

Please note: An update has been published for this report. To view the update, please click here

Centers for Disease Control and Prevention
MMWR

Morbidity and Mortality Weekly Report

Recommendations and Reports / Vol. 65 / No. 4

July 29, 2016

U.S. Selected Practice Recommendations for Contraceptive Use, 2016

Continuing Education Examination available at <http://www.cdc.gov/mmwr/cme/conted.html>.



U.S. Department of Health and Human Services
Centers for Disease Control and Prevention

CONTENTS

Introduction	1
Summary of Changes from the 2013 U.S. SPR	2
Methods.....	2
Maintaining Updated Guidance	3
How To Use This Document.....	3
Contraceptive Method Choice	4
How To Be Reasonably Certain that a Woman Is Not Pregnant.....	5
Intrauterine Contraception.....	7
Implants.....	14
Injectables.....	18
Combined Hormonal Contraceptives.....	22
Progestin-Only Pills.....	31
Standard Days Method.....	34
Emergency Contraception.....	34
Female Sterilization	36
Male Sterilization.....	37
When Women Can Stop Using Contraceptives	38
Conclusion	39
References.....	41

CONTENTS (Continued)

Appendix A: Summary Chart of U.S. Medical Eligibility Criteria for Contraceptive Use, 2016.....	53
Appendix B: When To Start Using Specific Contraceptive Methods	62
Appendix C: Examinations and Tests Needed Before Initiation of Contraceptive Methods.....	63
Appendix D: Routine Follow-Up After Contraceptive Initiation.....	64
Appendix E: Management of Women with Bleeding Irregularities While Using Contraception	65
Appendix F: Management of Intrauterine Devices When Users are Found To Have Pelvic Inflammatory Disease	66

Disclosure of Relationship

CDC, our planners, and our content experts wish to disclose they have no financial interest or other relationships with the manufacturers of commercial products, suppliers of commercial services, or commercial supporters. Planners have reviewed content to ensure there is no bias. This document will not include any discussion of the unlabeled use of a product or a product under investigational use, with the exception that some of the recommendations in this document might be inconsistent with package labeling.

The *MMWR* series of publications is published by the Center for Surveillance, Epidemiology, and Laboratory Services, Centers for Disease Control and Prevention (CDC), U.S. Department of Health and Human Services, Atlanta, GA 30329-4027.

Suggested citation: [Author names; first three, then et al., if more than six.] [Title]. *MMWR Recomm Rep* 2016;65(No. RR-#):[inclusive page numbers].

Centers for Disease Control and Prevention

Thomas R. Frieden, MD, MPH, *Director*
 Harold W. Jaffe, MD, MA, *Associate Director for Science*
 Joanne Cono, MD, ScM, *Director, Office of Science Quality*
 Chesley L. Richards, MD, MPH, *Deputy Director for Public Health Scientific Services*
 Michael F. Iademarco, MD, MPH, *Director, Center for Surveillance, Epidemiology, and Laboratory Services*

MMWR Editorial and Production Staff (Serials)

Sonja A. Rasmussen, MD, MS, *Editor-in-Chief*
 Charlotte K. Kent, PhD, MPH, *Executive Editor*
 Christine G. Casey, MD, *Editor*
 Teresa F. Rutledge, *Managing Editor*
 David C. Johnson, *Lead Technical Writer-Editor*
 Catherine B. Lansdowne, MS, *Project Editor*

Martha F. Boyd, *Lead Visual Information Specialist*
 Maureen A. Leahy, Julia C. Martinroe,
 Stephen R. Spriggs, Moua Yang, Tong Yang,
Visual Information Specialists
 Quang M. Doan, MBA, Phyllis H. King, Terraye M. Starr,
Information Technology Specialists

MMWR Editorial Board

Timothy F. Jones, MD, *Chairman*
 Matthew L. Boulton, MD, MPH
 Virginia A. Caine, MD
 Katherine Lyon Daniel, PhD
 Jonathan E. Fielding, MD, MPH, MBA
 David W. Fleming, MD

William E. Halperin, MD, DrPH, MPH
 King K. Holmes, MD, PhD
 Robin Ikeda, MD, MPH
 Rima F. Khabbaz, MD
 Phyllis Meadows, PhD, MSN, RN
 Jewel Mullen, MD, MPH, MPA

Jeff Niederdeppe, PhD
 Patricia Quinlisk, MD, MPH
 Patrick L. Remington, MD, MPH
 Carlos Roig, MS, MA
 William L. Roper, MD, MPH
 William Schaffner, MD

U.S. Selected Practice Recommendations for Contraceptive Use, 2016

Kathryn M. Curtis, PhD¹
 Tara C. Jattaoui, MD¹
 Naomi K. Tepper, MD¹
 Lauren B. Zapata, PhD¹
 Leah G. Horton, MSPH¹
 Denise J. Jamieson, MD¹
 Maura K. Whiteman, PhD¹

¹Division of Reproductive Health, National Center for Chronic Disease Prevention and Health Promotion

Summary

The 2016 U.S. Selected Practice Recommendations for Contraceptive Use (U.S. SPR) addresses a select group of common, yet sometimes controversial or complex, issues regarding initiation and use of specific contraceptive methods. These recommendations for health care providers were updated by CDC after review of the scientific evidence and consultation with national experts who met in Atlanta, Georgia, during August 26–28, 2015. The information in this report updates the 2013 U.S. SPR (CDC. U.S. selected practice recommendations for contraceptive use, 2013. MMWR 2013;62[No. RR-5]). Major updates include 1) revised recommendations for starting regular contraception after the use of emergency contraceptive pills and 2) new recommendations for the use of medications to ease insertion of intrauterine devices. The recommendations in this report are intended to serve as a source of clinical guidance for health care providers and provide evidence-based guidance to reduce medical barriers to contraception access and use. Health care providers should always consider the individual clinical circumstances of each person seeking family planning services. This report is not intended to be a substitute for professional medical advice for individual patients. Persons should seek advice from their health care providers when considering family planning options.

Introduction

Unintended pregnancy rates remain high in the United States; approximately 45% of all pregnancies are unintended, with higher proportions among adolescent and young women, women who are racial/ethnic minorities, and women with lower levels of education and income (1). Unintended pregnancies increase the risk for poor maternal and infant outcomes (2) and in 2010, resulted in U.S. government health care expenditures of \$21 billion (3). Approximately half of unintended pregnancies are among women who were not using contraception at the time they became pregnant; the other half are among women who became pregnant despite reported use of contraception (4). Strategies to prevent unintended pregnancy include assisting women at risk for unintended pregnancy and their partners with choosing appropriate contraceptive methods and helping them use methods correctly and consistently to prevent pregnancy.

In 2013, CDC published the first *U.S. Selected Practice Recommendations for Contraceptive Use* (U.S. SPR), adapted from global guidance developed by the World Health Organization (WHO SPR), which provided evidence-based guidance on how to use contraceptive methods safely and effectively once they are deemed to be medically appropriate.

U.S. SPR is a companion document to *U.S. Medical Eligibility Criteria for Contraceptive Use* (U.S. MEC) (<http://www.cdc.gov/reproductivehealth/unintendedpregnancy/usmec.htm>), which provides recommendations on safe use of contraceptive methods for women with various medical conditions and other characteristics (5). WHO intended for the global guidance to be used by local or national policy makers, family planning program managers, and the scientific community as a reference when they develop family planning guidance at the country or program level. During 2012–2013, CDC went through a formal process to adapt the global guidance for best implementation in the United States, which included rigorous identification and critical appraisal of the scientific evidence through systematic reviews, and input from national experts on how to translate that evidence into recommendations for U.S. health care providers (6). At that time, CDC committed to keeping this guidance up to date and based on the best available evidence, with full review every few years (6).

This document updates the 2013 U.S. SPR (6) with new evidence and input from experts. Major updates include 1) revised recommendations for starting regular contraception after the use of emergency contraceptive pills and 2) new recommendations for the use of medications to ease insertion of intrauterine devices (IUDs). Recommendations are provided for health care providers on the safe and effective use of contraceptive methods and address provision of contraceptive methods and management of side effects and other problems

Corresponding author: Kathryn M. Curtis, PhD, Division of Reproductive Health, National Center for Chronic Disease Prevention and Health Promotion, CDC. Telephone: 770-488-5200; E-mail: kmc6@cdc.gov.

with contraceptive method use, within the framework of removing unnecessary medical barriers to accessing and using contraception. These recommendations are meant to serve as a source of clinical guidance for health care providers; health care providers should always consider the individual clinical circumstances of each person seeking family planning services. This report is not intended to be a substitute for professional medical advice for individual patients, who should seek advice from their health care providers when considering family planning options.

Summary of Changes from the 2013 U.S. SPR

Updated Recommendations

Recommendations have been updated regarding when to start regular contraception after ulipristal acetate (UPA) emergency contraceptive pills:

- Advise the woman to start or resume hormonal contraception no sooner than 5 days after use of UPA, and provide or prescribe the regular contraceptive method as needed. For methods requiring a visit to a health care provider, such as depo-medroxyprogesterone acetate (DMPA), implants, and IUDs, starting the method at the time of UPA use may be considered; the risk that the regular contraceptive method might decrease the effectiveness of UPA must be weighed against the risk of not starting a regular hormonal contraceptive method.
- The woman needs to abstain from sexual intercourse or use barrier contraception for the next 7 days after starting or resuming regular contraception or until her next menses, whichever comes first.
- Any nonhormonal contraceptive method can be started immediately after the use of UPA.
- Advise the woman to have a pregnancy test if she does not have a withdrawal bleed within 3 weeks.

New Recommendations

New recommendations have been made for medications to ease IUD insertion:

- Misoprostol is not recommended for routine use before IUD insertion. Misoprostol might be helpful in select circumstances (e.g., in women with a recent failed insertion).
- Paracervical block with lidocaine might reduce patient pain during IUD insertion.

Methods

Since publication of the 2013 U.S. SPR, CDC has monitored the literature for new evidence relevant to the recommendations through the WHO/CDC continuous identification of research evidence (CIRE) system (7). This system identifies new evidence as it is published and allows WHO and CDC to update systematic reviews and facilitate updates to recommendations as new evidence warrants. Automated searches are run in PubMed weekly, and the results are reviewed. Abstracts that meet specific criteria are added to the web-based CIRE system, which facilitates coordination and peer review of systematic reviews for both WHO and CDC. In 2014, CDC reviewed all of the existing recommendations in the 2013 U.S. SPR for new evidence identified by CIRE that had the potential to lead to a changed recommendation. During August 27–28, 2014, CDC held a meeting in Atlanta, Georgia, of 11 family planning experts and representatives from partner organizations to solicit their input on the scope of and process for updating both the 2010 U.S. MEC and the 2013 U.S. SPR. The participants were experts in family planning and represented different provider types and organizations that represent health care providers. A list of participants is provided at the end of this report. The meeting related to topics to be addressed in the update of U.S. SPR based on new scientific evidence published since 2013 (identified through the CIRE system), topics addressed at a 2014 WHO meeting to update global guidance, and suggestions CDC received from providers for the addition of recommendations not included in the 2013 U.S. SPR (e.g., from provider feedback through e-mail, public inquiry, and questions received at conferences). CDC identified one topic to consider adding to the guidance: the use of medications to ease IUD insertion (evidence question: “Among women of reproductive age, does use of medications before IUD insertion improve the safety or effectiveness of the procedure [ease of insertion, need for adjunctive insertion measures, or insertion success] or affect patient outcomes [pain or side effects] compared with nonuse of these medications?”). CDC also identified one topic for which new evidence warranted a review of an existing recommendation: initiation of regular contraception after emergency contraceptive pills (evidence question: “Does ulipristal acetate for emergency contraception interact with regular use of hormonal contraception leading to decreased effectiveness of either contraceptive method?”). CDC determined that all other recommendations in the 2013 U.S. SPR were up to date and consistent with the current body of evidence for that recommendation.

In preparation for a subsequent expert meeting August 26–28, 2015, to review the scientific evidence

for potential recommendations, CDC staff conducted independent systematic reviews for each of the topics being considered. The purpose of these systematic reviews was to identify direct evidence related to the common clinical challenges associated with the recommendations. Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines were followed for reporting systematic reviews (8,9), and strength and quality of the evidence were assigned using the system of the U.S. Preventive Services Task Force (10). When direct evidence was limited or not available, indirect evidence (e.g., evidence on surrogate outcomes) and theoretical issues were considered and either added to direct evidence within a systematic review or separately compiled for presentation to the meeting participants. Completed systematic reviews were peer reviewed by two or three experts and then provided to participants before the expert meeting. Reviews are referenced throughout this document; the full reviews have been published and contain the details of each review, including systematic review question, literature search protocol, inclusion and exclusion criteria, evidence tables, and quality assessment. CDC staff continued to monitor new evidence identified through the CIRE system during the preparation for the August 2015 meeting.

During August 26–28, 2015, CDC held a meeting in Atlanta, Georgia, of 29 participants who were invited to provide their individual perspectives on the scientific evidence presented and to discuss potential recommendations that followed. Participants represented a wide range of expertise in family planning provision and research and included obstetrician/gynecologists, pediatricians, family physicians, nurse practitioners, epidemiologists, and others with research and clinical practice expertise in contraceptive safety, effectiveness, and management. Lists of participants and any potential conflicts of interest are provided at the end of this report. During the meeting, the evidence from the systematic review for each topic was presented, including direct evidence and any indirect evidence or theoretical concerns. Participants provided their perspectives on using the evidence to develop the recommendations that would meet the needs of U.S. health care providers. After the meeting, CDC determined the recommendations in this report, taking into consideration the perspectives provided by the meeting participants. Feedback also was received from four external reviewers, composed of health care providers and researchers who had not participated in the update meetings. These providers were asked to provide comments on the accuracy, feasibility, and clarity of the recommendations. Areas of research that need additional investigation also were considered during the meeting (11).

Maintaining Updated Guidance

As with any evidence-based guidance document, a key challenge is keeping the recommendations up to date as new scientific evidence becomes available. Working with WHO, CDC uses the CIRE system to ensure that WHO and CDC guidance is based on the best available evidence and that a mechanism is in place to update guidance when new evidence becomes available (7). CDC will continue to work with WHO to identify and assess all new relevant evidence and determine whether changes in the recommendations are warranted. In most cases, U.S. SPR will follow any updates in the WHO guidance, which typically occurs every 5 years (or sooner if warranted by new data). In addition, CDC will review any interim WHO updates for their application in the United States. CDC also will identify and assess any new literature for the recommendations that are not included in the WHO guidance and will completely review U.S. SPR every 5 years. Updates to the guidance can be found on the U.S. SPR website (<http://www.cdc.gov/reproductivehealth/UnintendedPregnancy/USSPR.htm>).

How To Use This Document

The recommendations in this report are intended to help health care providers address issues related to use of contraceptives, such as how to help a woman initiate use of a contraceptive method, which examinations and tests are needed before initiating use of a contraceptive method, what regular follow-up is needed, and how to address problems that often arise during use, including missed pills and side effects such as unscheduled bleeding. Each recommendation addresses what a woman or health care provider can do in specific situations. For situations in which certain groups of women might be medically ineligible to follow the recommendations, comments and reference to U.S. MEC are provided (5). The full U.S. MEC recommendations and the evidence supporting those recommendations have been updated in 2016 (5) and are summarized (Appendix A).

The information in this document is organized by contraceptive method, and the methods generally are presented in order of effectiveness, from highest to lowest. However, the recommendations are not intended to provide guidance on every aspect of provision and management of contraceptive method use. Instead, they incorporate the best available evidence to address specific issues regarding common, yet sometimes complex, clinical issues. Each contraceptive method section generally includes information about initiation of the method, regular follow-up, and management of problems with use (e.g., usage errors and side effects). Each section first

provides the recommendation and then includes comments and a brief summary of the scientific evidence on which the recommendation is based. The level of evidence from the systematic reviews for each evidence summary are provided based on the U.S. Preventive Services Task Force system, which includes ratings for study design (I: randomized controlled trials; II-1: controlled trials without randomization; II-2: observational studies; and II-3: multiple time series or descriptive studies), ratings for internal validity (good, fair, or poor), and categorization of the evidence as direct or indirect for the specific review question (10).

Recommendations in this document are provided for permanent methods of contraception, such as vasectomy and female sterilization, as well as for reversible methods of contraception, including the copper-containing intrauterine device (Cu-IUD); levonorgestrel-releasing IUDs (LNG-IUDs); the etonogestrel implant; progestin-only injectables; progestin-only pills (POPs); combined hormonal contraceptive methods that contain both estrogen and a progestin, including combined oral contraceptives (COCs), a transdermal contraceptive patch, and a vaginal contraceptive ring; and the standard days method (SDM). Recommendations also are provided for emergency use of the Cu-IUD and emergency contraceptive pills (ECPs).

For each contraceptive method, recommendations are provided on the timing for initiation of the method and indications for when and for how long additional contraception, or a back-up method, is needed. Many of these recommendations include guidance that a woman can start a contraceptive method at any time during her menstrual cycle if it is reasonably certain that she is not pregnant. Guidance for health care providers on how to be reasonably certain that a woman is not pregnant also is provided.

For each contraceptive method, recommendations include the examinations and tests needed before initiation of the method. These recommendations apply to persons who are presumed to be healthy. Those with known medical problems or other special conditions might need additional examinations or tests before being determined to be appropriate candidates for a particular method of contraception. U.S. MEC might be useful in such circumstances (5). Most women need no or very few examinations or tests before initiating a contraceptive method although they might be needed to address other noncontraceptive health needs (12). Any additional screening needed for preventive health care can be performed at the time of contraception initiation, and initiation should not be delayed for test results. The following classification system was developed by WHO and adopted by CDC to categorize the applicability of the various examinations or tests before initiation of contraceptive methods (13):

Class A: These tests and examinations are essential and mandatory in all circumstances for safe and effective use of the contraceptive method.

Class B: These tests and examinations contribute substantially to safe and effective use, although implementation can be considered within the public health context, service context, or both. The risk for not performing an examination or test should be balanced against the benefits of making the contraceptive method available.

Class C: These tests and examinations do not contribute substantially to safe and effective use of the contraceptive method.

These classifications focus on the relation of the examinations or tests to safe initiation of a contraceptive method. They are not intended to address the appropriateness of these examinations or tests in other circumstances. For example, some of the examinations or tests that are not deemed necessary for safe and effective contraceptive use might be appropriate for good preventive health care or for diagnosing or assessing suspected medical conditions. Systematic reviews were conducted for several different types of examinations and tests to assess whether a screening test was associated with safe use of contraceptive methods. Because no single convention exists for screening panels for certain diseases, including diabetes, lipid disorders, and liver diseases, the search strategies included broad terms for the tests and diseases of interest.

Summary charts and clinical algorithms that summarize the guidance for the various contraceptive methods have been developed for many of the recommendations, including when to start using specific contraceptive methods (Appendix B), examinations and tests needed before initiating the various contraceptive methods (Appendix C), routine follow-up after initiating contraception (Appendix D), management of bleeding irregularities (Appendix E), and management of IUDs when users are found to have pelvic inflammatory disease (PID) (Appendix F). These summaries might be helpful to health care providers when managing family planning patients. Additional tools are available on the U.S. SPR website (<http://www.cdc.gov/reproductivehealth/UnintendedPregnancy/USSPR.htm>).

Contraceptive Method Choice

Many elements need to be considered individually by a woman, man, or couple when choosing the most appropriate contraceptive method. Some of these elements include safety, effectiveness, availability (including accessibility and affordability), and acceptability. Although most contraceptive methods are safe for use by most women, U.S. MEC provides recommendations on the safety of specific contraceptive methods for women with certain characteristics and medical conditions (5); a U.S. MEC summary (Appendix A) and the categories of medical eligibility criteria for contraceptive use (Box 1) are provided.

Voluntary informed choice of contraceptive methods is an essential guiding principle, and contraceptive counseling, where applicable, might be an important contributor to the successful use of contraceptive methods.

Contraceptive method effectiveness is critically important in minimizing the risk for unintended pregnancy, particularly

BOX 1. Categories of medical eligibility criteria for contraceptive use

- U.S. MEC 1 = A condition for which there is no restriction for the use of the contraceptive method.
- U.S. MEC 2 = A condition for which the advantages of using the method generally outweigh the theoretical or proven risks.
- U.S. MEC 3 = A condition for which the theoretical or proven risks usually outweigh the advantages of using the method.
- U.S. MEC 4 = A condition that represents an unacceptable health risk if the contraceptive method is used.

Source: Curtis KM, Jatlaoui TC, Tepper NK, et al. U.S. medical eligibility criteria for contraceptive use. *MMWR* 2016;65(No. RR-3).

Abbreviation: U.S. MEC = U.S. Medical Eligibility Criteria for Contraceptive Use.

among women for whom an unintended pregnancy would pose additional health risks. The effectiveness of contraceptive methods depends both on the inherent effectiveness of the method itself and on how consistently and correctly it is used (Figure 1). Both consistent and correct use can vary greatly with characteristics such as age, income, desire to prevent or delay pregnancy, and culture. Methods that depend on consistent and correct use by clients have a wide range of effectiveness between typical use (actual use, including incorrect or inconsistent use) and perfect use (correct and consistent use according to directions) (14). IUDs and implants are considered long-acting, reversible contraception (LARC); these methods are highly effective because they do not depend on regular compliance from the user. LARC methods are appropriate for most women, including adolescents and nulliparous women. All women should be counseled about the full range and effectiveness of contraceptive options for which they are medically eligible so that they can identify the optimal method.

In choosing a method of contraception, dual protection from the simultaneous risk for human immunodeficiency virus (HIV) and other sexually transmitted diseases (STDs) also should be considered. Although hormonal contraceptives and IUDs are highly effective at preventing pregnancy, they do not protect against STDs, including HIV. Consistent and correct use of the male latex condom reduces the risk for HIV

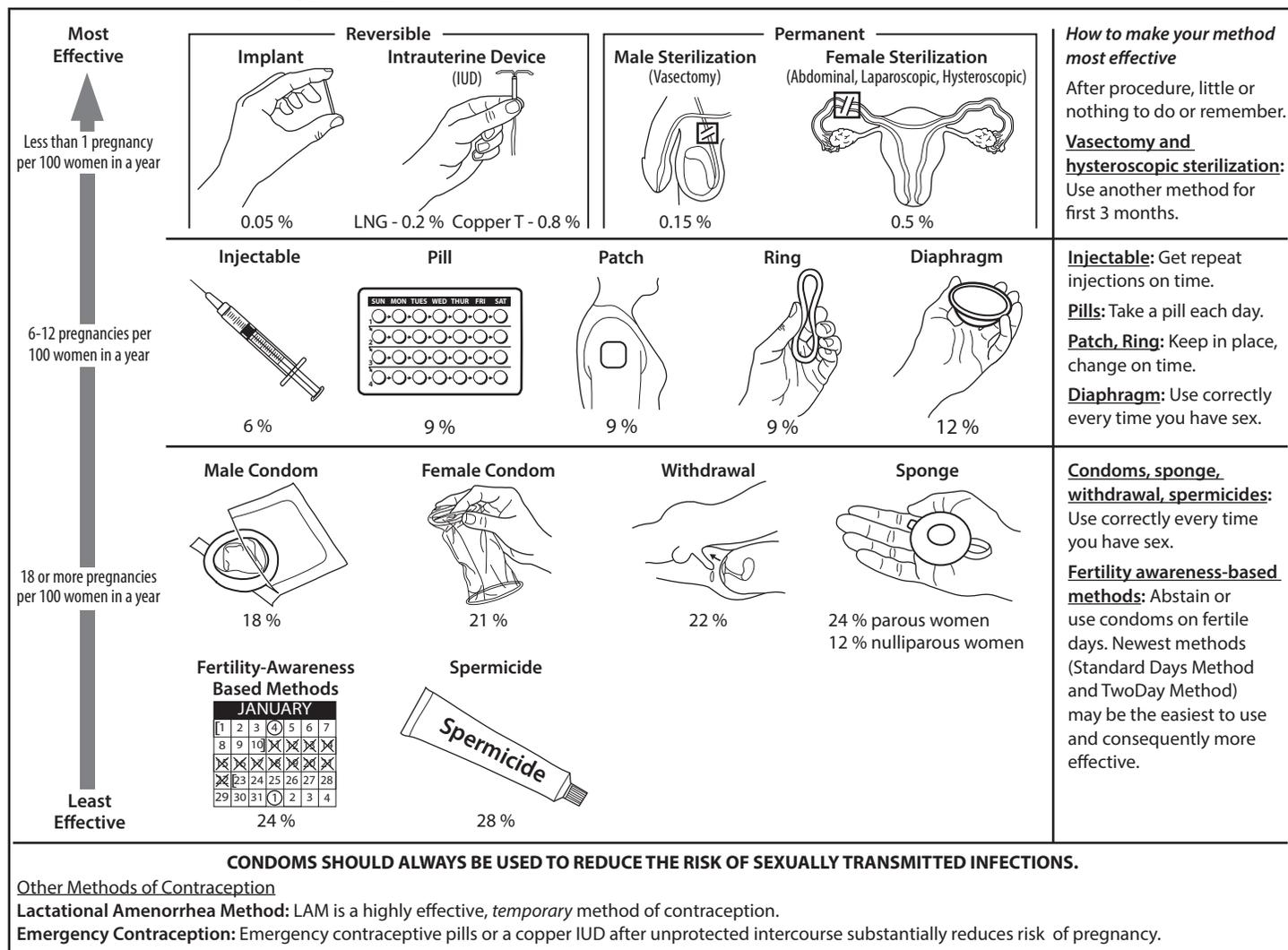
infection and other STDs, including chlamydial infection, gonococcal infection, and trichomoniasis (15). Although evidence is limited, use of female condoms can provide protection from acquisition and transmission of STDs (15). All patients, regardless of contraceptive choice, should be counseled about the use of condoms and the risk for STDs, including HIV infection (15). Additional information about prevention and treatment of STDs is available from the CDC *Sexually Transmitted Diseases Treatment Guidelines* (<http://www.cdc.gov/std/treatment>) (15).

Women, men, and couples have increasing numbers of safe and effective choices for contraceptive methods, including LARC methods such as IUDs and implants, to reduce the risk for unintended pregnancy. However, with these expanded options comes the need for evidence-based guidance to help health care providers offer quality family planning care to their patients, including assistance in choosing the most appropriate contraceptive method for individual circumstances and using that method correctly, consistently, and continuously to maximize effectiveness. Removing unnecessary barriers can help patients access and successfully use contraceptive methods. Several medical barriers to initiating and continuing contraceptive methods might exist, such as unnecessary screening examinations and tests before starting the method (e.g., a pelvic examination before initiation of COCs), inability to receive the contraceptive on the same day as the visit (e.g., waiting for test results that might not be needed or waiting until the woman's next menstrual cycle to start use), and difficulty obtaining continued contraceptive supplies (e.g., restrictions on number of pill packs dispensed at one time). Removing unnecessary steps, such as providing prophylactic antibiotics at the time of IUD insertion or requiring unnecessary follow-up procedures, also can help patients access and successfully use contraception.

How To Be Reasonably Certain that a Woman Is Not Pregnant

In most cases, a detailed history provides the most accurate assessment of pregnancy risk in a woman who is about to start using a contraceptive method. Several criteria for assessing pregnancy risk are listed in the recommendation that follows. These criteria are highly accurate (i.e., a negative predictive value of 99%–100%) in ruling out pregnancy among women who are not pregnant (16–19). Therefore, CDC recommends that health care providers use these criteria to assess pregnancy status in a woman who is about to start using contraceptives (Box 2). If a woman meets one of these criteria (and therefore the health care provider can be reasonably certain that she is not pregnant), a urine pregnancy

FIGURE 1. Effectiveness of family planning methods*



Sources: Adapted from World Health Organization (WHO) Department of Reproductive Health and Research, Johns Hopkins Bloomberg School of Public Health/Center for Communication Programs (CCP). Knowledge for health project. Family planning: a global handbook for providers (2011 update). Baltimore, MD; Geneva, Switzerland: CCP and WHO; 2011; and Trussell J. Contraceptive failure in the United States. Contraception 2011;83:397-404.

* The percentages indicate the number out of every 100 women who experienced an unintended pregnancy within the first year of typical use of each contraceptive method.

test might be considered in addition to these criteria (based on clinical judgment), bearing in mind the limitations of the accuracy of pregnancy testing. If a woman does not meet any of these criteria, then the health care provider cannot be reasonably certain that she is not pregnant, even with a negative pregnancy test. Routine pregnancy testing for every woman is not necessary.

On the basis of clinical judgment, health care providers might consider the addition of a urine pregnancy test; however, they should be aware of the limitations, including accuracy of the test relative to the time of last sexual intercourse, recent delivery, or spontaneous or induced abortion. Routine pregnancy testing for every woman is not necessary. If a woman has had recent (i.e., within the last 5 days) unprotected sexual

intercourse, consider offering emergency contraception (either a Cu-IUD or ECPs) if pregnancy is not desired.

Comments and Evidence Summary. The criteria for determining whether a woman is pregnant depend on the assurance that she has not ovulated within a certain amount of time after her last menses, spontaneous or induced abortion, or delivery. Among menstruating women, the timing of ovulation can vary widely. During an average 28-day cycle, ovulation generally occurs during days 9-20 (20). In addition, the likelihood of ovulation is low from days 1-7 of the menstrual cycle (21). After a spontaneous or an induced abortion, ovulation can occur within 2-3 weeks and has been found to occur as early as 8-13 days after the end of the pregnancy. Therefore, the likelihood of ovulation is low ≤7 days after

BOX 2. How to be reasonably certain that a woman is not pregnant

A health care provider can be reasonably certain that a woman is not pregnant if she has no symptoms or signs of pregnancy and meets any one of the following criteria:

- is ≤ 7 days after the start of normal menses
- has not had sexual intercourse since the start of last normal menses.
- has been correctly and consistently using a reliable method of contraception
- is ≤ 7 days after spontaneous or induced abortion
- is within 4 weeks postpartum
- is fully or nearly fully breastfeeding (exclusively breastfeeding or the vast majority [$\geq 85\%$] of feeds are breastfeeds), amenorrheic, and < 6 months postpartum

an abortion (22–24). A systematic review reported that the mean day of first ovulation among postpartum nonlactating women occurred 45–94 days after delivery (25). In one study, the earliest ovulation was reported at 25 days after delivery. Among women who are within 6 months postpartum, are fully or nearly fully breastfeeding (exclusively breastfeeding or the vast majority [$\geq 85\%$] of feeds are breastfeeds), and are amenorrheic, the risk for pregnancy is $< 2\%$ (26,27).

Although pregnancy tests often are performed before initiating contraception, the accuracy of qualitative urine pregnancy tests varies depending on the timing of the test relative to missed menses, recent sexual intercourse, or recent pregnancy. The sensitivity of a pregnancy test is defined as the concentration of human chorionic gonadotropin (hCG) at which 95% of tests are positive. Most qualitative pregnancy tests approved by the U.S. Food and Drug Administration (FDA) report a sensitivity of 20–25 mIU/mL in urine (28–31). However, pregnancy detection rates can vary widely because of differences in test sensitivity and the timing of testing relative to missed menses (30,32). Some studies have shown that an additional 11 days past the day of expected menses are needed to detect 100% of pregnancies using qualitative tests (29). In addition, pregnancy tests cannot detect a pregnancy resulting from recent sexual intercourse. Qualitative tests also might have positive results for several weeks after termination of pregnancy because hCG can be present for several weeks after delivery or abortion (spontaneous or induced) (33–35).

For contraceptive methods other than IUDs, the benefits of starting to use a contraceptive method likely exceed any risk, even in situations in which the health care provider is uncertain whether the woman is pregnant. Therefore, the health care provider can consider having patients start using

contraceptive methods other than IUDs at any time, with a follow-up pregnancy test in 2–4 weeks. The risks of not starting to use contraception should be weighed against the risks of initiating contraception use in a woman who might be already pregnant. Most studies have shown no increased risk for adverse outcomes, including congenital anomalies or neonatal or infant death, among infants exposed in utero to COCs (36–38). Studies also have shown no increased risk for neonatal or infant death or developmental abnormalities among infants exposed in utero to DMPA (37,39,40).

In contrast, for women who want to begin using an IUD (Cu-IUD or LNG-IUD), in situations in which the health care provider is uncertain whether the woman is pregnant, the woman should be provided with another contraceptive method to use until the health care provider is reasonably certain that she is not pregnant and can insert the IUD. Pregnancies among women with IUDs are at higher risk for complications such as spontaneous abortion, septic abortion, preterm delivery, and chorioamnionitis (41).

A systematic review identified four analyses of data from three diagnostic accuracy studies that evaluated the performance of the listed criteria (Box 2) through use of a pregnancy checklist compared with a urine pregnancy test conducted concurrently (42). The performance of the checklist to diagnose or exclude pregnancy varied, with sensitivity of 55%–100% and specificity of 39%–89%. The negative predictive value was consistent across studies at 99%–100%; the pregnancy checklist correctly ruled out women who were not pregnant. One of the studies assessed the added usefulness of signs and symptoms of pregnancy and found that these criteria did not substantially improve the performance of the pregnancy checklist, although the number of women with signs and symptoms was small (16) (Level of evidence: Diagnostic accuracy studies, fair, direct).

Intrauterine Contraception

Four IUDs are available in the United States, the copper-containing IUD and three levonorgestrel-releasing IUDs (containing a total of either 13.5 mg or 52 mg levonorgestrel). Fewer than 1 woman out of 100 becomes pregnant in the first year of using IUDs (with typical use) (14). IUDs are long-acting, are reversible, and can be used by women of all ages, including adolescents, and by parous and nulliparous women. IUDs do not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

Initiation of Cu-IUDs

Timing

- The Cu-IUD can be inserted at any time if it is reasonably certain that the woman is not pregnant (Box 2).
- The Cu-IUD also can be inserted within 5 days of the first act of unprotected sexual intercourse as an emergency contraceptive. If the day of ovulation can be estimated, the Cu-IUD also can be inserted >5 days after sexual intercourse as long as insertion does not occur >5 days after ovulation.

Need for Back-Up Contraception

- No additional contraceptive protection is needed after Cu-IUD insertion.

Special Considerations

Amenorrhea (Not Postpartum)

- **Timing:** The Cu-IUD can be inserted at any time if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** No additional contraceptive protection is needed.

Postpartum (Including After Cesarean Delivery)

- **Timing:** The Cu-IUD can be inserted at any time postpartum, including immediately postpartum (U.S. MEC 1 or 2) (Box 1), if it is reasonably certain that the woman is not pregnant (Box 2). The Cu-IUD should not be inserted in a woman with postpartum sepsis (e.g., chorioamnionitis or endometritis) (U.S. MEC 4).
- **Need for back-up contraception:** No additional contraceptive protection is needed.

Postabortion (Spontaneous or Induced)

- **Timing:** The Cu-IUD can be inserted within the first 7 days, including immediately postabortion (U.S. MEC 1 for first-trimester abortion and U.S. MEC 2 for second-trimester abortion). The Cu-IUD should not be inserted immediately after a septic abortion (U.S. MEC 4).
- **Need for back-up contraception:** No additional contraceptive protection is needed.

Switching from Another Contraceptive Method

- **Timing:** The Cu-IUD can be inserted immediately if it is reasonably certain that the woman is not pregnant (Box 2). Waiting for her next menstrual cycle is unnecessary.
- **Need for back-up contraception:** No additional contraceptive protection is needed.

Comments and Evidence Summary. In situations in which the health care provider is not reasonably certain that the woman is not pregnant, the woman should be provided with another contraceptive method to use until the health care provider can be reasonably certain that she is not pregnant and can insert the Cu-IUD.

A systematic review identified eight studies that suggested that timing of Cu-IUD insertion in relation to the menstrual cycle in non-postpartum women had little effect on long-term outcomes (rates of continuation, removal, expulsion, or pregnancy) or on short-term outcomes (pain at insertion, bleeding at insertion, or immediate expulsion) (43) (Level of evidence: II-2, fair, direct).

Initiation of LNG-IUDs

Timing of LNG-IUD Insertion

- The LNG-IUD can be inserted at any time if it is reasonably certain that the woman is not pregnant (Box 2).

Need for Back-Up Contraception

- If the LNG-IUD is inserted within the first 7 days since menstrual bleeding started, no additional contraceptive protection is needed.
- If the LNG-IUD is inserted >7 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Special Considerations

Amenorrhea (Not Postpartum)

- **Timing:** The LNG-IUD can be inserted at any time if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postpartum (Including After Cesarean Delivery)

- **Timing:** The LNG-IUD can be inserted at any time, including immediately postpartum (U.S. MEC 1 or 2) if it is reasonably certain that the woman is not pregnant (Box 2). The LNG-IUD should not be inserted in a woman with postpartum sepsis (e.g., chorioamnionitis or endometritis) (U.S. MEC 4).
- **Need for back-up contraception:** If the woman is <6 months postpartum, amenorrheic, and fully or nearly fully breastfeeding (exclusively breastfeeding or the vast majority [$\geq 85\%$] of feeds are breastfeeds) (27), no

additional contraceptive protection is needed. Otherwise, a woman who is ≥ 21 days postpartum and has not experienced return of her menstrual cycle needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days. If her menstrual cycles have returned and it has been >7 days since menstrual bleeding began, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postabortion (Spontaneous or Induced)

- **Timing:** The LNG-IUD can be inserted within the first 7 days, including immediately postabortion (U.S. MEC 1 for first-trimester abortion and U.S. MEC 2 for second-trimester abortion). The LNG-IUD should not be inserted immediately after a septic abortion (U.S. MEC 4).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days unless the IUD is placed at the time of a surgical abortion.

Switching from Another Contraceptive Method

- **Timing:** The LNG-IUD can be inserted immediately if it is reasonably certain that the woman is not pregnant (Box 2). Waiting for her next menstrual cycle is unnecessary.
- **Need for back-up contraception:** If it has been >7 days since menstrual bleeding began, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.
- **Switching from a Cu-IUD:** If the woman has had sexual intercourse since the start of her current menstrual cycle and it has been >5 days since menstrual bleeding started, theoretically, residual sperm might be in the genital tract, which could lead to fertilization if ovulation occurs. A health care provider may consider providing any type of ECPs at the time of LNG-IUD insertion.

Comments and Evidence Summary. In situations in which the health care provider is uncertain whether the woman might be pregnant, the woman should be provided with another contraceptive method to use until the health care provider can be reasonably certain that she is not pregnant and can insert the LNG-IUD. If a woman needs to use additional contraceptive protection when switching to an LNG-IUD from another contraceptive method, consider continuing her previous method for 7 days after LNG-IUD insertion. No direct evidence was found regarding the effects of inserting LNG-IUDs on different days of the cycle on short- or long-term outcomes (43).

