no

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bleeding could have been a recording error, and we have excluded her from this analysis. Of the remaining women, 36 provided complete data on bleed length and pad-and-tampon use for at least one regular menses in addition to a pregnancy loss bleeding episode. One non-conception cycle was excluded because no bleeding was reported. All but one woman in this sub-sample were white; most were college-educated (72%). The median age was 29 years (range 24–36 years), and 61% were parous at enrollment.

These 36 women in the analysis sample had 38 pregnancy losses prior to six weeks gestation (two women had two losses). Information on bleeding after an ovulatory non-conception cycle was available for 1 to 7 menses per woman (median = 3) for a total of 198 non-conception menses. Analyses were based on differences between the within-woman averages. For each woman, a value for each bleed characteristic was calculated for both the non-conception and the pregnancy loss bleeds. If a woman had more than one non-conception or pregnancy loss bleed, then the mean value of the bleed characteristic was used. A bleed characteristic difference was then computed by subtracting the non-conception bleed value from the pregnancy loss bleed value, and differences were tested with a Wilcoxon signed rank test. Non-parametric correlation statistics and tests for difference were used due to the inherent non-normality of some of the bleed characteristic variables. The 25<sup>th</sup> and 75<sup>th</sup> percentile values for the distributions are presented, but means and standard errors are also presented for ease of comparison with other published bleed length values. All *P*-values cited are two-sided.

The pregnancy losses represented a range of durations and amounts of hCG production, which we hypothesized might also be related to the pattern of bleeding. In order to explore this, we used two markers: maximum level of hCG obtained during the pregnancy, and the length of the failed pregnancy (days from ovulation to onset of bleeding).

## Results

The overall pattern of bleeding after a pregnancy loss of less than six weeks gestation was remarkably similar to the pattern seen with regular menses (Figure 1). There was a high correlation between the length of a woman's bleeding with pregnancy loss and her usual bleeding with menses (Spearman correlation coefficient = 0.78; P < 0.001) (Figure 2). However, a closer look at within-women comparisons revealed some differences. Bleeding after a pregnancy loss tended to be slightly longer, by 0.4 days (P = 0.01) (Figure 2, Table I). This increased duration of bleeding was due primarily to an increase in light bleeding was 0.5 days (P = 0.004). Consistent with the additional light bleeding with pregnancy loss, the mean daily number of pads and tampons was slightly decreased in the pregnancy loss bleeding (Table 1). Pregnancy-loss bleeding showed no increase in either the total number of pads and tampons or the maximum number of pads and tampons in one day (Table I).

Pregnancies ending in losses prior to six weeks gestation are not all the same, either in their duration or in the amount of hCG they produce. Among the 38 pregnancy losses in this analysis, the time from conception (i.e. ovulation) to onset of bleeding ranged from 11 to 28 days. Similarly, maximal hCG concentration varied more than a hundred-fold, from 0.035 to 46 ng per milliliter. We explored whether the bleeding patterns after a loss varied according to the pregnancy duration and the level of hCG measured (Table II).

We found no relation between maximal hCG concentration or pregnancy duration and the number of days of bleeding. However, the total number of pads and tampons used in association with the pregnancy loss did correlate with both maximum hCG (Spearman correlation coefficient = 0.37; P = 0.02) and days from ovulation to bleeding (Spearman correlation coefficient = 0.48; P = 0.003). As one might expect, women with higher maximal hCG levels

and longer pregnancy duration used more pads and tampons for the bleeding associated with their pregnancy loss than they did for their typical menses. However, we unexpectedly also found that women whose pregnancies had lowest maximum hCG values and were of shortest duration tended to use *fewer* pads and tampons than they did for ordinary menses (Figure 3). This difference was most notable between women in the lowest and highest quartiles of maximum hCG (Table III).

## Discussion

In the Early Pregnancy Study, the bleeding accompanying a woman's pregnancy loss after less than six weeks gestation was, on the whole, not remarkably different from her typical menses. Overall, we saw no evidence of heavier bleeding after the pregnancy loss, either on a given day or over the entire bleeding episode. There was a slight tendency for women to bleed longer following pregnancy loss than they did for their typical menses, but this difference was small (0.4 days) and due entirely to light bleeding. The evidence suggests that most women would be unlikely to distinguish the bleeding episode that follows an early pregnancy loss from their ordinary menses.

