

UC Davis

UC Davis Previously Published Works

Title

Return of fertility in nulliparous and parous women after levonorgestrel 52 mg intrauterine system discontinuation

Permalink

<https://escholarship.org/uc/item/1s1994f3>

Journal

Fertility and Sterility, 110(4)

ISSN

0015-0282

Authors

Carr, BR
Thomas, MA
Gangestad, A
et al.

Publication Date

2018-09-01

DOI

10.1016/j.fertnstert.2018.07.145

Peer reviewed

status, income quartile, patient urban-rural designation, hospital geographic region and hospital teaching status. We compared data to visits by women in the same age range without an EC diagnosis. Z-tests were performed to assess any significant differences between groups.

RESULTS: The estimated number of encounters for EC in the ED decreased significantly during the study period. The steepest decline occurred from 2006 (15,039 visits; 95%CI: 11,270 - 18,902) to 2007 (4,370 visits; 95%CI: 3,523 - 5,217). After 2007, visits decreased steadily, reaching 685 visits in 2014 (95%CI: 488 - 881). Overall, this represented a 95% decrease in visits over time. Annual total ED charges decreased from \$6.2M in 2006 to \$0.65M in 2014. When compared to all other ED visits by women aged 15-44 yo in 2014, ED visits for EC were more likely to be by women aged 15-19 yo (30.7% vs 13.3%; $p < .001$), who lived in the Northeast (53.7% vs 17.7%; $p < .001$), lived in large metropolitan areas (64.2% vs 48.1%; $p = 0.010$), or presented to metropolitan teaching hospitals (64.0% vs 54.4%; $p = 0.111$). The proportion of visits for EC by patients who lived in zip codes with average incomes in the lowest and second income quartiles was 29.9% (95%CI: 20.7% - 39.2%), and 36.2% (95%CI: 23.4% - 49.0%), respectively in 2014. Trends by payer showed that patients with private insurance or self-pay status experienced steeper declines in ED visit counts compared to patients with Medicaid, with estimated percent changes of -98%, -98% and -83% between 2006-2014 respectively.

CONCLUSIONS: ED utilization for emergency contraception decreased substantially between 2006-2014, demonstrating the influence of policy on these care seeking patterns. This decrease also led to cost-savings in the ED by reducing utilization. Nonetheless, trends by payer indicate a potential disparity in how women currently access (ED vs other outlets) and can pay for EC. Future policies should ensure that patients seeking care for unintended pregnancy are provided timely and cost-effective care.

Supported by: University of Michigan.

O-104 Tuesday, October 9, 2018 11:00 AM

RETURN OF FERTILITY IN NULLIPAROUS AND PAROUS WOMEN AFTER LEVONORGESTREL 52 MG INTRAUTERINE SYSTEM DISCONTINUATION.

B. R. Carr,^a M. A. Thomas,^b A. Gangestad,^c D. L. Eisenberg,^d A. I. Olariu,^e M. D. Creinin.^f ^aUTSWMC, Dallas, TX; ^bObstetrics and Gynecology, University of Cincinnati, Cincinnati, OH; ^cObstetrics and Gynecology, Case Western University, Cleveland, OH; ^dObstetrics and Gynecology, Washington University in St. Louis, St. Louis, MO; ^eMedicines360, San Francisco, CA; ^fObstetrics and Gynecology, University of California, Davis, Sacramento, CA.

OBJECTIVE: Evaluate reproductive function in nulliparous and parous women after levonorgestrel (LNG) 52mg intrauterine system (IUS) discontinuation based on time to pregnancy.

DESIGN: Prospective clinical trial.

MATERIALS AND METHODS: Eligible nulliparous and parous women 16-45 years old received the Liletta® IUS in a multicenter trial evaluating efficacy and safety for up to 10 years. Participants who discontinued the IUS within the first 5 years of use and desired pregnancy were contacted every three months for up to 12 months to determine whether pregnancy occurred; pregnancy outcomes were not assessed. We evaluated fertility endpoints in women 16-35 years at study entry. We compared outcomes by parity, gravidity and age using Fisher's exact test, student's t-tests and chi-square test for trend as indicated.

RESULTS: Overall, 132 (86.8%) of 152 women 16-35 years old at study entry who attempted to conceive did so within 12 months. The 12-month conception rates did not differ between nulliparous (70/80 [87.5%]) and par-

ous (62/72 [86.1%]) women ($p = 0.82$) or between nulligravid (60/68 [88.2%]) and gravid (72/84 [85.7%]) women ($p = 0.81$). Pregnancy by the end of 3 and 6 months occurred in 66 (43.4%) and 106 (69.7%), respectively, with a median time to conception of 91.5 days. Median use before discontinuation for the evaluated population was 34 months (range 1.3-59.8 months). Pregnancy rates at 12 months post-discontinuation did not differ by length of IUS use for nulliparous or parous women (Table). The age at IUS discontinuation of women who successfully conceived (29.8±4.0 years) did not differ from women who did not conceive (29.6±4.3 years), $p = 0.84$.