Examinations and Tests Needed Before Initiation of a Cu-IUD or an LNG-IUD

Among healthy women, few examinations or tests are needed before initiation of an IUD (Table 1). Bimanual examination and cervical inspection are necessary before IUD insertion. A baseline weight and BMI measurement might be useful for monitoring IUD users over time. If a woman has not been screened for STDs according to STD screening guidelines, screening can be performed at the time of insertion. Women with known medical problems or other special conditions might need additional examinations or tests before being determined to be appropriate candidates for a particular method of contraception. U.S. MEC might be useful in such circumstances (5).

Comments and Evidence Summary. Weight (BMI): Obese women can use IUDs (U.S. MEC 1) (5); therefore, screening for obesity is not necessary for the safe initiation of IUDs. However, measuring weight and calculating

TABLE 1. Classification of examinations and tests needed before IUD insertion

Examination or test	Class*	
	Copper-containing IUD	Levonorgestrel-releasing IUD
Examination		
Blood pressure	C	C
Weight (BMI) (weight [kg] / height [m] ²)	—†	—†
Clinical breast examination	C	C
Bimanual examination and cervical inspection	A	A
Laboratory test		
Glucose	C	C
Lipids	C	C
Liver enzymes	C	C
Hemoglobin	C	C
Thrombogenic mutations	C	C
Cervical cytology (Papanicolaou smear)	C	C
STD screening with laboratory tests	—§	—§
HIV screening with laboratory tests	C	C

Abbreviations: BMI = body mass index; HIV = human immunodeficiency virus; IUD = intrauterine device; STD = sexually transmitted disease; U.S. MEC = U.S. Medical Eligibility Criteria for Contraceptive Use.

* **Class A:** essential and mandatory in all circumstances for safe and effective use of the contraceptive method. **Class B:** contributes substantially to safe and effective use, but implementation may be considered within the public health and/or service context; the risk of not performing an examination or test should be balanced against the benefits of making the contraceptive method available. **Class C:** does not contribute substantially to safe and effective use of the contraceptive method.

† Weight (BMI) measurement is not needed to determine medical eligibility for any methods of contraception because all methods can be used (U.S. MEC 1) or generally can be used (U.S. MEC 2) among obese women (Box 1). However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

§ Most women do not require additional STD screening at the time of IUD insertion. If a woman with risk factors for STDs has not been screened for gonorrhea and chlamydia according to CDC's *STD Treatment Guidelines* (<http://www.cdc.gov/std/treatment>), screening can be performed at the time of IUD insertion, and insertion should not be delayed. Women with current purulent cervicitis or chlamydial infection or gonococcal infection should not undergo IUD insertion (U.S. MEC 4).

BMI (weight [kg] / height [m²]) at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

Bimanual examination and cervical inspection: Bimanual examination and cervical inspection are necessary before IUD insertion to assess uterine size and position and to detect any cervical or uterine abnormalities that might indicate infection or otherwise prevent IUD insertion (44,45).

STDs: Women should be routinely screened for chlamydial infection and gonorrhea according to national screening guidelines. The CDC *Sexually Transmitted Diseases Treatment Guidelines* provide information on screening eligibility, timing, and frequency of screening and on screening for persons with risk factors (15) (<http://www.cdc.gov/std/treatment>). If STD screening guidelines have been followed, most women do not need additional STD screening at the time of IUD insertion, and insertion should not be delayed. If a woman with risk factors for STDs has not been screened for gonorrhea and chlamydia according to CDC STD treatment guidelines, screening can be performed at the time of IUD insertion, and insertion should not be delayed. Women with current purulent cervicitis or chlamydial infection or gonococcal infection should not undergo IUD insertion (U.S. MEC 4). A systematic review identified two studies that demonstrated no differences in PID rates among women who screened positive for gonorrhea or chlamydia and underwent concurrent IUD insertion compared with women who screened positive and initiated other contraceptive methods (46). Indirect evidence demonstrates women who undergo same-day STD screening and IUD insertion have similar PID rates compared with women who have delayed IUD insertion. Women who undergo same-day STD screening and IUD insertion have low incidence rates of PID. Algorithms for predicting PID among women with risk factors for STDs have poor predictive value. Risk for PID among women with risk factors for STDs is low (15,47–57). Although women with STDs at the time of IUD insertion have a higher risk for PID, the overall rate of PID among all IUD users is low (51,54).

Hemoglobin: Women with iron-deficiency anemia can use the LNG-IUD (U.S. MEC 1) (5); therefore, screening for anemia is not necessary for safe initiation of the LNG-IUD. Women with iron-deficiency anemia generally can use Cu-IUDs (U.S. MEC 2) (5). Measurement of hemoglobin before initiation of Cu-IUDs is not necessary because of the minimal change in hemoglobin among women with and without anemia using Cu-IUDs. A systematic review identified four studies that provided direct evidence for changes in hemoglobin among women with anemia who received Cu-IUDs (58). Evidence from one randomized trial (59)

and one prospective cohort study (60) showed no significant changes in hemoglobin among Cu-IUD users with anemia, whereas two prospective cohort studies (61,62) showed a statistically significant decrease in hemoglobin levels during 12 months of follow-up; however, the magnitude of the decrease was small and most likely not clinically significant. The systematic review also identified 21 studies that provided indirect evidence by examining changes in hemoglobin among healthy women receiving Cu-IUDs (63–83), which generally showed no clinically significant changes in hemoglobin levels with up to 5 years of follow up (Level of evidence: I to II-2, fair, direct).

Lipids: Screening for dyslipidemias is not necessary for the safe initiation of Cu-IUD or LNG-IUD because of the low prevalence of undiagnosed disease in women of reproductive age and the low likelihood of clinically significant changes with use of hormonal contraceptives. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with lipid measurement before initiation of hormonal contraceptives (57). During 2009–2012 among women aged 20–44 years in the United States, 7.6% had high cholesterol, defined as total serum cholesterol ≥ 240 mg/dL (84). During 1999–2008, the prevalence of undiagnosed hypercholesterolemia among women aged 20–44 years was approximately 2% (85). Studies have shown mixed results about the effects of hormonal methods on lipid levels among both healthy women and women with baseline lipid abnormalities, and the clinical significance of these changes is unclear (86–89).

Liver enzymes: Women with liver disease can use the Cu-IUD (U.S. MEC 1) (5); therefore, screening for liver disease is not necessary for the safe initiation of the Cu-IUD. Although women with certain liver diseases generally should not use the LNG-IUD (U.S. MEC 3) (5), screening for liver disease before initiation of the LNG-IUD is not necessary because of the low prevalence of these conditions and the high likelihood that women with liver disease already would have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with liver enzyme tests before initiation of hormonal contraceptive use (57). In 2012, among U.S. women, the percentage with liver disease (not further specified) was 1.3% (90). In 2013, the incidence of acute hepatitis A, B, or C was ≤ 1 per 100,000 U.S. population (91). During 2002–2011, the incidence of liver carcinoma among U.S. women was approximately 3.7 per 100,000 population (92). Because estrogen and progestins are metabolized in the liver, the use of hormonal contraceptives among women with liver disease might, theoretically, be a concern. The use of hormonal contraceptives, specifically COCs and POPs,

does not affect disease progression or severity in women with hepatitis, cirrhosis, or benign focal nodular hyperplasia (93,94), although evidence is limited, and no evidence exists for the LNG-IUD.

Clinical breast examination: Women with breast disease can use the Cu-IUD (U.S. MEC 1) (5); therefore, screening for breast disease is not necessary for the safe initiation of the Cu-IUD. Although women with current breast cancer should not use the LNG-IUD (U.S. MEC 4) (5), screening asymptomatic women with a clinical breast examination before inserting an IUD is not necessary because of the low prevalence of breast cancer among women of reproductive age. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with a breast examination before initiation of hormonal contraceptives (95). The incidence of breast cancer among women of reproductive age in the United States is low. In 2012, the incidence of breast cancer among women aged 20–49 years was approximately 70.7 per 100,000 women (96).

Cervical cytology: Although women with cervical cancer should not undergo IUD insertion (U.S. MEC 4) (5), screening asymptomatic women with cervical cytology before IUD insertion is not necessary because of the high rates of cervical screening, low incidence of cervical cancer in the United States, and high likelihood that a woman with cervical cancer already would have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with cervical cytology before initiation of IUDs (57). Cervical cancer is rare in the United States, with an incidence rate of 9.8 per 100,000 women during 2012 (96). The incidence and mortality rates from cervical cancer have declined dramatically in the United States, largely because of cervical cytology screening (97). Overall screening rates for cervical cancer in the United States are high; in 2013 among women aged 18–44 years, approximately 77% reported having cervical cytology screening within the last 3 years (98).

HIV screening: Women with HIV infection can use (U.S. MEC 1) or generally can use (U.S. MEC 2) IUDs (5). Therefore, HIV screening is not necessary before IUD insertion. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened for HIV infection before IUD insertion (57). Limited evidence suggests that IUDs are not associated with disease progression, increased infection, or other adverse health effects among women with HIV infection (99–114).

Other screening: Women with hypertension, diabetes, or thrombogenic mutations can use (U.S. MEC 1) or generally can use (U.S. MEC 2) IUDs (5). Therefore, screening for these conditions is not necessary for the safe initiation of IUDs.

Provision of Medications to Ease IUD Insertion

- Misoprostol is not recommended for routine use before IUD insertion. Misoprostol might be helpful in select circumstances (e.g., in women with a recent failed insertion).
- Paracervical block with lidocaine might reduce patient pain during IUD insertion.

Comments and Evidence Summary. Potential barriers to IUD use include anticipated pain with insertion and provider concerns about difficult insertion. Identifying effective approaches to ease IUD insertion might increase IUD initiation.

Evidence for misoprostol from two systematic reviews, including a total of 10 randomized controlled trials, suggests that misoprostol does not improve provider ease of insertion, reduce the need for adjunctive insertion measures, or improve insertion success (Level of evidence: I, good to fair, direct) and might increase patient pain and side effects (Level of evidence: I, high quality) (115,116). However, one randomized controlled trial examined women with a recent failed IUD insertion and found significantly higher insertion success with second insertion attempt among women pretreated with misoprostol versus placebo (Level of evidence: I, good, direct) (117).

Limited evidence for paracervical block with lidocaine from one systematic review suggests that it might reduce patient pain (115). In this review, two randomized controlled trials found significantly reduced pain at either tenaculum placement or IUD insertion among women receiving paracervical block with 1% lidocaine 3–5 minutes before IUD insertion (118,119). Neither trial found differences in side effects among women receiving paracervical block compared with controls (Level of evidence: I, moderate to low quality) (118,119).

Limited evidence on nonsteroidal antiinflammatory drugs (NSAIDs) and nitric oxide donors generally suggested no positive effect; evidence on lidocaine with administration other than paracervical block was limited and inconclusive (Level of evidence for provider ease of insertion: I, good to poor, direct; Level of evidence for need for adjunctive insertion measures: I, fair, direct; Level of evidence for patient pain: I, high to low quality; Level of evidence for side effects: I, high to low quality) (115,116).

Provision of Prophylactic Antibiotics at the Time of IUD Insertion

- Prophylactic antibiotics are generally not recommended for Cu-IUD or LNG-IUD insertion.

Comments and Evidence Summary. Theoretically, IUD insertion could induce bacterial spread and lead to

complications such as PID or infective endocarditis. A metaanalysis was conducted of randomized controlled trials examining antibiotic prophylaxis versus placebo or no treatment for IUD insertion (120). Use of prophylaxis reduced the frequency of unscheduled return visits but did not significantly reduce the incidence of PID or premature IUD discontinuation. Although the risk for PID was higher within the first 20 days after insertion, the incidence of PID was low among all women who had IUDs inserted (51). In addition, the American Heart Association recommends that the use of prophylactic antibiotics solely to prevent infective endocarditis is not needed for genitourinary procedures (121). Studies have not demonstrated a conclusive link between genitourinary procedures and infective endocarditis or a preventive benefit of prophylactic antibiotics during such procedures (121).

Routine Follow-Up After IUD Insertion

These recommendations address when routine follow-up is needed for safe and effective continued use of contraception for healthy women. The recommendations refer to general situations and might vary for different users and different situations. Specific populations who might benefit from more frequent follow-up visits include adolescents, persons with certain medical conditions or characteristics, and persons with multiple medical conditions.

- Advise the woman to return at any time to discuss side effects or other problems, if she wants to change the method being used, and when it is time to remove or replace the contraceptive method. No routine follow-up visit is required.
- At other routine visits, health care providers who see IUD users should do the following:
 - Assess the woman's satisfaction with her contraceptive method and whether she has any concerns about method use.
 - Assess any changes in health status, including medications, that would change the appropriateness of the IUD for safe and effective continued use on the basis of U.S. MEC (e.g., category 3 and 4 conditions and characteristics).
 - Consider performing an examination to check for the presence of the IUD strings.
 - Consider assessing weight changes and counseling women who are concerned about weight changes perceived to be associated with their contraceptive method.

Comments and Evidence Summary. Evidence from a systematic review about the effect of a specific follow-up visit schedule on IUD continuation is very limited and of poor quality. The evidence did not suggest that greater frequency of

visits or earlier timing of the first follow-up visit after insertion improves continuation of use (122) (Level of evidence: II-2, poor, direct). Evidence from four studies from a systematic review on the incidence of PID among IUD initiators, or IUD removal as a result of PID, suggested that the incidence of PID did not differ between women using Cu-IUDs and those using DMPA, COCs, or LNG-IUDs (123) (Level of evidence: I to II-2, good, indirect). Evidence on the timing of PID after IUD insertion is mixed. Although the rate of PID generally was low, the largest study suggested that the rate of PID was significantly higher in the first 20 days after insertion (51) (Level of evidence: I to II-3, good to poor, indirect).

Bleeding Irregularities with Cu-IUD Use

- Before Cu-IUD insertion, provide counseling about potential changes in bleeding patterns during Cu-IUD use. Unscheduled spotting or light bleeding, as well as heavy or prolonged bleeding, is common during the first 3–6 months of Cu-IUD use, is generally not harmful, and decreases with continued Cu-IUD use.
- If clinically indicated, consider an underlying gynecological problem, such as Cu-IUD displacement, an STD, pregnancy, or new pathologic uterine conditions (e.g., polyps or fibroids), especially in women who have already been using the Cu-IUD for a few months or longer and who have developed a new onset of heavy or prolonged bleeding. If an underlying gynecological problem is found, treat the condition or refer for care.
- If an underlying gynecological problem is not found and the woman requests treatment, the following treatment option can be considered during days of bleeding:
 - NSAIDs for short-term treatment (5–7 days)
- If bleeding persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Comments and Evidence Summary. During contraceptive counseling and before insertion of the Cu-IUD, information about common side effects such as unscheduled spotting or light bleeding or heavy or prolonged menstrual bleeding, especially during the first 3–6 months of use, should be discussed (64). These bleeding irregularities are generally not harmful. Enhanced counseling about expected bleeding patterns and reassurance that bleeding irregularities are generally not harmful has been shown to reduce method discontinuation in clinical trials with other contraceptives (i.e., DMPA) (124,125).

Evidence is limited on specific drugs, doses, and durations of use for effective treatments for bleeding irregularities with Cu-IUD use. Therefore, although this report includes general

recommendations for treatments to consider, evidence for specific regimens is lacking.

A systematic review identified 11 studies that examined various therapeutic treatments for heavy menstrual bleeding, prolonged menstrual bleeding, or both among women using Cu-IUDs (126). Nine studies examined the use of various oral NSAIDs for the treatment of heavy or prolonged menstrual bleeding among Cu-IUD users and compared them with either a placebo or a baseline cycle. Three of these trials examined the use of indomethacin (127–129), three examined mefenamic acid (130–132), and three examined flufenamic acid (127,128,133). Other NSAIDs used in the reported trials included alclufenac (127,128), suprofen (134), and diclofenac sodium (135). All but one NSAID study (131) demonstrated statistically significant or notable reductions in mean total menstrual blood loss with NSAID use. One study among 19 Cu-IUD users with heavy bleeding suggested that treatment with oral tranexamic acid can significantly reduce mean blood loss during treatment compared with placebo (135). Data regarding the overall safety of tranexamic acid are limited; an FDA warning states that tranexamic acid is contraindicated in women with active thromboembolic disease or with a history or intrinsic risk for thrombosis or thromboembolism (136,137). Treatment with aspirin demonstrated no statistically significant change in mean blood loss among women whose pretreatment menstrual blood loss was >80 ml or 60–80 mL; treatment resulted in a significant increase among women whose pretreatment menstrual blood loss was <60 mL (138). One study examined the use of a synthetic form of vasopressin, intranasal desmopressin (300 µg/day), for the first 5 days of menses for three treatment cycles and found a significant reduction in mean blood loss compared with baseline (130) (Level of evidence: I to II-3, poor to fair, direct). Only one small study examined treatment of spotting with three separate NSAIDs and did not observe improvements in spotting in any of the groups (127) (Level of evidence: I, poor, direct).

Bleeding Irregularities (Including Amenorrhea) with LNG-IUD Use

- Before LNG-IUD insertion, provide counseling about potential changes in bleeding patterns during LNG-IUD use. Unscheduled spotting or light bleeding is expected during the first 3–6 months of LNG-IUD use, is generally not harmful, and decreases with continued LNG-IUD use. Over time, bleeding generally decreases with LNG-IUD use, and many women experience only light menstrual bleeding or amenorrhea. Heavy or prolonged bleeding, either unscheduled or menstrual, is uncommon during LNG-IUD use.

Irregular Bleeding (Spotting, Light Bleeding, or Heavy or Prolonged Bleeding)

- If clinically indicated, consider an underlying gynecological problem, such as LNG-IUD displacement, an STD, pregnancy, or new pathologic uterine conditions (e.g., polyps or fibroids). If an underlying gynecological problem is found, treat the condition or refer for care.
- If bleeding persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Amenorrhea

- Amenorrhea does not require any medical treatment. Provide reassurance.
 - If a woman's regular bleeding pattern changes abruptly to amenorrhea, consider ruling out pregnancy if clinically indicated.
- If amenorrhea persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Comments and Evidence Summary. During contraceptive counseling and before insertion of the LNG-IUD, information about common side effects such as unscheduled spotting or light bleeding, especially during the first 3–6 months of use, should be discussed. Approximately half of LNG-IUD users are likely to experience amenorrhea or oligomenorrhea by 2 years of use (139). These bleeding irregularities are generally not harmful. Enhanced counseling about expected bleeding patterns and reassurance that bleeding irregularities are generally not harmful has been shown to reduce method discontinuation in clinical trials with other hormonal contraceptives (i.e., DMPA) (124,125). No direct evidence was found regarding therapeutic treatments for bleeding irregularities during LNG-IUD use.

Management of the IUD when a Cu-IUD or an LNG-IUD User Is Found To Have PID

- Treat the PID according to the CDC *Sexually Transmitted Diseases Treatment Guidelines* (15).
- Provide comprehensive management for STDs, including counseling about condom use.
- The IUD does not need to be removed immediately if the woman needs ongoing contraception.
- Reassess the woman in 48–72 hours. If no clinical improvement occurs, continue antibiotics and consider removal of the IUD.
- If the woman wants to discontinue use, remove the IUD sometime after antibiotics have been started to avoid the

potential risk for bacterial spread resulting from the removal procedure.

- If the IUD is removed, consider ECPs if appropriate. Counsel the woman on alternative contraceptive methods, and offer another method if it is desired.
- A summary of IUD management in women with PID is provided (Appendix F).

Comments and Evidence Summary. Treatment outcomes do not generally differ between women with PID who retain the IUD and those who have the IUD removed; however, appropriate antibiotic treatment and close clinical follow-up are necessary.

A systematic review identified four studies that included women using copper or nonhormonal IUDs who developed PID and compared outcomes between women who had the IUD removed or did not (140). One randomized trial showed that women with IUDs removed had longer hospitalizations than those who did not, although no differences in PID recurrences or subsequent pregnancies were observed (141). Another randomized trial showed no differences in laboratory findings among women who removed the IUD compared with those who did not (142). One prospective cohort study showed no differences in clinical or laboratory findings during hospitalization; however, the IUD removal group had longer hospitalizations (143). One randomized trial showed that the rate of recovery for most clinical signs and symptoms was higher among women who had the IUD removed than among women who did not (144). No evidence was found regarding women using LNG-IUDs (Level of evidence: I to II-2, fair, direct.)

Management of the IUD when a Cu-IUD or an LNG-IUD User is Found To Be Pregnant

- Evaluate for possible ectopic pregnancy.
- Advise the woman that she has an increased risk for spontaneous abortion (including septic abortion that might be life threatening) and for preterm delivery if the IUD is left in place. The removal of the IUD reduces these risks but might not decrease the risk to the baseline level of a pregnancy without an IUD.
 - If she does not want to continue the pregnancy, counsel her about options.
 - If she wants to continue the pregnancy, advise her to seek care promptly if she has heavy bleeding, cramping, pain, abnormal vaginal discharge, or fever.

IUD Strings Are Visible or Can Be Retrieved Safely from the Cervical Canal

- Advise the woman that the IUD should be removed as soon as possible.

- If the IUD is to be removed, remove it by pulling on the strings gently.
- Advise the woman that she should return promptly if she has heavy bleeding, cramping, pain, abnormal vaginal discharge, or fever.
- If she chooses to keep the IUD, advise her to seek care promptly if she has heavy bleeding, cramping, pain, abnormal vaginal discharge, or fever.

IUD Strings Are Not Visible and Cannot Be Safely Retrieved

- If ultrasonography is available, consider performing or referring for ultrasound examination to determine the location of the IUD. If the IUD cannot be located, it might have been expelled or have perforated the uterine wall.
- If ultrasonography is not possible or the IUD is determined by ultrasound to be inside the uterus, advise the woman to seek care promptly if she has heavy bleeding, cramping, pain, abnormal vaginal discharge, or fever.

Comments and Evidence Summary. Removing the IUD improves the pregnancy outcome if the IUD strings are visible or the device can be retrieved safely from the cervical canal. Risks for spontaneous abortion, preterm delivery, and infection are substantial if the IUD is left in place.

Theoretically, the fetus might be affected by hormonal exposure from an LNG-IUD. However, whether this exposure increases the risk for fetal abnormalities is unknown.

A systematic review identified nine studies suggesting that women who did not remove their IUDs during pregnancy were at greater risk for adverse pregnancy outcomes (including spontaneous abortion, septic abortion, preterm delivery, and chorioamnionitis) compared with women who had their IUDs removed or who did not have an IUD (41). Cu-IUD removal decreased risks but not to the baseline risk for pregnancies without an IUD. One case series examined LNG-IUDs. When they were not removed, 8 out of 10 pregnancies ended in spontaneous abortions (Level of evidence: II-2, fair, direct).

Implants

The etonogestrel implant, a single rod with 68 mg of etonogestrel, is available in the United States. Fewer than 1 woman out of 100 become pregnant in the first year of use of the etonogestrel implant with typical use (14). The implant is long acting, is reversible, and can be used by women of all ages, including adolescents. The implant does not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

Initiation of Implants

Timing

- The implant can be inserted at any time if it is reasonably certain that the woman is not pregnant (Box 2).

Need for Back-Up Contraception

- If the implant is inserted within the first 5 days since menstrual bleeding started, no additional contraceptive protection is needed.
- If the implant is inserted >5 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Special Considerations

Amenorrhea (Not Postpartum)

- **Timing:** The implant can be inserted at any time if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postpartum (Breastfeeding)

- **Timing:** The implant can be inserted at any time (U.S. MEC 2 if <1 month postpartum and U.S. MEC 1 if ≥1 month postpartum) if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** If the woman is <6 months postpartum, amenorrheic, and fully or nearly fully breastfeeding (exclusively breastfeeding or the vast majority [≥85%] of feeds are breastfeeds) (27), no additional contraceptive protection is needed. Otherwise, a woman who is ≥21 days postpartum and has not experienced return of her menstrual cycle needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days. If her menstrual cycles have returned and it has been >5 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postpartum (Not Breastfeeding)

- **Timing:** The implant can be inserted at any time, including immediately postpartum (U.S. MEC 1) if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** If a woman is <21 days postpartum, no additional contraceptive protection is needed. A woman who is ≥21 days postpartum and has not experienced return of her menstrual cycle needs to abstain from sexual intercourse or use additional

contraceptive protection for the next 7 days. If her menstrual cycles have returned and it has been >5 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postabortion (Spontaneous or Induced)

- **Timing:** The implant can be inserted within the first 7 days, including immediately after the abortion (U.S. MEC 1).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days unless the implant is placed at the time of a surgical abortion.

Switching from Another Contraceptive Method

- **Timing:** The implant can be inserted immediately if it is reasonably certain that the woman is not pregnant (Box 2). Waiting for her next menstrual cycle is unnecessary.
- **Need for back-up contraception:** If it has been >5 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days after insertion.
- **Switching from an IUD:** If the woman has had sexual intercourse since the start of her current menstrual cycle and it has been >5 days since menstrual bleeding started, theoretically, residual sperm might be in the genital tract, which could lead to fertilization if ovulation occurs. A health care provider may consider any of the following options:
 - Advise the woman to retain the IUD for at least 7 days after the implant is inserted and return for IUD removal.
 - Advise the woman to abstain from sexual intercourse or use barrier contraception for 7 days before removing the IUD and switching to the new method.
 - If the woman cannot return for IUD removal and has not abstained from sexual intercourse or used barrier contraception for 7 days, advise the woman to use ECPs (with the exception of UPA) at the time of IUD removal.

Comments and Evidence Summary. In situations in which the health care provider is uncertain whether the woman might be pregnant, the benefits of starting the implant likely exceed any risk; therefore, starting the implant should be considered at any time, with a follow-up pregnancy test in 2–4 weeks.

If a woman needs to use additional contraceptive protection when switching to an implant from another contraceptive method, consider continuing her previous method for 7 days after implant insertion. No direct evidence was found regarding the effects of starting the etonogestrel implant at different times of the cycle.

Examinations and Tests Needed Before Implant Insertion

Among healthy women, no examinations or tests are needed before initiation of an implant, although a baseline weight and BMI measurement might be useful for monitoring implant users over time (Table 2). Women with known medical problems or other special conditions might need additional examinations or tests before being determined to be appropriate candidates for a particular method of contraception. U.S. MEC might be useful in such circumstances (5).

Comments and Evidence Summary. Weight (BMI): Obese women can use implants (U.S. MEC 1) (5); therefore, screening for obesity is not necessary for the safe initiation of implants. However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

Bimanual examination and cervical inspection: A pelvic examination is not necessary before initiation of implants because it would not facilitate detection of conditions for which implant use would be unsafe. Women with current breast cancer should not use implants (U.S. MEC 4); women with certain liver diseases generally should not (U.S. MEC 3) use implants (5). However, none of these conditions are likely to be detected

by pelvic examination (145). A systematic review identified two case-control studies that compared delayed and immediate pelvic examination before initiation of hormonal contraceptives, specifically oral contraceptives or DMPA (95). No differences in risk factors for cervical neoplasia, incidence of STDs, incidence of abnormal Papanicolaou smears, or incidence of abnormal wet mounts were observed. No evidence was found regarding implants (Level of evidence: II-2 fair, direct).

Lipids: Screening for dyslipidemias is not necessary for the safe initiation of implants because of the low prevalence of undiagnosed disease in women of reproductive age and the low likelihood of clinically significant changes with use of hormonal contraceptives. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with lipid measurement before initiation of hormonal contraceptives (57). During 2009–2012 among women aged 20–44 years in the United States, 7.6% had high cholesterol, defined as total serum cholesterol ≥ 240 mg/dL (84). During 1999–2008, the prevalence of undiagnosed hypercholesterolemia among women aged 20–44 years was approximately 2% (85). Studies have shown mixed results regarding the effects of hormonal methods on lipid levels among both healthy women and women with baseline lipid abnormalities, and the clinical significance of these changes is unclear (86–89).

Liver enzymes: Although women with certain liver diseases generally should not use implants (U.S. MEC 3) (5), screening for liver disease before initiation of implants is not necessary because of the low prevalence of these conditions and the high likelihood that women with liver disease already would have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with liver enzyme tests before initiation of hormonal contraceptives (57). In 2012, the percentage of U.S. women with liver disease (not further specified) was 1.3% (90). In 2013, the incidence of acute hepatitis A, B, or C was ≤ 1 per 100,000 U.S. population (91). During 2002–2011, the incidence of liver carcinoma among U.S. women was approximately 3.7 per 100,000 population (92). Because estrogen and progestins are metabolized in the liver, the use of hormonal contraceptives among women with liver disease might, theoretically, be a concern. The use of hormonal contraceptives, specifically COCs and POPs, does not affect disease progression or severity in women with hepatitis, cirrhosis, or benign focal nodular hyperplasia (93,94), although evidence is limited and no evidence exists for implants.

Clinical breast examination: Although women with current breast cancer should not use implants (U.S. MEC 4) (5), screening asymptomatic women with a clinical breast

TABLE 2. Classification of examinations and tests needed before implant insertion

Examination or test	Class*
Examination	
Blood pressure	C
Weight (BMI) (weight [kg] / height [m] ²)	—†
Clinical breast examination	C
Bimanual examination and cervical inspection	C
Laboratory test	
Glucose	C
Lipids	C
Liver enzymes	C
Hemoglobin	C
Thrombogenic mutations	C
Cervical cytology (Papanicolaou smear)	C
STD screening with laboratory tests	C
HIV screening with laboratory tests	C

Abbreviations: BMI = body mass index; HIV = human immunodeficiency virus; STD = sexually transmitted disease; U.S. MEC = U.S. Medical Eligibility Criteria for Contraceptive Use.

* **Class A:** essential and mandatory in all circumstances for safe and effective use of the contraceptive method. **Class B:** contributes substantially to safe and effective use, but implementation may be considered within the public health and/or service context; the risk of not performing an examination or test should be balanced against the benefits of making the contraceptive method available. **Class C:** does not contribute substantially to safe and effective use of the contraceptive method.

† Weight (BMI) measurement is not needed to determine medical eligibility for any methods of contraception because all methods can be used (U.S. MEC 1) or generally can be used (U.S. MEC 2) among obese women (Box 1). However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

examination before initiation of implants is not necessary because of the low prevalence of breast cancer among women of reproductive age (15–49 years). A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with a breast examination before initiation of hormonal contraceptives (95). The incidence of breast cancer among women of reproductive age in the United States is low. In 2012, the incidence of breast cancer among women aged 20–49 years was approximately 70.7 per 100,000 women (96).

Other screening: Women with hypertension, diabetes, anemia, thrombogenic mutations, cervical intraepithelial neoplasia, cervical cancer, STDs, or HIV infection can use (U.S. MEC 1) or generally can use (U.S. MEC 2) implants (5); therefore, screening for these conditions is not necessary for the safe initiation of implants.

Routine Follow-Up After Implant Insertion

These recommendations address when routine follow-up is needed for safe and effective continued use of contraception for healthy women. The recommendations refer to general situations and might vary for different users and different situations. Specific populations who might benefit from more frequent follow-up visits include adolescents, those with certain medical conditions or characteristics, and those with multiple medical conditions.

- Advise the woman to return at any time to discuss side effects or other problems, if she wants to change the method being used, and when it is time to remove or replace the contraceptive method. No routine follow-up visit is required.
- At other routine visits, health care providers seeing implant users should do the following:
 - Assess the woman's satisfaction with her contraceptive method and whether she has any concerns about method use.
 - Assess any changes in health status, including medications, that would change the appropriateness of the implant for safe and effective continued use based on U.S. MEC (e.g., category 3 and 4 conditions and characteristics).
 - Consider assessing weight changes and counseling women who are concerned about weight change perceived to be associated with their contraceptive method.

Comments and Evidence Summary. A systematic review did not identify any evidence regarding whether a routine follow-up visit after initiating an implant improves correct or continued use (122).

Bleeding Irregularities (Including Amenorrhea) During Implant Use

- Before implant insertion, provide counseling about potential changes in bleeding patterns during implant use. Unscheduled spotting or light bleeding is common with implant use, and some women experience amenorrhea. These bleeding changes are generally not harmful and might or might not decrease with continued implant use. Heavy or prolonged bleeding, unscheduled or menstrual, is uncommon during implant use.

Irregular Bleeding (Spotting, Light Bleeding, or Heavy or Prolonged Bleeding)

- If clinically indicated, consider an underlying gynecological problem, such as interactions with other medications, an STD, pregnancy, or new pathologic uterine conditions (e.g., polyps or fibroids). If an underlying gynecological problem is found, treat the condition or refer for care.
- If an underlying gynecologic problem is not found and the woman wants treatment, the following treatment options during days of bleeding can be considered:
 - NSAIDs for short-term treatment (5–7 days)
 - Hormonal treatment (if medically eligible) with low-dose COCs or estrogen for short-term treatment (10–20 days)
- If irregular bleeding persists and the woman finds it unacceptable, counsel her on alternative methods, and offer another method if it is desired.

Amenorrhea

- Amenorrhea does not require any medical treatment. Provide reassurance.
 - If a woman's regular bleeding pattern changes abruptly to amenorrhea, consider ruling out pregnancy if clinically indicated.
- If amenorrhea persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Comments and Evidence Summary. During contraceptive counseling and before insertion of the implant, information about common side effects, such as unscheduled spotting or light bleeding and amenorrhea, especially during the first year of use, should be discussed. A pooled analysis of data from 11 clinical trials indicates that a significant proportion of etonogestrel implant users had relatively little bleeding: 22% of women experienced amenorrhea and 34% experienced infrequent spotting, although 7% reported frequent bleeding

and 18% reported prolonged bleeding (146). Unscheduled bleeding or amenorrhea is generally not harmful. Enhanced counseling about expected bleeding patterns and reassurance that bleeding irregularities are generally not harmful has been shown to reduce discontinuation in clinical trials with other hormonal contraceptives (i.e., DMPA) (124,125).

A systematic review and four newly published studies examined several medications for the treatment of bleeding irregularities with primarily levonorgestrel contraceptive implants (147–151). Two small studies found significant cessation of bleeding within 7 days of start of treatment among women taking oral celecoxib (200 mg) daily for 5 days or oral mefenamic acid (500 mg) 3 times daily for 5 days compared with placebo (149,150). Differences in bleeding cessation were not found among women with etonogestrel implants taking mifepristone but were found when women with the implants combined mifepristone with either ethinyl estradiol or doxycycline (151,152). Doxycycline alone or in combination with ethinyl estradiol did not improve bleeding cessation among etonogestrel implant users (151). Among LNG implant users, mifepristone reduced the number of bleeding or spotting days but only after 6 months of treatment (153). Evidence also suggests that estrogen (154–156), daily COCs (154), LNG pills (155), tamoxifen (157), or tranexamic acid (158) can reduce the number of bleeding or spotting days during treatment among LNG implant users. In one small study, vitamin E was found to significantly reduce the mean number of bleeding days after the first treatment cycle; however, another larger study reported no significant differences in length of bleeding and spotting episodes with vitamin E treatment (159,160). Use of aspirin did not result in a significant difference in median length of bleeding or bleeding and spotting episodes after treatment (159). One study among implant users reported a reduction in number of bleeding days after initiating ibuprofen; however, another trial did not demonstrate any significant differences in the number of spotting and bleeding episodes with ibuprofen compared with placebo (148,155).

Injectables

Progestin-only injectable contraceptives (DMPA, 150 mg intramuscularly or 104 mg subcutaneously) are available in the United States; the only difference between these two formulations is the route of administration. Approximately 6 out of 100 women will become pregnant in the first year of use of DMPA with typical use (14). DMPA is reversible and can be used by women of all ages, including adolescents. DMPA does not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

Initiation of Injectables

Timing

- The first DMPA injection can be given at any time if it is reasonably certain that the woman is not pregnant (Box 2).

Need for Back-Up Contraception

- If DMPA is started within the first 7 days since menstrual bleeding started, no additional contraceptive protection is needed.
- If DMPA is started >7 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Special Considerations

Amenorrhea (Not Postpartum)

- **Timing:** The first DMPA injection can be given at any time if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postpartum (Breastfeeding)

- **Timing:** The first DMPA injection can be given at any time, including immediately postpartum (U.S. MEC 2 if <1 month postpartum; U.S. MEC 1 if ≥1 month postpartum) if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** If the woman is <6 months postpartum, amenorrheic, and fully or nearly fully breastfeeding (exclusively breastfeeding or the vast majority [≥85%] of feeds are breastfeeds) (27), no additional contraceptive protection is needed. Otherwise, a woman who is ≥21 days postpartum and has not experienced return of her menstrual cycle needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days. If her menstrual cycles have returned and it has been >7 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postpartum (Not Breastfeeding)

- **Timing:** The first DMPA injection can be given at any time, including immediately postpartum (U.S. MEC 1) if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** If a woman is <21 days postpartum, no additional contraceptive protection is needed. A woman who is ≥21 days postpartum and has

not experienced return of her menstrual cycle needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days. If her menstrual cycles have returned and it has been >7 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postabortion (Spontaneous or Induced)

- **Timing:** The first DMPA injection can be given within the first 7 days, including immediately after the abortion (U.S. MEC 1).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days unless the injection is given at the time of a surgical abortion.

Switching from Another Contraceptive Method

- **Timing:** The first DMPA injection can be given immediately if it is reasonably certain that the woman is not pregnant (Box 2). Waiting for her next menstrual cycle is unnecessary.
- **Need for back-up contraception:** If it has been >7 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.
- **Switching from an IUD:** If the woman has had sexual intercourse since the start of her current menstrual cycle and it has been >5 days since menstrual bleeding started, theoretically, residual sperm might be in the genital tract, which could lead to fertilization if ovulation occurs. A health care provider may consider any of the following options:
 - Advise the women to retain the IUD for at least 7 days after the injection and return for IUD removal.
 - Advise the woman to abstain from sexual intercourse or use barrier contraception for 7 days before removing the IUD and switching to the new method.
 - If the woman cannot return for IUD removal and has not abstained from sexual intercourse or used barrier contraception for 7 days, advise the woman to use ECPs (with the exception of UPA) at the time of IUD removal.

Comments and Evidence Summary. In situations in which the health care provider is uncertain whether the woman might be pregnant, the benefits of starting DMPA likely exceed any risk; therefore, starting DMPA should be considered at any time, with a follow-up pregnancy test in 2–4 weeks. If a woman needs to use additional contraceptive protection when switching to DMPA from another contraceptive method, consider continuing her previous method for 7 days after DMPA injection.