Our results are consistent with those from a study of highland Bolivian women. In this study of 189 women, gestational age of the pregnancy losses was not reported, but for the 14 losses that were likely to have happened at very early gestational ages, the mean bleed duration of 3.3 days did not differ markedly from the study population's mean non-conception menses duration of 3.6 days (Vitzthum *et al.* 2001).

Most information on bleeding with pregnancy loss comes from studies of expectant management of women seeking medical care for miscarriage. The majority of these miscarriages occurred in the latter half of the first trimester. Mean length of bleeding with these miscarriages was in the range of 9–11 days (Haines *et al.* 1994;Nielsen and Hahlin 1995;Chung *et al.* 1998;Nielsen *et al.* 1999;Bagratee *et al.* 2004). For comparison, the mean duration of menstrual bleeding among a large cohort of predominantly white, college-educated, United States women has been reported to be 5.3 days (Cooper *et al.* 1996), virtually the same as the 5.4 days seen for non-conception cycles in our study (Table I).

While variations in maximal hCG production and pregnancy duration were unrelated to the length of the bleeding associated with the loss in Early Pregnancy Study women, they were correlated with intensity of bleeding (compared with usual menses). An increase in bleeding for longer lasting pregnancies is not surprising; such pregnancies will generally have reached a later stage of development, with more advanced decidualization and trophoblast invasion (Cunningham *et al.* 2005). However, the reduction in bleeding seen with pregnancy losses of shorter duration and lower hCG production was surprising. It may be that a menstrual cycle with poorer endometrial development may increase the risk of very early pregnancy loss. A related hypothesis is that a hormonally suboptimal cycle predisposes the woman to both suboptimal endometrial development and very early pregnancy loss, producing an association of the two. Alternatively, impending early loss may produce a change in the endometrium that results in reduced shedding of tissue.

In conclusion, we found that the bleeding after a pregnancy loss of less than six weeks gestation was only slightly longer than the woman's typical menstrual bleeding. This is counter to expectations based on observations of later miscarriages. Overall, there was no meaningful difference in the number of pads and tampons women used after a pregnancy loss compared to what they used in a typical menstrual period (either as recorded per day or in total over the course of the bleeding episode). In general, it seems unlikely that bleeding after an early

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pregnancy loss is different enough from a woman's usual menstrual bleeding to trigger suspicion of pregnancy loss.

Maximal hCG production and pregnancy duration were, however, correlated with the amount of bleeding associated with the loss. Unexpectedly, women bled less than usual following those pregnancy losses with the shortest duration and lowest maximal hCG production. This apparent reduction in bleeding after the very earliest losses bears further investigation, since it could indicate the presence of possible physiologic changes in the earliest days after conception and implantation that may be relevant to fecundity and the risk of early loss.