CONCLUSIONS: After Liletta discontinuation, women have rapid return of normal fertility as demonstrated by time to pregnancy in the year post-removal. Fertility rates after IUS removal do not vary based on parity, gravidity, duration of IUS use or age at discontinuation.

Supported by: Medicines360.

O-105 Tuesday, October 9, 2018 11:15 AM

PLASMA LEVONORGESTREL LEVELS IN NON-OBESE AND OBESE WOMEN USING A LEVONORGESTREL 52 MG INTRAUTERINE SYSTEM FOR UP TO 7 YEARS.

M. D. Creinin,^a A. Gangestad,^b T. D. Kimble,^c B. Carr,^d A. Olariu,^e C. L. Westhoff.^f ^aObstetrics and Gynecology, University of California, Davis, Sacramento, CA; ^bObstetrics and Gynecology, Case Western Reserve University, Cleveland, OH; ^cObstetrics and Gynecology, Eastern Virginia Medical School, Norfolk, VA; ^dUTSWMC, Dallas, TX; ^eMedicines360, San Francisco, CA; ^fObstetrics and Gynecology, Columbia University, New York, NY.

OBJECTIVE: To evaluate levonorgestrel plasma concentrations for up to seven years in non-obese and obese women using a levonorgestrel (LNG) 52mg intrauterine system (IUS).

DESIGN: Prospective clinical trial.

MATERIALS AND METHODS: Eligible women 16-45 years old received an LNG 52mg IUS (Liletta®) in a multicenter trial evaluating efficacy and safety for up to 10 years. A planned sub-study enrolled 40 participants (19 obese, 21 non-obese) to evaluate LNG concentrations over time at baseline, weeks 1 and 2, and months 1, 3, 6, 9, 12, 18, 24, 30 and 36. Additionally, all study subjects (219 obese and 670 non-obese) began blood sampling every 6 months at month 36. The liquid chromatography-tandem mass spectrometry assay had a lower limit of LNG detection of 25 pg/mL. We compared LNG concentrations in obese and non-obese women at each time point through 84 months (7 years) using an independent-samples t-test.

RESULTS: Plasma LNG concentration was lower in obese compared with non-obese subjects at all time-points through 84 months (Table). All obese users had maximum levels <300 pg/mL at day 30 and thereafter. Maximum LNG concentrations in non-obese LNG 52mg IUS users were 603 pg/mL at week 1, 492 pg/mL at week 14, and <300 pg/mL by 1 year. From 3 to 84 months, LNG concentrations were 21-41% lower in obese subjects ($p < 0.01$ for all months). Average BMI in the obese and non-obese sub-study participants were $38.6 \pm 5.8 \text{ kg/m}^2$ and $24.6 \pm 2.9 \text{ kg/m}^2$, respectively, with ranges of 30.4-49.2 kg/m^2 and 18.9-29.5 kg/m^2 , respectively. Average BMI in the obese and non-obese study subjects beginning sampling at 36 months were similar, with 23.2% of obese subjects having a BMI $\geq 40 \text{ kg/m}^2$.

CONCLUSIONS: Obese women demonstrate lower plasma LNG concentrations throughout seven years of LNG 52mg IUS use. The LNG concentrations in obese and non-obese women may be helpful for patient education

Mean Plasma Levonorgestrel Levels in Levonorgestrel 52mg IUS Users

Time Point (months)	Obese		Non-Obese		P-value
	N	LNG (pg/mL)	N	LNG (pg/mL)	
0.25 (1 week)	19	188 ± 53	21	310 ± 140	<.001
1	19	180 ± 45	21	248 ± 83	.006
6	16	151 ± 40	20	230 ± 67	<.001
12	15	144 ± 52	18	192 ± 36	.006
24	15	116 ± 27	15	178 ± 38	.002
36	219	104 ± 53	670	141 ± 51	<.001
48	182	90 ± 44	550	121 ± 52	<.001
60	95	83 ± 46	250	114 ± 45	<.001
72	65	69 ± 29	133	102 ± 39	<.001
84	47	64 ± 29	94	94 ± 32	<.001

Pregnancy Rates by Parity After Levonorgestrel 52mg IUS Discontinuation					
Total		Nulliparous		Parous	
Years of IUS Use	p- value*	Pregnant	p- value*	Pregnant	p- value*
≤1	0.43	11/12 (92%)	0.62	3/3 (100%)	0.29
1+ to 2		27/30 (90%)		11/12 (92%)	
2+ to 3		36/41 (88%)		17/21 (81%)	
3+ to 4		33/42 (79%)		21/25 (84%)	
4+ to 5		25/27 (93%)		18/19 (95%)	

*Chi-square test for trend