A systematic review identified eight articles examining DMPA initiation on different days of the menstrual cycle (161). Evidence from two studies with small sample sizes indicated that DMPA injections given up to day 7 of the menstrual cycle inhibited ovulation; when DMPA was administered after day 7, ovulation occurred in some women. Cervical mucus was of poor quality (i.e., not favorable for sperm penetration) in 90% of women within 24 hours of the injection (Level of evidence: II-2, fair) (162–164). Studies found that use of another contraceptive method until DMPA could be initiated (bridging option) did not help women initiate DMPA and was associated with more unintended pregnancies than immediate receipt of DMPA (165–169) (Level of evidence: I to II-3, fair to poor, indirect).

Examinations and Tests Needed Before Initiation of an Injectable

Among healthy women, no examinations or tests are needed before initiation of DMPA, although a baseline weight and BMI measurement might be useful to monitor DMPA users over time (Table 3). Women with known medical problems or other special conditions might need additional examinations or tests before being determined to be appropriate candidates for a particular method of contraception. U.S. MEC might be useful in such circumstances (5).

Comments and Evidence Summary. Weight (BMI): Obese women can use (U.S. MEC 1) or generally can use (U.S. MEC 2) DMPA (5); therefore, screening for obesity is not necessary for the safe initiation of DMPA. However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method. (See guidance on follow-up for DMPA users for evidence on weight gain with DMPA use).

Bimanual examination and cervical inspection: Pelvic examination is not necessary before initiation of DMPA because it does not facilitate detection of conditions for which DMPA would be unsafe. Although women with current breast cancer should not use DMPA (U.S. MEC 4), and women with severe hypertension, heart disease, vascular disease, or certain liver diseases generally should not use DMPA (U.S. MEC 3) (5), none of these conditions are likely to be detected by pelvic examination (145). A systematic review identified two case-control studies that compared delayed versus immediate pelvic examination before initiation of hormonal contraceptives, specifically oral contraceptives or DMPA (95). No differences in risk factors for cervical neoplasia, incidence of STDs, incidence of abnormal Papanicolaou smears, or incidence of

TABLE 3. Classification of examinations and tests needed before depo-medroxyprogesterone acetate initiation

Examination or test	Class*
Examination	
Blood pressure	C
Weight (BMI) (weight [kg]/height [m] ²)	—†
Clinical breast examination	C
Bimanual examination and cervical inspection	C
Laboratory test	
Glucose	C
Lipids	C
Liver enzymes	C
Hemoglobin	C
Thrombogenic mutations	C
Cervical cytology (Papanicolaou smear)	C
STD screening with laboratory tests	C
HIV screening with laboratory tests	C

Abbreviations: BMI = body mass index; HIV = human immunodeficiency virus; STD = sexually transmitted disease; U.S. MEC = U.S. Medical Eligibility Criteria for Contraceptive Use.

* **Class A:** essential and mandatory in all circumstances for safe and effective use of the contraceptive method. **Class B:** contributes substantially to safe and effective use, but implementation may be considered within the public health and/or service context; the risk of not performing an examination or test should be balanced against the benefits of making the contraceptive method available. **Class C:** does not contribute substantially to safe and effective use of the contraceptive method.

† Weight (BMI) measurement is not needed to determine medical eligibility for any methods of contraception because all methods can be used (U.S. MEC 1) or generally can be used (U.S. MEC 2) among obese women (Box 1). However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

abnormal wet mounts were observed (Level of evidence: II-2, fair, direct).

Blood pressure: Women with hypertension generally can use DMPA (U.S. MEC 2), with the exception of women with severe hypertension or vascular disease, who generally should not use DMPA (U.S. MEC 3) (5). Screening for hypertension before initiation of DMPA is not necessary because of the low prevalence of undiagnosed severe hypertension and the high likelihood that women with these conditions already would have had them diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with a blood pressure measurement before initiation of progestin-only contraceptives (170). The prevalence of undiagnosed hypertension among women of reproductive age is low. During 2009–2012 among women aged 20–44 years in the United States, the prevalence of hypertension was 8.7% (84). During 1999–2008, the percentage of women aged 20–44 years with undiagnosed hypertension was 1.9% (85).

Glucose: Although women with complicated diabetes generally should not use DMPA (U.S. MEC 3) (5), screening for diabetes before initiation of DMPA is not necessary because of the low prevalence of undiagnosed diabetes and the high likelihood that women with complicated diabetes would

already have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with glucose measurement before initiation of hormonal contraceptives (57). The prevalence of diabetes among women of reproductive age is low. During 2009–2012 among women aged 20–44 years in the United States, the prevalence of diabetes was 3.3% (84). During 1999–2008, the percentage of women aged 20–44 years with undiagnosed diabetes was 0.5% (85). Although hormonal contraceptives can have some adverse effects on glucose metabolism in healthy and diabetic women, the overall clinical effect is minimal (171–177).

Lipids: Screening for dyslipidemias is not necessary for the safe initiation of injectables because of the low prevalence of undiagnosed disease in women of reproductive age and the low likelihood of clinically significant changes with use of hormonal contraceptives. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with lipid measurement before initiation of hormonal contraceptives (57). During 2009–2012 among women aged 20–44 years in the United States, 7.6% had high cholesterol, defined as total serum cholesterol ≥240 mg/dL (84). During 1999–2008, the prevalence of undiagnosed hypercholesterolemia among women aged 20–44 years was approximately 2% (85). Studies have shown mixed results about the effects of hormonal methods on lipid levels among both healthy women and women with baseline lipid abnormalities, and the clinical significance of these changes is unclear (86–89).

Liver enzymes: Although women with certain liver diseases generally should not use DMPA (U.S. MEC 3) (5), screening for liver disease before initiation of DMPA is not necessary because of the low prevalence of these conditions and the high likelihood that women with liver disease already would have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with liver enzyme tests before initiation of hormonal contraceptives (57). In 2012, among U.S. women, the percentage with liver disease (not further specified) was 1.3% (90). In 2013, the incidence of acute hepatitis A, B, or C was ≤1 per 100,000 U.S. population (91). During 2002–2011, the incidence of liver carcinoma among U.S. women was approximately 3.7 per 100,000 population (92). Because estrogen and progestins are metabolized in the liver, the use of hormonal contraceptives among women with liver disease might, theoretically, be a concern. The use of hormonal contraceptives, specifically COCs and POPs, does not affect disease progression or severity in women with hepatitis, cirrhosis, or benign focal nodular hyperplasia (93,94), although evidence is limited and no evidence exists for DMPA.

Clinical breast examination: Although women with current breast cancer should not use DMPA (U.S. MEC 4) (5), screening asymptomatic women with a clinical breast examination before initiating DMPA is not necessary because of the low prevalence of breast cancer among women of reproductive age. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with a clinical breast examination before initiation of hormonal contraceptives (95). The incidence of breast cancer among women of reproductive age in the United States is low. In 2012, the incidence of breast cancer among women aged 20–49 years was approximately 70.7 per 100,000 women (96).

Other screening: Women with anemia, thrombogenic mutations, cervical intraepithelial neoplasia, cervical cancer, HIV infection, or other STDs can use (U.S. MEC 1) or generally can use (U.S. MEC 2) DMPA (5); therefore, screening for these conditions is not necessary for the safe initiation of DMPA.

Routine Follow-Up After Injectable Initiation

These recommendations address when routine follow-up is recommended for safe and effective continued use of contraception for healthy women. The recommendations refer to general situations and might vary for different users and different situations. Specific populations who might benefit from frequent follow-up visits include adolescents, those with certain medical conditions or characteristics, and those with multiple medical conditions.

- Advise the woman to return at any time to discuss side effects or other problems, if she wants to change the method being used, and when it is time for reinjection. No routine follow-up visit is required.
- At other routine visits, health care providers seeing injectable users should do the following:
 - Assess the woman's satisfaction with her contraceptive method and whether she has any concerns about method use.
 - Assess any changes in health status, including medications, that would change the appropriateness of the injectable for safe and effective continued use based on U.S. MEC (e.g., category 3 and 4 conditions and characteristics).
 - Consider assessing weight changes and counseling women who are concerned about weight change perceived to be associated with their contraceptive method.

Comments and Evidence Summary. Although no evidence exists regarding whether a routine follow-up visit after initiating DMPA improves correct or continued use, monitoring weight or BMI change over time is important for DMPA users.

A systematic review identified a limited body of evidence that examined whether weight gain in the few months after DMPA initiation predicted future weight gain (123). Two studies found significant differences in weight gain or BMI at follow-up periods ranging from 12 to 36 months between early weight gainers (i.e., those who gained >5% of their baseline body weight within 6 months after initiation) and those who were not early weight gainers (178,179). The differences between groups were more pronounced at 18, 24, and 36 months than at 12 months. One study found that most adolescent DMPA users who had gained >5% of their baseline weight by 3 months gained even more weight by 12 months (180) (Level of evidence: II-2, fair, to II-3, fair, direct).

Timing of Repeat Injections

Reinjection Interval

- Provide repeat DMPA injections every 3 months (13 weeks).

Special Considerations

Early Injection

- The repeat DMPA injection can be given early when necessary.

Late Injection

- The repeat DMPA injection can be given up to 2 weeks late (15 weeks from the last injection) without requiring additional contraceptive protection.
- If the woman is >2 weeks late (>15 weeks from the last injection) for a repeat DMPA injection, she can have the injection if it is reasonably certain that she is not pregnant (Box 2). She needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days. She might consider the use of emergency contraception (with the exception of UPA) if appropriate.

Comments and Evidence Summary. No time limits exist for early injections; injections can be given when necessary (e.g., when a woman cannot return at the routine interval). WHO has extended the time that a woman can have a late reinjection (i.e., grace period) for DMPA use from 2 weeks to 4 weeks on the basis of data from one study showing low pregnancy rates through 4 weeks; however, the CDC expert group did not consider the data to be generalizable to the United States because a large proportion of women in the study were breastfeeding. Therefore, U.S. SPR recommends a grace period of 2 weeks.

A systematic review identified 12 studies evaluating time to pregnancy or ovulation after the last injection of DMPA (181). Although pregnancy rates were low during the 2-week interval following the reinjection date and for 4 weeks following the

reinjection date, data were sparse, and one study included a large proportion of breastfeeding women (182–184). Studies also indicated a wide variation in time to ovulation after the last DMPA injection, with the majority ranging from 15 to 49 weeks from the last injection (185–193) (Level of evidence: level II-2, fair, direct).

Bleeding Irregularities (Including Amenorrhea) During Injectable Use

- Before DMPA initiation, provide counseling about potential changes in bleeding patterns during DMPA use. Amenorrhea and unscheduled spotting or light bleeding is common with DMPA use, and heavy or prolonged bleeding can occur with DMPA use. These bleeding irregularities are generally not harmful and might decrease with continued DMPA use.

Unscheduled Spotting or Light Bleeding

- If clinically indicated, consider an underlying gynecological problem, such as interactions with other medications, an STD, pregnancy, or new pathologic uterine conditions (e.g., polyps or fibroids). If an underlying gynecological problem is found, treat the condition or refer for care.
- If an underlying gynecologic problem is not found and the woman wants treatment, the following treatment option during days of bleeding can be considered:
 - NSAIDs for short-term treatment (5–7 days)
- If unscheduled spotting or light bleeding persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Heavy or Prolonged Bleeding

- If clinically indicated, consider an underlying gynecological problem, such as interactions with other medications, an STD, pregnancy, or new pathologic uterine conditions (such as fibroids or polyps). If an underlying gynecologic problem is identified, treat the condition or refer for care.
- If an underlying gynecologic problem is not found and the woman wants treatment, the following treatment options during days of bleeding can be considered:
 - NSAIDs for short-term treatment (5–7 days)
 - Hormonal treatment (if medically eligible) with low-dose COCs or estrogen for short-term treatment (10–20 days)
- If heavy or prolonged bleeding persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Amenorrhea

- Amenorrhea does not require any medical treatment. Provide reassurance.
 - If a woman's regular bleeding pattern changes abruptly to amenorrhea, consider ruling out pregnancy if clinically indicated.
- If amenorrhea persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Comments and Evidence Summary. During contraceptive counseling and before initiation of DMPA, information about common side effects such as irregular bleeding should be discussed. Unscheduled bleeding or spotting is common with DMPA use (194). In addition, amenorrhea is common after ≥ 1 years of continuous use (194,195). These bleeding irregularities are generally not harmful. Enhanced counseling among DMPA users detailing expected bleeding patterns and reassurance that these irregularities generally are not harmful has been shown to reduce DMPA discontinuation in clinical trials (124,125).

A systematic review, as well as two additional studies, examined the treatment of bleeding irregularities during DMPA use (195–197). Two small studies found significant cessation of bleeding within 7 days of starting treatment among women taking valdecoxib for 5 days or mefenamic acid for 5 days compared with placebo (198,199). Treatment with ethinyl estradiol was found to stop bleeding better than placebo during the treatment period, although rates of discontinuation were high and safety outcomes were not examined (200). In one small study among DMPA users who had been experiencing amenorrhea for 2 months, treatment with COCs was found to alleviate amenorrhea better than placebo (201). No studies examined the effects of aspirin on bleeding irregularities among DMPA users.

Combined Hormonal Contraceptives

Combined hormonal contraceptives contain both estrogen and a progestin and include 1) COCs (various formulations), 2) a transdermal contraceptive patch (which releases 150 μg of norelgestromin and 20 μg ethinyl estradiol daily), and 3) a vaginal contraceptive ring (which releases 120 μg etonogestrel and 15 μg ethinyl estradiol daily). Approximately 9 out of 100 women become pregnant in the first year of use with combined hormonal contraceptives with typical use (14). These methods are reversible and can be used by women of all ages. Combined hormonal contraceptives are generally used for

21–24 consecutive days, followed by 4–7 hormone-free days (either no use or placebo pills). These methods are sometimes used for an extended period with infrequent or no hormone-free days. Combined hormonal contraceptives do not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

Initiation of Combined Hormonal Contraceptives

Timing

- Combined hormonal contraceptives can be initiated at any time if it is reasonably certain that the woman is not pregnant (Box 2).

Need for Back-Up Contraception

- If combined hormonal contraceptives are started within the first 5 days since menstrual bleeding started, no additional contraceptive protection is needed.
- If combined hormonal contraceptives are started >5 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Special Considerations

Amenorrhea (Not Postpartum)

- **Timing:** Combined hormonal contraceptives can be started at any time if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postpartum (Breastfeeding)

- **Timing:** Combined hormonal contraceptives can be started when the woman is medically eligible to use the method (5) and if it is reasonably certain that she is not pregnant. (Box 2).
- Postpartum women who are breastfeeding should not use combined hormonal contraceptives during the first 3 weeks after delivery (U.S. MEC 4) because of concerns about increased risk for venous thromboembolism and generally should not use combined hormonal contraceptives during the fourth week postpartum (U.S. MEC 3) because of concerns about potential effects on breastfeeding performance. Postpartum breastfeeding women with other risk factors for venous thromboembolism generally should not use combined hormonal contraceptives 4–6 weeks after delivery (U.S. MEC 3).

- **Need for back-up contraception:** If the woman is <6 months postpartum, amenorrheic, and fully or nearly fully breastfeeding (exclusively breastfeeding or the vast majority [$\geq 85\%$] of feeds are breastfeeds) (27), no additional contraceptive protection is needed. Otherwise, a woman who is ≥ 21 days postpartum and has not experienced return of her menstrual cycle needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days. If her menstrual cycles have returned and it has been >5 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postpartum (Not Breastfeeding)

- **Timing:** Combined hormonal contraceptives can be started when the woman is medically eligible to use the method (5) and if it is reasonably certain that she is not pregnant (Box 2).
- Postpartum women should not use combined hormonal contraceptives during the first 3 weeks after delivery (U.S. MEC 4) because of concerns about increased risk for venous thromboembolism. Postpartum women with other risk factors for venous thromboembolism generally should not use combined hormonal contraceptives 3–6 weeks after delivery (U.S. MEC 3).
- **Need for back-up contraception:** If a woman is <21 days postpartum, no additional contraceptive protection is needed. A woman who is ≥ 21 days postpartum and whose menstrual cycles have not returned needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days. If her menstrual cycles have returned and it has been >5 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.

Postabortion (Spontaneous or Induced)

- **Timing:** Combined hormonal contraceptives can be started within the first 7 days following first-trimester or second-trimester abortion, including immediately postabortion (U.S. MEC 1).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days unless combined hormonal contraceptives are started at the time of a surgical abortion.

Switching from Another Contraceptive Method

- **Timing:** Combined hormonal contraceptives can be started immediately if it is reasonably certain that the

woman is not pregnant (Box 2). Waiting for her next menstrual cycle is unnecessary.

- **Need for back-up contraception:** If it has been >5 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 7 days.
- **Switching from an IUD:** If the woman has had sexual intercourse since the start of her current menstrual cycle and it has been >5 days since menstrual bleeding started, theoretically, residual sperm might be in the genital tract, which could lead to fertilization if ovulation occurs. A health care provider may consider any of the following options:
 - Advise the women to retain the IUD for at least 7 days after combined hormonal contraceptives are initiated and return for IUD removal.
 - Advise the woman to abstain from sexual intercourse or use barrier contraception for 7 days before removing the IUD and switching to the new method.
 - If the woman cannot return for IUD removal and has not abstained from sexual intercourse or used barrier contraception for 7 days, advise the woman to use ECPs at the time of IUD removal. Combined hormonal contraceptives can be started immediately after use of ECPs (with the exception of UPA). Combined hormonal contraceptives can be started no sooner than 5 days after use of UPA.

Comments and Evidence Summary. In situations in which the health care provider is uncertain whether the woman might be pregnant, the benefits of starting combined hormonal contraceptives likely exceed any risk; therefore, starting combined hormonal contraceptives should be considered at any time, with a follow-up pregnancy test in 2–4 weeks. If a woman needs to use additional contraceptive protection when switching to combined hormonal contraceptives from another contraceptive method, consider continuing her previous method for 7 days after starting combined hormonal contraceptives.

A systematic review of 18 studies examined the effects of starting combined hormonal contraceptives on different days of the menstrual cycle (202). Overall, the evidence suggested that pregnancy rates did not differ by the timing of combined hormonal contraceptive initiation (169,203–205) (Level of evidence: I to II-3, fair, indirect). The more follicular activity that occurred before starting COCs, the more likely ovulation was to occur; however, no ovulations occurred when COCs were started at a follicle diameter of 10 mm (mean cycle day 7.6) or when the ring was started at 13 mm (median cycle day 11) (206–215) (Level of evidence: I to II-3, fair, indirect). Bleeding patterns and other side effects did not vary with the timing of combined hormonal contraceptive initiation (204,205,216–220) (Level of evidence: I to II-2,

good to poor, direct). Although continuation rates of combined hormonal contraceptives were initially improved by the “quick start” approach (i.e., starting on the day of the visit), the advantage disappeared over time (203,204,216–221) (Level of evidence: I to II-2, good to poor, direct).

Examinations and Test Needed Before Initiation of Combined Hormonal Contraceptives

Among healthy women, few examinations or tests are needed before initiation of combined hormonal contraceptives (Table 4). Blood pressure should be measured before initiation of combined hormonal contraceptives. Baseline weight and BMI measurements might be useful for monitoring combined hormonal contraceptive users over time. Women with known medical problems or other special conditions might need additional examinations or tests before being determined to be appropriate candidates for a particular method of contraception. U.S. MEC might be useful in such circumstances (5).

Comments and Evidence Summary. Blood pressure: Women who have more severe hypertension (systolic pressure of ≥ 160 mmHg or diastolic pressure of ≥ 100 mm Hg) or vascular disease should not use combined hormonal contraceptives (U.S. MEC 4), and women who have less severe hypertension (systolic pressure of 140–159 mm Hg or diastolic pressure of 90–99 mm Hg) or adequately controlled hypertension generally should not use combined hormonal contraceptives (U.S. MEC 3) (5). Therefore, blood pressure should be evaluated before initiating combined hormonal contraceptives. In instances in which blood pressure cannot be measured by a provider, blood pressure measured in other settings can be reported by the woman to her provider. Evidence suggests that cardiovascular outcomes are worse among women who did not have their blood pressure measured before initiating COCs. A systematic review identified six articles from three studies that reported cardiovascular outcomes among women who had blood pressure measurements and women who did not have blood pressure measurements before initiating COCs (170). Three case-control studies showed that women who did not have blood pressure measurements before initiating COCs had a higher risk for acute myocardial infarction than women who did have blood pressure measurements (222–224). Two case-control studies showed that women who did not have blood pressure measurements before initiating COCs had a higher risk for ischemic stroke than women who did have blood pressure measurements (225,226). One case-control study showed no difference in the risk for hemorrhagic stroke among women who initiated COCs regardless of whether their

TABLE 4. Classification of examinations and tests needed before combined hormonal contraceptive initiation

Examination or test	Class*
Examination	
Blood pressure	A [†]
Weight (BMI) (weight [kg]/height [m] ²)	— [§]
Clinical breast examination	C
Bimanual examination and cervical inspection	C
Laboratory test	
Glucose	C
Lipids	C
Liver enzymes	C
Hemoglobin	C
Thrombogenic mutations	C
Cervical cytology (Papanicolaou smear)	C
STD screening with laboratory tests	C
HIV screening with laboratory tests	C

Abbreviations: BMI = body mass index; HIV = human immunodeficiency virus; STD = sexually transmitted disease; U.S. MEC = U.S. Medical Eligibility Criteria for Contraceptive Use.

* **Class A:** essential and mandatory in all circumstances for safe and effective use of the contraceptive method. **Class B:** contributes substantially to safe and effective use, but implementation may be considered within the public health and/or service context; the risk of not performing an examination or test should be balanced against the benefits of making the contraceptive method available. **Class C:** does not contribute substantially to safe and effective use of the contraceptive method.

[†] In instances in which blood pressure cannot be measured by a provider, blood pressure measured in other settings can be reported by the woman to her provider.

[§] Weight (BMI) measurement is not needed to determine medical eligibility for any methods of contraception because all methods can be used (U.S. MEC 1) or generally can be used (U.S. MEC 2) among obese women (Box 1). However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

blood pressure was measured (227). Studies that examined hormonal contraceptive methods other than COCs were not identified (Level of evidence: II-2, fair, direct).

Weight (BMI): Obese women generally can use combined hormonal contraceptives (U.S. MEC 2) (5); therefore, screening for obesity is not necessary for the safe initiation of combined hormonal contraceptives. However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

Bimanual examination and cervical inspection: Pelvic examination is not necessary before initiation of combined hormonal contraceptives because it does not facilitate detection of conditions for which hormonal contraceptives would be unsafe. Women with certain conditions such as current breast cancer, severe hypertension or vascular disease, heart disease, migraine headaches with aura, and certain liver diseases, as well as women aged ≥ 35 years and who smoke ≥ 15 cigarettes per day, should not use (U.S. MEC 4) or generally should not use (U.S. MEC 3) combined hormonal contraceptives (5); however, none of these conditions are likely to be detected by pelvic examination (145). A systematic review

identified two case-control studies that compared delayed and immediate pelvic examination before initiation of hormonal contraceptives, specifically oral contraceptives or DMPA (95). No differences in risk factors for cervical neoplasia, incidence of STDs, incidence of abnormal Papanicolaou smears, or incidence of abnormal wet mounts were found (Level of evidence: Level II-2 fair, direct).

Glucose: Although women with complicated diabetes should not use (U.S. MEC 4) or generally should not use (U.S. MEC 3) combined hormonal contraceptives, depending on the severity of the condition (5), screening for diabetes before initiation of hormonal contraceptives is not necessary because of the low prevalence of undiagnosed diabetes and the high likelihood that women with complicated diabetes already would have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with glucose measurement before initiation of hormonal contraceptives (57). The prevalence of diabetes among women of reproductive age is low. During 2009–2012 among women aged 20–44 years in the United States, the prevalence of diabetes was 3.3% (84). During 1999–2008, the percentage of women aged 20–44 years with undiagnosed diabetes was 0.5% (85). Although hormonal contraceptives can have some adverse effects on glucose metabolism in healthy and diabetic women, the overall clinical effect is minimal (171–177).

Lipids: Screening for dyslipidemias is not necessary for the safe initiation of combined hormonal contraceptives because of the low prevalence of undiagnosed disease in women of reproductive age and the low likelihood of clinically significant changes with use of hormonal contraceptives. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with lipid measurement before initiation of hormonal contraceptives (57). During 2009–2012 among women aged 20–44 years in the United States, 7.6% had high cholesterol, defined as total serum cholesterol ≥ 240 mg/dL (84). During 1999–2008, the prevalence of undiagnosed hypercholesterolemia among women aged 20–44 years was approximately 2% (85). A systematic review identified few studies, all of poor quality, that suggest that women with known dyslipidemias using combined hormonal contraceptives might be at increased risk for myocardial infarction, cerebrovascular accident, or venous thromboembolism compared with women without dyslipidemias; no studies were identified that examined risk for pancreatitis among women with known dyslipidemias using combined hormonal contraceptives (89). Studies have shown mixed results regarding the effects of hormonal contraceptives on lipid levels among both healthy women and women with

baseline lipid abnormalities, and the clinical significance of these changes is unclear (86–89).

Liver enzymes: Although women with certain liver diseases should not use (U.S. MEC 4) or generally should not use (U.S. MEC 3) combined hormonal contraceptives (5), screening for liver disease before initiation of combined hormonal contraceptives is not necessary because of the low prevalence of these conditions and the high likelihood that women with liver disease already would have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with liver enzyme tests before initiation of hormonal contraceptives (57). In 2012, among U.S. women, the percentage with liver disease (not further specified) was 1.3% (90). In 2013, the incidence of acute hepatitis A, B, or C was ≤ 1 per 100,000 U.S. population (91). During 2002–2011, the incidence of liver carcinoma among U.S. women was approximately 3.7 per 100,000 population (92). Because estrogen and progestins are metabolized in the liver, the use of hormonal contraceptives among women with liver disease might, theoretically, be a concern. The use of hormonal contraceptives, specifically COCs and POPs, does not affect disease progression or severity in women with hepatitis, cirrhosis, or benign focal nodular hyperplasia (93,94), although evidence is limited; no evidence exists for other types of combined hormonal contraceptives.

Thrombogenic mutations: Women with thrombogenic mutations should not use combined hormonal contraceptives (U.S. MEC 4) (5) because of the increased risk for venous thromboembolism (228). However, studies have shown that universal screening for thrombogenic mutations before initiating COCs is not cost-effective because of the rarity of the conditions and the high cost of screening (229–231).

Clinical breast examination: Although women with current breast cancer should not use combined hormonal contraceptives (U.S. MEC 4) (5), screening asymptomatic women with a clinical breast examination before initiating combined hormonal contraceptives is not necessary because of the low prevalence of breast cancer among women of reproductive age. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with a breast examination before initiation of hormonal contraceptives (95). The incidence of breast cancer among women of reproductive age in the United States is low. In 2012, the incidence of breast cancer among women aged 20–49 years was approximately 70.7 per 100,000 women (96).

Other screening: Women with anemia, cervical intraepithelial neoplasia, cervical cancer, HIV infection, or other STDs can use (U.S. MEC 1) or generally can use (U.S. MEC 2) combined

hormonal contraceptives (5); therefore, screening for these conditions is not necessary for the safe initiation of combined hormonal contraceptives.

Number of Pill Packs that Should Be Provided at Initial and Return Visits

- At the initial and return visits, provide or prescribe up to a 1-year supply of COCs (e.g., 13 28-day pill packs), depending on the woman's preferences and anticipated use.
- A woman should be able to obtain COCs easily in the amount and at the time she needs them.

Comments and Evidence Summary. The more pill packs given up to 13 cycles, the higher the continuation rates. Restricting the number of pill packs distributed or prescribed can result in unwanted discontinuation of the method and increased risk for pregnancy.

A systematic review of the evidence suggested that providing a greater number of pill packs was associated with increased continuation (232). Studies that compared provision of one versus 12 packs, one versus 12 or 13 packs, or three versus seven packs found increased continuation of pill use among women provided with more pill packs (233–235). However, one study found no difference in continuation when patients were provided one and then three packs versus four packs all at once (236). In addition to continuation, a greater number of pill packs provided was associated with fewer pregnancy tests, fewer pregnancies, and lower cost per client. However, a greater number of pill packs (i.e., 13 packs versus three packs) also was associated with increased pill wastage in one study (234) (Level of evidence: I to II-2, fair, direct).

Routine Follow-Up After Combined Hormonal Contraceptive Initiation

These recommendations address when routine follow-up is recommended for safe and effective continued use of contraception for healthy women. The recommendations refer to general situations and might vary for different users and different situations. Specific populations who might benefit from more frequent follow-up visits include adolescents, those with certain medical conditions or characteristics, and those with multiple medical conditions.

- Advise the woman to return at any time to discuss side effects or other problems or if she wants to change the method being used. No routine follow-up visit is required.
- At other routine visits, health care providers seeing combined hormonal contraceptive users should do the following:

- Assess the woman's satisfaction with her contraceptive method and whether she has any concerns about method use.
- Assess any changes in health status, including medications, that would change the appropriateness of combined hormonal contraceptives for safe and effective continued use based on U.S. MEC (e.g., category 3 and 4 conditions and characteristics).
- Assess blood pressure.
- Consider assessing weight changes and counseling women who are concerned about weight change perceived to be associated with their contraceptive method.

Comments and Evidence Summary. No evidence exists regarding whether a routine follow-up visit after initiating combined hormonal contraceptives improves correct or continued use. Monitoring blood pressure is important for combined hormonal contraceptive users. Health care providers might consider recommending women obtain blood pressure measurements in other settings.

A systematic review identified five studies that examined the incidence of hypertension among women who began using a COC versus those who started a nonhormonal method of contraception or a placebo (123). Few women developed hypertension after initiating COCs, and studies examining increases in blood pressure after COC initiation found mixed results. No studies were identified that examined changes in blood pressure among patch or vaginal ring users (Level of evidence: I, fair, to II-2, fair, indirect).

Late or Missed Doses and Side Effects from Combined Hormonal Contraceptive Use

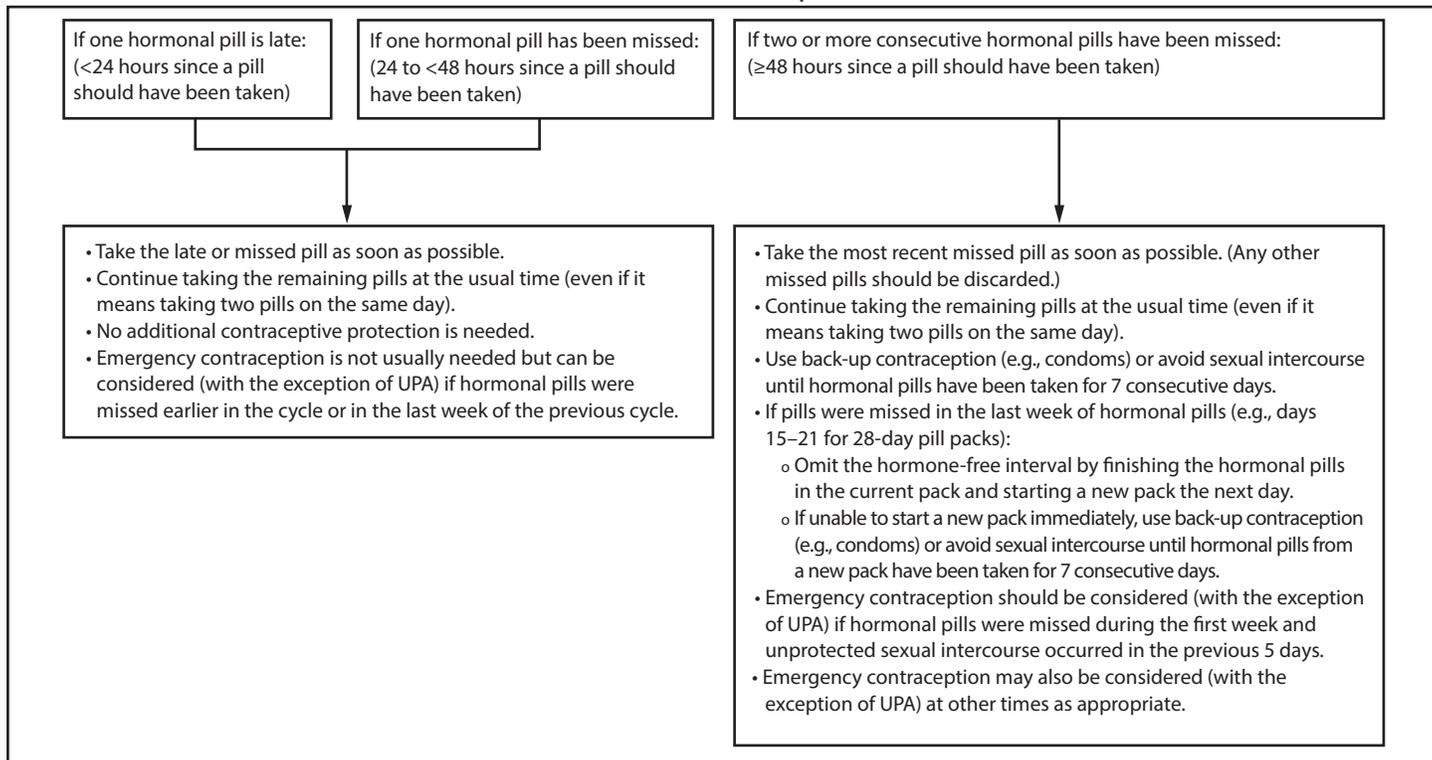
For the following recommendations, a dose is considered late when <24 hours have elapsed since the dose should have been taken. A dose is considered missed if ≥ 24 hours have elapsed since the dose should have been taken. For example, if a COC pill was supposed to have been taken on Monday at 9:00 a.m. and is taken at 11:00 a.m., the pill is late; however, by Tuesday morning at 11:00 a.m., Monday's 9:00 a.m. pill has been missed and Tuesday's 9:00 a.m. pill is late. For COCs, the recommendations only apply to late or missed hormonally active pills and not to placebo pills. Recommendations are provided for late or missed pills (Figure 2), the patch (Figure 3), and the ring (Figure 4).

Comments and Evidence Summary. Inconsistent or incorrect use of combined hormonal contraceptives is a major cause of combined hormonal contraceptive failure. Extending the hormone-free interval is considered to be a particularly risky time to miss combined hormonal contraceptives. Seven days of continuous combined hormonal contraceptive use is deemed

necessary to reliably prevent ovulation. The recommendations reflect a balance between simplicity and precision of science. Women who frequently miss COCs or experience other usage errors with combined hormonal patch or combined vaginal ring should consider an alternative contraceptive method that is less dependent on the user to be effective (e.g., IUD, implant, or injectable).

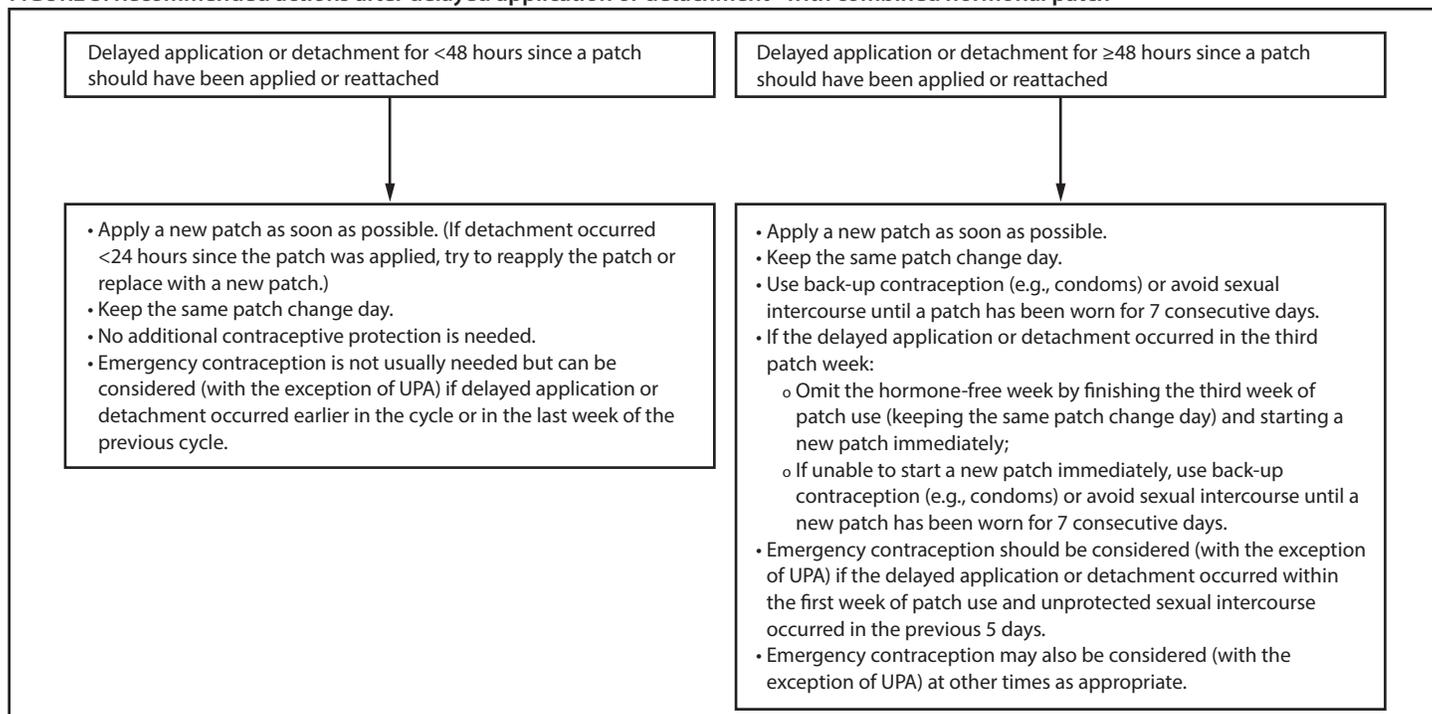
A systematic review identified 36 studies that examined measures of contraceptive effectiveness of combined hormonal contraceptives during cycles with extended hormone-free intervals, shortened hormone-free intervals, or deliberate nonadherence on days not adjacent to the hormone-free interval (237). Most of the studies examined COCs (215,238–265), two examined the combined hormonal patch (259,266), and six examined the combined vaginal ring (211,267–271). No direct evidence on the effect of missed pills on the risk for pregnancy was found. Studies of women deliberately extending the hormone-free interval up to 14 days found wide variability in the amount of follicular development and occurrence of ovulation (241,244,246,247,249,250,252–255); in general, the risk for ovulation was low, and among women who did ovulate, cycles were usually abnormal. In studies of women who deliberately missed pills on various days during the cycle not adjacent to the hormone-free interval, ovulation occurred infrequently (239,245–247,255,256,258,259). Studies comparing 7-day hormone-free intervals with shorter hormone-free intervals found lower rates of pregnancy (238,242,251,257) and significantly greater suppression of ovulation (240,250,261–263,265) among women with shorter intervals in all but one study (260), which found no difference. Two studies that compared 30- μg ethinyl estradiol pills with 20- μg ethinyl estradiol pills showed more follicular activity when 20- μg ethinyl estradiol pills were missed (241,244). In studies examining the combined vaginal ring, three studies found that nondeliberate extension of the hormone-free interval for 24 to <48 hours from the scheduled period did not increase the risk for pregnancy (267,268,270); one study found that ring insertion after a deliberately extended hormone-free interval that allowed a 13-mm follicle to develop interrupted ovarian function and further follicular growth (211); and one study found that inhibition of ovulation was maintained after deliberately forgetting to remove the ring for up to 2 weeks after normal ring use (271). In studies examining the combined hormonal patch, one study found that missing 1–3 consecutive days before patch replacement (either wearing one patch 3 days longer before replacement or going 3 days without a patch before replacing the next patch) on days not adjacent to the patch-free interval resulted in little follicular activity and low risk for ovulation (259), and one pharmacokinetic study found that serum levels of

FIGURE 2. Recommended actions after late or missed combined oral contraceptives



Abbreviation: UPA = ulipristal acetate.