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### References

- Armstrong EG, Ehrlich PH, Birken S, Schlatterer JP, Siris E, Hembree W, Canfield RE. Use of a highly sensitive and specific immunoradiometric assay for detection of human chorionic gonadotropin in urine of normal, nonpregnant and pregnant individuals. J Clin Endocrinol Metab 1984;59:867–874. [PubMed: 6480810]
- Bagratee JS, Khullar V, Regan L, Moodley J, Kagoro H. A randomized controlled trial comparing medical and expectant management of first trimester miscarriage. Hum Reprod 2004;19:266–271. [PubMed: 14747165]
- Baird DD, Weinberg CR, Wilcox AJ, McConnaughey DR, Musey PI. Using the ratio of urinary oestrogen and progesterone metabolites to estimate day of ovulation. Stat Med 1991;10:255–266. [PubMed: 2052803]
- Chung TKH, Cheung LP, Sahota DS, Haines CJ, Chang AMZ. Spontaneous abortion: Short-term complications following either conservative or surgical management. Aust NZ J Obstet Gynaecol 1998;38:61–64.
- Cooper GS, Sandler DP, Whelan EA, Smith KR. Association of physical and behavioral characteristics with menstrual cycle patterns in women age 29–31 years. Epidemiol 1996;7:624–628.
- Cunningham, FG.; Hauth, JC.; Leveno, KJ.; Gilstrap, L., III; Bloom, SL.; Wenstrom, KD., editors. Williams Obstetrics. New York: McGraw-Hill; 2005.
- Fraser IS, Pearse C, Shearman RP, Elliott PM, McIlveen J, Markham R. Efficacy of mefenamic acid in patients with a complaint of menorrhagia. Obstet Gynecol 1981;58:543–551. [PubMed: 7029369]
- Haines CJ, Chung T, Leung DYL. Transvaginal sonography and the conservative management of spontaneous abortion. Gynecol Obstet Invest 1994;37:14–17. [PubMed: 8125401]
- Higham JM, O'Brien PM, Shaw RW. Assessment of menstrual blood loss using a pictorial chart. Br J Obstet Gynaecol 1990;97:734–739. [PubMed: 2400752]
- Higham JM, Shaw RW. Clinical associations with objective menstrual blood volume. Eur J Obstet Gnecol Reprod Biol 1999;82:73–76.
- McChesney R, Wilcox AJ, O'Connor JF, Weinberg CR, Baird DD, Schlatterer JP, McConnaughey DR, Birken S, Canfield RE. Intact HCG, free HCG β subunit and HCG β core fragment: longitudinal patterns in urine during early pregnancy. Hum Reprod 2005;20:928–935. [PubMed: 15665026]
- Nielsen S, Hahlin M. Expectant management of first-trimester spontaneous abortion. Lancet 1995;345:84–86. [PubMed: 7815886]
- Nielsen S, Hahlin M, Platz-Christensen J. Randomised trial comparing expectant with medical management for first trimester miscarriages. Br J Obstet Gynaecol 1999;106:804–807. [PubMed: 10453830]
- Oats, J.; Abraham, S. Llewellyn-Jones Obstetrics and Gynaecology. Edinburgh: Elsevier Mosby; 2005.
- Vitzthum VJ, Spielvogel H, Caceres E, Miller A. Vaginal bleeding patterns among rural highland Bolivian women: Relationship to fecundity and fetal loss. Contraception 2001;64:319–325. [PubMed: 11777494]

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- Wang X, Chen C, Wang L, Chen D, Guang W, French J. Conception, early pregnancy loss, and time to clinical pregnancy: a population-based prospective study. Fertil Steril 2003;79:577–584. [PubMed: 12620443]
- Warner PE, Critchley HOD, Lumsden M, Campbell-Brown M, Douglas A, Murray GD. Menorrhagia I: Measured blood loss, clinical features, and outcome in women with heavy periods: A survey with follow-up data. Am J Obstet Gynecol 2004;190:1216–1223. [PubMed: 15167821]
- Wilcox AJ, Weinberg CR, Baird DD. Risk factors for early pregnancy loss. Epidemiol 1990;1:382–385.
- Wilcox AJ, Weinberg CR, O'Connor JF, Baird DD, Schlatterer JP, Canfield RE, Armstrong EG, Nisula BC. Incidence of early loss of pregnancy. N Engl J Med 1988;319:189–194. [PubMed: 3393170]
- World Health Organization Task Force on Adolescent Reproductive Health. World Health Organization multicenter study on menstrual and ovulatory patterns in adolescent girls. J Adol Health 1986;7:236–244.
- Zinaman MJ, Clegg ED, Brown CC, O'Connor J, Selevan SG. Estimates of human fertility and pregnancy loss. Fertil Steril 1996;65:503–509. [PubMed: 8774277]



#### Figure 1.

Mean daily pad plus tampon use total by day of bleed period for 36 women for bleeds following their early pregnancy losses (solid line) versus their cycles with no detected conception (dashed line). Mean daily pad plus tampon totals are the across women means of each woman's mean values. Every menses contributes data for all 12 days (zero values after bleeding ends).



#### Figure 2.

Plot of each woman's bleed length (days) for the bleed period following her early pregnancy loss (mean value used for the two women with two losses each) versus her mean bleed length (days) for her bleed periods following cycles with no detected conception (N = 36 women). Dashed line is line of equality.



Figure 3.

Plot of the within-woman difference in total number of pads plus tampons used during bleed period (total number used for bleed following early pregnancy loss minus mean of totals for bleeds following cycles with no detected conception) versus the maximum hCG value (shown on a log scale) obtained for the pregnancy that was lost (N = 36 women). Spearman correlation coefficient = 0.37 (*P*-value = 0.02).

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Table I

n versus menstrual bleed characteristics	Early Loss – No Conception
a pregnancy lost prior to six weeks gestation	No Conception
eed characteristics associated with conception (N = $36$ women)	Early Loss
Within-woman comparison of ble following cycles with no detected	Characteristics of Pad +

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Characteristics of Pad +		Early	' Loss			No Con	ception		Early	Loss – No Co	nception
	Mean	SE <sup>a</sup>	$25^{\mathrm{th}}b$	$75^{\mathrm{th}b}$	Mean	SE	$25^{\text{th}}$	75 <sup>th</sup>	Mean	SE	<i>p</i> -value <sup><i>c</i></sup>
Length of use (days)	5.8	0.3	5.0	6.8	5.4	0.2	4.9	6.0	0.4	0.2	0.01
Total no. used during bleed period	18.2	1.2	12.5	23.0	18.4	1.3	13.0	24.3	-0.2	0.9	0.94
Max. no. used in one day	5.2	0.4	3.0	6.0	5.4	0.4	3.6	7.3	-0.3	0.3	0.53
Mean no. per bleeding day	3.2	0.2	2.2	4.1	3.5	0.2	2.4	4.4	-0.2	0.1	0.06
Period day of heaviest use	2.3	0.1	1.9	3.0	2.2	0.1	1.8	2.5	0.0	0.1	0.78
No. of light days $(0-3)^{d}$ , e	3.8	0.3	3.0	5.0	3.3	0.3	2.0	5.0	0.5	0.2	0.004
No. of medium days (4–6)	1.6	0.2	0.0	3.0	1.6	0.2	0.8	2.0	0.1	0.2	0.90
No. of heavy days $(7+)$	0.4	0.1	0.0	0.0	0.6	0.1	0.0	1.0	-0.2	0.1	0.15
a Standard Error											
-											

 $b_{25}$ th, 75th percentile values

 $^{c}$ Wilcoxon signed rank test

 $d_{\rm N}$  umbers in parentheses indicate daily total for pad + tampon use.

 $^{e}$  One bleed episode has a single day with pads + tampons = 0.

## Table II

## Correlation of bleed characteristics with markers of the strength of the lost pregnancy (N = 36 women)

(Characteristics of Pad + Tampon Use Within-		Pregnar	ey Marker	
Woman Difference) <sup>w</sup>	Maximum hCG		Days from ovul	ation to bleeding
	$\rho^b$	<i>P</i> -value	ρ <sup>b</sup>	<i>P</i> -value
Length of use (days)	0.09	0.59	0.07	0.66
Total no. used during bleed period	0.37	0.02	0.48	0.003
Max. no. used in one day	0.16	0.35	0.16	0.34
Mean no. per bleeding day	0.29	0.08	0.47	0.004
Period day of heaviest use	-0.11	0.53	-0.29	0.09
No. of light days $(0-3)^c d$	-0.38	0.02	-0.42	0.01
No. of medium days $(4-6)$	0.32	0.05	0.39	0.02
No. of heavy days (7+)	0.05	0.79	0.08	0.66

 $^{a}$ Value for woman's bleed following her early loss minus mean value for her bleeds following cycles with no detected conception

<sup>b</sup>Spearman correlation coefficient

 $^{C}$ Numbers in parentheses indicate daily total for pad + tampon use.

 $^{d}$ One bleed episode has a single day with pads + tampons = 0.

#### Table III

Within-woman difference in total number of pads plus tampons used per bleed by quartile of Ln (Maximum hCG).

Maximum hCG	No.	Total No. of Pads + Tampons: Within Conce	Total No. of Pads + Tampons: Within Woman Difference (Early Loss – No Conception)		
		Mean	SE		
$\leq 0.12$ 0.12 to 0.27 0.27 to 1.65 > 1.65	9 9 9 9	-3.12 -0.23 -1.34 3.87	2.01 1.07 1.64 1.48		

<sup>*a*</sup> Overall *p*-value based on rank correlation = 0.01.