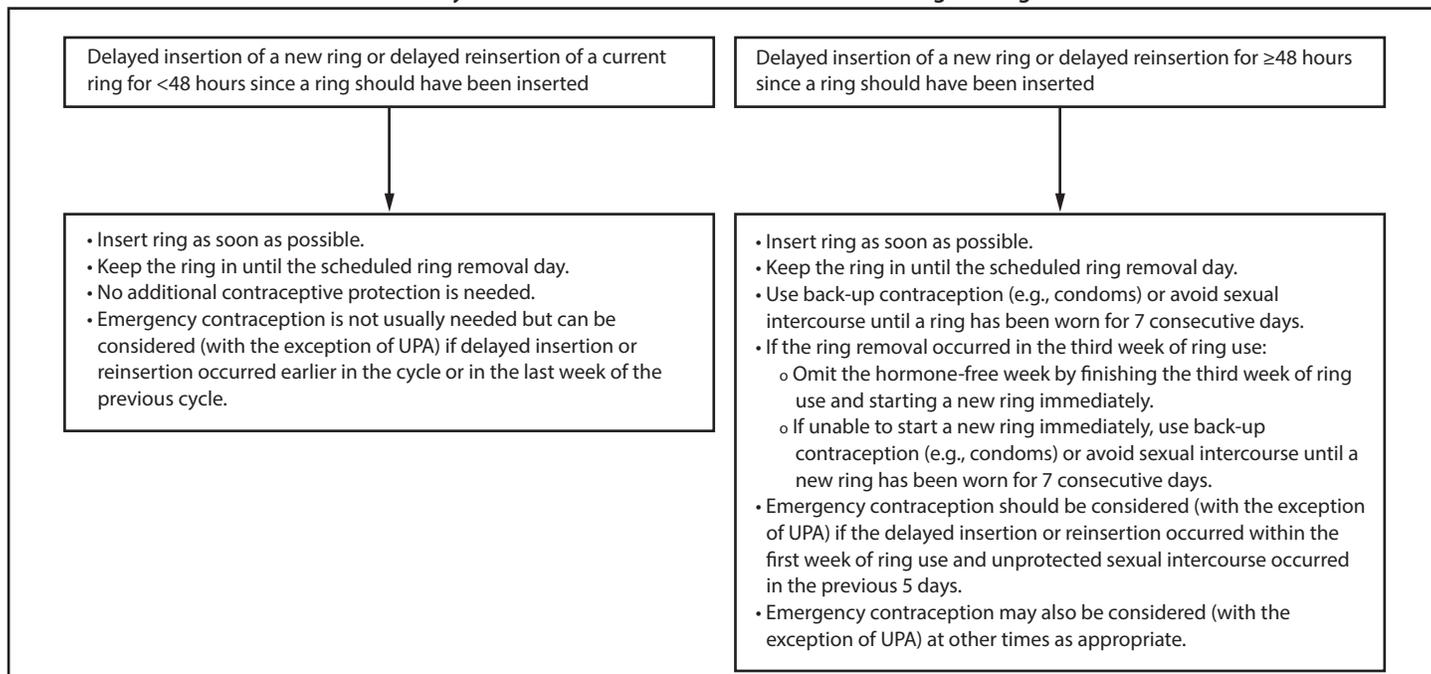
FIGURE 3. Recommended actions after delayed application or detachment* with combined hormonal patch



Abbreviation: UPA = ulipristal acetate.

* If detachment takes place but the woman is unsure when the detachment occurred, consider the patch to have been detached for ≥48 hours since a patch should have been applied or reattached.

FIGURE 4. Recommended actions after delayed insertion or reinsertion* with combined vaginal ring



Abbreviation: UPA = ulipristal acetate.

* If removal takes place but the woman is unsure of how long the ring has been removed, consider the ring to have been removed for ≥48 hours since a ring should have been inserted or reinserted.

ethinyl estradiol and progestin norelgestromin remained within reference ranges after extending patch wear for 3 days (266). No studies were found on extending the patch-free interval. In studies that provide indirect evidence on the effects of missed combined hormonal contraception on surrogate measures of pregnancy, how differences in surrogate measures correspond to pregnancy risk is unclear (Level of evidence: I, good, indirect to II-3, poor, direct).

Vomiting or Severe Diarrhea While Using COCs

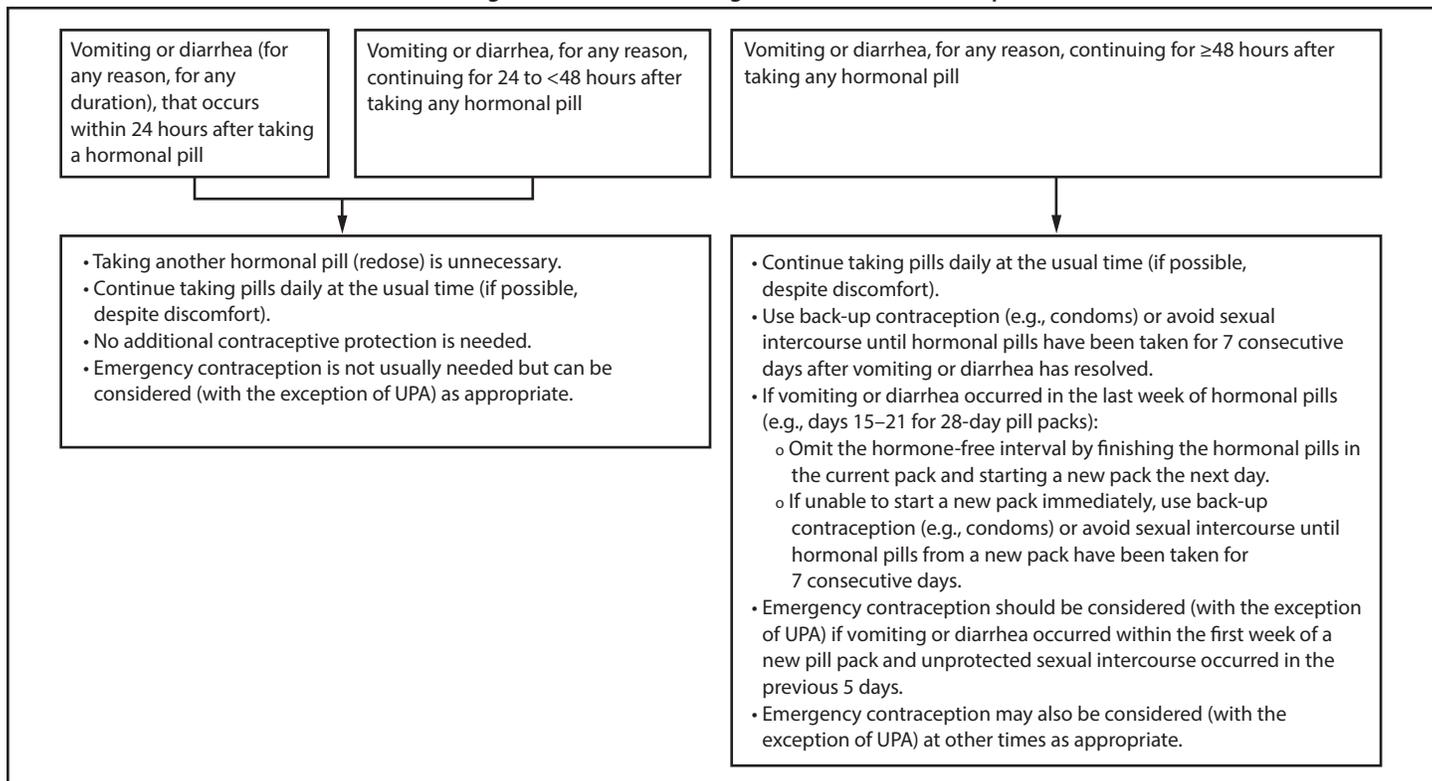
Certain steps should be taken by women who experience vomiting or severe diarrhea while using COCs (Figure 5).

Comments and Evidence Summary. Theoretically, the contraceptive effectiveness of COCs might be decreased because of vomiting or severe diarrhea. Because of the lack of evidence that addresses vomiting or severe diarrhea while using COCs, these recommendations are based on the recommendations for missed COCs. No evidence was found on the effects of vomiting or diarrhea on measures of contraceptive effectiveness including pregnancy, follicular development, hormone levels, or cervical mucus quality.

Unscheduled Bleeding with Extended or Continuous Use of Combined Hormonal Contraceptives

- Before initiation of combined hormonal contraceptives, provide counseling about potential changes in bleeding patterns during extended or continuous combined hormonal contraceptive use. (Extended contraceptive use is defined as a planned hormone-free interval after at least two contiguous cycles. Continuous contraceptive use is defined as uninterrupted use of hormonal contraception without a hormone-free interval) (272).
- Unscheduled spotting or bleeding is common during the first 3–6 months of extended or continuous combined hormonal contraceptive use. It is generally not harmful and decreases with continued combined hormonal contraceptive use.
- If clinically indicated, consider an underlying gynecological problem, such as inconsistent use, interactions with other medications, cigarette smoking, an STD, pregnancy, or new pathologic uterine conditions (e.g., polyps or fibroids). If an underlying gynecological problem is found, treat the condition or refer for care.
- If an underlying gynecological problem is not found and the woman wants treatment, the following treatment option can be considered:

FIGURE 5. Recommended actions after vomiting or diarrhea while using combined oral contraceptives



Abbreviation: UPA = ulipristal acetate.

- Advise the woman to discontinue combined hormonal contraceptive use (i.e., a hormone-free interval) for 3–4 consecutive days; a hormone-free interval is not recommended during the first 21 days of using the continuous or extended combined hormonal contraceptive method. A hormone-free interval also is not recommended more than once per month because contraceptive effectiveness might be reduced.
- If unscheduled spotting or bleeding persists and the woman finds it unacceptable, counsel her on alternative contraceptive methods, and offer another method if it is desired.

Comments and Evidence Summary. During contraceptive counseling and before initiating extended or continuous combined hormonal contraceptives, information about common side effects such as unscheduled spotting or bleeding, especially during the first 3–6 months of use, should be discussed (273). These bleeding irregularities are generally not harmful and usually improve with persistent use of the hormonal method. To avoid unscheduled spotting or bleeding, counseling should emphasize the importance of correct use and timing; for users of contraceptive pills, emphasize consistent pill use. Enhanced counseling about expected bleeding patterns

and reassurance that bleeding irregularities are generally not harmful has been shown to reduce method discontinuation in clinical trials with DMPA (124,125,274).

A systematic review identified three studies with small study populations that addressed treatments for unscheduled bleeding among women using extended or continuous combined hormonal contraceptives (275). In two separate randomized clinical trials in which women were taking either contraceptive pills or using the contraceptive ring continuously for 168 days, women assigned to a hormone-free interval of 3 or 4 days reported improved bleeding. Although they noted an initial increase in flow, this was followed by an abrupt decrease 7–8 days later with eventual cessation of flow 11–12 days later. These findings were compared with women who continued to use their method without a hormone-free interval, in which a greater proportion reported either treatment failure or fewer days of amenorrhea (276,277). In another randomized trial of 66 women with unscheduled bleeding among women using 84 days of hormonally active contraceptive pills, oral doxycycline (100 mg twice daily) initiated the first day of bleeding and taken for 5 days did not result in any improvement in bleeding compared with placebo (278) (Level of evidence: I, fair, direct).

Progestin-Only Pills

POPs contain only a progestin and no estrogen and are available in the United States. Approximately 9 out of 100 women become pregnant in the first year of use with POPs with typical use (14). POPs are reversible and can be used by women of all ages. POPs do not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

Initiation of POPs

Timing

- POPs can be started at any time if it is reasonably certain that the woman is not pregnant (Box 2).

Need for Back-Up Contraception

- If POPs are started within the first 5 days since menstrual bleeding started, no additional contraceptive protection is needed.
- If POPs are started >5 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days.

Special Considerations

Amenorrhea (Not Postpartum)

- **Timing:** POPs can be started at any time if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days.

Postpartum (Breastfeeding)

- **Timing:** POPs can be started at any time, including immediately postpartum (U.S. MEC 2 if <1 month postpartum; U.S. MEC 1 if ≥1 month postpartum) if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** If the woman is <6 months postpartum, amenorrheic, and fully or nearly fully breastfeeding (exclusively breastfeeding or the vast majority [≥85%] of feeds are breastfeeds) (27), no additional contraceptive protection is needed. Otherwise, a woman who is ≥21 days postpartum and has not experienced return of her menstrual cycles, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days. If her menstrual cycles have returned and it has been >5 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days.

Postpartum (Not Breastfeeding)

- **Timing:** POPs can be started at any time, including immediately postpartum (U.S. MEC 1), if it is reasonably certain that the woman is not pregnant (Box 2).
- **Need for back-up contraception:** If a woman is <21 days postpartum, no additional contraceptive protection is needed. Women who are ≥21 days postpartum and whose menstrual cycles have not returned need to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days. If her menstrual cycles have returned and it has been >5 days since menstrual bleeding started, she needs to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days.

Postabortion (Spontaneous or Induced)

- **Timing:** POPs can be started within the first 7 days, including immediately postabortion (U.S. MEC 1).
- **Need for back-up contraception:** The woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days unless POPs are started at the time of a surgical abortion.

Switching from Another Contraceptive Method

- **Timing:** POPs can be started immediately if it is reasonably certain that the woman is not pregnant (Box 2). Waiting for her next menstrual cycle is unnecessary.
- **Need for back-up contraception:** If it has been >5 days since menstrual bleeding started, the woman needs to abstain from sexual intercourse or use additional contraceptive protection for the next 2 days.
- **Switching from an IUD:** If the woman has had sexual intercourse since the start of her current menstrual cycle and it has been >5 days since menstrual bleeding started, theoretically, residual sperm might be in the genital tract, which could lead to fertilization if ovulation occurs. A health care provider may consider any of the following options:
 - Advise the women to retain the IUD for at least 2 days after POPs are initiated and return for IUD removal.
 - Advise the woman to abstain from sexual intercourse or use barrier contraception for 7 days before removing the IUD and switching to the new method.
 - If the woman cannot return for IUD removal and has not abstained from sexual intercourse or used barrier contraception for 7 days, advise the woman to use ECPs at the time of IUD removal. POPs can be started immediately after use of ECPs (with the exception of UPA). POPs can be started no sooner than 5 days after use of UPA.

Comments and Evidence Summary. In situations in which the health care provider is uncertain whether the woman might

be pregnant, the benefits of starting POPs likely exceed any risk; therefore, starting POPs should be considered at any time, with a follow-up pregnancy test in 2–4 weeks.

Unlike COCs, POPs inhibit ovulation in about half of cycles, although the rates vary widely by individual (279). Peak serum steroid levels are reached about 2 hours after administration, followed by rapid distribution and elimination, such that by 24 hours after administration, serum steroid levels are near baseline (279). Therefore, taking POPs at approximately the same time each day is important. An estimated 48 hours of POP use has been deemed necessary to achieve the contraceptive effects on cervical mucus (279). If a woman needs to use additional contraceptive protection when switching to POPs from another contraceptive method, consider continuing her previous method for 2 days after starting POPs. No direct evidence was found regarding the effects of starting POPs at different times of the cycle.

Examinations and Tests Needed Before Initiation of POPs

Among healthy women, no examinations or tests are needed before initiation of POPs, although a baseline weight and BMI measurement might be useful for monitoring POP users over time (Table 5). Women with known medical problems or other special conditions might need additional examinations or tests before being determined to be appropriate candidates for a particular method of contraception. The U.S. MEC might be useful in such circumstances (5).

Comments and Evidence Summary. Weight (BMI): Obese women can use POPs (U.S. MEC 1) (5); therefore, screening for obesity is not necessary for the safe initiation of POPs. However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

Bimanual examination and cervical inspection: Pelvic examination is not necessary before initiation of POPs because it does not facilitate detection of conditions for which POPs would be unsafe. Women with current breast cancer should not use POPs (U.S. MEC 4), and women with certain liver diseases generally should not use POPs (U.S. MEC 3) (5); however, neither of these conditions are likely to be detected by pelvic examination (145). A systematic review identified two case-control studies that compared delayed versus immediate pelvic examination before initiation of hormonal contraceptives, specifically oral contraceptives or DMPA (95). No differences in risk factors for cervical neoplasia, incidence of STDs, incidence of abnormal Papanicolaou smears, or incidence of

TABLE 5. Classification of examinations and tests needed before progestin-only pill initiation

Examination or test	Class*
Examination	
Blood pressure	C
Weight (BMI) (weight [kg]/height [m] ²)	—†
Clinical breast examination	C
Bimanual examination and cervical inspection	C
Laboratory test	
Glucose	C
Lipids	C
Liver enzymes	C
Hemoglobin	C
Thrombogenic mutations	C
Cervical cytology (Papanicolaou smear)	C
STD screening with laboratory tests	C
HIV screening with laboratory tests	C

Abbreviations: BMI = body mass index; HIV = human immunodeficiency virus; STD = sexually transmitted disease; U.S. MEC = U.S. Medical Eligibility Criteria for Contraceptive Use.

* **Class A:** essential and mandatory in all circumstances for safe and effective use of the contraceptive method. **Class B:** contributes substantially to safe and effective use, but implementation may be considered within the public health and/or service context; the risk of not performing an examination or test should be balanced against the benefits of making the contraceptive method available. **Class C:** does not contribute substantially to safe and effective use of the contraceptive method.

† Weight (BMI) measurement is not needed to determine medical eligibility for any methods of contraception because all methods can be used (U.S. MEC 1) or generally can be used (U.S. MEC 2) among obese women (Box 1). However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

abnormal findings from wet mounts were observed (Level of evidence: II-2 fair, direct).

Lipids: Screening for dyslipidemias is not necessary for the safe initiation of POPs because of the low prevalence of undiagnosed disease in women of reproductive age and the low likelihood of clinically significant changes with use of hormonal contraceptives. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with lipid measurement before initiation of hormonal contraceptives (57). During 2009–2012 among women aged 20–44 years in the United States, 7.6% had high cholesterol, defined as total serum cholesterol ≥ 240 mg/dL (84). During 1999–2008, the prevalence of undiagnosed hypercholesterolemia among women aged 20–44 years was approximately 2% (85). Studies have shown mixed results about the effects of hormonal methods on lipid levels among both healthy women and women with baseline lipid abnormalities, and the clinical significance of these changes is unclear (86–89).

Liver enzymes: Although women with certain liver diseases generally should not use POPs (U.S. MEC 3) (5), screening for liver disease before initiation of POPs is not necessary because of the low prevalence of these conditions and the high likelihood that women with liver disease already would

have had the condition diagnosed. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with liver enzyme tests before initiation of hormonal contraceptives (57). In 2012, among U.S. women, the percentage with liver disease (not further specified) was 1.3% (90). In 2013, the incidence of acute hepatitis A, B, or C was ≤ 1 per 100,000 U.S. population (91). During 2002–2011, the incidence of liver carcinoma among U.S. women was approximately 3.7 per 100,000 population (92). Because estrogen and progestins are metabolized in the liver, the use of hormonal contraceptives among women with liver disease might, theoretically, be a concern. The use of hormonal contraceptives, specifically COCs and POPs, does not affect disease progression or severity in women with hepatitis, cirrhosis, or benign focal nodular hyperplasia (93,94).

Clinical breast examination: Although women with current breast cancer should not use POPs (U.S. MEC 4) (5), screening asymptomatic women with a clinical breast examination before initiating POPs is not necessary because of the low prevalence of breast cancer among women of reproductive age. A systematic review did not identify any evidence regarding outcomes among women who were screened versus not screened with a clinical breast examination before initiation of hormonal contraceptives (95). The incidence of breast cancer among women of reproductive age in the United States is low. In 2012, the incidence of breast cancer among women aged 20–49 years was approximately 70.7 per 100,000 women (96).

Other screening: Women with hypertension, diabetes, anemia, thrombogenic mutations, cervical intraepithelial neoplasia, cervical cancer, STDs, or HIV infection can use (U.S. MEC 1) or generally can use (U.S. MEC 2) POPs (5); therefore, screening for these conditions is not necessary for the safe initiation of POPs.

Number of Pill Packs that Should Be Provided at Initial and Return Visits

- At the initial and return visit, provide or prescribe up to a 1-year supply of POPs (e.g., 13 28-day pill packs), depending on the woman's preferences and anticipated use.
- A woman should be able to obtain POPs easily in the amount and at the time she needs them.

Comments and Evidence Summary. The more pill packs given up to 13 cycles, the higher the continuation rates. Restricting the number of pill packs distributed or prescribed can result in unwanted discontinuation of the method and increased risk for pregnancy.

A systematic review of the evidence suggested that providing a greater number of pill packs was associated with increased continuation (232). Studies that compared provision of one

versus 12 packs, one versus 12 or 13 packs, or three versus seven packs found increased continuation of pill use among women provided with more pill packs (233–235). However, one study found no difference in continuation when patients were provided one and then three packs versus four packs all at once (236). In addition to continuation, a greater number of pill packs provided was associated with fewer pregnancy tests, fewer pregnancies, and lower cost per client. However, a greater number of pill packs (13 packs versus three packs) also was associated with increased pill wastage in one study (234) (Level of evidence: I to II-2, fair, direct).

Routine Follow-Up After POP Initiation

These recommendations address when routine follow-up is recommended for safe and effective continued use of contraception for healthy women. The recommendations refer to general situations and might vary for different users and different situations. Specific populations who might benefit from more frequent follow-up visits include adolescents, those with certain medical conditions or characteristics, and those with multiple medical conditions.

- Advise the woman to return at any time to discuss side effects or other problems or if she wants to change the method being used. No routine follow-up visit is required.
- At other routine visits, health care providers seeing POP users should do the following:
 - Assess the woman's satisfaction with her contraceptive method and whether she has any concerns about method use.
 - Assess any changes in health status, including medications, that would change the appropriateness of POPs for safe and effective continued use based on U.S. MEC (e.g., category 3 and 4 conditions and characteristics).
 - Consider assessing weight changes and counseling women who are concerned about weight change perceived to be associated with their contraceptive method.

Comments and Evidence Summary. No evidence was found regarding whether a routine follow-up visit after initiating POPs improves correct and continued use.

Missed POPs

For the following recommendations, a dose is considered missed if it has been >3 hours since it should have been taken.

- Take one pill as soon as possible.
- Continue taking pills daily, one each day, at the same time each day, even if it means taking two pills on the same day.
- Use back-up contraception (e.g., condoms) or avoid sexual intercourse until pills have been taken correctly, on time, for 2 consecutive days.

- Emergency contraception should be considered (with the exception of UPA) if the woman has had unprotected sexual intercourse.

Comments and Evidence Summary. Inconsistent or incorrect use of oral contraceptive pills is a major reason for oral contraceptive failure. Unlike COCs, POPs inhibit ovulation in about half of cycles, although this rate varies widely by individual (279). Peak serum steroid levels are reached about 2 hours after administration, followed by rapid distribution and elimination, such that by 24 hours after administration, serum steroid levels are near baseline (279). Therefore, taking POPs at approximately the same time each day is important. An estimated 48 hours of POP use was deemed necessary to achieve the contraceptive effects on cervical mucus (279). Women who frequently miss POPs should consider an alternative contraceptive method that is less dependent on the user to be effective (e.g., IUD, implant, or injectable). No evidence was found regarding the effects of missed POPs available in the United States on measures of contraceptive effectiveness including pregnancy, follicular development, hormone levels, or cervical mucus quality.

Vomiting or Diarrhea (for any Reason or Duration) that Occurs Within 3 Hours After Taking a Pill

- Take another pill as soon as possible (if possible, despite discomfort).
- Continue taking pills daily, one each day, at the same time each day.
- Use back-up contraception (e.g., condoms) or avoid sexual intercourse until 2 days after vomiting or diarrhea has resolved.
- Emergency contraception should be considered (with the exception of UPA) if the woman has had unprotected sexual intercourse.

Comments and Evidence Summary. Theoretically, the contraceptive effectiveness of POPs might be decreased because of vomiting or severe diarrhea. Because of the lack of evidence to address this question, these recommendations are based on the recommendations for missed POPs. No evidence was found regarding the effects of vomiting or diarrhea on measures of contraceptive effectiveness, including pregnancy, follicular development, hormone levels, or cervical mucus quality.

Standard Days Method

SDM is a method based on fertility awareness; users must avoid unprotected sexual intercourse on days 8–19 of the menstrual cycle (280). Approximately 5 out of 100 women

become pregnant in the first year of use with perfect (i.e., correct and consistent) use of SDM (280); effectiveness based on typical use is not available for this method but is expected to be lower than that for perfect use. SDM is reversible and can be used by women of all ages. SDM does not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

Use of SDM Among Women with Various Durations of the Menstrual Cycle

Menstrual Cycles of 26–32 Days

- The woman may use the method.
- Provide a barrier method of contraception for protection on days 8–19 if she wants one.
- If she has unprotected sexual intercourse during days 8–19, consider the use of emergency contraception if appropriate.

Two or More Cycles of <26 or >32 Days Within Any 1 Year of SDM Use

- Advise the woman that the method might not be appropriate for her because of a higher risk for pregnancy. Help her consider another method.

Comments and Evidence Summary. The probability of pregnancy is increased when the menstrual cycle is outside the range of 26–32 days, even if unprotected sexual intercourse is avoided on days 8–19. A study of 7,600 menstrual cycles, including information on cycle length and signs of ovulation, concluded that the theoretical effectiveness of SDM is greatest for women with cycles of 26–32 days, that the method is still effective for women who occasionally have a cycle outside this range, and that it is less effective for women who consistently have cycles outside this range. Information from daily hormonal measurements shows that the timing of the 6-day fertile window varies greatly, even among women with regular cycles (21,281,282).

Emergency Contraception

Emergency contraception consists of methods that can be used by women after sexual intercourse to prevent pregnancy. Emergency contraception methods have varying ranges of effectiveness depending on the method and timing of administration. Four options are available in the United States: the Cu-IUD and three types of ECPs.

Types of Emergency Contraception

Intrauterine Device

- Cu-IUD

ECPs

- UPA in a single dose (30 mg)
- Levonorgestrel in a single dose (1.5 mg) or as a split dose (1 dose of 0.75 mg of levonorgestrel followed by a second dose of 0.75 mg of levonorgestrel 12 hours later)
- Combined estrogen and progestin in 2 doses (Yuzpe regimen: 1 dose of 100 μ g of ethinyl estradiol plus 0.50 mg of levonorgestrel followed by a second dose of 100 μ g of ethinyl estradiol plus 0.50 mg of levonorgestrel 12 hours later)

Initiation of Emergency Contraception

Timing

Cu-IUD

- The Cu-IUD can be inserted within 5 days of the first act of unprotected sexual intercourse as an emergency contraceptive.
- In addition, when the day of ovulation can be estimated, the Cu-IUD can be inserted beyond 5 days after sexual intercourse, as long as insertion does not occur >5 days after ovulation.

ECPs

- ECPs should be taken as soon as possible within 5 days of unprotected sexual intercourse.

Comments and Evidence Summary. Cu-IUDs are highly effective as emergency contraception (283) and can be continued as regular contraception. UPA and levonorgestrel ECPs have similar effectiveness when taken within 3 days after unprotected sexual intercourse; however, UPA has been shown to be more effective than the levonorgestrel formulation 3–5 days after unprotected sexual intercourse (284). The combined estrogen and progestin regimen is less effective than UPA or levonorgestrel and also is associated with more frequent occurrence of side effects (nausea and vomiting) (285). The levonorgestrel formulation might be less effective than UPA among obese women (286).

Two studies of UPA use found consistent decreases in pregnancy rates when administered within 120 hours of unprotected sexual intercourse (284,287). Five studies found that the levonorgestrel and combined regimens decreased risk for pregnancy through the fifth day after unprotected sexual intercourse; however, rates of pregnancy were slightly higher

when ECPs were taken after 3 days (288–292). A meta-analysis of levonorgestrel ECPs found that pregnancy rates were low when administered within 4 days after unprotected sexual intercourse but increased at 4–5 days (293) (Level of evidence: I to II-2, good to poor, direct).

Advance Provision of ECPs

- An advance supply of ECPs may be provided so that ECPs will be available when needed and can be taken as soon as possible after unprotected sexual intercourse.

Comments and Evidence Summary. A systematic review identified 17 studies that reported on safety or effectiveness of advance ECPs in adult or adolescent women (294). Any use of ECPs was two to seven times greater among women who received an advance supply of ECPs. However, a summary estimate (relative risk = 0.97; 95% confidence interval = 0.77–1.22) of five randomized controlled trials did not indicate a significant reduction in unintended pregnancies at 12 months with advance provision of ECPs. In the majority of studies among adults or adolescents, patterns of regular contraceptive use, pregnancy rates, and incidence of STDs did not vary between those who received advance ECPs and those who did not. Although available evidence supports the safety of advance provision of ECPs, effectiveness of advance provision of ECPs in reducing pregnancy rates at the population level has not been demonstrated (Level of evidence: I to II-3, good to poor, direct).

Initiation of Regular Contraception After ECPs

UPA

- Advise the woman to start or resume hormonal contraception no sooner than 5 days after use of UPA, and provide or prescribe the regular contraceptive method as needed. For methods requiring a visit to a health care provider, such as DMPA, implants, and IUDs, starting the method at the time of UPA use may be considered; the risk that the regular contraceptive method might decrease the effectiveness of UPA must be weighed against the risk of not starting a regular hormonal contraceptive method.
- The woman needs to abstain from sexual intercourse or use barrier contraception for the next 7 days after starting or resuming regular contraception or until her next menses, whichever comes first.
- Any nonhormonal contraceptive method can be started immediately after the use of UPA.
- Advise the woman to have a pregnancy test if she does not have a withdrawal bleed within 3 weeks.

Levonorgestrel and Combined Estrogen and Progestin ECPs

- Any regular contraceptive method can be started immediately after the use of levonorgestrel or combined estrogen and progestin ECPs.
- The woman needs to abstain from sexual intercourse or use barrier contraception for 7 days.
- Advise the woman to have a pregnancy test if she does not have a withdrawal bleed within 3 weeks.

Comments and Evidence Summary. The resumption or initiation of regular hormonal contraception after ECP use involves consideration of the risk for pregnancy if ECPs fail and the risks for unintended pregnancy if contraception initiation is delayed until the subsequent menstrual cycle. A health care provider may provide or prescribe pills, the patch, or the ring for a woman to start no sooner than 5 days after use of UPA. For methods requiring a visit to a health care provider, such as DMPA, implants, and IUDs, starting the method at the time of UPA use may be considered; the risk that the regular contraceptive method might decrease the effectiveness of UPA must be weighed against the risk of not starting a regular hormonal contraceptive method.

Data on when a woman can start regular contraception after ECPs are limited to pharmacodynamic data and expert opinion (295–297). In one pharmacodynamic study of women who were randomly assigned to either UPA or placebo groups mid-cycle followed by a 21-day course of combined hormonal contraception found no difference between UPA and placebo groups in the time for women's ovaries to reach quiescence by ultrasound and serum estradiol (296); this finding suggests that UPA did not have an effect on the combined hormonal contraception. In another pharmacodynamic study with a crossover design, women were randomly assigned to one of three groups: 1) UPA followed by desogestrel for 20 days started 1 day later; 2) UPA plus placebo; or 3) placebo plus desogestrel for 20 days (295). Among women taking UPA followed by desogestrel, a higher incidence of ovulation in the first 5 days was found compared with UPA alone (45% versus 3%, respectively), suggesting desogestrel might decrease the effectiveness of UPA. No concern exists that administering combined estrogen and progestin or levonorgestrel formulations of ECPs concurrently with systemic hormonal contraception decreases the effectiveness of either emergency or regular contraceptive methods because these formulations do not have antiprogestin properties like UPA. If a woman is planning to initiate contraception after the next menstrual bleeding after ECP use, the cycle in which ECPs are used might be shortened, prolonged, or involve unscheduled bleeding.

Prevention and Management of Nausea and Vomiting with ECP Use

Nausea and Vomiting

- Levonorgestrel and UPA ECPs cause less nausea and vomiting than combined estrogen and progestin ECPs.
- Routine use of antiemetics before taking ECPs is not recommended. Pretreatment with antiemetics may be considered depending on availability and clinical judgment.

Vomiting Within 3 Hours of Taking ECPs

- Another dose of ECP should be taken as soon as possible. Use of an antiemetic should be considered.

Comments and Evidence Summary. Many women do not experience nausea or vomiting when taking ECPs, and predicting which women will experience nausea or vomiting is difficult. Although routine use of antiemetics before taking ECPs is not recommended, antiemetics are effective in some women and can be offered when appropriate. Health care providers who are deciding whether to offer antiemetics to women taking ECPs should consider the following: 1) women taking combined estrogen and progestin ECPs are more likely to experience nausea and vomiting than those who take levonorgestrel or UPA ECPs; 2) evidence indicates that antiemetics reduce the occurrence of nausea and vomiting in women taking combined estrogen and progestin ECPs; and 3) women who take antiemetics might experience other side effects from the antiemetics.

A systematic review examined incidence of nausea and vomiting with different ECP regimens and effectiveness of antinausea drugs in reducing nausea and vomiting with ECP use (298). The levonorgestrel regimen was associated with significantly less nausea than a nonstandard dose of UPA (50 mg) and the standard combined estrogen and progestin regimen (299–301). Use of the split-dose levonorgestrel showed no differences in nausea and vomiting compared with the single-dose levonorgestrel (288,290,292,302) (Level of evidence: I, good-fair, indirect). Two trials of antinausea drugs, meclizine and metoclopramide, taken before combined estrogen and progestin ECPs, reduced the severity of nausea (303,304). Significantly less vomiting occurred with meclizine but not metoclopramide (Level of evidence: I, good-fair, direct). No direct evidence was found regarding the effects of vomiting after taking ECPs.

Female Sterilization

Laparoscopic, abdominal, and hysteroscopic methods of female sterilization are available in the United States, and

some of these procedures can be performed in an outpatient procedure or office setting. Fewer than 1 out of 100 women become pregnant in the first year after female sterilization (14). Because these methods are intended to be irreversible, all women should be appropriately counseled about the permanency of sterilization and the availability of highly effective, long-acting, reversible methods of contraception. Female sterilization does not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

When Hysteroscopic Sterilization is Reliable for Contraception

- Before a woman can rely on hysteroscopic sterilization for contraception, a hysterosalpingogram (HSG) must be performed 3 months after the sterilization procedure to confirm bilateral tubal occlusion.
- The woman should be advised that she needs to abstain from sexual intercourse or use additional contraceptive protection until she has confirmed bilateral tubal occlusion.

When Laparoscopic and Abdominal Approaches are Reliable for Contraception

- A woman can rely on sterilization for contraception immediately after laparoscopic and abdominal approaches. No additional contraceptive protection is needed.

Comments and Evidence Summary. HSG confirmation is necessary to confirm bilateral tubal occlusion after hysteroscopic sterilization. The inserts for the hysteroscopic sterilization system available in the United States are placed bilaterally into the fallopian tubes and require 3 months for adequate fibrosis and scarring leading to bilateral tubal occlusion. After hysteroscopic sterilization, advise the woman to correctly and consistently use an effective method of contraception while awaiting confirmation. If compliance with another method might be a problem, a woman and her health care provider may consider DMPA injection at the time of sterilization to ensure adequate contraception for 3 months. Unlike laparoscopic and abdominal sterilizations, pregnancy risk beyond 7 years of follow-up has not been studied among women who received hysteroscopic sterilization.

Pregnancy risk with at least 10 years of follow-up has been studied among women who received laparoscopic and abdominal sterilizations (305,306). Although these methods are highly effective, pregnancies can occur many years after the procedure, and the risk for pregnancy is higher among younger women (306,307).

A systematic review was conducted to identify studies that reported whether pregnancies occurred after hysteroscopic sterilization (308). Twenty-four studies were identified that reported whether pregnancies occurred after hysteroscopic sterilization and found that very few pregnancies occurred among women with confirmed bilateral tubal occlusion; however, few studies include long-term follow-up, and none with follow up for >7 years. Among women who had successful bilateral placement, most pregnancies that occurred after hysteroscopic sterilization were in women who did not have confirmed bilateral tubal occlusion at 3 months, either because of lack of follow up or misinterpretation of HSG results (309–311). Some pregnancies occurred within 3 months of placement, including among women who were already pregnant at the time of the procedure, women who did not use alternative contraception, or women who had failures of alternative contraception (310–315). Although these studies generally demonstrated high rates of bilateral placement, some pregnancies occurred as a result of lack of bilateral placement identified on later imaging (310,311,313–316). Most pregnancies occurred after deviations from FDA directions, which include placement in the early follicular phase of the menstrual cycle, imaging at 3 months to document proper placement, and use of effective alternative contraception until documented occlusion (Level of evidence: II-3, fair, direct).

Male Sterilization

Male sterilization, or vasectomy, is one of the few contraceptive methods available to men and can be performed in an outpatient procedure or office setting. Fewer than 1 woman out of 100 becomes pregnant in the first year after her male partner undergoes sterilization (14). Because male sterilization is intended to be irreversible, all men should be appropriately counseled about the permanency of sterilization and the availability of highly effective, long-acting, reversible methods of contraception for women. Male sterilization does not protect against STDs; consistent and correct use of male latex condoms reduces the risk for STDs, including HIV.

When Vasectomy is Reliable for Contraception

- A semen analysis should be performed 8–16 weeks after a vasectomy to ensure the procedure was successful.
- The man should be advised that he should use additional contraceptive protection or abstain from sexual intercourse until he has confirmation of vasectomy success by postvasectomy semen analysis.

Other Postprocedure Recommendations

- The man should refrain from ejaculation for approximately 1 week after the vasectomy to allow for healing of surgical sites and, after certain methods of vasectomy, occlusion of the vas.

Comments and Evidence Summary. The Vasectomy Guideline Panel of the American Urological Association performed a systematic review of key issues concerning the practice of vasectomy (317). All English-language publications on vasectomy published during 1949–2011 were reviewed. For more information, see the American Urological Association *Vasectomy Guidelines* (<https://www.auanet.org/common/pdf/education/clinical-guidance/Vasectomy.pdf>).

Motile sperm disappear within a few weeks after vasectomy (318–321). The time to azoospermia varies widely in different studies; however, by 12 weeks after the vasectomy, 80% of men have azoospermia, and almost all others have rare nonmotile sperm (defined as $\leq 100,000$ nonmotile sperm per milliliter) (317). The number of ejaculations after vasectomy is not a reliable indicator of when azoospermia or rare nonmotile sperm will be achieved (317). Once azoospermia or rare nonmotile sperm has been achieved, patients can rely on the vasectomy for contraception, although not with 100% certainty. The risk for pregnancy after a man has achieved postvasectomy azoospermia is approximately one in 2,000 (322–326).

A median of 78% (range 33%–100%) of men return for a single postvasectomy semen analysis (317). In the largest cohorts that appear typical of North American vasectomy practice, approximately two thirds of men (55%–71%) return for at least one postvasectomy semen analysis (322,327–331). Assigning men an appointment after their vasectomy might improve compliance with follow-up (332).

When Women Can Stop Using Contraceptives

- Contraceptive protection is still needed for women aged >44 years if the woman wants to avoid pregnancy.

Comments and Evidence Summary. The age at which a woman is no longer at risk for pregnancy is not known. Although uncommon, spontaneous pregnancies occur among women aged >44 years. Both the American College of Obstetricians and Gynecologists and the North American Menopause Society recommend that women continue contraceptive use until menopause or age 50–55 years (333,334). The median age of menopause is approximately 51 years in North America (333)

but can vary from ages 40–60 years (335). The median age of definitive loss of natural fertility is 41 years but can range up to age 51 years (336,337). No reliable laboratory tests are available to confirm definitive loss of fertility in a woman. The assessment of follicle-stimulating hormone levels to determine when a woman is no longer fertile might not be accurate (333).

Health care providers should consider the risks for becoming pregnant in a woman of advanced reproductive age, as well as any risks of continuing contraception until menopause. Pregnancies among women of advanced reproductive age are at higher risk for maternal complications, such as hemorrhage, venous thromboembolism, and death, and fetal complications, such as spontaneous abortion, stillbirth, and congenital anomalies (338–340). Risks associated with continuing contraception, in particular risks for acute cardiovascular events (venous thromboembolism, myocardial infarction, or stroke) or breast cancer, also are important to consider. U.S. MEC states that on the basis of age alone, women aged >45 years can use POPs, implants, the LNG-IUD, or the Cu-IUD (U.S. MEC 1) (5). Women aged >45 years generally can use combined hormonal contraceptives and DMPA (U.S. MEC 2) (5). However, women in this age group might have chronic conditions or other risk factors that might render use of hormonal contraceptive methods unsafe; U.S. MEC might be helpful in guiding the safe use of contraceptives in these women.

In two studies, the incidence of venous thromboembolism was higher among oral contraceptive users aged ≥ 45 years compared with younger oral contraceptive users (341–343); however, an interaction between hormonal contraception and increased age compared with baseline risk was not demonstrated (341,342) or was not examined (343). The relative risk for myocardial infarction was higher among all oral contraceptive users than in nonusers, although a trend of increased relative risk with increasing age was not demonstrated (344,345). No studies were found regarding the risk for stroke in COC users aged ≥ 45 years (Level of evidence: II-2, good to poor, direct).

A pooled analysis by the Collaborative Group on Hormonal Factors and Breast Cancer in 1996 (346) found small increased relative risks for breast cancer among women aged ≥ 45 years whose last use of combined hormonal contraceptives was <5 years previously and for those whose last use was 5–9 years previously. Seven more recent studies suggested small but nonsignificant increased relative risks for breast carcinoma in situ or breast cancer among women who had used oral contraceptives or DMPA when they were aged ≥ 40 years compared with those who had never used either method (347–353) (Level of evidence: II-2, fair, direct).

Conclusion

Most women can start most contraceptive methods at any time, and few examinations or tests, if any, are needed before starting a contraceptive method. Routine follow-up for most women includes assessment of her satisfaction with the contraceptive method, concerns about method use, and changes in health status or medications that could affect medical eligibility for continued use of the method. Because changes in bleeding patterns are one of the major reasons for discontinuation of contraception, recommendations are provided for the management of bleeding irregularities with various contraceptive methods. In addition, because women and health care providers can be confused about the procedures for missed pills and dosing errors with the contraceptive patch and ring, the instructions are streamlined for easier use. ECPs and emergency use of the Cu-IUD are important options for women, and recommendations on using these methods, as well as starting regular contraception after use of emergency contraception, are provided. Male and female sterilization are highly effective methods of contraception for men, women, and couples who have completed childbearing; for men undergoing vasectomy and women undergoing a hysteroscopic sterilization procedure, additional contraceptive protection is needed until the success of the procedure can be confirmed.

CDC is committed to working with partners at the federal, national, and local levels to disseminate, implement, and evaluate U.S. SPR recommendations so that the information reaches health care providers. Strategies for dissemination and implementation include collaborating with other federal agencies and professional and service organizations to widely distribute the recommendations through presentations, electronic distribution, newsletters, and other publications; development of provider tools and job aids to assist providers in implementing the new recommendations; and training activities for students, as well as for continuing education. CDC conducts surveys of family planning health care providers to assess attitudes and practices related to contraceptive use. Results from these surveys will assist CDC in evaluating the impact of these recommendations on the provision of contraceptives in the United States. Finally, CDC will continually monitor new scientific evidence and will update

these recommendations as warranted by new evidence. Updates to the recommendations, as well as provider tools and other resources, are available on the CDC U.S. SPR website: <http://www.cdc.gov/reproductivehealth/UnintendedPregnancy/USSPR.htm>.

Acknowledgments

This report is based, in part, on the work of the Promoting Family Planning Team, Department of Reproductive Health and Research, World Health Organization, and its development of the *Selected Practice Recommendations for Contraceptive Use, 3rd edition*.

U.S. Selected Practice Recommendations for Contraceptive Use Participants

CDC Guideline Development Group for U.S. Medical Eligibility Criteria for Contraceptive Use and U.S. Selected Practice Recommendations for Contraceptive Use

Kathryn M. Curtis, PhD (Chair), Erin Berry-Bibee, MD, Suzanne G. Folger, PhD, Leah G. Horton, MSPH, Denise J. Jamieson, MD, Tara C. Jatlaoui, MD, Polly A. Marchbanks, PhD, H. Pamela Pagano, MPH, Halley E.M. Riley, MPH, Mirelys Rodriguez, Katharine B. Simmons, MD, Naomi K. Tepper, MD, Maura K. Whiteman, PhD, Lauren B. Zapata, PhD, CDC, Atlanta, Georgia.

Invited Meeting Participants

August 27–28, 2014, Atlanta, Georgia

Herbert Peterson, MD, University of North Carolina, Chapel Hill, North Carolina (Chair); Gale Burstein, MD, Erie County Department of Health, Buffalo, New York; Alison Edelman, MD, Oregon Health and Science University, Portland, Oregon; Eve Espey, MD, University of New Mexico, Albuquerque, New Mexico; Emily Godfrey, MD, University of Washington, Seattle, WA; Andrew Kaunitz, MD, University of Florida, Jacksonville, Florida; Susan Moskosky, MS, U.S. Department of Health and Human Services, Rockville, Maryland; Kavita Nanda, MD, FHI360 and American College of Obstetricians and Gynecologists, Durham, NC; Deborah Nucatola, MD, Planned Parenthood Federation of America, New York, New York; Michael Policar, MD, University of California, San Francisco, California; Carolyn Westhoff, MD, Columbia University, New York, New York.

Systematic Review Presenters and Authors

Meeting, August 26–28, 2015, Atlanta, Georgia

Kathryn M. Curtis, PhD, Tara C. Jatlaoui, MD, Polly A. Marchbanks, PhD, Lauren B. Zapata, PhD, CDC, Atlanta, Georgia; Nathalie Kapp, MD, HRA Pharma, Paris, France.

Invited Meeting Participants**August 26–28, 2015, Atlanta, Georgia**

Herbert Peterson, MD, University of North Carolina, Chapel Hill, North Carolina (Chair); Rebecca Allen, MD, American Society for Reproductive Medicine and Women and Infants Hospital, Providence, Rhode Island; Abbey Berenson, MD, University of Texas Medical Branch, Galveston, Texas; Amanda Black, MD, University of Ottawa, Ontario, Canada; Cora Collette Breuner, MD, American Academy of Pediatrics and Seattle Children's Hospital, Seattle, Washington; Gale Burstein, MD, Erie County Department of Health, Buffalo, New York; Alicia Christy, MD, National Institutes of Health, Rockville, Maryland; Mitchell D. Creinin, MD, University of California, Davis, California; Linda Dominguez, Southwest Women's Health, Albuquerque, New Mexico; Alison Edelman, MD, Oregon Health and Science University, Portland, Oregon; Eve Espey, MD, University of New Mexico, Albuquerque, New Mexico; Emily Godfrey, MD, University of Washington, Seattle, WA; Marji Gold, MD, Albert Einstein College of Medicine, Bronx, New York; Robert Hatcher, MD, Emory University, Atlanta, Georgia; Mark Hathaway, MD, JHPIEGO and Unity Healthcare, Washington, DC; Stephen Heartwell, MD, Susan Thompson Buffett Foundation, Omaha, Nebraska; Paula Hillard, MD, Stanford University, Palo Alto, California; Andrew Kaunitz, MD, University of Florida, Jacksonville, Florida; Reagan McDonald-Mosley, MD, Planned Parenthood Federation of America, New York, New York; Susan Moskosky, MS, U.S. Department of Health and Human Services, Rockville, Maryland; Kavita Nanda, MD, FHI360, Durham, North Carolina; Jeffrey Peipert, MD, Washington University, St. Louis, Missouri; Michael Policar, MD, University of California, San Francisco, California; Sarah Prager, MD, University of Washington, Seattle Washington; David Soper, MD, Medical University of South Carolina, Charleston, South Carolina; Lisa Soule, MD, Food and Drug Administration, Rockville, Maryland; James Trussell, PhD, Princeton University, Princeton, New Jersey; Carolyn Westhoff, MD, Columbia University, New York, New York; Christopher Zahn, MD, American College of Obstetricians and Gynecologists, Washington, DC.

CDC Attendees

Wanda D. Barfield, MD, Peter Briss, MD, W. Craig Hooper, PhD, Jill Huppert, MD, Caroline King, PhD, Michele Mandel, Titilope Oduyebo, MD, Cria Perrine, MD, Sam Posner, PhD.

External Reviewers

Claude Burnett, MD, Northeast Health District, Georgia Department of Public Health, Athens, Georgia; Carrie Cwiak, MD, Emory University, Atlanta, Georgia; David Klein, MD, Uniformed Services University, Bethesda, Maryland; Seema Menon, MD, Medical College of Wisconsin, Milwaukee, Wisconsin.

Conflicts of Interest for Invited Meeting Participants**August 26–28, 2015, Atlanta, Georgia**

Rebecca Allen, Nexplanon trainer for Merck and Liletta trainer for Actavis, consultant, advisory board, and education grant from Bayer; Mitchell D. Creinin, Nexplanon trainer for Merck, litigation consultant for Bayer, advisory board for Merck and Teva Pharmaceutical Industries Ltd., consultant for Lemonaid — PolkaDoc app, research support to University of California, Davis from Medicines360, ContraMed, Merck, Eunice Kennedy Shriver National Institute of Child Health and Human Development, and Society of Family Planning; Linda Dominguez, speaker for Bayer, Merck, and Actavis; Alison Edelman, royalties from Up to Date, Inc., consultant for Genzyme, grant support from National Institutes of Health and Gates Foundation, travel funds from World Health Organization, grant support and honorarium from Society of Family Planning, honorarium and travel funds from Contemporary Forum, trainer for Merck, consultant for Gynuity Health Projects, honorarium from CDC, Projects In Knowledge, and American Congress of Obstetricians and Gynecologists, advisory board for Agile Therapeutics; Eve Espey, travel funds from the American Congress of Obstetricians and Gynecologists, Society for Family Planning, and U.S. Food and Drug Administration, Reproductive and Drug Advisory Committee for U.S. Food and Drug Administration, travel funds and honoraria from Wayne State University, Telluride Conference, New Mexico Department of Health Clinician Conference, Planned Parenthood National Medical Conference and Society of Family Planning, British Columbia Contraception Access Research Team Conference, and American Congress of Obstetricians and Gynecologists annual meeting; Emily Godfrey, research funding from Bayer Women's Health, Prima-Temp, and Teva Pharmaceutical Industries Ltd., trainer for Merck and Upstream USA, grant reviewer for Fellowship of Family Planning and Society of Family Planning Research Fund; Mark Hathaway, Liletta trainer and speaker for Actavis and Medicines360, Nexplanon trainer for Merck, advisory board for ContraMed LLC and Afaxys Pharmaceuticals; Paula Hillard consultant for American Civil Liberties Union, Advanced Health Media, CMEology, National Sleep Foundation, and Planned Parenthood Federation of America, honoraria from National Sleep Foundation, Dignity Health, CMEology, Advance Health Media, and Medscape, editorial board for Advanstar — Contemporary OB/GYN, board examiner for the American Board of Obstetrics and Gynecology, contract reviewer for the Department of Health and Human Services, editorial board for EBSCO — PEMSsoft, Nexplanon trainer for Merck, scientific advisor to Proctor and Gamble, publication royalties from Wiley Blackwell Publishing; Nathalie Kapp, employee of HRA Pharma; Andrew Kaunitz, advisory board participant of Allergan, Bayer, Merck, and Pfizer, clinical trial funding to University of Florida from Agile Therapeutics, Bayer, Merck; Jeffrey Peipert, research funding from Bayer and Teva Pharmaceutical Industries Ltd., advisory board for Perrigo; Michael Policar, litigation consultant for Bayer; James Trussell, advisory board for Merck and Teva Pharmaceutical Industries Ltd., consultant for Bayer; Carolyn Westhoff, data and safety monitoring board for Merck and Bayer, advisory board for Agile Therapeutics, MicroChips Biotech, and Actavis, research support to Columbia University from Medicines360, León Farma, and ContraMed.

Handling Conflict of Interest

To promote transparency, all participants were asked to disclose any potential conflicts of interest to CDC prior to the expert meeting and to report any potential conflicts of interest during the introductory portion of the expert meeting. All potential conflicts of interest are listed above. No participants were excluded from discussion based on potential conflicts of interest. One presenter was an employee of a pharmaceutical company and participated by teleconference; after the presentation and questions related to the presentation, the presenter was excused from the discussion. CDC staff who ultimately decided and developed these recommendations have no financial interests or other relationships with the manufacturers of commercial products, suppliers of commercial services, or commercial supporters relevant to these recommendations.

References

- Finer LB, Zolna MR. Declines in unintended pregnancy in the United States, 2008–2011. *N Engl J Med* 2016;374:843–52. <http://dx.doi.org/10.1056/NEJMsa1506575>
- Gipson JD, Koenig MA, Hindin MJ. The effects of unintended pregnancy on infant, child, and parental health: a review of the literature. *Stud Fam Plann* 2008;39:18–38. <http://dx.doi.org/10.1111/j.1728-4465.2008.00148.x>
- Sonfield A, Kost K. Public costs from unintended pregnancies and the role of public insurance programs in paying for pregnancy-related care: national and state estimates for 2010. New York, NY: The Guttmacher Institute; 2015.
- Finer LB, Henshaw SK. Disparities in rates of unintended pregnancy in the United States, 1994 and 2001. *Perspect Sex Reprod Health* 2006;38:90–6. <http://dx.doi.org/10.1363/3809006>
- CDC. U.S. medical eligibility criteria for contraceptive use, 2016. *MMWR Recomm Rep* (No. RR-3);2016.
- CDC. U.S. selected practice recommendations for contraceptive use, 2013: adapted from the World Health Organization selected practice recommendations for contraceptive use, 2nd edition. *MMWR Recomm Rep* 2013;62(No. RR-5).
- Mohllajee AP, Curtis KM, Flanagan RG, Rinehart W, Gaffield ML, Peterson HB. Keeping up with evidence: a new system for WHO's evidence-based family planning guidance. *Am J Prev Med* 2005;28:483–90. <http://dx.doi.org/10.1016/j.amepre.2005.02.008>
- Liberati A, Altman DG, Tetzlaff J, et al. The PRISMA statement for reporting systematic reviews and meta-analyses of studies that evaluate health care interventions: explanation and elaboration. *J Clin Epidemiol* 2009;62:e1–34. <http://dx.doi.org/10.1016/j.jclinepi.2009.06.006>
- Moher D, Liberati A, Tetzlaff J, Altman DG; PRISMA Group. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *BMJ* 2009;339:b2535.
- Harris RP, Helfand M, Woolf SH, et al; Methods Work Group, Third US Preventive Services Task Force. Current methods of the U.S. Preventive Services Task Force: a review of the process. *Am J Prev Med* 2001;20(Suppl):21–35. [http://dx.doi.org/10.1016/S0749-3797\(01\)00261-6](http://dx.doi.org/10.1016/S0749-3797(01)00261-6)
- Horton LG, Folger SG, Berry-Bibee E, Jatlaoui TC, Tepper NK, Curtis KM. Research gaps from evidence-based contraceptives guidance: the U.S. Medical Eligibility Criteria for Contraceptive Use, 2016, and the U.S. Selected Practice Recommendations for Contraceptive Use, 2016. *Contraception*. In press 2016.
- Gavin L, Moskosky S, Carter M, et al. Providing quality family planning services: Recommendations of CDC and the U.S. Office of Population Affairs. *MMWR Recomm Rep* 2014;63(No. RR-4).
- World Health Organization. Selected practice recommendations for contraceptive use. 2nd ed. Geneva, Switzerland: WHO Press; 2004.
- Trussell J. Contraceptive failure in the United States. *Contraception* 2011;83:397–404. <http://dx.doi.org/10.1016/j.contraception.2011.01.021>
- Workowski KA, Bolan GA. Sexually transmitted diseases treatment guidelines, 2015. *MMWR Recomm Rep* 2015;64(No. RR-3).
- Stanback J, Nakintu N, Qureshi Z, Nasution M. Does assessment of signs and symptoms add to the predictive value of an algorithm to rule out pregnancy? *J Fam Plann Reprod Health Care* 2006;32:27–9. <http://dx.doi.org/10.1783/147118906775275370>
- Stanback J, Nanda K, Ramirez Y, Rountree W, Cameron SB. Validation of a job aid to rule out pregnancy among family planning clients in Nicaragua. *Rev Panam Salud Publica* 2008;23:116–8. <http://dx.doi.org/10.1590/S1020-49892008000200007>
- Stanback J, Qureshi Z, Sekadde-Kigundu C, Gonzalez B, Nutley T. Checklist for ruling out pregnancy among family-planning clients in primary care. *Lancet* 1999;354:566. [http://dx.doi.org/10.1016/S0140-6736\(99\)01578-0](http://dx.doi.org/10.1016/S0140-6736(99)01578-0)
- Torpey K, Mwenda L, Kabaso M, et al. Excluding pregnancy among women initiating antiretroviral therapy: efficacy of a family planning job aid. *BMC Public Health* 2010;10:249. <http://dx.doi.org/10.1186/1471-2458-10-249>
- Cole LA, Ladner DG, Byrn FW. The normal variabilities of the menstrual cycle. *Fertil Steril* 2009;91:522–7. <http://dx.doi.org/10.1016/j.fertnstert.2007.11.073>
- Wilcox AJ, Dunson D, Baird DD. The timing of the “fertile window” in the menstrual cycle: day specific estimates from a prospective study. *BMJ* 2000;321:1259–62. <http://dx.doi.org/10.1136/bmj.321.7271.1259>
- Donnet ML, Howie PW, Marnie M, Cooper W, Lewis M. Return of ovarian function following spontaneous abortion. *Clin Endocrinol (Oxf)* 1990;33:13–20. <http://dx.doi.org/10.1111/j.1365-2265.1990.tb00460.x>
- Lähteenmäki P. Postabortal contraception. *Ann Med* 1993;25:185–9. <http://dx.doi.org/10.3109/07853899309164166>
- Stoddard A, Eisenberg DL. Controversies in family planning: timing of ovulation after abortion and the conundrum of postabortion intrauterine device insertion. *Contraception* 2011;84:119–21. <http://dx.doi.org/10.1016/j.contraception.2010.12.010>
- Jackson E, Glasier A. Return of ovulation and menses in postpartum nonlactating women: a systematic review. *Obstet Gynecol* 2011;117:657–62. <http://dx.doi.org/10.1097/AOG.0b013e31820ce18c>
- Kennedy KI, Rivera R, McNeilly AS. Consensus statement on the use of breastfeeding as a family planning method. *Contraception* 1989;39:477–96. [http://dx.doi.org/10.1016/0010-7824\(89\)90103-0](http://dx.doi.org/10.1016/0010-7824(89)90103-0)
- Labbok MH, Perez A, Valdes V, et al. The lactational amenorrhea method (LAM): a postpartum introductory family planning method with policy and program implications. *Adv Contracept* 1994;10:93–109. <http://dx.doi.org/10.1007/BF01978103>
- US Food and Drug Administration. 510(k) premarket notification. Silver Spring, MD; 2015. <https://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfpmn/pmn.cfm>
- Cervinski MA, Gronowski AM. Reproductive-endocrine point-of-care testing: current status and limitations. *Clin Chem Lab Med* 2010;48:935–42. <http://dx.doi.org/10.1515/CCLM.2010.183>
- Cole LA. Human chorionic gonadotropin tests. *Expert Rev Mol Diagn* 2009;9:721–47. <http://dx.doi.org/10.1586/erm.09.51>
- Eichner SF, Timpe EM. Urinary-based ovulation and pregnancy: point-of-care testing. *Ann Pharmacother* 2004;38:325–31. <http://dx.doi.org/10.1345/aph.1D210>
- Wilcox AJ, Baird DD, Dunson D, McChesney R, Weinberg CR. Natural limits of pregnancy testing in relation to the expected menstrual period. *JAMA* 2001;286:1759–61. <http://dx.doi.org/10.1001/jama.286.14.1759>
- Korhonen J, Alfthan H, Ylöstalo P, Veldhuis J, Stenman UH. Disappearance of human chorionic gonadotropin and its alpha- and beta-subunits after term pregnancy. *Clin Chem* 1997;43:2155–63.

34. Reyes FI, Winter JS, Faiman C. Postpartum disappearance of chorionic gonadotropin from the maternal and neonatal circulations. *Am J Obstet Gynecol* 1985;153:486–9. [http://dx.doi.org/10.1016/0002-9378\(85\)90458-2](http://dx.doi.org/10.1016/0002-9378(85)90458-2)
35. Steier JA, Bergsjø P, Myking OL. Human chorionic gonadotropin in maternal plasma after induced abortion, spontaneous abortion, and removed ectopic pregnancy. *Obstet Gynecol* 1984;64:391–4.
36. Bracken MB. Oral contraception and congenital malformations in offspring: a review and meta-analysis of the prospective studies. *Obstet Gynecol* 1990;76:552–7.
37. Gray RH, Pardthaisong T. In utero exposure to steroid contraceptives and survival during infancy. *Am J Epidemiol* 1991;134:804–11.
38. Pardthaisong T, Gray RH. In utero exposure to steroid contraceptives and outcome of pregnancy. *Am J Epidemiol* 1991;134:795–803.
39. Jaffe B, Harlap S, Baras M, et al. Long-term effects of MPA on human progeny: intellectual development. *Contraception* 1988;37:607–19. [http://dx.doi.org/10.1016/0010-7824\(88\)90007-8](http://dx.doi.org/10.1016/0010-7824(88)90007-8)
40. Pardthaisong T, Yencht C, Gray R. The long-term growth and development of children exposed to Depo-Provera during pregnancy or lactation. *Contraception* 1992;45:313–24. [http://dx.doi.org/10.1016/0010-7824\(92\)90053-V](http://dx.doi.org/10.1016/0010-7824(92)90053-V)
41. Brahmi D, Steenland MW, Renner RM, Gaffield ME, Curtis KM. Pregnancy outcomes with an IUD in situ: a systematic review. *Contraception* 2012;85:131–9. <http://dx.doi.org/10.1016/j.contraception.2011.06.010>
42. Tepper NK, Marchbanks PA, Curtis KM. Use of a checklist to rule out pregnancy: a systematic review. *Contraception* 2013;87:661–5. <http://dx.doi.org/10.1016/j.contraception.2012.08.007>
43. Whiteman MK, Tyler CP, Folger SG, Gaffield ME, Curtis KM. When can a woman have an intrauterine device inserted? A systematic review. *Contraception* 2013;87:666–73. <http://dx.doi.org/10.1016/j.contraception.2012.08.015>
44. Teva Women's Health Inc. ParaGard T 380a intrauterine copper contraceptive [Prescribing information]. Sellersville, PA; 2013. <http://paragard.com/Pdf/ParaGard-PI.pdf>
45. Bayer HealthCare Pharmaceuticals Inc. Mirena (levonorgestrel-releasing intrauterine system) [Prescribing information]. Whippany, NJ; 2014. http://labeling.bayerhealthcare.com/html/products/pi/Mirena_PI.pdf
46. Jatlaoui TC, Simmons KB, Curtis KM. The safety of intrauterine contraception initiation among women with current asymptomatic cervical infections or at increased risk of sexually transmitted infections. *Contraception* 2016. Epub June 1, 2016. <http://dx.doi.org/10.1016/j.contraception.2016.05.013>
47. Avonts D, Sercu M, Heyerick P, Vandermeeren I, Meheus A, Piot P. Incidence of uncomplicated genital infections in women using oral contraception or an intrauterine device: a prospective study. *Sex Transm Dis* 1990;17:23–9. <http://dx.doi.org/10.1097/00007435-199017010-00006>
48. Birgisson NE, Zhao Q, Secura GM, Madden T, Peipert JF. Positive testing for *Neisseria gonorrhoeae* and *Chlamydia trachomatis* and the risk of pelvic inflammatory disease in IUD users. *J Womens Health (Larchmt)* 2015;24:354–9. <http://dx.doi.org/10.1089/jwh.2015.5190>
49. Campbell SJ, Cropsey KL, Matthews CA. Intrauterine device use in a high-risk population: experience from an urban university clinic. *Am J Obstet Gynecol* 2007;197: 193.e1–6; discussion 193.e6–7.
50. Cropsey KL, Matthews C, Campbell S, Ivey S, Adawadkar S. Long-term, reversible contraception use among high-risk women treated in a university-based gynecology clinic: comparison between IUD and Depo-Provera. *J Womens Health (Larchmt)* 2010;19:349–53. <http://dx.doi.org/10.1089/jwh.2009.1518>
51. Farley TMM, Rosenberg MJ, Rowe PJ, Chen JH, Meirik O. Intrauterine devices and pelvic inflammatory disease: an international perspective. *Lancet* 1992;339:785–8. [http://dx.doi.org/10.1016/0140-6736\(92\)91904-M](http://dx.doi.org/10.1016/0140-6736(92)91904-M)
52. Grimes DA, Schulz KF. Antibiotic prophylaxis for intrauterine contraceptive device insertion. *Cochrane Database Syst Rev* 2001;(2):CD001327.
53. LeFevre ML; US Preventive Services Task Force. Screening for chlamydia and gonorrhea: U.S. Preventive Services Task Force recommendation statement. *Ann Intern Med* 2014;161:902–10. <http://dx.doi.org/10.7326/M14-1981>
54. Mohllajee AP, Curtis KM, Peterson HB. Does insertion and use of an intrauterine device increase the risk of pelvic inflammatory disease among women with sexually transmitted infection? A systematic review. *Contraception* 2006;73:145–53. <http://dx.doi.org/10.1016/j.contraception.2005.08.007>
55. Morrison CS, Turner AN, Jones LB. Highly effective contraception and acquisition of HIV and other sexually transmitted infections. *Best Pract Res Clin Obstet Gynaecol* 2009;23:263–84. <http://dx.doi.org/10.1016/j.bpobgyn.2008.11.004>
56. Sufrin CB, Postlethwaite D, Armstrong MA, Merchant M, Wendt JM, Steinauer JE. *Neisseria gonorrhoea* and *Chlamydia trachomatis* screening at intrauterine device insertion and pelvic inflammatory disease. *Obstet Gynecol* 2012;120:1314–21.
57. Tepper NK, Steenland MW, Marchbanks PA, Curtis KM. Laboratory screening prior to initiating contraception: a systematic review. *Contraception* 2013;87:645–9. <http://dx.doi.org/10.1016/j.contraception.2012.08.009>
58. Tepper NK, Steenland MW, Marchbanks PA, Curtis KM. Hemoglobin measurement prior to initiating copper intrauterine devices: a systematic review. *Contraception* 2013;87:639–44. <http://dx.doi.org/10.1016/j.contraception.2012.08.008>
59. Rivera R, Almonte H, Arreola M, et al. The effects of three different regimens of oral contraceptives and three different intrauterine devices on the levels of hemoglobin, serum iron and iron binding capacity in anemic women. *Contraception* 1983;27:311–27. [http://dx.doi.org/10.1016/0010-7824\(83\)90009-4](http://dx.doi.org/10.1016/0010-7824(83)90009-4)
60. Ursin G, Ross RK, Sullivan-Halley J, Hanisch R, Henderson B, Bernstein L. Use of oral contraceptives and risk of breast cancer in young women. *Breast Cancer Res Treat* 1998;50:175–84. <http://dx.doi.org/10.1023/A:1006037823178>
61. Hassan EO, el-Husseini M, el-Nahal N. The effect of 1-year use of the CuT 380A and oral contraceptive pills on hemoglobin and ferritin levels. *Contraception* 1999;60:101–5. [http://dx.doi.org/10.1016/S0010-7824\(99\)00065-7](http://dx.doi.org/10.1016/S0010-7824(99)00065-7)
62. Calzolari E, Guglielmo R, Viola F, Migliore L. Hematological parameters and iron therapy in women with IUDs. Experimental study [Italian]. *Minerva Ginecol* 1981;33:355–62.
63. Andersson K, Odland V, Rybo G. Levonorgestrel-releasing and copper-releasing (Nova T) IUDs during five years of use: a randomized comparative trial. *Contraception* 1994;49:56–72. [http://dx.doi.org/10.1016/0010-7824\(94\)90109-0](http://dx.doi.org/10.1016/0010-7824(94)90109-0)
64. Andrade AT, Pizarro E, Shaw STJ Jr, Souza JP, Belsey EM, Rowe PJ; World Health Organization. Consequences of uterine blood loss caused by various intrauterine contraceptive devices in South American women. *Contraception* 1988;38:1–18. [http://dx.doi.org/10.1016/0010-7824\(88\)90091-1](http://dx.doi.org/10.1016/0010-7824(88)90091-1)
65. Blum M, Ariel J, Zacharowitch D. Ferritin, a faithful reflection of iron deficiency in IUD wearers with mild vaginal spotting. *Adv Contracept* 1991;7:39–42. <http://dx.doi.org/10.1007/BF01850717>
66. El-sheikha Z, Hamza A, Mahmoud M. Menstrual blood loss of TCu-380 A and TCu-200 B IUDs. *Popul Sci* 1990;July:55–62.
67. Gallegos AJ, Aznar R, Merino G, Guizer E. Intrauterine devices and menstrual blood loss. A comparative study of eight devices during the first six months of use. *Contraception* 1978;17:153–61. [http://dx.doi.org/10.1016/0010-7824\(78\)90071-9](http://dx.doi.org/10.1016/0010-7824(78)90071-9)
68. Gao J, Zeng S, Sun BL, et al. Menstrual blood loss, haemoglobin and ferritin concentration of Beijing women wearing steel ring, VCu 200, and TCu 220c IUDs. *Contraception* 1986;34:559–71. [http://dx.doi.org/10.1016/S0010-7824\(86\)80012-9](http://dx.doi.org/10.1016/S0010-7824(86)80012-9)

69. Goh TH, Hariharan M. Effect of laparoscopic sterilization and insertion of Multiload Cu 250 and Progestasert IUDs on serum ferritin levels. *Contraception* 1983;28:329–36. [http://dx.doi.org/10.1016/0010-7824\(83\)90034-3](http://dx.doi.org/10.1016/0010-7824(83)90034-3)
70. Goh TH, Hariharan M, Tan CH. A longitudinal study of serum iron indices and haemoglobin concentration following copper-IUD insertion. *Contraception* 1980;22:389–95. [http://dx.doi.org/10.1016/0010-7824\(80\)90024-4](http://dx.doi.org/10.1016/0010-7824(80)90024-4)
71. Guillebaud J, Bonnar J, Morehead J, Matthews A. Menstrual blood-loss with intrauterine devices. *Lancet* 1976;1:387–90. [http://dx.doi.org/10.1016/S0140-6736\(76\)90216-6](http://dx.doi.org/10.1016/S0140-6736(76)90216-6)
72. Haugan T, Skjeldestad FE, Halvorsen LE, Kahn H. A randomized trial on the clinical performance of Nova T380 and Gyne T380 Slimline copper IUDs. *Contraception* 2007;75:171–6. <http://dx.doi.org/10.1016/j.contraception.2006.09.005>
73. Kivijärvi A, Timonen H, Rajamäki A, Grönroos M. Iron deficiency in women using modern copper intrauterine devices. *Obstet Gynecol* 1986;67:95–8.
74. Larsson B, Hamberger L, Rybo G. Influence of copper intrauterine contraceptive devices (Cu-7-IUD) on the menstrual blood-loss. *Acta Obstet Gynecol Scand* 1975;54:315–8. <http://dx.doi.org/10.3109/00016347509156760>
75. Larsson G, Milsom I, Jonasson K, Lindstedt G, Rybo G. The long-term effects of copper surface area on menstrual blood loss and iron status in women fitted with an IUD. *Contraception* 1993;48:471–80. [http://dx.doi.org/10.1016/0010-7824\(93\)90136-U](http://dx.doi.org/10.1016/0010-7824(93)90136-U)
76. Malmqvist R, Petersohn L, Bengtsson LP. Menstrual bleeding with copper-covered intrauterine contraceptive devices. *Contraception* 1974;9:627–33. [http://dx.doi.org/10.1016/0010-7824\(74\)90048-1](http://dx.doi.org/10.1016/0010-7824(74)90048-1)
77. Milsom I, Rybo G, Lindstedt G. The influence of copper surface area on menstrual blood loss and iron status in women fitted with an IUD. *Contraception* 1990;41:271–81. [http://dx.doi.org/10.1016/0010-7824\(90\)90068-7](http://dx.doi.org/10.1016/0010-7824(90)90068-7)
78. Piedras J, Córdova MS, Pérez-Toral MC, Lince E, Garza-Flores J. Predictive value of serum ferritin in anemia development after insertion of T Cu 220 intrauterine device. *Contraception* 1983;27:289–97. [http://dx.doi.org/10.1016/0010-7824\(83\)90007-0](http://dx.doi.org/10.1016/0010-7824(83)90007-0)
79. Sivin I, Alvarez F, Diaz J, et al. Intrauterine contraception with copper and with levonorgestrel: a randomized study of the TCu 380Ag and levonorgestrel 20 mcg/day devices. *Contraception* 1984;30:443–56. [http://dx.doi.org/10.1016/0010-7824\(84\)90036-2](http://dx.doi.org/10.1016/0010-7824(84)90036-2)
80. Sivin I, Stern J, Diaz J, et al. Two years of intrauterine contraception with levonorgestrel and with copper: a randomized comparison of the TCu 380Ag and levonorgestrel 20 mcg/day devices. *Contraception* 1987;35:245–55. [http://dx.doi.org/10.1016/0010-7824\(87\)90026-6](http://dx.doi.org/10.1016/0010-7824(87)90026-6)
81. Tchai BS, Kim SW, Han JH, Im MW. Menstrual blood loss, iron nutrition, and the effects of Alza-T IPCS 52, T-Cu 220C and Lippes Loop D in Korean women. *Seoul J Med* 1987;28:51–9.
82. Wright EA, Kapu MM, Isichei UP. Zinc depletion and menorrhagia in Nigerians using copper T-200 intrauterine device. *Trace Elem Med* 1989;6:147–9.
83. Milsom I, Andersson K, Jonasson K, Lindstedt G, Rybo G. The influence of the Gyne-T 380S IUD on menstrual blood loss and iron status. *Contraception* 1995;52:175–9. [http://dx.doi.org/10.1016/0010-7824\(95\)00163-5](http://dx.doi.org/10.1016/0010-7824(95)00163-5)
84. CDC. Safe motherhood at a glance: Hyattsville, MD: CDC; 2015. <http://www.cdc.gov/chronicdisease/resources/publications/aag/pdf/2015/safe-motherhood-aag-2015.pdf>
85. Cox S, Dietz P, Bowman B, Posner S. Prevalence of chronic conditions among women of reproductive age. Third National Summit on Preconception Health and Health Care: Tampa, FL; 2011.
86. Berenson AB, Rahman M, Wilkinson G. Effect of injectable and oral contraceptives on serum lipids. *Obstet Gynecol* 2009;114:786–94. <http://dx.doi.org/10.1097/AOG.0b013e3181b76bea>
87. Dilbaz B, Ozdegirmenci O, Caliskan E, Dilbaz S, Haberal A. Effect of etonogestrel implant on serum lipids, liver function tests and hemoglobin levels. *Contraception* 2010;81:510–4. <http://dx.doi.org/10.1016/j.contraception.2010.01.014>
88. Nelson AL, Cwiak C. Combined oral contraceptives. In: Hatcher RA, Trussel J, Nelson A, Cates W Jr, Kowal D, Policar M, editors. *Contraceptive technology*. 20th ed. New York, NY: Ardent Media; 2011:249–342.
89. Dragoman M, Curtis KM, Gaffield ME. Combined hormonal contraceptive use among women with known dyslipidemias: a systematic review of critical safety outcomes. *Contraception* 2015;S0010-7824(15)00509-0.
90. US Department of Health and Human Services. Summary health statistics for U.S. adults: National Health Interview Survey, 2012. Report No. 0083–1972. Hyattsville, MD: US Department of Health and Human Services; 2014.
91. CDC. Surveillance for viral hepatitis—United States, 2013. Atlanta, GA: CDC; 2013. <http://www.cdc.gov/hepatitis/statistics/2013surveillance/index.htm>
92. Kohler BA, Sherman RL, Howlader N, et al. Annual report to the nation on the status of cancer, 1975–2011, featuring incidence of breast cancer subtypes by race/ethnicity, poverty, and state. *J Natl Cancer Inst* 2015;107:djv048. <http://dx.doi.org/10.1093/jnci/djv048>
93. Kapp N, Tilley IB, Curtis KM. The effects of hormonal contraceptive use among women with viral hepatitis or cirrhosis of the liver: a systematic review. *Contraception* 2009;80:381–6. <http://dx.doi.org/10.1016/j.contraception.2009.04.007>
94. Kapp N, Curtis KM. Hormonal contraceptive use among women with liver tumors: a systematic review. *Contraception* 2009;80:387–90. <http://dx.doi.org/10.1016/j.contraception.2009.01.021>
95. Tepper NK, Curtis KM, Steenland MW, Marchbanks PA. Physical examination prior to initiating hormonal contraception: a systematic review. *Contraception* 2013;87:650–4. <http://dx.doi.org/10.1016/j.contraception.2012.08.010>
96. National Cancer Institute. SEER Fast Stats. Washington DC: Surveillance Research Program, National Cancer Institute; 2015. <http://seer.cancer.gov/faststats>
97. Watson M, Saraiya M, Benard V, et al. Burden of cervical cancer in the United States, 1998–2003. *Cancer* 2008;113(Suppl):2855–64. <http://dx.doi.org/10.1002/cncr.23756>
98. National Center for Health Statistics, CDC. Health, United States, 2014: with special feature on adults. Hyattsville, MD: CDC; 2015.
99. European Study Group on Heterosexual Transmission of HIV. Comparison of female to male and male to female transmission of HIV in 563 stable couples. *BMJ* 1992;304:809–13. <http://dx.doi.org/10.1136/bmj.304.6830.809>
100. Coleman JS, Mwachari C, Balkus J, et al. Effect of the levonorgestrel intrauterine device on genital HIV-1 RNA shedding among HIV-1-infected women not taking antiretroviral therapy in Nairobi, Kenya. *J Acquir Immune Defic Syndr* 2013;63:245–8. <http://dx.doi.org/10.1097/QAI.0b013e31828decf8>
101. Haddad LB, Cwiak C, Jamieson DJ, et al. Contraceptive adherence among HIV-infected women in Malawi: a randomized controlled trial of the copper intrauterine device and depot medroxyprogesterone acetate. *Contraception* 2013;88:737–43. <http://dx.doi.org/10.1016/j.contraception.2013.08.006>

102. Heikinheimo O, Lehtovirta P, Aho I, Ristola M, Paavonen J. The levonorgestrel-releasing intrauterine system in human immunodeficiency virus-infected women: a 5-year follow-up study. *Am J Obstet Gynecol* 2011;204:126 e1–4.
103. Heikinheimo O, Lehtovirta P, Suni J, Paavonen J. The levonorgestrel-releasing intrauterine system (LNG-IUS) in HIV-infected women—effects on bleeding patterns, ovarian function and genital shedding of HIV. *Hum Reprod* 2006;21:2857–61. <http://dx.doi.org/10.1093/humrep/del264>
104. Kakaire O, Byamugisha JK, Tumwesigye NM, Gemzell-Danielsson K. Clinical versus laboratory screening for sexually transmitted infections prior to insertion of intrauterine contraception among women living with HIV/AIDS: a randomized controlled trial. *Hum Reprod* 2015;30:1573–9. <http://dx.doi.org/10.1093/humrep/dev109>
105. Kovacs A, Wasserman SS, Burns D, et al; DATRI Study Group; WIHS Study Group. Determinants of HIV-1 shedding in the genital tract of women. *Lancet* 2001;358:1593–601. [http://dx.doi.org/10.1016/S0140-6736\(01\)06653-3](http://dx.doi.org/10.1016/S0140-6736(01)06653-3)
106. Landolt NK, Phanuphak N, Teeratakulpisarn N, et al. Uptake and continuous use of copper intrauterine device in a cohort of HIV-positive women. *AIDS Care* 2013;25:710–4. <http://dx.doi.org/10.1080/09540121.2012.752786>
107. Lehtovirta P, Paavonen J, Heikinheimo O. Experience with the levonorgestrel-releasing intrauterine system among HIV-infected women. *Contraception* 2007;75:37–9. <http://dx.doi.org/10.1016/j.contraception.2006.09.006>
108. Morrison CS, Sekadde-Kigonde C, Sinei SK, Weiner DH, Kwok C, Kokonya D. Is the intrauterine device appropriate contraception for HIV-1-infected women? *BJOG* 2001;108:784–90. <http://dx.doi.org/10.1111/j.1471-0528.2001.00204.x>
109. Mostad SB, Overbaugh J, DeVange DM, et al. Hormonal contraception, vitamin A deficiency, and other risk factors for shedding of HIV-1 infected cells from the cervix and vagina. *Lancet* 1997;350:922–7. [http://dx.doi.org/10.1016/S0140-6736\(97\)04240-2](http://dx.doi.org/10.1016/S0140-6736(97)04240-2)
110. Richardson BA, Morrison CS, Sekadde-Kigonde C, et al. Effect of intrauterine device use on cervical shedding of HIV-1 DNA. *AIDS* 1999;13:2091–7. <http://dx.doi.org/10.1097/00002030-199910220-00012>
111. Sinei SK, Morrison CS, Sekadde-Kigonde C, Allen M, Kokonya D. Complications of use of intrauterine devices among HIV-1-infected women. *Lancet* 1998;351:1238–41. [http://dx.doi.org/10.1016/S0140-6736\(97\)10319-1](http://dx.doi.org/10.1016/S0140-6736(97)10319-1)
112. Stringer EM, Kaseba C, Levy J, Sinkala M, Goldenberg RL, Chi BH, et al. A randomized trial of the intrauterine contraceptive device vs hormonal contraception in women who are infected with the human immunodeficiency virus. *Am J Obstet Gynecol* 2007;197:144 e1–8.
113. Stringer EM, Levy J, Sinkala M, et al. HIV disease progression by hormonal contraceptive method: secondary analysis of a randomized trial. *AIDS* 2009;23:1377–82. <http://dx.doi.org/10.1097/QAD.0b013e32832cbca8>
114. Tepper NK, Curtis KM, Nanda K, Jamieson DJ. Safety of intrauterine devices among women with HIV: a systematic review. *Contraception* 2016. Epub June 22, 2016. <http://dx.doi.org/10.1016/j.contraception.2016.06.011>
115. Lopez LM, Bernholc A, Zeng Y, et al. Interventions for pain with intrauterine device insertion. *Cochrane Database Syst Rev* 2015;7:CD007373.
116. Zapata LB, Jataoui TC, Marchbanks PA, Curtis KM. Medications to ease intrauterine device insertion: a systematic review. *Contraception* 2016. Epub June 29, 2016. <http://dx.doi.org/10.1016/j.contraception.2016.06.014>
117. Bahamondes MV, Espejo-Arce X, Bahamondes L. Effect of vaginal administration of misoprostol before intrauterine contraceptive insertion following previous insertion failure: a double blind RCT. *Hum Reprod* 2015;30:1861–6. <http://dx.doi.org/10.1093/humrep/dev137>
118. Cirik D, Taskin E, Tuglu A, Ortac A, Dai O. Paracervical block with 1% lidocaine for pain control during intrauterine device insertion: a prospective, single-blinded, controlled study. *Int J Reprod Contracept Obstet Gynecol* 2013;2:263–7.
119. Mody SK, Kiley J, Rademaker A, Gawron L, Stika C, Hammond C. Pain control for intrauterine device insertion: a randomized trial of 1% lidocaine paracervical block. *Contraception* 2012;86:704–9. <http://dx.doi.org/10.1016/j.contraception.2012.06.004>
120. Grimes DA, Schulz KF. Prophylactic antibiotics for intrauterine device insertion: a metaanalysis of the randomized controlled trials. *Contraception* 1999;60:57–63. [http://dx.doi.org/10.1016/S0010-7824\(99\)00071-2](http://dx.doi.org/10.1016/S0010-7824(99)00071-2)
121. Wilson W, Taubert KA, Gewitz M, et al; American Heart Association Rheumatic Fever, Endocarditis, and Kawasaki Disease Committee; American Heart Association Council on Cardiovascular Disease in the Young; American Heart Association Council on Clinical Cardiology; American Heart Association Council on Cardiovascular Surgery and Anesthesia; Quality of Care and Outcomes Research Interdisciplinary Working Group. Prevention of infective endocarditis: guidelines from the American Heart Association: a guideline from the American Heart Association Rheumatic Fever, Endocarditis, and Kawasaki Disease Committee, Council on Cardiovascular Disease in the Young, and the Council on Clinical Cardiology, Council on Cardiovascular Surgery and Anesthesia, and the Quality of Care and Outcomes Research Interdisciplinary Working Group. *Circulation* 2007;116:1736–54. <http://dx.doi.org/10.1161/CIRCULATIONAHA.106.183095>
122. Steenland MW, Zapata LB, Brahma D, Marchbanks PA, Curtis KM. The effect of follow-up visits or contacts after contraceptive initiation on method continuation and correct use. *Contraception* 2013;87:625–30. <http://dx.doi.org/10.1016/j.contraception.2012.09.018>
123. Steenland MW, Zapata LB, Brahma D, Marchbanks PA, Curtis KM. Appropriate follow up to detect potential adverse events after initiation of select contraceptive methods: a systematic review. *Contraception* 2013;87:611–24. <http://dx.doi.org/10.1016/j.contraception.2012.09.017>
124. Canto De Cetina TE, Canto P, Ordoñez Luna M. Effect of counseling to improve compliance in Mexican women receiving depot-medroxyprogesterone acetate. *Contraception* 2001;63:143–6. [http://dx.doi.org/10.1016/S0010-7824\(01\)00181-0](http://dx.doi.org/10.1016/S0010-7824(01)00181-0)
125. Lei ZW, Wu SC, Garceau RJ, et al. Effect of pretreatment counseling on discontinuation rates in Chinese women given depot-medroxyprogesterone acetate for contraception. *Contraception* 1996;53:357–61. [http://dx.doi.org/10.1016/0010-7824\(96\)00085-6](http://dx.doi.org/10.1016/0010-7824(96)00085-6)
126. Godfrey EM, Folger SG, Jeng G, Jamieson DJ, Curtis KM. Treatment of bleeding irregularities in women with copper-containing IUDs: a systematic review. *Contraception* 2013;87:549–66. <http://dx.doi.org/10.1016/j.contraception.2012.09.006>
127. Topozada M, Anwar M, Abdel Rahman H, Gaweesh S. Control of IUD-induced bleeding by three non-steroidal anti-inflammatory drugs. *Contracept Deliv Syst* 1982;3:117–25.
128. Topozada M, El-Attar A, El-Ayyat MA, Khamis Y. Management of uterine bleeding by PGs or their synthesis inhibitors. *Adv Prostaglandin Thromboxane Res* 1980;8:1459–63.

129. Wu S, Wang C, Cheng W, et al. Randomized multi-center study of baofuxin for treatment of bleeding side-effect induced by IUD. *Reprod Contracept* 2000;11:152–7.
130. Mercorio F, De Simone R, Di Carlo C, et al. Effectiveness and mechanism of action of desmopressin in the treatment of copper intrauterine device-related menorrhagia: a pilot study. *Hum Reprod* 2003;18:2319–22. <http://dx.doi.org/10.1093/humrep/deg449>
131. Pedron N, Lozano M, Aznar R. Treatment of hypermenorrhea with mefenamic acid in women using IUDs. *Contracept Deliv Syst* 1982;3:135–9.
132. Pizarro E, Mehech G, Hidalgo M, Muñoz G, Romero C. Effect of meclofenamic acid on menstruation in hypermenorrhagic women using intrauterine devices [Spanish]. *Rev Chil Obstet Ginecol* 1988;53:43–56.
133. Chinese National IUD Research Working Group. Prevention and treatment of IUD-induced menorrhagia with antifibrinolytic and antiprostaglandin drugs [Chinese]. *Zhonghua Fu Chan Ke Za Zhi* 1987;22:291–4, 312.
134. Di Lieto A, Catalano D, Miranda L, Paladini A. Action of a prostaglandin synthetase inhibitor on IUD associated uterine bleeding. *Clin Exp Obstet Gynecol* 1987;14:41–4.
135. Ylikorkala O, Viinikka L. Comparison between antifibrinolytic and antiprostaglandin treatment in the reduction of increased menstrual blood loss in women with intrauterine contraceptive devices. *Br J Obstet Gynaecol* 1983;90:78–83. <http://dx.doi.org/10.1111/j.1471-0528.1983.tb06751.x>
136. US Food and Drug Administration. *Cyklokapron: tranexamic acid injection [Prescribing information]*. New York, NY; 2011.
137. US Food and Drug Administration. *Lysteda: tranexamic acid tablets [Prescribing information]*. Parsippany, NJ. 2011.
138. Pedrón N, Lozano M, Gallegos AJ. The effect of acetylsalicylic acid on menstrual blood loss in women with IUDs. *Contraception* 1987;36:295–303. [http://dx.doi.org/10.1016/0010-7824\(87\)90099-0](http://dx.doi.org/10.1016/0010-7824(87)90099-0)
139. Hidalgo M, Bahamondes L, Perrotti M, Diaz J, Dantas-Monteiro C, Petta C. Bleeding patterns and clinical performance of the levonorgestrel-releasing intrauterine system (Mirena) up to two years. *Contraception* 2002;65:129–32. [http://dx.doi.org/10.1016/S0010-7824\(01\)00302-X](http://dx.doi.org/10.1016/S0010-7824(01)00302-X)
140. Tepper NK, Steenland MW, Gaffield ME, Marchbanks PA, Curtis KM. Retention of intrauterine devices in women who acquire pelvic inflammatory disease: a systematic review. *Contraception* 2013;87:655–60. <http://dx.doi.org/10.1016/j.contraception.2012.08.011>
141. Larsson B, Wennergren M. Investigation of a copper-intrauterine device (Cu-IUD) for possible effect on frequency and healing of pelvic inflammatory disease. *Contraception* 1977;15:143–9. [http://dx.doi.org/10.1016/0010-7824\(77\)90012-9](http://dx.doi.org/10.1016/0010-7824(77)90012-9)
142. Söderberg G, Lindgren S. Influence of an intrauterine device on the course of an acute salpingitis. *Contraception* 1981;24:137–43. [http://dx.doi.org/10.1016/0010-7824\(81\)90086-X](http://dx.doi.org/10.1016/0010-7824(81)90086-X)
143. Teisala K. Removal of an intrauterine device and the treatment of acute pelvic inflammatory disease. *Ann Med* 1989;21:63–5. <http://dx.doi.org/10.3109/07853898909149184>
144. Altunyurt S, Demir N, Posaci C. A randomized controlled trial of coil removal prior to treatment of pelvic inflammatory disease. *Eur J Obstet Gynecol Reprod Biol* 2003;107:81–4. [http://dx.doi.org/10.1016/S0301-2115\(02\)00342-1](http://dx.doi.org/10.1016/S0301-2115(02)00342-1)
145. Stewart FH, Harper CC, Ellertson CE, Grimes DA, Sawaya GF, Trussell J. Clinical breast and pelvic examination requirements for hormonal contraception: current practice vs evidence. *JAMA* 2001;285:2232–9. <http://dx.doi.org/10.1001/jama.285.17.2232>
146. Mansour D, Korver T, Marintcheva-Petrova M, Fraser IS. The effects of Implanon on menstrual bleeding patterns. *Eur J Contracept Reprod Health Care* 2008;13(Suppl 1):13–28. <http://dx.doi.org/10.1080/13625180801959931>
147. Abdel-Aleem H, d’Arcangues C, Vogelsong K, Gülmezoglu AM. Treatment of vaginal bleeding irregularities induced by progestin only contraceptives. *Cochrane Database Syst Rev* 2007;(2):CD003449.
148. Archer DF, Philput CB, Levine AS, et al. Effects of ethinyl estradiol and ibuprofen compared to placebo on endometrial bleeding, cervical mucus and the postcoital test in levonorgestrel subcutaneous implant users. *Contraception* 2008;78:106–12. <http://dx.doi.org/10.1016/j.contraception.2008.04.003>
149. Buasang K, Taneepanichskul S. Efficacy of celecoxib on controlling irregular uterine bleeding secondary to Jadelle use. *J Med Assoc Thai* 2009;92:301–7.
150. Phaliwong P, Taneepanichskul S. The effect of mefenamic acid on controlling irregular uterine bleeding second to Implanon use. *J Med Assoc Thai* 2004;87(Suppl 3):S64–8.
151. Weisberg E, Hickey M, Palmer D, et al. A randomized controlled trial of treatment options for troublesome uterine bleeding in Implanon users. *Hum Reprod* 2009;24:1852–61. <http://dx.doi.org/10.1093/humrep/dep081>
152. Weisberg E, Hickey M, Palmer D, et al. A pilot study to assess the effect of three short-term treatments on frequent and/or prolonged bleeding compared to placebo in women using Implanon. *Hum Reprod* 2006;21:295–302. <http://dx.doi.org/10.1093/humrep/dei273>
153. Cheng L, Zhu H, Wang A, Ren F, Chen J, Glasier A. Once a month administration of mifepristone improves bleeding patterns in women using subdermal contraceptive implants releasing levonorgestrel. *Hum Reprod* 2000;15:1969–72. <http://dx.doi.org/10.1093/humrep/15.9.1969>
154. Alvarez-Sanchez F, Brache V, Thevenin F, Cochon L, Faundes A. Hormonal treatment for bleeding irregularities in Norplant implant users. *Am J Obstet Gynecol* 1996;174:919–22. [http://dx.doi.org/10.1016/S0002-9378\(96\)70326-5](http://dx.doi.org/10.1016/S0002-9378(96)70326-5)
155. Díaz S, Croxatto HB, Pavez M, Belhadj H, Stern J, Sivin I. Clinical assessment of treatments for prolonged bleeding in users of Norplant implants. *Contraception* 1990;42:97–109. [http://dx.doi.org/10.1016/0010-7824\(90\)90094-C](http://dx.doi.org/10.1016/0010-7824(90)90094-C)
156. Witjaksono J, Lau TM, Affandi B, Rogers PA. Oestrogen treatment for increased bleeding in Norplant users: preliminary results. *Hum Reprod* 1996;11(Suppl 2):109–14. http://dx.doi.org/10.1093/humrep/11.suppl_2.109
157. Abdel-Aleem H, Shaaban OM, Amin AF, Abdel-Aleem AM. Tamoxifen treatment of bleeding irregularities associated with Norplant use. *Contraception* 2005;72:432–7. <http://dx.doi.org/10.1016/j.contraception.2005.05.015>
158. Phupong V, Sophonsritsuk A, Taneepanichskul S. The effect of tranexamic acid for treatment of irregular uterine bleeding secondary to Norplant use. *Contraception* 2006;73:253–6. <http://dx.doi.org/10.1016/j.contraception.2005.09.012>
159. d’Arcangues C, Piaggio G, Brache V, et al; Study Group on Progestogen-induced Vaginal Bleeding Disturbances. Effectiveness and acceptability of vitamin E and low-dose aspirin, alone or in combination, on Norplant-induced prolonged bleeding. *Contraception* 2004;70:451–62. <http://dx.doi.org/10.1016/j.contraception.2004.05.012>

160. Subakir SB, Setiadi E, Affandi B, Pringgoutomo S, Freisleben HJ. Benefits of vitamin E supplementation to Norplant users—in vitro and in vivo studies. *Toxicology* 2000;148:173–8. [http://dx.doi.org/10.1016/S0300-483X\(00\)00208-0](http://dx.doi.org/10.1016/S0300-483X(00)00208-0)
161. Kapp N, Gaffield ME. Initiation of progestogen-only injectables on different days of the menstrual cycle and its effect on contraceptive effectiveness and compliance: a systematic review. *Contraception* 2013;87:576–82. <http://dx.doi.org/10.1016/j.contraception.2012.08.017>
162. Petta CA, Faúndes A, Dunson TR, et al. Timing of onset of contraceptive effectiveness in Depo-Provera users. II. Effects on ovarian function. *Fertil Steril* 1998;70:817–20. [http://dx.doi.org/10.1016/S0015-0282\(98\)00309-4](http://dx.doi.org/10.1016/S0015-0282(98)00309-4)
163. Petta CA, Faundes A, Dunson TR, et al. Timing of onset of contraceptive effectiveness in Depo-Provera users: Part I. Changes in cervical mucus. *Fertil Steril* 1998;69:252–7. [http://dx.doi.org/10.1016/S0015-0282\(97\)00477-9](http://dx.doi.org/10.1016/S0015-0282(97)00477-9)
164. Siritwongse T, Snidvongs W, Tantayaporn P, Leepipatpaiboon S. Effect of depo-medroxyprogesterone acetate on serum progesterone levels when administered on various cycle days. *Contraception* 1982;26:487–93. [http://dx.doi.org/10.1016/0010-7824\(82\)90147-0](http://dx.doi.org/10.1016/0010-7824(82)90147-0)
165. Balkus J, Miller L. Same-day administration of depot-medroxyprogesterone acetate injection: a retrospective chart review. *Contraception* 2005;71:395–8. <http://dx.doi.org/10.1016/j.contraception.2004.10.014>
166. Morroni C, Grams M, Tiezzi L, Westhoff C. Immediate monthly combination contraception to facilitate initiation of the depot medroxyprogesterone acetate contraceptive injection. *Contraception* 2004;70:19–23. <http://dx.doi.org/10.1016/j.contraception.2004.02.007>
167. Nelson AL, Katz T. Initiation and continuation rates seen in 2-year experience with Same Day injections of DMPA. *Contraception* 2007;75:84–7. <http://dx.doi.org/10.1016/j.contraception.2006.09.007>
168. Rickert VI, Tiezzi L, Lipshutz J, León J, Vaughan RD, Westhoff C. Depo Now: preventing unintended pregnancies among adolescents and young adults. *J Adolesc Health* 2007;40:22–8. <http://dx.doi.org/10.1016/j.jadohealth.2006.10.018>
169. Sneed R, Westhoff C, Morroni C, Tiezzi L. A prospective study of immediate initiation of depo medroxyprogesterone acetate contraceptive injection. *Contraception* 2005;71:99–103. <http://dx.doi.org/10.1016/j.contraception.2004.08.014>
170. Tepper NK, Curtis KM, Steenland MW, Marchbanks PA. Blood pressure measurement prior to initiating hormonal contraception: a systematic review. *Contraception* 2013;87:631–8. <http://dx.doi.org/10.1016/j.contraception.2012.08.025>
171. Diab KM, Zaki MM. Contraception in diabetic women: comparative metabolic study of Norplant, depot medroxyprogesterone acetate, low dose oral contraceptive pill and CuT380A. *J Obstet Gynaecol Res* 2000;26:17–26. <http://dx.doi.org/10.1111/j.1447-0756.2000.tb01195.x>
172. Garg SK, Chase HP, Marshall G, Hoops SL, Holmes DL, Jackson WE. Oral contraceptives and renal and retinal complications in young women with insulin-dependent diabetes mellitus. *JAMA* 1994;271:1099–102. <http://dx.doi.org/10.1001/jama.1994.03510380055037>
173. Grigoryan OR, Grodnitskaya EE, Andreeva EN, Chebotnikova TV, Melnichenko GA. Use of the NuvaRing hormone-releasing system in late reproductive-age women with type 1 diabetes mellitus. *Gynecol Endocrinol* 2008;24:99–104. <http://dx.doi.org/10.1080/09513590701708795>
174. Kahn HS, Curtis KM, Marchbanks PA. Effects of injectable or implantable progestin-only contraceptives on insulin-glucose metabolism and diabetes risk. *Diabetes Care* 2003;26:216–25. <http://dx.doi.org/10.2337/diacare.26.1.216>
175. Lopez LM, Grimes DA, Schulz KF. Steroidal contraceptives: effect on carbohydrate metabolism in women without diabetes mellitus. *Cochrane Database Syst Rev* 2009; (4):CD006133.
176. Rogovskaya S, Rivera R, Grimes DA, et al. Effect of a levonorgestrel intrauterine system on women with type 1 diabetes: a randomized trial. *Obstet Gynecol* 2005;105:811–5. <http://dx.doi.org/10.1097/01.AOG.0000156301.11939.56>
177. Troisi RJ, Cowie CC, Harris MI. Oral contraceptive use and glucose metabolism in a national sample of women in the United States. *Am J Obstet Gynecol* 2000;183:389–95. <http://dx.doi.org/10.1067/mob.2000.105909>
178. Bonny AE, Secic M, Cromer B. Early weight gain related to later weight gain in adolescents on depot medroxyprogesterone acetate. *Obstet Gynecol* 2011;117:793–7. <http://dx.doi.org/10.1097/AOG.0b013e31820f387c>
179. Le YC, Rahman M, Berenson AB. Early weight gain predicting later weight gain among depot medroxyprogesterone acetate users. *Obstet Gynecol* 2009;114:279–84. <http://dx.doi.org/10.1097/AOG.0b013e3181af68b2>
180. Risser WL, Gefter LR, Barratt MS, Risser JM. Weight change in adolescents who used hormonal contraception. *J Adolesc Health* 1999;24:433–6. [http://dx.doi.org/10.1016/S1054-139X\(98\)00151-7](http://dx.doi.org/10.1016/S1054-139X(98)00151-7)
181. Paulen ME, Curtis KM. When can a woman have repeat progestogen-only injectables—depot medroxyprogesterone acetate or norethisterone enantate? *Contraception* 2009;80:391–408. <http://dx.doi.org/10.1016/j.contraception.2009.03.023>
182. Pardthaisong T. Return of fertility after use of the injectable contraceptive Depo Provera: up-dated data analysis. *J Biosoc Sci* 1984;16:23–34. <http://dx.doi.org/10.1017/S0021932000014760>
183. Schwallie PC, Assenzo JR. The effect of depo-medroxyprogesterone acetate on pituitary and ovarian function, and the return of fertility following its discontinuation: a review. *Contraception* 1974;10:181–202. [http://dx.doi.org/10.1016/0010-7824\(74\)90073-0](http://dx.doi.org/10.1016/0010-7824(74)90073-0)
184. Steiner MJ, Kwok C, Stanback J, et al. Injectable contraception: what should the longest interval be for reinjections? *Contraception* 2008;77:410–4. <http://dx.doi.org/10.1016/j.contraception.2008.01.017>
185. Bassol S, Garza-Flores J, Cravioto MC, et al. Ovarian function following a single administration of depo-medroxyprogesterone acetate (DMPA) at different doses. *Fertil Steril* 1984;42:216–22.
186. Fotherby K, Koetsawang S, Mathrubutham M. Pharmacokinetic study of different doses of Depo Provera. *Contraception* 1980;22:527–36. [http://dx.doi.org/10.1016/0010-7824\(80\)90105-5](http://dx.doi.org/10.1016/0010-7824(80)90105-5)
187. Fotherby K, Saxena BN, Shrimanker K, et al. A preliminary pharmacokinetic and pharmacodynamic evaluation of depot-medroxyprogesterone acetate and norethisterone oenanthate. *Fertil Steril* 1980;34:131–9.
188. Garza-Flores J, Cardenas S, Rodríguez V, Cravioto MC, Diaz-Sanchez V, Perez-Palacios G. Return to ovulation following the use of long-acting injectable contraceptives: a comparative study. *Contraception* 1985;31:361–6. [http://dx.doi.org/10.1016/0010-7824\(85\)90004-6](http://dx.doi.org/10.1016/0010-7824(85)90004-6)
189. Jain J, Dutton C, Nicosia A, Wajszczuk C, Bode FR, Mishell DR Jr. Pharmacokinetics, ovulation suppression and return to ovulation following a lower dose subcutaneous formulation of Depo-Provera. *Contraception* 2004;70:11–8. <http://dx.doi.org/10.1016/j.contraception.2004.01.011>
190. Lan PT, Aedo AR, Landgren BM, Johannisson E, Diczfalusy E. Return of ovulation following a single injection of depo-medroxyprogesterone acetate: a pharmacokinetic and pharmacodynamic study. *Contraception* 1984;29:1–18. [http://dx.doi.org/10.1016/0010-7824\(84\)90054-4](http://dx.doi.org/10.1016/0010-7824(84)90054-4)

191. Ortiz A, Hirol M, Stanczyk FZ, Goebelsmann U, Mishell DR Jr. Serum medroxyprogesterone acetate (MPA) concentrations and ovarian function following intramuscular injection of depo-MPA. *J Clin Endocrinol Metab* 1977;44:32–8. <http://dx.doi.org/10.1210/jcem-44-1-32>
192. Saxena BN, Dusitsin N, Tankeyoon M, Chaudhury RR. Return of ovulation after the cessation of depot-medroxy progesterone acetate treatment in Thai women. *J Med Assoc Thai* 1980;63:66–9.
193. Toh YC, Jain J, Rahny MH, Bode FR, Ross D. Suppression of ovulation by a new subcutaneous depot medroxyprogesterone acetate (104 mg/0.65 mL) contraceptive formulation in Asian women. *Clin Ther* 2004;26:1845–54. <http://dx.doi.org/10.1016/j.clinthera.2004.11.013>
194. Arias RD, Jain JK, Brucker C, Ross D, Ray A. Changes in bleeding patterns with depot medroxyprogesterone acetate subcutaneous injection 104 mg. *Contraception* 2006;74:234–8. <http://dx.doi.org/10.1016/j.contraception.2006.03.008>
195. Dempsey A, Roca C, Westhoff C. Vaginal estrogen supplementation during Depo-Provera initiation: a randomized controlled trial. *Contraception* 2010;82:250–5. <http://dx.doi.org/10.1016/j.contraception.2010.04.003>
196. Abdel-Aleem H, d’Arcangues C, Vogelzang KM, Gülmezoglu AM. Treatment of vaginal bleeding irregularities induced by progestin only contraceptives. *Cochrane Database Syst Rev* 2007; (4):CD003449.
197. Harel Z, Biro F, Kollar L, Riggs S, Flanagan P, Vaz R. Supplementation with vitamin C and/or vitamin B(6) in the prevention of Depo-Provera side effects in adolescents. *J Pediatr Adolesc Gynecol* 2002;15:153–8. [http://dx.doi.org/10.1016/S1083-3188\(02\)00148-1](http://dx.doi.org/10.1016/S1083-3188(02)00148-1)
198. Nathirojanakun P, Taneepanichskul S, Sappakitkumjorn N. Efficacy of a selective COX-2 inhibitor for controlling irregular uterine bleeding in DMPA users. *Contraception* 2006;73:584–7. <http://dx.doi.org/10.1016/j.contraception.2005.09.013>
199. Tantiwattanakul P, Taneepanichskul S. Effect of mefenamic acid on controlling irregular uterine bleeding in DMPA users. *Contraception* 2004;70:277–9. <http://dx.doi.org/10.1016/j.contraception.2004.04.003>
200. Said S, Sadek W, Rocca M, et al. Clinical evaluation of the therapeutic effectiveness of ethinyl oestradiol and oestrone sulphate on prolonged bleeding in women using depot medroxyprogesterone acetate for contraception. World Health Organization, Special Programme of Research, Development and Research Training in Human Reproduction, Task Force on Long-acting Systemic Agents for Fertility Regulation. *Hum Reprod* 1996;11(Suppl 2):1–13. http://dx.doi.org/10.1093/humrep/11.suppl_2.1
201. Sadeghi-Bazargani H, Ehdacivand F, Arshi S, Eftekhar H, Sezavar H, Amanati L. Low-dose oral contraceptive to re-induce menstrual bleeding in amenorrheic women on DMPA treatment: a randomized clinical trial. *Med Sci Monit* 2006;12:CR420–5.
202. Brahmi D, Curtis KM. When can a woman start combined hormonal contraceptives (CHCs)? A systematic review. *Contraception* 2013;87:524–38. <http://dx.doi.org/10.1016/j.contraception.2012.09.010>
203. Edwards SM, Ziemann M, Jones K, Diaz A, Robilotto C, Westhoff C. Initiation of oral contraceptives—start now! *J Adolesc Health* 2008;43:432–6. <http://dx.doi.org/10.1016/j.jadohealth.2008.06.008>
204. Westhoff C, Heartwell S, Edwards S, et al. Initiation of oral contraceptives using a quick start compared with a conventional start: a randomized controlled trial. *Obstet Gynecol* 2007;109:1270–6. <http://dx.doi.org/10.1097/01.AOG.0000264550.41242.f2>
205. Westhoff C, Morroni C, Kerns J, Murphy PA. Bleeding patterns after immediate vs. conventional oral contraceptive initiation: a randomized, controlled trial. *Fertil Steril* 2003;79:322–9. [http://dx.doi.org/10.1016/S0015-0282\(02\)04680-0](http://dx.doi.org/10.1016/S0015-0282(02)04680-0)
206. Baerwald AR, Olatunbosun OA, Pierson RA. Ovarian follicular development is initiated during the hormone-free interval of oral contraceptive use. *Contraception* 2004;70:371–7. <http://dx.doi.org/10.1016/j.contraception.2004.05.006>
207. Baerwald AR, Pierson RA. Ovarian follicular development during the use of oral contraception: a review. *J Obstet Gynaecol Can* 2004;26:19–24. [http://dx.doi.org/10.1016/S1701-2163\(16\)30692-2](http://dx.doi.org/10.1016/S1701-2163(16)30692-2)
208. Duijkers IJM, Klipping C, Verhoeven CHJ, Dieben TOM. Ovarian function with the contraceptive vaginal ring or an oral contraceptive: a randomized study. *Hum Reprod* 2004;19:2668–73. <http://dx.doi.org/10.1093/humrep/deh493>
209. Killick S, Eyoung E, Elstein M. Ovarian follicular development in oral contraceptive cycles. *Fertil Steril* 1987;48:409–13.
210. Molloy BG, Coulson KA, Lee JM, Watters JK. “Missed pill” conception: fact or fiction? *Br Med J (Clin Res Ed)* 1985;290:1474–5. <http://dx.doi.org/10.1136/bmj.290.6480.1474-a>
211. Mulders TM, Dieben TO, Bennink HJ. Ovarian function with a novel combined contraceptive vaginal ring. *Hum Reprod* 2002;17:2594–9. <http://dx.doi.org/10.1093/humrep/17.10.2594>
212. Schwartz JL, Creinin MD, Pymar HC, Reid L. Predicting risk of ovulation in new start oral contraceptive users. *Obstet Gynecol* 2002;99:177–82.
213. Sitavarin S, Jaisamrarn U, Taneepanichskul S. A randomized trial on the impact of starting day on ovarian follicular activity in very low dose oral contraceptive pills users. *J Med Assoc Thai* 2003;86:442–8.
214. Taylor DR, Anthony FW, Dennis KJ. Suppression of ovarian function by Microgynon 30 in day 1 and day 5 “starters.” *Contraception* 1986;33:463–71. [http://dx.doi.org/10.1016/S0010-7824\(86\)80005-1](http://dx.doi.org/10.1016/S0010-7824(86)80005-1)
215. Smith SK, Kirkman RJ, Arce BB, McNeilly AS, Loudon NB, Baird DT. The effect of deliberate omission of Trinordiol or Microgynon on the hypothalamo-pituitary-ovarian axis. *Contraception* 1986;34:513–22. [http://dx.doi.org/10.1016/0010-7824\(86\)90060-0](http://dx.doi.org/10.1016/0010-7824(86)90060-0)
216. Lara-Torre E, Schroeder B. Adolescent compliance and side effects with Quick Start initiation of oral contraceptive pills. *Contraception* 2002;66:81–5. [http://dx.doi.org/10.1016/S0010-7824\(02\)00326-8](http://dx.doi.org/10.1016/S0010-7824(02)00326-8)
217. Murthy AS, Creinin MD, Harwood B, Schreiber CA. Same-day initiation of the transdermal hormonal delivery system (contraceptive patch) versus traditional initiation methods. *Contraception* 2005;72:333–6. <http://dx.doi.org/10.1016/j.contraception.2005.05.009>
218. Westhoff C, Osborne LM, Schafer JE, Morroni C. Bleeding patterns after immediate initiation of an oral compared with a vaginal hormonal contraceptive. *Obstet Gynecol* 2005;106:89–96. <http://dx.doi.org/10.1097/01.AOG.0000164483.13326.59>
219. Yeshaya A, Orvieto R, Kaplan B, et al. Flexible starting schedule for oral contraception: effect on the incidence of breakthrough bleeding and compliance. *Eur J Contracept Reprod Health Care* 1998;3:121–3. <http://dx.doi.org/10.3109/13625189809051414>
220. Yeshaya A, Orvieto R, Kauschansky A, et al. A delayed starting schedule of oral contraception: the effect on the incidence of breakthrough bleeding and compliance in women. *Eur J Contracept Reprod Health Care* 1996;1:263–5. <http://dx.doi.org/10.3109/13625189609150668>
221. Westhoff C, Kerns J, Morroni C, Cushman LF, Tiezzi L, Murphy PA. Quick start: novel oral contraceptive initiation method. *Contraception* 2002;66:141–5. [http://dx.doi.org/10.1016/S0010-7824\(02\)00351-7](http://dx.doi.org/10.1016/S0010-7824(02)00351-7)
222. WHO Collaborative Study of Cardiovascular Disease and Steroid Hormone Contraception. Acute myocardial infarction and combined oral contraceptives: results of an international multicentre case-control study. *Lancet* 1997;349:1202–9. [http://dx.doi.org/10.1016/S0140-6736\(97\)02358-1](http://dx.doi.org/10.1016/S0140-6736(97)02358-1)

223. Dunn N, Thorogood M, Faragher B, et al. Oral contraceptives and myocardial infarction: results of the MICA case-control study. *BMJ* 1999;318:1579–83. <http://dx.doi.org/10.1136/bmj.318.7198.1579>
224. Lewis MA, Heinemann LA, Spitzer WO, MacRae KD, Bruppacher R. The use of oral contraceptives and the occurrence of acute myocardial infarction in young women. Results from the Transnational Study on Oral Contraceptives and the Health of Young Women. *Contraception* 1997;56:129–40. [http://dx.doi.org/10.1016/S0010-7824\(97\)00118-2](http://dx.doi.org/10.1016/S0010-7824(97)00118-2)
225. WHO Collaborative Study of Cardiovascular Disease and Steroid Hormone Contraception. Ischaemic stroke and combined oral contraceptives: results of an international, multicentre, case-control study. *Lancet* 1996;348:498–505. [http://dx.doi.org/10.1016/S0140-6736\(95\)12393-8](http://dx.doi.org/10.1016/S0140-6736(95)12393-8)
226. Heinemann LA, Lewis MA, Spitzer WO, Thorogood M, Guggenmoos-Holzmann I, Bruppacher R; Transnational Research Group on Oral Contraceptives and the Health of Young Women. Thromboembolic stroke in young women. A European case-control study on oral contraceptives. *Contraception* 1998;57:29–37. [http://dx.doi.org/10.1016/S0010-7824\(97\)00204-7](http://dx.doi.org/10.1016/S0010-7824(97)00204-7)
227. WHO Collaborative Study of Cardiovascular Disease and Steroid Hormone Contraception. Haemorrhagic stroke, overall stroke risk, and combined oral contraceptives: results of an international, multicentre, case-control study. *Lancet* 1996;348:505–10. [http://dx.doi.org/10.1016/S0140-6736\(95\)12394-6](http://dx.doi.org/10.1016/S0140-6736(95)12394-6)
228. Mohllajee AP, Curtis KM, Martins SL, Peterson HB. Does use of hormonal contraceptives among women with thrombogenic mutations increase their risk of venous thromboembolism? A systematic review. *Contraception* 2006;73:166–78. <http://dx.doi.org/10.1016/j.contraception.2005.08.011>
229. Wu O, Robertson L, Twaddle S, et al. Screening for thrombophilia in high-risk situations: systematic review and cost-effectiveness analysis. The Thrombosis: Risk and Economic Assessment of Thrombophilia Screening (TREATS) study. *Health Technol Assess* 2006;10:1–110. <http://dx.doi.org/10.3310/hta10110>
230. Wu O, Greer IA. Is screening for thrombophilia cost-effective? *Curr Opin Hematol* 2007;14:500–3. <http://dx.doi.org/10.1097/MOH.0b013e32825f5318>
231. Blickstein D, Blickstein I. Oral contraception and thrombophilia. *Curr Opin Obstet Gynecol* 2007;19:370–6. <http://dx.doi.org/10.1097/GCO.0b013e32821642e6>
232. Steenland MW, Rodriguez MI, Marchbanks PA, Curtis KM. How does the number of oral contraceptive pill packs dispensed or prescribed affect continuation and other measures of consistent and correct use? A systematic review. *Contraception* 2013;87:605–10. <http://dx.doi.org/10.1016/j.contraception.2012.08.004>
233. White KO, Westhoff C. The effect of pack supply on oral contraceptive pill continuation: a randomized controlled trial. *Obstet Gynecol* 2011;118:615–22. <http://dx.doi.org/10.1097/AOG.0b013e3282289eab>
234. Foster DG, Parvataneni R, de Bocanegra HT, Lewis C, Bradsberry M, Darney P. Number of oral contraceptive pill packages dispensed, method continuation, and costs. *Obstet Gynecol* 2006;108:1107–14. <http://dx.doi.org/10.1097/01.AOG.0000239122.98508.39>
235. Foster DG, Hulett D, Bradsberry M, Darney P, Policar M. Number of oral contraceptive pill packages dispensed and subsequent unintended pregnancies. *Obstet Gynecol* 2011;117:566–72. <http://dx.doi.org/10.1097/AOG.0b013e3282056309>
236. Chin-Quee DS, Cuthbertson C, Janowitz B. Over-the-counter pill provision: evidence from Jamaica. *Stud Fam Plann* 2006;37:99–110. <http://dx.doi.org/10.1111/j.1728-4465.2006.00089.x>
237. Zapata LB, Steenland MW, Brahma D, Marchbanks PA, Curtis KM. Effect of missed combined hormonal contraceptives on contraceptive effectiveness: a systematic review. *Contraception* 2013;87:685–700. <http://dx.doi.org/10.1016/j.contraception.2012.08.035>
238. Anttila L, Kunz M, Marr J. Bleeding pattern with drospirenone 3 mg+ethinyl estradiol 20 mcg 24/4 combined oral contraceptive compared with desogestrel 150 mcg+ethinyl estradiol 20 mcg 21/7 combined oral contraceptive. *Contraception* 2009;80:445–51. <http://dx.doi.org/10.1016/j.contraception.2009.03.013>
239. Chowdhury V, Joshi UM, Gopalkrishna K, Betrabet S, Mehta S, Saxena BN. ‘Escape’ ovulation in women due to the missing of low dose combination oral contraceptive pills. *Contraception* 1980;22:241–7. [http://dx.doi.org/10.1016/S0010-7824\(80\)80003-5](http://dx.doi.org/10.1016/S0010-7824(80)80003-5)
240. Christin-Maitre S, Serfaty D, Chabbert-Buffet N, Ochslein E, Chassard D, Thomas JL. Comparison of a 24-day and a 21-day pill regimen for the novel combined oral contraceptive, norgestrel acetate and 17 β -estradiol (NOMAC/E2): a double-blind, randomized study. *Hum Reprod* 2011;26:1338–47. <http://dx.doi.org/10.1093/humrep/der058>
241. Creinin MD, Lippman JS, Eder SE, Godwin AJ, Olson W. The effect of extending the pill-free interval on follicular activity: triphasic norgestimate/35 μ g ethinyl estradiol versus monophasic levonorgestrel/20 μ g ethinyl estradiol. *Contraception* 2002;66:147–52. [http://dx.doi.org/10.1016/S0010-7824\(02\)00344-X](http://dx.doi.org/10.1016/S0010-7824(02)00344-X)
242. Dinger J, Minh TD, Buttman N, Bardenheuer K. Effectiveness of oral contraceptive pills in a large U.S. cohort comparing progestogen and regimen. *Obstet Gynecol* 2011;117:33–40. <http://dx.doi.org/10.1097/AOG.0b013e32820095a2>
243. Elomaa K, Lähteenmäki P. Ovulatory potential of preovulatory sized follicles during oral contraceptive treatment. *Contraception* 1999;60:275–9. [http://dx.doi.org/10.1016/S0010-7824\(99\)00094-3](http://dx.doi.org/10.1016/S0010-7824(99)00094-3)
244. Elomaa K, Rolland R, Brosens I, et al. Omitting the first oral contraceptive pills of the cycle does not automatically lead to ovulation. *Am J Obstet Gynecol* 1998;179:41–6. [http://dx.doi.org/10.1016/S0002-9378\(98\)70249-2](http://dx.doi.org/10.1016/S0002-9378(98)70249-2)
245. Endrikat J, Wessel J, Rosenbaum P, Düsterberg B. Plasma concentrations of endogenous hormones during one regular treatment cycle with a low-dose oral contraceptive and during two cycles with deliberate omission of two tablets. *Gynecol Endocrinol* 2004;18:318–26. <http://dx.doi.org/10.1080/0951359042000199869>
246. Hamilton CJ, Hoogland HJ. Longitudinal ultrasonographic study of the ovarian suppressive activity of a low-dose triphasic oral contraceptive during correct and incorrect pill intake. *Am J Obstet Gynecol* 1989;161:1159–62. [http://dx.doi.org/10.1016/0002-9378\(89\)90655-8](http://dx.doi.org/10.1016/0002-9378(89)90655-8)
247. Hedon B, Cristol P, Plauchut A, et al. Ovarian consequences of the transient interruption of combined oral contraceptives. *Int J Fertil* 1992;37:270–6.
248. Killick SR. Ovarian follicles during oral contraceptive cycles: their potential for ovulation. *Fertil Steril* 1989;52:580–2.
249. Killick SR, Bancroft K, Oelbaum S, Morris J, Elstein M. Extending the duration of the pill-free interval during combined oral contraception. *Adv Contracept* 1990;6:33–40. <http://dx.doi.org/10.1007/BF01849485>
250. Klipping C, Duijkers I, Trummer D, Marr J. Suppression of ovarian activity with a drospirenone-containing oral contraceptive in a 24/4 regimen. *Contraception* 2008;78:16–25. <http://dx.doi.org/10.1016/j.contraception.2008.02.019>

251. Klipping C, Marr J. Effects of two combined oral contraceptives containing ethinyl estradiol 20 microg combined with either drospirenone or desogestrel on lipids, hemostatic parameters and carbohydrate metabolism. *Contraception* 2005;71:409–16. <http://dx.doi.org/10.1016/j.contraception.2004.12.005>
252. Landgren BM, Csemiczky G. The effect of follicular growth and luteal function of “missing the pill.” A comparison between a monophasic and a triphasic combined oral contraceptive. *Contraception* 1991;43:149–59. [http://dx.doi.org/10.1016/0010-7824\(91\)90042-E](http://dx.doi.org/10.1016/0010-7824(91)90042-E)
253. Landgren BM, Diczfalusy E. Hormonal consequences of missing the pill during the first two days of three consecutive artificial cycles. *Contraception* 1984;29:437–46. [http://dx.doi.org/10.1016/0010-7824\(84\)90017-9](http://dx.doi.org/10.1016/0010-7824(84)90017-9)
254. Letterie GS. A regimen of oral contraceptives restricted to the periovulatory period may permit folliculogenesis but inhibit ovulation. *Contraception* 1998;57:39–44. [http://dx.doi.org/10.1016/S0010-7824\(97\)00205-9](http://dx.doi.org/10.1016/S0010-7824(97)00205-9)
255. Letterie GS, Chow GE. Effect of “missed” pills on oral contraceptive effectiveness. *Obstet Gynecol* 1992;79:979–82.
256. Morris SE, Groom GV, Cameron ED, Buckingham MS, Everitt JM, Elstein M. Studies on low dose oral contraceptives: plasma hormone changes in relation to deliberate pill (“Microgynon 30”) omission. *Contraception* 1979;20:61–9. [http://dx.doi.org/10.1016/0010-7824\(79\)90045-3](http://dx.doi.org/10.1016/0010-7824(79)90045-3)
257. Nakajima ST, Archer DF, Ellman H. Efficacy and safety of a new 24-day oral contraceptive regimen of norethindrone acetate 1 mg/ethinyl estradiol 20 µg (Loestrin 24 Fe). *Contraception* 2007;75:16–22. <http://dx.doi.org/10.1016/j.contraception.2006.08.004>
258. Nuttall ID, Elstein M, McCafferty E, Seth J, Cameron ED. The effect of ethinyl estradiol 20 mcg and levonorgestrel 250 mcg on the pituitary-ovarian function during normal tablet-taking and when tablets are missed. *Contraception* 1982;26:121–35. [http://dx.doi.org/10.1016/0010-7824\(82\)90081-6](http://dx.doi.org/10.1016/0010-7824(82)90081-6)
259. Pierson RA, Archer DF, Moreau M, Shangold GA, Fisher AC, Creasy GW. Ortho Evra/Evra versus oral contraceptives: follicular development and ovulation in normal cycles and after an intentional dosing error. *Fertil Steril* 2003;80:34–42. [http://dx.doi.org/10.1016/S0015-0282\(03\)00556-9](http://dx.doi.org/10.1016/S0015-0282(03)00556-9)
260. Rible RD, Taylor D, Wilson ML, Stanczyk FZ, Mishell DR Jr. Follicular development in a 7-day versus 4-day hormone-free interval with an oral contraceptive containing 20 mcg ethinyl estradiol and 1 mg norethindrone acetate. *Contraception* 2009;79:182–8. <http://dx.doi.org/10.1016/j.contraception.2008.10.005>
261. Schlaff WD, Lynch AM, Hughes HD, Cedars MI, Smith DL. Manipulation of the pill-free interval in oral contraceptive pill users: the effect on follicular suppression. *Am J Obstet Gynecol* 2004;190:943–51. <http://dx.doi.org/10.1016/j.ajog.2004.02.012>
262. Spona J, Elstein M, Feichtinger W, et al. Shorter pill-free interval in combined oral contraceptives decreases follicular development. *Contraception* 1996;54:71–7. [http://dx.doi.org/10.1016/0010-7824\(96\)00137-0](http://dx.doi.org/10.1016/0010-7824(96)00137-0)
263. Sullivan H, Furniss H, Spona J, Elstein M. Effect of 21-day and 24-day oral contraceptive regimens containing gestodene (60 µg) and ethinyl estradiol (15 µg) on ovarian activity. *Fertil Steril* 1999;72:115–20. [http://dx.doi.org/10.1016/S0015-0282\(99\)00205-8](http://dx.doi.org/10.1016/S0015-0282(99)00205-8)
264. Wang E, Shi S, Cekan SZ, Landgren BM, Diczfalusy E. Hormonal consequences of “missing the pill.” *Contraception* 1982;26:545–66. [http://dx.doi.org/10.1016/0010-7824\(82\)90131-7](http://dx.doi.org/10.1016/0010-7824(82)90131-7)
265. Willis SA, Kuehl TJ, Spiekerman AM, Sulak PJ. Greater inhibition of the pituitary—ovarian axis in oral contraceptive regimens with a shortened hormone-free interval. *Contraception* 2006;74:100–3. <http://dx.doi.org/10.1016/j.contraception.2006.02.006>
266. Abrams LS, Skee DM, Natarajan J, et al. Pharmacokinetics of norelgestromin and ethinyl estradiol delivered by a contraceptive patch (Ortho Evra/Evra) under conditions of heat, humidity, and exercise. *J Clin Pharmacol* 2001;41:1301–9. <http://dx.doi.org/10.1177/00912700122012887>
267. Ahrendt HJ, Nisand I, Bastianelli C, et al. Efficacy, acceptability and tolerability of the combined contraceptive ring, NuvaRing, compared with an oral contraceptive containing 30 microg of ethinyl estradiol and 3 mg of drospirenone. *Contraception* 2006;74:451–7. <http://dx.doi.org/10.1016/j.contraception.2006.07.004>
268. Bjarnadóttir RI, Tuppurainen M, Killick SR. Comparison of cycle control with a combined contraceptive vaginal ring and oral levonorgestrel/ethinyl estradiol. *Am J Obstet Gynecol* 2002;186:389–95. <http://dx.doi.org/10.1067/mob.2002.121103>
269. Brucker C, Karck U, Merkle E. Cycle control, tolerability, efficacy and acceptability of the vaginal contraceptive ring, NuvaRing: results of clinical experience in Germany. *Eur J Contracept Reprod Health Care* 2008;13:31–8. <http://dx.doi.org/10.1080/13625180701577122>
270. Dieben TO, Roumen FJ, Apter D. Efficacy, cycle control, and user acceptability of a novel combined contraceptive vaginal ring. *Obstet Gynecol* 2002;100:585–93.
271. Mulders TM, Dieben TO. Use of the novel combined contraceptive vaginal ring NuvaRing for ovulation inhibition. *Fertil Steril* 2001;75:865–70. [http://dx.doi.org/10.1016/S0015-0282\(01\)01689-2](http://dx.doi.org/10.1016/S0015-0282(01)01689-2)
272. Guilbert E, Boroditsky R, Black A, et al; Society of Obstetricians and Gynaecologists of Canada. Canadian Consensus Guideline on Continuous and Extended Hormonal Contraception, 2007. *J Obstet Gynaecol Can* 2007;29(Suppl 2):S1–32. [http://dx.doi.org/10.1016/S1701-2163\(16\)32526-9](http://dx.doi.org/10.1016/S1701-2163(16)32526-9)
273. Wiegratz I, Stahlberg S, Manthey T, et al. Effect of extended-cycle regimen with an oral contraceptive containing 30 mcg ethinylestradiol and 2 mg dienogest on bleeding patterns, safety, acceptance and contraceptive efficacy. *Contraception* 2011;84:133–43. <http://dx.doi.org/10.1016/j.contraception.2011.01.002>
274. Hubacher D, Fortney J. Follow-up visits after IUD insertion. Are more better? *J Reprod Med* 1999;44:801–6.
275. Godfrey EM, Whiteman MK, Curtis KM. Treatment of unscheduled bleeding in women using extended- or continuous-use combined hormonal contraception: a systematic review. *Contraception* 2013;87:567–75. <http://dx.doi.org/10.1016/j.contraception.2012.08.005>
276. Sulak PJ, Kuehl TJ, Coffee A, Willis S. Prospective analysis of occurrence and management of breakthrough bleeding during an extended oral contraceptive regimen. *Am J Obstet Gynecol* 2006;195:935–41. <http://dx.doi.org/10.1016/j.ajog.2006.02.048>
277. Sulak PJ, Smith V, Coffee A, Witt I, Kuehl AL, Kuehl TJ. Frequency and management of breakthrough bleeding with continuous use of the transvaginal contraceptive ring: a randomized controlled trial. *Obstet Gynecol* 2008;112:563–71. <http://dx.doi.org/10.1097/AOG.0b013e3181842071>
278. Kaneshiro B, Edelman A, Carlson N, Morgan K, Nichols M, Jensen J. Treatment of unscheduled bleeding in continuous oral contraceptive users with doxycycline: a randomized controlled trial. *Obstet Gynecol* 2010;115:1141–9. <http://dx.doi.org/10.1097/AOG.0b013e3181e0119c>

279. McCann MF, Potter LS. Progestin-only oral contraception: a comprehensive review. *Contraception* 1994;50(Suppl 1):S1–195. [http://dx.doi.org/10.1016/0010-7824\(94\)90124-4](http://dx.doi.org/10.1016/0010-7824(94)90124-4)
280. Arévalo M, Jennings V, Sinai I. Efficacy of a new method of family planning: the Standard Days Method. *Contraception* 2002;65:333–8. [http://dx.doi.org/10.1016/S0010-7824\(02\)00288-3](http://dx.doi.org/10.1016/S0010-7824(02)00288-3)
281. Arévalo M, Sinai I, Jennings V. A fixed formula to define the fertile window of the menstrual cycle as the basis of a simple method of natural family planning. *Contraception* 1999;60:357–60. [http://dx.doi.org/10.1016/S0010-7824\(99\)00106-7](http://dx.doi.org/10.1016/S0010-7824(99)00106-7)
282. Wilcox AJ, Dunson DB, Weinberg CR, Trussell J, Baird DD. Likelihood of conception with a single act of intercourse: providing benchmark rates for assessment of post-coital contraceptives. *Contraception* 2001;63:211–5. [http://dx.doi.org/10.1016/S0010-7824\(01\)00191-3](http://dx.doi.org/10.1016/S0010-7824(01)00191-3)
283. Cleland K, Zhu H, Goldstuck N, Cheng L, Trussell J. The efficacy of intrauterine devices for emergency contraception: a systematic review of 35 years of experience. *Hum Reprod* 2012;27:1994–2000. <http://dx.doi.org/10.1093/humrep/des140>
284. Glasier AF, Cameron ST, Fine PM, et al. Ulipristal acetate versus levonorgestrel for emergency contraception: a randomised non-inferiority trial and meta-analysis. *Lancet* 2010;375:555–62. [http://dx.doi.org/10.1016/S0140-6736\(10\)60101-8](http://dx.doi.org/10.1016/S0140-6736(10)60101-8)
285. Raymond E, Taylor D, Trussell J, Steiner MJ. Minimum effectiveness of the levonorgestrel regimen of emergency contraception. *Contraception* 2004;69:79–81. <http://dx.doi.org/10.1016/j.contraception.2003.09.013>
286. Jataoui TC, Curtis KM. Safety and effectiveness data for emergency contraceptive pills among women with obesity: a systematic review. *Contraception* 2016. Epub May 24, 2016. <http://dx.doi.org/10.1016/j.contraception.2016.05.002>
287. Fine P, Mathé H, Ginde S, Cullins V, Morfesis J, Gainer E. Ulipristal acetate taken 48–120 hours after intercourse for emergency contraception. *Obstet Gynecol* 2010;115:257–63. <http://dx.doi.org/10.1097/AOG.0b013e3181c8e2aa>
288. Dada OA, Godfrey EM, Piaggio G, von Hertzen H; Nigerian Network for Reproductive Health Research and Training. A randomized, double-blind, noninferiority study to compare two regimens of levonorgestrel for emergency contraception in Nigeria. *Contraception* 2010;82:373–8. <http://dx.doi.org/10.1016/j.contraception.2010.06.004>
289. Ellertson C, Evans M, Ferden S, et al. Extending the time limit for starting the Yuzpe regimen of emergency contraception to 120 hours. *Obstet Gynecol* 2003;101:1168–71.
290. Ngai SW, Fan S, Li S, et al. A randomized trial to compare 24 h versus 12 h double dose regimen of levonorgestrel for emergency contraception. *Hum Reprod* 2005;20:307–11. <http://dx.doi.org/10.1093/humrep/deh583>
291. Rodrigues I, Grou F, Joly J. Effectiveness of emergency contraceptive pills between 72 and 120 hours after unprotected sexual intercourse. *Am J Obstet Gynecol* 2001;184:531–7. <http://dx.doi.org/10.1067/mob.2001.111102>
292. von Hertzen H, Piaggio G, Ding J, et al; WHO Research Group on Post-ovulatory Methods of Fertility Regulation. Low dose mifepristone and two regimens of levonorgestrel for emergency contraception: a WHO multicentre randomised trial. *Lancet* 2002;360:1803–10. [http://dx.doi.org/10.1016/S0140-6736\(02\)11767-3](http://dx.doi.org/10.1016/S0140-6736(02)11767-3)
293. Piaggio G, Kapp N, von Hertzen H. Effect on pregnancy rates of the delay in the administration of levonorgestrel for emergency contraception: a combined analysis of four WHO trials. *Contraception* 2011;84:35–9. <http://dx.doi.org/10.1016/j.contraception.2010.11.010>
294. Rodriguez MI, Curtis KM, Gaffield ML, Jackson E, Kapp N. Advance supply of emergency contraception: a systematic review. *Contraception* 2013;87:590–601. <http://dx.doi.org/10.1016/j.contraception.2012.09.011>
295. Brache V, Cochon L, Duijkers IJ, et al. A prospective, randomized, pharmacodynamic study of quick-starting a desogestrel progestin-only pill following ulipristal acetate for emergency contraception. *Hum Reprod* 2015;30:2785–93.
296. Cameron ST, Berger C, Michie L, Klipping C, Gemzell-Danielsson K. The effects on ovarian activity of ulipristal acetate when ‘quickstarting’ a combined oral contraceptive pill: a prospective, randomized, double-blind parallel-arm, placebo-controlled study. *Hum Reprod* 2015;30:1566–72. <http://dx.doi.org/10.1093/humrep/dev115>
297. Salcedo J, Rodriguez MI, Curtis KM, Kapp N. When can a woman resume or initiate contraception after taking emergency contraceptive pills? A systematic review. *Contraception* 2013;87:602–4. <http://dx.doi.org/10.1016/j.contraception.2012.08.013>
298. Rodriguez MI, Godfrey EM, Warden M, Curtis KM. Prevention and management of nausea and vomiting with emergency contraception: a systematic review. *Contraception* 2013;87:583–9. <http://dx.doi.org/10.1016/j.contraception.2012.09.031>
299. Creinin MD, Schlaff W, Archer DF, et al. Progesterone receptor modulator for emergency contraception: a randomized controlled trial. *Obstet Gynecol* 2006;108:1089–97. <http://dx.doi.org/10.1097/01.AOG.0000239440.02284.45>
300. Farajkhoda T, Khoshbin A, Enjebab B, Bokaei M, Karimi Zarchi M. Assessment of two emergency contraceptive regimens in Iran: levonorgestrel versus the Yuzpe. *Niger J Clin Pract* 2009;12:450–2.
301. Ho PC, Kwan MSW. A prospective randomized comparison of levonorgestrel with the Yuzpe regimen in post-coital contraception. *Hum Reprod* 1993;8:389–92.
302. Arowojolu AO, Okewole IA, Adekunle AO. Comparative evaluation of the effectiveness and safety of two regimens of levonorgestrel for emergency contraception in Nigerians. *Contraception* 2002;66:269–73. [http://dx.doi.org/10.1016/S0010-7824\(02\)00337-2](http://dx.doi.org/10.1016/S0010-7824(02)00337-2)
303. Ragan RE, Rock RW, Buck HW. Metoclopramide pretreatment attenuates emergency contraceptive-associated nausea. *Am J Obstet Gynecol* 2003;188:330–3. <http://dx.doi.org/10.1067/mob.2003.90>
304. Raymond EG, Creinin MD, Barnhart KT, Lovvorn AE, Rountree RW, Trussell J. Meclizine for prevention of nausea associated with use of emergency contraceptive pills: a randomized trial. *Obstet Gynecol* 2000;95:271–7.
305. Peterson HB. Sterilization. *Obstet Gynecol* 2008;111:189–203. <http://dx.doi.org/10.1097/01.AOG.0000298621.98372.62>
306. Peterson HB, Xia Z, Hughes JM, Wilcox LS, Tylor LR, Trussell J. The risk of pregnancy after tubal sterilization: findings from the U.S. Collaborative Review of Sterilization. *Am J Obstet Gynecol* 1996;174:1161–8, discussion 1168–70. [http://dx.doi.org/10.1016/S0002-9378\(96\)70658-0](http://dx.doi.org/10.1016/S0002-9378(96)70658-0)
307. Lawrie TA, Nardin JM, Kulier R, Boulvain M. Techniques for the interruption of tubal patency for female sterilisation. *Cochrane Database Syst Rev* 2011; (2):CD003034.
308. Cleary TP, Tepper NK, Cwiak C, et al. Pregnancies after hysteroscopic sterilization: a systematic review. *Contraception* 2013;87:539–48. <http://dx.doi.org/10.1016/j.contraception.2012.08.006>
309. Legendre G, Levallant JM, Faivre E, Deffieux X, Gervaise A, Fernandez H. 3D ultrasound to assess the position of tubal sterilization microinserts. *Hum Reprod* 2011;26:2683–9. <http://dx.doi.org/10.1093/humrep/der242>
310. Levy B, Levie MD, Childers ME. A summary of reported pregnancies after hysteroscopic sterilization. *J Minim Invasive Gynecol* 2007;14:271–4. <http://dx.doi.org/10.1016/j.jmig.2006.11.007>

311. Veersema S, Vleugels MB, Moolenaar LM, Janssen CA, Brölmann HA. Unintended pregnancies after Essure sterilization in the Netherlands. *Fertil Steril* 2010;93:35–8. <http://dx.doi.org/10.1016/j.fertnstert.2008.10.005>
312. Arjona JE, Miño M, Cordon J, Povedano B, Pelegrin B, Castelo-Branco C. Satisfaction and tolerance with office hysteroscopic tubal sterilization. *Fertil Steril* 2008;90:1182–6. <http://dx.doi.org/10.1016/j.fertnstert.2007.08.007>
313. Duffy S, Marsh F, Rogerson L, et al. Female sterilisation: a cohort controlled comparative study of ESSURE versus laparoscopic sterilisation. *BJOG* 2005;112:1522–8. <http://dx.doi.org/10.1111/j.1471-0528.2005.00726.x>
314. Grosdemouge I, Engrand JB, Dhainault C, et al. Essure implants for tubal sterilisation in France [French]. *Gynecol Obstet Fertil* 2009;37:389–95. <http://dx.doi.org/10.1016/j.gyobfe.2009.03.024>
315. Levie MD, Chudnoff SG. Prospective analysis of office-based hysteroscopic sterilization. *J Minim Invasive Gynecol* 2006;13:98–101. <http://dx.doi.org/10.1016/j.jmig.2005.11.010>
316. Shavell VI, Abdallah ME, Diamond MP, Berman JM. Placement of a permanent birth control device at a university medical center. *J Reprod Med* 2009;54:218–22.
317. Teisala K. Removal of an intrauterine device and the treatment of acute pelvic inflammatory disease. *Ann Med* 1989;21:63–5. <http://dx.doi.org/10.3109/07853898909149184>
318. Bedford JM, Zelikovskiy G. Viability of spermatozoa in the human ejaculate after vasectomy. *Fertil Steril* 1979;32:460–3.
319. Edwards IS. Earlier testing after vasectomy, based on the absence of motile sperm. *Fertil Steril* 1993;59:431–6.
320. Jouannet P, David G. Evolution of the properties of semen immediately following vasectomy. *Fertil Steril* 1978;29:435–41.
321. Labrecque M, Hays M, Chen-Mok M, Barone MA, Sokal D. Frequency and patterns of early recanalization after vasectomy. *BMC Urol* 2006;6:25. <http://dx.doi.org/10.1186/1471-2490-6-25>
322. Alderman PM. The lurking sperm. A review of failures in 8879 vasectomies performed by one physician. *JAMA* 1988;259:3142–4. <http://dx.doi.org/10.1001/jama.1988.03720210032024>
323. Black T, Francome C. The evolution of the Marie Stopes electrocautery no-scalpel vasectomy procedure. *J Fam Plann Reprod Health Care* 2002;28:137–8. <http://dx.doi.org/10.1783/147118902101196270>
324. Davies AH, Sharp RJ, Cranston D, Mitchell RG. The long-term outcome following “special clearance” after vasectomy. *Br J Urol* 1990;66:211–2. <http://dx.doi.org/10.1111/j.1464-410X.1990.tb14907.x>
325. Philp T, Guillebaud J, Budd D. Late failure of vasectomy after two documented analyses showing azoospermic semen. *Br Med J (Clin Res Ed)* 1984;289:77–9. <http://dx.doi.org/10.1136/bmj.289.6437.77>
326. Philp T, Guillebaud J, Budd D. Complications of vasectomy: review of 16,000 patients. *Br J Urol* 1984;56:745–8. <http://dx.doi.org/10.1111/j.1464-410X.1984.tb06161.x>
327. Belker AM, Sexter MS, Sweitzer SJ, Raff MJ. The high rate of noncompliance for post-vasectomy semen examination: medical and legal considerations. *J Urol* 1990;144:284–6.
328. Chawla A, Bowles B, Zini A. Vasectomy follow-up: clinical significance of rare nonmotile sperm in postoperative semen analysis. *Urology* 2004;64:1212–5. <http://dx.doi.org/10.1016/j.urology.2004.07.007>
329. Labrecque M, Bédard L, Laperrière L. Efficacy and complications associated with vasectomies in two clinics in the Quebec region [French]. *Can Fam Physician* 1998;44:1860–6.
330. Labrecque M, Nazerali H, Mondor M, Fortin V, Nasution M. Effectiveness and complications associated with 2 vasectomy occlusion techniques. *J Urol* 2002;168:2495–8, discussion 2498. [http://dx.doi.org/10.1016/S0022-5347\(05\)64176-6](http://dx.doi.org/10.1016/S0022-5347(05)64176-6)
331. Maatman TJ, Aldrin L, Carothers GG. Patient noncompliance after vasectomy. *Fertil Steril* 1997;68:552–5. [http://dx.doi.org/10.1016/S0015-0282\(97\)00251-3](http://dx.doi.org/10.1016/S0015-0282(97)00251-3)
332. Dhar NB, Jones JS, Bhatt A, Babineau D. A prospective evaluation of the impact of scheduled follow-up appointments with compliance rates after vasectomy. *BJU Int* 2007;99:1094–7. <http://dx.doi.org/10.1111/j.1464-410X.2006.06725.x>
333. Hillard PJ, Berek JS, Barss VA, et al. Guidelines for women’s health care: A resource manual. 3rd ed. Washington, DC: American College of Obstetricians and Gynecologists; 2007.
334. Shifren J, Gass M; NAMS Recommendations for Clinical Care of Midlife Women Working Group. The North American Menopause Society recommendation for clinical care of midlife women. *Menopause* 2014;21:1038–62.
335. te Velde ER, Pearson PL. The variability of female reproductive ageing. *Hum Reprod Update* 2002;8:141–54. <http://dx.doi.org/10.1093/humupd/8.2.141>
336. Broekmans FJ, Soules MR, Fauser BC. Ovarian aging: mechanisms and clinical consequences. *Endocr Rev* 2009;30:465–93. <http://dx.doi.org/10.1210/er.2009-0006>
337. Wood JW. Fecundity and natural fertility in humans. *Oxf Rev Reprod Biol* 1989;11:61–109.
338. Balasch J, Gratacós E. Delayed childbearing: effects on fertility and the outcome of pregnancy. *Curr Opin Obstet Gynecol* 2012;24:187–93. <http://dx.doi.org/10.1097/GCO.0b013e3283517908>
339. Bateman BT, Simpson LL. Higher rate of stillbirth at the extremes of reproductive age: a large nationwide sample of deliveries in the United States. *Am J Obstet Gynecol* 2006;194:840–5. <http://dx.doi.org/10.1016/j.ajog.2005.08.038>
340. Berg CJ, Callaghan WM, Syverson C, Henderson Z. Pregnancy-related mortality in the United States, 1998 to 2005. *Obstet Gynecol* 2010;116:1302–9. <http://dx.doi.org/10.1097/AOG.0b013e3181fdff11>
341. Lidegaard Ø, Løkkegaard E, Svendsen AL, Agger C. Hormonal contraception and risk of venous thromboembolism: national follow-up study. *BMJ* 2009;339:b2890.
342. Lidegaard Ø, Nielsen LH, Skovlund CW, Skjeldestad FE, Løkkegaard E. Risk of venous thromboembolism from use of oral contraceptives containing different progestogens and oestrogen doses: Danish cohort study, 2001–9. *BMJ* 2011;343:d6423.
343. Nightingale AL, Lawrenson RA, Simpson EL, Williams TJ, MacRae KD, Farmer RD. The effects of age, body mass index, smoking and general health on the risk of venous thromboembolism in users of combined oral contraceptives. *Eur J Contracept Reprod Health Care* 2000;5:265–74. <http://dx.doi.org/10.1080/13625180008500402>
344. Slone D, Shapiro S, Kaufman DW, Rosenberg L, Miettinen OS, Stolley PD. Risk of myocardial infarction in relation to current and discontinued use of oral contraceptives. *N Engl J Med* 1981;305:420–4. <http://dx.doi.org/10.1056/NEJM198108203050802>
345. Tanis BC, van den Bosch MA, Kemmeren JM, et al. Oral contraceptives and the risk of myocardial infarction. *N Engl J Med* 2001;345:1787–93. <http://dx.doi.org/10.1056/NEJMoa003216>
346. Collaborative Group on Hormonal Factors in Breast Cancer. Breast cancer and hormonal contraceptives: collaborative reanalysis of individual data on 53 297 women with breast cancer and 100 239 women without breast cancer from 54 epidemiological studies. *Lancet* 1996;347:1713–27. [http://dx.doi.org/10.1016/S0140-6736\(96\)90806-5](http://dx.doi.org/10.1016/S0140-6736(96)90806-5)
347. Gill JK, Press MF, Patel AV, Bernstein L. Oral contraceptive use and risk of breast carcinoma in situ (United States). *Cancer Causes Control* 2006;17:1155–62. <http://dx.doi.org/10.1007/s10552-006-0056-0>

Recommendations and Reports

348. Kumle M, Weiderpass E, Braaten T, Persson I, Adami HO, Lund E. Use of oral contraceptives and breast cancer risk: The Norwegian-Swedish Women's Lifestyle and Health Cohort Study. *Cancer Epidemiol Biomarkers Prev* 2002;11:1375–81.
349. Marchbanks PA, McDonald JA, Wilson HG, et al. Oral contraceptives and the risk of breast cancer. *N Engl J Med* 2002;346:2025–32. <http://dx.doi.org/10.1056/NEJMoa013202>
350. Newcomb PA, Longnecker MP, Storer BE, et al. Recent oral contraceptive use and risk of breast cancer (United States). *Cancer Causes Control* 1996;7:525–32. <http://dx.doi.org/10.1007/BF00051885>
351. Rosenberg L, Palmer JR, Rao RS, et al. Case-control study of oral contraceptive use and risk of breast cancer. *Am J Epidemiol* 1996;143:25–37. <http://dx.doi.org/10.1093/oxfordjournals.aje.a008654>
352. Rosenberg L, Zhang Y, Coogan PF, Strom BL, Palmer JR. A case-control study of oral contraceptive use and incident breast cancer. *Am J Epidemiol* 2009;169:473–9. <http://dx.doi.org/10.1093/aje/kwn360>
353. Shapiro S, Rosenberg L, Hoffman M, et al. Risk of breast cancer in relation to the use of injectable progestogen contraceptives and combined estrogen/progestogen contraceptives. *Am J Epidemiol* 2000;151:396–403. <http://dx.doi.org/10.1093/oxfordjournals.aje.a010219>

Appendix A

Summary Chart of U.S. Medical Eligibility Criteria for Contraceptive Use, 2016

Health-care providers can use the summary table as a quick reference guide to the classifications for hormonal contraceptive methods and intrauterine contraception to compare classifications across these methods (Box A1) (Table A1). For

complete guidance, see the 2016 *U.S. Medical Eligibility Criteria for Contraceptive Use* (U.S. MEC) (Curtis KM, Tepper NK, Jatlaoui TC, et al. U.S. medical eligibility criteria for contraceptive use, 2016. MMWR Recomm Rep 2016;65[No. RR-3]) for clarifications to the numeric categories, as well as for summaries of the evidence and additional comments. Hormonal contraceptives and intrauterine devices do not protect against sexually transmitted diseases (STDs), including human immunodeficiency virus (HIV), and women using these methods should be counseled that consistent and correct use of the male latex condom reduces the risk for transmission of HIV and other STDs. Use of female condoms can provide protection from transmission of STDs, although data are limited.

BOX A1. Categories for classifying hormonal contraceptives and intrauterine devices

- 1 = A condition for which there is no restriction for the use of the contraceptive method.
- 2 = A condition for which the advantages of using the method generally outweigh the theoretical or proven risks.
- 3 = A condition for which the theoretical or proven risks usually outweigh the advantages of using the method.
- 4 = A condition that represents an unacceptable health risk if the contraceptive method is used.

TABLE A1. Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD	LNG-IUD	Implants	DMPA	POP	CHCs
Personal Characteristics and Reproductive History						
Pregnancy	4*	4*	NA*	NA*	NA*	NA*
Age	Menarche to <20 years: 2 ≥20 years: 1	Menarche to <20 years: 2 ≥20 years: 1	Menarche to <18 years: 1 18–45 years: 1 >45 years: 1	Menarche to <18 years: 2 18–45 years: 1 >45 years: 2	Menarche to <18 years: 1 18–45 years: 1 >45 years: 1	Menarche to <40 years: 1 ≥40 years: 2
Parity						
a. Nulliparous	2	2	1	1	1	1
b. Parous	1	1	1	1	1	1
Breastfeeding						
a. <21 days postpartum	—	—	2*	2*	2*	4*
b. 21 to <30 days postpartum						
i. With other risk factors for VTE (e.g., age ≥35 years, previous VTE, thrombophilia, immobility, transfusion at delivery, peripartum cardiomyopathy, BMI ≥30 kg/m ² , postpartum hemorrhage, postcesarean delivery, preeclampsia, or smoking)	—	—	2*	2*	2*	3*
ii. Without other risk factors for VTE	—	—	2*	2*	2*	3*
c. 30–42 days postpartum						
i. With other risk factors for VTE (e.g., age ≥35 years, previous VTE, thrombophilia, immobility, transfusion at delivery, peripartum cardiomyopathy, BMI ≥30 kg/m ² , postpartum hemorrhage, postcesarean delivery, preeclampsia, or smoking)	—	—	1*	1*	1*	3*

See table footnotes on page 61.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD	LNG-IUD	Implants	DMPA	POP	CHCs
ii. Without other risk factors for VTE	—	—	1*	1*	1*	2*
d. >42 days postpartum	—	—	1*	1*	1*	2*
Postpartum (nonbreastfeeding women)						
a. <21 days postpartum	—	—	1	1	1	4
b. 21–42 days postpartum						
i. With other risk factors for VTE (e.g., age ≥35 years, previous VTE, thrombophilia, immobility, transfusion at delivery, peripartum cardiomyopathy, BMI ≥30 kg/m ² , postpartum hemorrhage, postcesarean delivery, preeclampsia, or smoking)	—	—	1	1	1	3*
ii. Without other risk factors for VTE	—	—	1	1	1	2
c. >42 days postpartum	—	—	1	1	1	1
Postpartum (including cesarean delivery)						
a. <10 minutes after delivery of the placenta						
i. Breastfeeding	1*	2*	—	—	—	—
ii. Nonbreastfeeding	1*	1*	—	—	—	—
b. 10 minutes after delivery of the placenta to <4 weeks (breastfeeding or nonbreastfeeding)	2*	2*	—	—	—	—
c. ≥4 weeks (breastfeeding or nonbreastfeeding)	1*	1*	—	—	—	—
d. Postpartum sepsis	4	4	—	—	—	—
Postabortion						
a. First trimester	1*	1*	1*	1*	1*	1*
b. Second trimester	2*	2*	1*	1*	1*	1*
c. Immediate postseptic abortion	4	4	1*	1*	1*	1*
Past ectopic pregnancy	1	1	1	1	2	1
History of pelvic surgery (see Postpartum [Including Cesarean Delivery] section)	1	1	1	1	1	1
Smoking						
a. Age <35 years	1	1	1	1	1	2
b. Age ≥35 years						
i. <15 cigarettes/day	1	1	1	1	1	3
ii. ≥15 cigarettes/day	1	1	1	1	1	4
Obesity						
a. BMI ≥30 kg/m ²	1	1	1	1	1	2
b. Menarche to <18 years and BMI ≥30 kg/m ²	1	1	1	2	1	2
History of bariatric surgery This condition is associated with increased risk for adverse health events as a result of pregnancy.						
a. Restrictive procedures: decrease storage capacity of the stomach (vertical banded gastroplasty, laparoscopic adjustable gastric band, or laparoscopic sleeve gastrectomy)	1	1	1	1	1	1
b. Malabsorptive procedures: decrease absorption of nutrients and calories by shortening the functional length of the small intestine (Roux-en-Y gastric bypass or biliopancreatic diversion)	1	1	1	1	3	COCs: 3 Patch and ring: 1

See table footnotes on page 61.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD	LNG-IUD	Implants	DMPA	POP	CHCs
Cardiovascular Disease						
Multiple risk factors for atherosclerotic cardiovascular disease (e.g., older age, smoking, diabetes, hypertension, low HDL, high LDL, or high triglyceride levels)	1	2	2*	3*	2*	3/4*
Hypertension Systolic blood pressure \geq 160 mm Hg or diastolic blood pressure \geq 100 mm Hg are associated with increased risk for adverse health events as a result of pregnancy.						
a. Adequately controlled hypertension	1*	1*	1*	2*	1*	3*
b. Elevated blood pressure levels (properly taken measurements)						
i. Systolic 140–159 mm Hg or diastolic 90–99 mm Hg	1*	1*	1*	2*	1*	3*
ii. Systolic \geq 160 mm Hg or diastolic \geq 100 mm Hg	1*	2*	2*	3*	2*	4*
c. Vascular disease	1*	2*	2*	3*	2*	4*
History of high blood pressure during pregnancy (when current blood pressure is measurable and normal)	1	1	1	1	1	2
Deep venous thrombosis/ Pulmonary embolism						
a. History of DVT/PE, not receiving anticoagulant therapy						
i. Higher risk for recurrent DVT/PE (one or more risk factors)	1	2	2	2	2	4
• History of estrogen-associated DVT/PE						
• Pregnancy-associated DVT/PE						
• Idiopathic DVT/PE						
• Known thrombophilia, including antiphospholipid syndrome						
• Active cancer (metastatic, receiving therapy, or within 6 months after clinical remission), excluding nonmelanoma skin cancer						
• History of recurrent DVT/PE						
ii. Lower risk for recurrent DVT/PE (no risk factors)	1	2	2	2	2	3
b. Acute DVT/PE	2	2	2	2	2	4
c. DVT/PE and established receiving anticoagulant therapy for at least 3 months						
i. Higher risk for recurrent DVT/PE (one or more risk factors)	2	2	2	2	2	4*
• Known thrombophilia, including antiphospholipid syndrome						
• Active cancer (metastatic, receiving therapy, or within 6 months after clinical remission), excluding nonmelanoma skin cancer						
• History of recurrent DVT/PE						
ii. Lower risk for recurrent DVT/PE (no risk factors)	2	2	2	2	2	3*

See table footnotes on page 61.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD		LNG-IUD		Implants		DMPA		POP		CHCs
d. Family history (first-degree relatives)	1		1		1		1		1		2
e. Major surgery											
i. With prolonged immobilization	1		2		2		2		2		4
ii. Without prolonged immobilization	1		1		1		1		1		2
f. Minor surgery without immobilization	1		1		1		1		1		1
Known thrombogenic mutations (e.g., factor V Leiden; prothrombin mutation; and protein S, protein C, and antithrombin deficiencies)	1*		2*		2*		2*		2*		4*
This condition is associated with increased risk for adverse health events as a result of pregnancy.											
Superficial venous disorders											
a. Varicose veins	1		1		1		1		1		1
b. Superficial venous thrombosis (acute or history)	1		1		1		1		1		3*
Current and history of ischemic heart disease	1		Initiation 2	Continuation 3	Initiation 2	Continuation 3	3		Initiation 2	Continuation 3	4
This condition is associated with increased risk for adverse health events as a result of pregnancy.											
Stroke (history of cerebrovascular accident)	1				Initiation 2	Continuation 3	3		Initiation 2	Continuation 3	4
This condition is associated with increased risk for adverse health events as a result of pregnancy.											
Valvular heart disease											
Complicated valvular heart disease is associated with increased risk for adverse health events as a result of pregnancy.											
a. Uncomplicated	1		1		1		1		1		2
b. Complicated (pulmonary hypertension, risk for atrial fibrillation, or history of subacute bacterial endocarditis)	1		1		1		1		1		4
Peripartum cardiomyopathy											
This condition is associated with increased risk for adverse health events as a result of pregnancy.											
a. Normal or mildly impaired cardiac function (New York Heart Association Functional Class I or II; patients with no limitation of activities or patients with slight, mild limitation of activity) (2)											
i. <6 months	2		2		1		1		1		4
ii. ≥6 months	2		2		1		1		1		3
b. Moderately or severely impaired cardiac function (New York Heart Association Functional Class III or IV; patients with marked limitation of activity or patients who should be at complete rest) (2).	2		2		2		2		2		4
Rheumatic Diseases											
Systemic lupus erythematosus	Initiation	Continuation					Initiation	Continuation			
This condition is associated with increased risk for adverse health events as a result of pregnancy.											
a. Positive (or unknown) antiphospholipid antibodies	1*	1*	3*		3*		3*	3*	3*		4*

See table footnotes on page 61.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD		LNG-IUD		Implants	DMPA		POP	CHCs
b. Severe thrombocytopenia	3*	2*	2*	2*	2*	3*	2*	2*	2*
c. Immunosuppressive therapy	2*	1*	2*	2*	2*	2*	2*	2*	2*
d. None of the above	1*	1*	2*	2*	2*	2*	2*	2*	2*
Rheumatoid arthritis	Initiation	Continuation	Initiation	Continuation					
a. Receiving immunosuppressive therapy	2	1	2	1	1	2/3*		1	2
b. Not receiving immunosuppressive therapy		1		1	1	2		1	2
Neurologic Conditions									
Headaches									
a. Nonmigraine (mild or severe)		1		1	1		1	1	1*
b. Migraine									
i. Without aura (This category of migraine includes menstrual migraine.)		1		1	1		1	1	2*
ii. With aura		1		1	1		1	1	4*
Epilepsy		1		1	1*		1*	1*	1*
This condition is associated with increased risk for adverse health events as a result of pregnancy.									
Multiple sclerosis									
a. With prolonged immobility		1		1	1		2	1	3
b. Without prolonged immobility		1		1	1		2	1	1
Depressive Disorders									
Depressive disorders		1*		1*	1*		1*	1*	1*
Reproductive Tract Infections and Disorders									
Vaginal bleeding patterns									
			Initiation	Continuation					
a. Irregular pattern without heavy bleeding		1	1	1	2		2	2	1
b. Heavy or prolonged bleeding (includes regular and irregular patterns)		2*	1*	2*	2*		2*	2*	1*
Unexplained vaginal bleeding (suspicious for serious condition) before evaluation	Initiation	Continuation	Initiation	Continuation					
	4*	2*	4*	2*	3*		3*	2*	2*
Endometriosis		2		1	1		1	1	1
Benign ovarian tumors (including cysts)		1		1	1		1	1	1
Severe dysmenorrhea		2		1	1		1	1	1
Gestational trophoblastic disease									
This condition is associated with increased risk for adverse health events as a result of pregnancy.									
a. Suspected gestational trophoblastic disease (immediate postevacuation)									
i. Uterine size first trimester		1*		1*	1*		1*	1*	1*
ii. Uterine size second trimester		2*		2*	1*		1*	1*	1*
b. Confirmed gestational trophoblastic disease (after initial evacuation and during monitoring)									
i. Undetectable/nonpregnant β -hCG levels		1*		1*	1*		1*	1*	1*
ii. Decreasing β -hCG levels		2*		2*	1*		1*	1*	1*
iii. Persistently elevated β -hCG levels or malignant disease, with no evidence or suspicion of intrauterine disease		2*		2*	1*		1*	1*	1*

See table footnotes on page 61.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD		LNG-IUD		Implants	DMPA	POP	CHCs
iv. Persistently elevated β -hCG levels or malignant disease, with evidence or suspicion of intrauterine disease	4*	2*	4*	2*	1*	1*	1*	1*
Cervical ectropion	1		1		1	1	1	1
Cervical intraepithelial neoplasia	1		2		2	2	1	2
Cervical cancer (awaiting treatment)	Initiation 4	Continuation 2	Initiation 4	Continuation 2	2	2	1	2
Breast disease Breast cancer is associated with increased risk of adverse health events as a result of pregnancy.								
a. Undiagnosed mass	1		2		2*	2*	2*	2*
b. Benign breast disease	1		1		1	1	1	1
c. Family history of cancer	1		1		1	1	1	1
d. Breast cancer								
i. Current	1		4		4	4	4	4
ii. Past and no evidence of current disease for 5 years	1		3		3	3	3	3
Endometrial hyperplasia	1		1		1	1	1	1
Endometrial cancer This condition is associated with increased risk for adverse health events as a result of pregnancy.	Initiation 4	Continuation 2	Initiation 4	Continuation 2	1	1	1	1
Ovarian cancer This condition is associated with increased risk for adverse health events as a result of pregnancy.	1		1		1	1	1	1
Uterine fibroids	2		2		1	1	1	1
Anatomical abnormalities								
a. Distorted uterine cavity (any congenital or acquired uterine abnormality distorting the uterine cavity in a manner that is incompatible with IUD insertion)	4		4		—	—	—	—
b. Other abnormalities (including cervical stenosis or cervical lacerations) not distorting the uterine cavity or interfering with IUD insertion	2		2		—	—	—	—
Pelvic inflammatory disease								
a. Past PID (assuming no current risk factors for STDs)								
i. With subsequent pregnancy	1	1	1	1	1	1	1	1
ii. Without subsequent pregnancy	2	2	2	2	1	1	1	1
b. Current PID	4	2*	4	2*	1	1	1	1
Sexually transmitted diseases	Initiation	Continuation	Initiation	Continuation				
a. Current purulent cervicitis or chlamydial infection or gonococcal infection	4	2*	4	2*	1	1	1	1
b. Vaginitis (including <i>Trichomonas vaginalis</i> and bacterial vaginosis)	2	2	2	2	1	1	1	1
c. Other factors related to STDs	2*	2	2*	2	1	1	1	1
HIV								
High risk for HIV	Initiation	Continuation	Initiation	Continuation				
HIV infection For women with HIV infection who are not clinically well or not receiving ARV therapy, this condition is associated with increased risk for adverse health events as a result of pregnancy.	2	2	2	2	1	1*	1	1
a. Clinically well receiving ARV therapy	—	—	—	—	1*	1*	1*	1*
	1	1	1	1	—	—	—	—

See table footnotes on page 61.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD		LNG-IUD		Implants	DMPA	POP	CHCs		
b. Not clinically well or not receiving ARV therapy	2	1	2	1	—	—	—	—		
Other Infections										
Schistosomiasis										
Schistosomiasis with fibrosis of the liver is associated with increased risk for adverse health events as a result of pregnancy.										
a. Uncomplicated	1		1		1	1	1	1		
b. Fibrosis of the liver (if severe, see Cirrhosis)	1		1		1	1	1	1		
Tuberculosis										
	Initiation	Continuation	Initiation	Continuation						
This condition is associated with increased risk for adverse health events as a result of pregnancy.										
a. Nonpelvic	1	1	1	1	1*	1*	1*	1*		
b. Pelvic	4	3	4	3	1*	1*	1*	1*		
Malaria	1		1		1	1	1	1		
Endocrine Conditions										
Diabetes										
Insulin-dependent diabetes; diabetes with nephropathy, retinopathy, neuropathy, or diabetes with other vascular disease; or diabetes of >20 years' duration are associated with increased risk of adverse health events as a result of pregnancy.										
a. History of gestational disease	1		1		1	1	1	1		
b. Nonvascular disease										
i. Non-insulin dependent	1		2		2	2	2	2		
ii. Insulin dependent	1		2		2	2	2	2		
c. Nephropathy, retinopathy, or neuropathy	1		2		2	3	2	3/4*		
d. Other vascular disease or diabetes of >20 years' duration	1		2		2	3	2	3/4*		
Thyroid disorders										
a. Simple goiter	1		1		1	1	1	1		
b. Hyperthyroid	1		1		1	1	1	1		
c. Hypothyroid	1		1		1	1	1	1		
Gastrointestinal Conditions										
Inflammatory bowel disease (ulcerative colitis or Crohn's disease)	1		1		1	2	2	2/3*		
Gallbladder disease										
a. Symptomatic										
i. Treated by cholecystectomy	1		2		2	2	2	2		
ii. Medically treated	1		2		2	2	2	3		
iii. Current	1		2		2	2	2	3		
b. Asymptomatic	1		2		2	2	2	2		
History of cholestasis										
a. Pregnancy related	1		1		1	1	1	2		
b. Past COC related	1		2		2	2	2	3		
Viral hepatitis										
a. Acute or flare	1		1		1	1	1	Initiation	Continuation	
b. Carrier	1		1		1	1	1	3/4*	2	
c. Chronic	1		1		1	1	1	1	1	
Cirrhosis										
Severe cirrhosis is associated with increased risk for adverse health events as a result of pregnancy.										
a. Mild (compensated)	1		1		1	1	1	1		
b. Severe (decompensated)	1		3		3	3	3	4		
Liver tumors										
Hepatocellular adenoma and malignant liver tumors are associated with increased risk for adverse health events as a result of pregnancy.										
a. Benign										

See table footnotes on page 61.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD		LNG-IUD		Implants	DMPA	POP	CHCs
i. Focal nodular hyperplasia	1		2		2	2	2	2
ii. Hepatocellular adenoma	1		3		3	3	3	4
b. Malignant (hepatoma)	1		3		3	3	3	4
Respiratory Conditions								
Cystic fibrosis	1*		1*		1*	2*	1*	1*
This condition is associated with increased risk for adverse health events as a result of pregnancy.								
Anemias								
Thalassemia	2		1		1	1	1	1
Sickle cell disease	2		1		1	1	1	2
This condition is associated with increased risk for adverse health events as a result of pregnancy.								
Iron-deficiency anemia	2		1		1	1	1	1
Solid Organ Transplantation								
Solid organ transplantation	Initiation	Continuation	Initiation	Continuation				
This condition is associated with increased risk for adverse health events as a result of pregnancy.								
a. Complicated: graft failure (acute or chronic), rejection, or cardiac allograft vasculopathy	3	2	3	2	2	2	2	4
b. Uncomplicated	2		2		2	2	2	2*
Drug Interactions								
Antiretroviral therapy	Initiation	Continuation	Initiation	Continuation				
a. Nucleoside reverse transcriptase inhibitors (NRTIs)								
i. Abacavir (ABC)	1/2*	1*	1/2*	1*	1	1	1	1
ii. Tenofovir (TDF)	1/2*	1*	1/2*	1*	1	1	1	1
iii. Zidovudine (AZT)	1/2*	1*	1/2*	1*	1	1	1	1
iv. Lamivudine (3TC)	1/2*	1*	1/2*	1*	1	1	1	1
v. Didanosine (DDI)	1/2*	1*	1/2*	1*	1	1	1	1
vi. Emtricitabine (FTC)	1/2*	1*	1/2*	1*	1	1	1	1
vii. Stavudine (D4T)	1/2*	1*	1/2*	1*	1	1	1	1
b. Nonnucleoside reverse transcriptase inhibitors (NNRTIs)								
i. Efavirenz (EFV)	1/2*	1*	1/2*	1*	2*	1*	2*	2*
ii. Etravirine (ETR)	1/2*	1*	1/2*	1*	1	1	1	1
iii. Nevirapine (NVP)	1/2*	1*	1/2*	1*	1	1	1	1
iv. Rilpivirine (RPV)	1/2*	1*	1/2*	1*	1	1	1	1
c. Ritonavir-boosted protease inhibitors								
i. Ritonavir-boosted atazanavir (ATV/r)	1/2*	1*	1/2*	1*	2*	1*	2*	2*
ii. Ritonavir-boosted darunavir (DRV/r)	1/2*	1*	1/2*	1*	2*	1*	2*	2*
iii. Ritonavir-boosted foseprenavir (FPV/r)	1/2*	1*	1/2*	1*	2*	1*	2*	2*
iv. Ritonavir-boosted lopinavir (LPV/r)	1/2*	1*	1/2*	1*	1	1	1	1
v. Ritonavir-boosted saquinavir (SQV/r)	1/2*	1*	1/2*	1*	2*	1*	2*	2*
vi. Ritonavir-boosted tipranavir (TPV/r)	1/2*	1*	1/2*	1*	2*	1*	2*	2*
d. Protease inhibitors without ritonavir								
i. Atazanavir (ATV)	1/2*	1*	1/2*	1*	1	1	1	2*
ii. Fosamprenavir (FPV)	1/2*	1*	1/2*	1*	2*	2*	2*	3*
iii. Indinavir (IDV)	1/2*	1*	1/2*	1*	1	1	1	1
iv. Nelfinavir (NFV)	1/2*	1*	1/2*	1*	2*	1*	2*	2*
e. CCR5 co-receptor antagonists								
i. Maraviroc (MVC)	1/2*	1*	1/2*	1*	1	1	1	1
f. HIV integrase strand transfer inhibitors								
i. Raltegravir (RAL)	1/2*	1*	1/2*	1*	1	1	1	1
ii. Dolutegravir (DTG)	1/2*	1*	1/2*	1*	1	1	1	1
iii. Elvitegravir (EVG)	1/2*	1*	1/2*	1*	1	1	1	1

See table footnotes on next page.

TABLE A1. (Continued) Summary of classifications for hormonal contraceptive methods and intrauterine devices

Condition	Cu-IUD		LNG-IUD		Implants	DMPA	POP	CHCs
g. Fusion inhibitors								
i. Enfuvirtide	1/2*	1*	1/2*	1*	1	1	1	1
Anticonvulsant therapy								
a. Certain anticonvulsants (phenytoin, carbamazepine, barbiturates, primidone, topiramate, and oxcarbazepine)	1		1		2*	1*	3*	3*
b. Lamotrigine	1		1		1	1	1	3*
Antimicrobial therapy								
a. Broad-spectrum antibiotics	1		1		1	1	1	1
b. Antifungals	1		1		1	1	1	1
c. Antiparasitics	1		1		1	1	1	1
d. Rifampin or rifabutin therapy	1		1		2*	1*	3*	3*
Psychotropic medications								
a. SSRIs	1		1		1	1	1	1
St. John's wort	1		1		2	1	2	2

Abbreviations: BMI = body mass index; COC = combined oral contraceptive; Cu-IUD = copper-containing IUD; DMPA = depot medroxyprogesterone acetate; DVT = deep venous thrombosis; hCG = human chorionic gonadotropin; HDL = high-density lipoprotein; HIV = human immunodeficiency virus; IUD = intrauterine device; LDL = low-density lipoprotein; LNG-IUD = levonorgestrel-releasing IUD; NA = not applicable; PE = pulmonary embolism; PID = pelvic inflammatory disease; POP = progestin-only pill; SSRI = selective serotonin reuptake inhibitor; STD = sexually transmitted disease.

* Consult the respective appendix for each contraceptive method in the 2016 *U.S. Medical Eligibility Criteria for Contraceptive Use (1)* for clarifications to the numeric categories.

References

1. Curtis KM, Tepper NK, Jatlaoui TC, et al. U.S. medical eligibility criteria for contraceptive use, 2016. *MMWR Recomm Rep* 2016;65(No. RR-3).
2. The Criteria Committee of the New York Heart Association. *Nomenclature and criteria for diagnosis of diseases of the heart and great vessels*. 9th ed. Boston, MA: Little, Brown & Co; 1994.

Appendix B

When To Start Using Specific Contraceptive Methods

Contraceptive method	When to start (if the provider is reasonably certain that the woman is not pregnant)	Additional contraception (i.e., back-up) needed	Examinations or tests needed before initiation*
Copper-containing IUD	Anytime	Not needed	Bimanual examination and cervical inspection [†]
Levonorgestrel-releasing IUD	Anytime	If >7 days after menses started, use back-up method or abstain for 7 days.	Bimanual examination and cervical inspection [†]
Implant	Anytime	If >5 days after menses started, use back-up method or abstain for 7 days.	None
Injectable	Anytime	If >7 days after menses started, use back-up method or abstain for 7 days.	None
Combined hormonal contraceptive	Anytime	If >5 days after menses started, use back-up method or abstain for 7 days.	Blood pressure measurement
Progestin-only pill	Anytime	If >5 days after menses started, use back-up method or abstain for 2 days.	None

Abbreviations: BMI = body mass index; HIV = human immunodeficiency virus; IUD = intrauterine device; STD = sexually transmitted disease; U.S. MEC = *U.S. Medical Eligibility Criteria for Contraceptive Use*.

* Weight (BMI) measurement is not needed to determine medical eligibility for any methods of contraception because all methods can be used (U.S. MEC 1) or generally can be used (U.S. MEC 2) among obese women (Box 1). However, measuring weight and calculating BMI (weight [kg] / height [m]²) at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

[†] Most women do not require additional STD screening at the time of IUD insertion. If a woman with risk factors for STDs has not been screened for gonorrhea and chlamydia according to CDC's *STD Treatment Guidelines* (<http://www.cdc.gov/std/treatment>), screening can be performed at the time of IUD insertion, and insertion should not be delayed. Women with current purulent cervicitis or chlamydial infection or gonococcal infection should not undergo IUD insertion (U.S. MEC 4).

Appendix C

Examinations and Tests Needed Before Initiation of Contraceptive Methods

The examinations or tests noted apply to women who are presumed to be healthy (Table C1). Those with known medical problems or other special conditions might need additional examinations or tests before being determined to be appropriate candidates for a particular method of contraception. The 2016 *U.S. Medical Eligibility Criteria for Contraceptive Use* (U.S. MEC) might be useful in such circumstances (1). The following classification was considered useful in differentiating the applicability of the various examinations or tests:

- **Class A:** essential and mandatory in all circumstances for safe and effective use of the contraceptive method.
- **Class B:** contributes substantially to safe and effective use, but implementation may be considered within the public health and/or service context; risk of not performing an examination or test should be balanced against the benefits of making the contraceptive method available.
- **Class C:** does not contribute substantially to safe and effective use of the contraceptive method.

These classifications focus on the relationship of the examinations or tests to safe initiation of a contraceptive method. They are not intended to address the appropriateness of these examinations or tests in other circumstances. For example, some of the examinations or tests that are not deemed necessary for safe and effective contraceptive use might be appropriate for good preventive health care or for diagnosing or assessing suspected medical conditions. Any additional screening needed for preventive health care can be performed at the time of contraception initiation and initiation should not be delayed for test results.

No examinations or tests are needed before initiating condoms or spermicides. A bimanual examination is necessary for diaphragm fitting. A bimanual examination and cervical inspection are needed for cervical cap fitting.

References

1. Curtis KM, Tepper NK, Jatlaoui TC, et al. U.S. medical eligibility criteria for contraceptive use, 2016. *MMWR Recomm Rep* 2016;65(No. RR-3).

TABLE C1. Examinations and tests needed before initiation of contraceptive methods

Examination or test	Contraceptive method and class							
	Cu-IUD and LNG-IUD	Implant	Injectable	CHC	POP	Condom	Diaphragm or cervical cap	Spermicide
Examination								
Blood pressure	C	C	C	A*	C	C	C	C
Weight (BMI) (weight [kg] / height [m] ²)	—†	—†	—†	—†	—†	C	C	C
Clinical breast examination	C	C	C	C	C	C	C	C
Bimanual examination and cervical inspection	A	C	C	C	C	C	A [§]	C
Laboratory test								
Glucose	C	C	C	C	C	C	C	C
Lipids	C	C	C	C	C	C	C	C
Liver enzymes	C	C	C	C	C	C	C	C
Hemoglobin	C	C	C	C	C	C	C	C
Thrombogenic mutations	C	C	C	C	C	C	C	C
Cervical cytology (Papanicolaou test)	C	C	C	C	C	C	C	C
STD screening with laboratory tests	—¶	C	C	C	C	C	C	C
HIV screening with laboratory tests	C	C	C	C	C	C	C	C

Abbreviations: BMI = body mass index; CHC = combined hormonal contraceptive; Cu-IUD = copper-containing intrauterine device; HIV = human immunodeficiency virus; LNG-IUD = levonorgestrel-releasing intrauterine device; POP = progestin-only pill; STD = sexually transmitted disease; U.S. MEC = *U.S. Medical Eligibility Criteria for Contraceptive Use*.

* In instances in which blood pressure cannot be measured by a provider, blood pressure measured in other settings can be reported by the woman to her provider.
 † Weight (BMI) measurement is not needed to determine medical eligibility for any methods of contraception because all methods can be used (U.S. MEC 1) or generally can be used (U.S. MEC 2) among obese women (Box 1). However, measuring weight and calculating BMI at baseline might be helpful for monitoring any changes and counseling women who might be concerned about weight change perceived to be associated with their contraceptive method.

§ A bimanual examination (not cervical inspection) is needed for diaphragm fitting.

¶ Most women do not require additional STD screening at the time of IUD insertion. If a woman with risk factors for STDs has not been screened for gonorrhea and chlamydia according to CDC's *STD Treatment Guidelines* (<http://www.cdc.gov/std/treatment>), screening can be performed at the time of IUD insertion, and insertion should not be delayed. Women with current purulent cervicitis or chlamydial infection or gonococcal infection should not undergo IUD insertion (U.S. MEC 4).

Appendix D

Routine Follow-Up After Contraceptive Initiation

These recommendations address when routine follow-up is recommended for safe and effective continued use of contraception for healthy women (Table D1). The recommendations refer to general situations and might

vary for different users and different situations. Specific populations who might benefit from frequent follow-up visits include adolescents, those with certain medical conditions or characteristics, and those with multiple medical conditions.

TABLE D1. Routine follow-up after contraceptive initiation

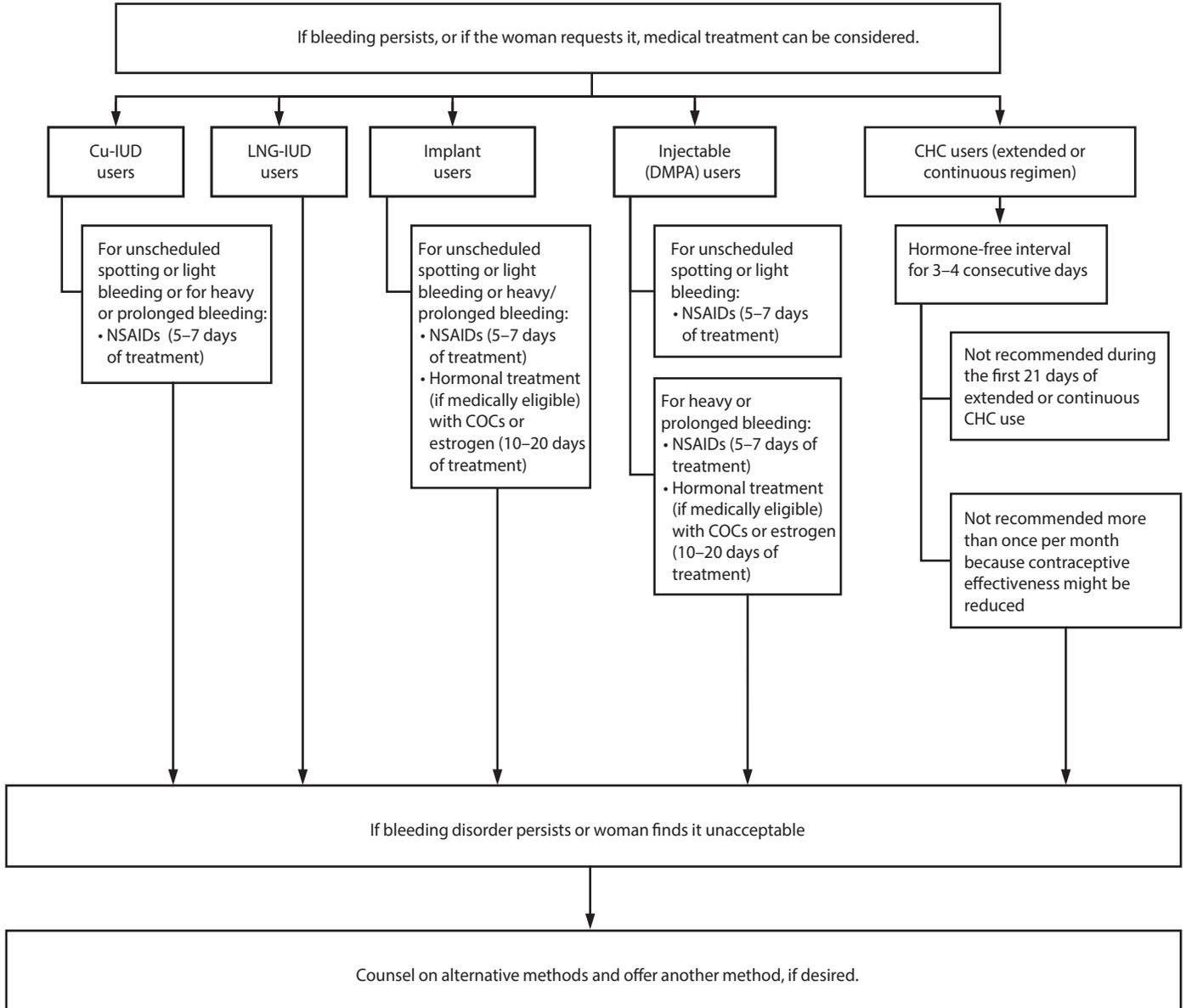
Action	Contraceptive method				
	Cu-IUD or LNG-IUD	Implant	Injectable	CHC	POP
General follow-up					
Advise women to return at any time to discuss side effects or other problems or if they want to change the method. Advise women using IUDs, implants, or injectables when the IUD or implant needs to be removed or when a reinjection is needed. No routine follow-up visit is required.	X	X	X	X	X
Other routine visits					
Assess the woman's satisfaction with her current method and whether she has any concerns about method use.	X	X	X	X	X
Assess any changes in health status, including medications, that would change the method's appropriateness for safe and effective continued use based on U.S. MEC (i.e., category 3 and 4 conditions and characteristics) (Box 1).	X	X	X	X	X
Consider performing an examination to check for the presence of IUD strings.	X	—	—	—	—
Consider assessing weight changes and counseling women who are concerned about weight change perceived to be associated with their contraceptive method.	X	X	X	X	X
Measure blood pressure.	—	—	—	X	—

Abbreviations: CHC = combined hormonal contraceptives; Cu-IUD = copper-containing intrauterine device; HIV = human immunodeficiency virus; IUD = intrauterine device; LNG-IUD = levonorgestrel-releasing intrauterine device; POP = progestin-only pills; U.S. MEC = *U.S. Medical Eligibility Criteria for Contraceptive Use*.

Appendix E

Management of Women with Bleeding Irregularities While Using Contraception*

Management of women with bleeding irregularities while using contraception



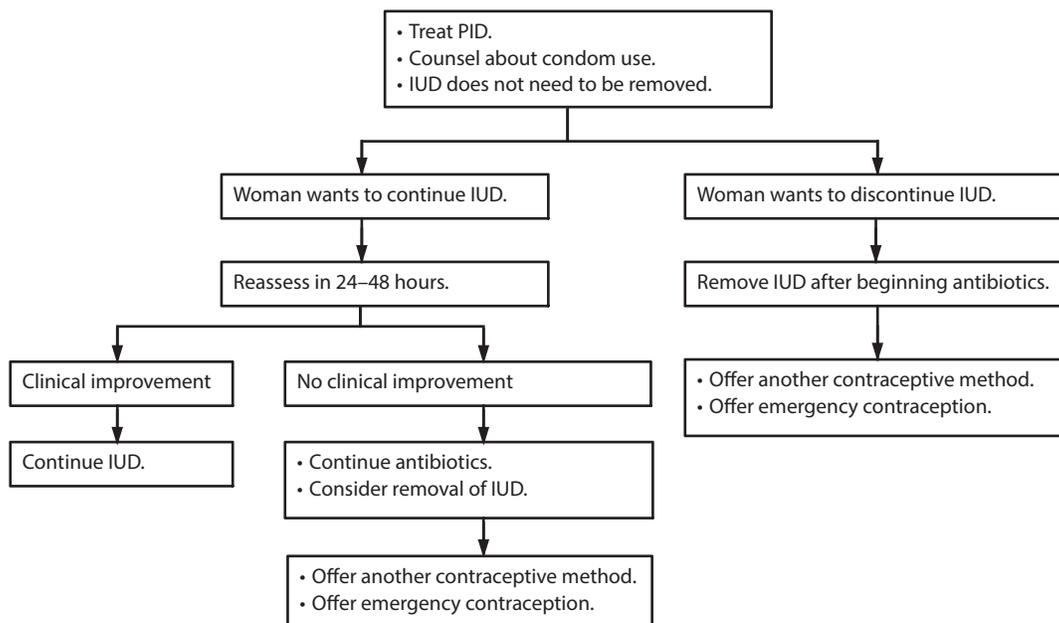
Abbreviations: CHC = combined hormonal contraceptive; COC = combined oral contraceptive; Cu-IUD = copper-containing intrauterine device; DMPA = depot medroxyprogesterone acetate; LNG-IUD = levonorgestrel-releasing intrauterine device; NSAIDs = nonsteroidal antiinflammatory drugs.

* If clinically warranted, evaluate for underlying condition. Treat the condition or refer for care. Heavy or prolonged bleeding, either unscheduled or menstrual, is uncommon among LNG-IUD users and implant users.

Appendix F

Management of Intrauterine Devices When Users are Found To Have Pelvic Inflammatory Disease*

Management of intrauterine devices when users of copper-containing intrauterine devices or levonorgestrel-releasing intrauterine devices are found to have pelvic inflammatory disease



Abbreviations: IUD = intrauterine device; PID = pelvic inflammatory disease.

*Treat according to the CDC *Sexually Transmitted Diseases Treatment Guidelines* (<http://www.cdc.gov/std/treatment>).

The *Morbidity and Mortality Weekly Report (MMWR)* Series is prepared by the Centers for Disease Control and Prevention (CDC) and is available free of charge in electronic format. To receive an electronic copy each week, visit *MMWR*'s free subscription page at <http://www.cdc.gov/mmwr/mmwrsubscribe.html>. Paper copy subscriptions are available through the Superintendent of Documents, U.S. Government Printing Office, Washington, DC 20402; telephone 202-512-1800.

Readers who have difficulty accessing this PDF file may access the HTML file at http://www.cdc.gov/mmwr/volumes/65/rr/rr6504a1.htm?s_cid=rr6504a1_w. Address all inquiries about the *MMWR* Series, including material to be considered for publication, to Executive Editor, *MMWR* Series, Mailstop E-90, CDC, 1600 Clifton Rd., N.E., Atlanta, GA 30329-4027 or to mmwrq@cdc.gov.

All material in the *MMWR* Series is in the public domain and may be used and reprinted without permission; citation as to source, however, is appreciated.

Use of trade names and commercial sources is for identification only and does not imply endorsement by the U.S. Department of Health and Human Services.

References to non-CDC sites on the Internet are provided as a service to *MMWR* readers and do not constitute or imply endorsement of these organizations or their programs by CDC or the U.S. Department of Health and Human Services. CDC is not responsible for the content of these sites. URL addresses listed in *MMWR* were current as of the date of publication.

ISSN: 1057-5987 (Print)