

ACCELERATING ACCESS TO POSTPARTUM FAMILY PLANNING

CHIANG-MAI, THAILAND
JUNE 8 -11



IN TECHNICAL
PARTNERSHIP
WITH



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innovating to save lives

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New Methods added to the MEC, 5th Edition

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Methods of contraception included in MEC, 4th Edition

- ❑ Combined oral contraceptives
- ❑ Combined hormonal contraceptives (1 month injectables, patch, vaginal ring)
- ❑ Progestogen-only contraceptives (pills, implants, 2-3 month injectables)
- ❑ Emergency contraceptive pills
- ❑ IUDs (copper bearing and levonorgestrel)
- ❑ Emergency IUD
- ❑ Barrier methods (condoms, spermicides & diaphragm)
- ❑ Fertility awareness-based methods
- ❑ Lactational amenorrhoea (LAM)
- ❑ Coitus Interruptus
- ❑ Sterilization (male and female)

New methods added to MEC 5th edition

- ❑ Subcutaneously-administered depot medroxyprogesterone acetate (DMPA-SC)
- ❑ Sino implant (II)
- ❑ Ulipristal acetate (UPA)
- ❑ Progesterone-releasing vaginal ring (PVR)

**SUBCUTANEOUSLY -ADMINISTERED
DEPOT MEDROXYPROGESTERONE
ACETATE (DMPA-SC)**

DMPA-SC

- Depot medroxyprogesterone acetate delivered subcutaneously (DMPA SC) at dose of 104 mg/0.65 mL
- Important service delivery implications:
 - Self-administration feasible and acceptable to both adolescents and adults

DMPA-SC: Summary of Evidence

- ❑ **Obesity (8 studies)**
 - Contraceptive efficacy, weight change and other adverse events similar across BMI groups
- ❑ **Age (3 studies)**
 - Declines in BMD, no differences in weight change or bleeding patterns by age group
- ❑ **Endometriosis (2 studies)**
 - Declines in BMD, few serious adverse events
- ❑ **HIV (1 study)**
 - No difference in serious adverse events (SC vs. IM)

- ❑ Among **healthy women (3 studies)**, evidence suggests DMPA-SC may be similar to DMPA-IM

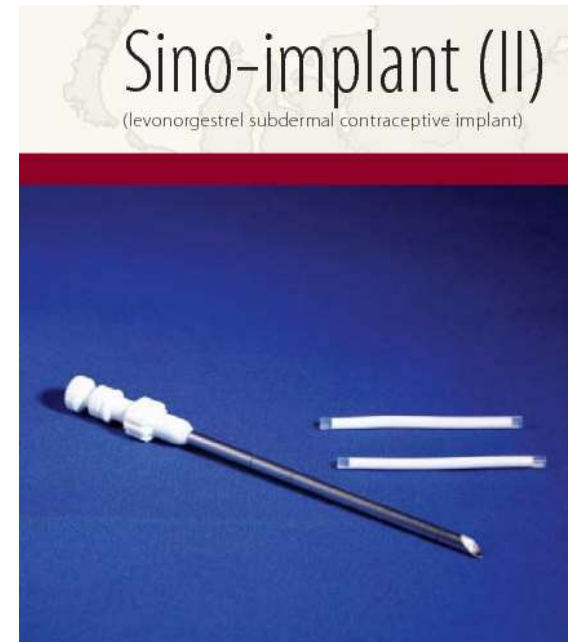
DMPA-SC

- ❑ No change in existing MEC recommendations for DMPA warranted with inclusion of DMPA-SC as a new method.

SINO-IMPLANT (II)

Background: Sino-implant (II)

- ❑ 2-rod, 150 mg LNG implant manufactured in China
- ❑ Currently used by millions of women worldwide (mostly Indonesia & China)



Sino-implant (II)

- ❑ No studies provided direct evidence for women with medical conditions
- ❑ Three RCTs comparing serious adverse events and other markers for disease in healthy women using Sino-implant (II) versus Norplant or Sino-implant (I)
- ❑ Noted similar safety profile

Sino-implant (II)

- ❑ No change in existing MEC recommendations for progestogen-only Implants warranted with inclusion of Sino-implant (II) as a new method.

ULIPRISTAL ACETATE

Ulipristal Acetate (UPA)

- ❑ Selective progesterone receptor modulator
- ❑ Single dose 30mg effective up to 120 hours
- ❑ Delays ovulation
- ❑ Approved in EU in 2009, US in 2010
- ❑ Currently registered in 72 countries

EMERGENCY CONTRACEPTIVE PILLS (ECPs)
 (including levonorgestrel contraceptive pills and combined oral contraceptive pills)

ECPs do not protect against STI/HIV. If there is risk of STI/HIV (including during pregnancy or postpartum), the correct and consistent use of condoms is recommended, either alone or with another contraceptive method. Male latex condoms are proven to protect against STI/HIV.

CONDITION * additional comments at end of table	CATEGORY	CLARIFICATIONS/EVIDENCE
PREGNANCY	NA	NA = not applicable Clarification: Although this method is not indicated for a woman with a known or suspected pregnancy, there is no known harm to the woman, the course of her pregnancy, or the fetus if ECPs are accidentally used.
BREASTFEEDING	1	
PAST ECTOPIC PREGNANCY	1	
HISTORY OF SEVERE CARDIOVASCULAR COMPLICATIONS* (ischaemic heart disease, cerebrovascular attack, or other thromboembolic conditions)	2	
ANGINA PECTORIS*	2	
MIGRAINE*	2	
SEVERE LIVER DISEASE* (including jaundice)	2	
REPEATED ECP USE	1	Clarification: Recurrence requires further counselling. Repeated ECP use may be considered as 2, 3 or 4 for CHC.
RAPE*	1	

MEC, 4th Edition ECP Recommendations

ADDITIONAL COMMENTS

HISTORY OF SEVERE CARDIOVASCULAR COMPLICATIONS

The duration of use of ECPs is less than that of regular use of COCs or POPs and thus would be expected to have less clinical impact.

ANGINA PECTORIS

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MIGRAINE

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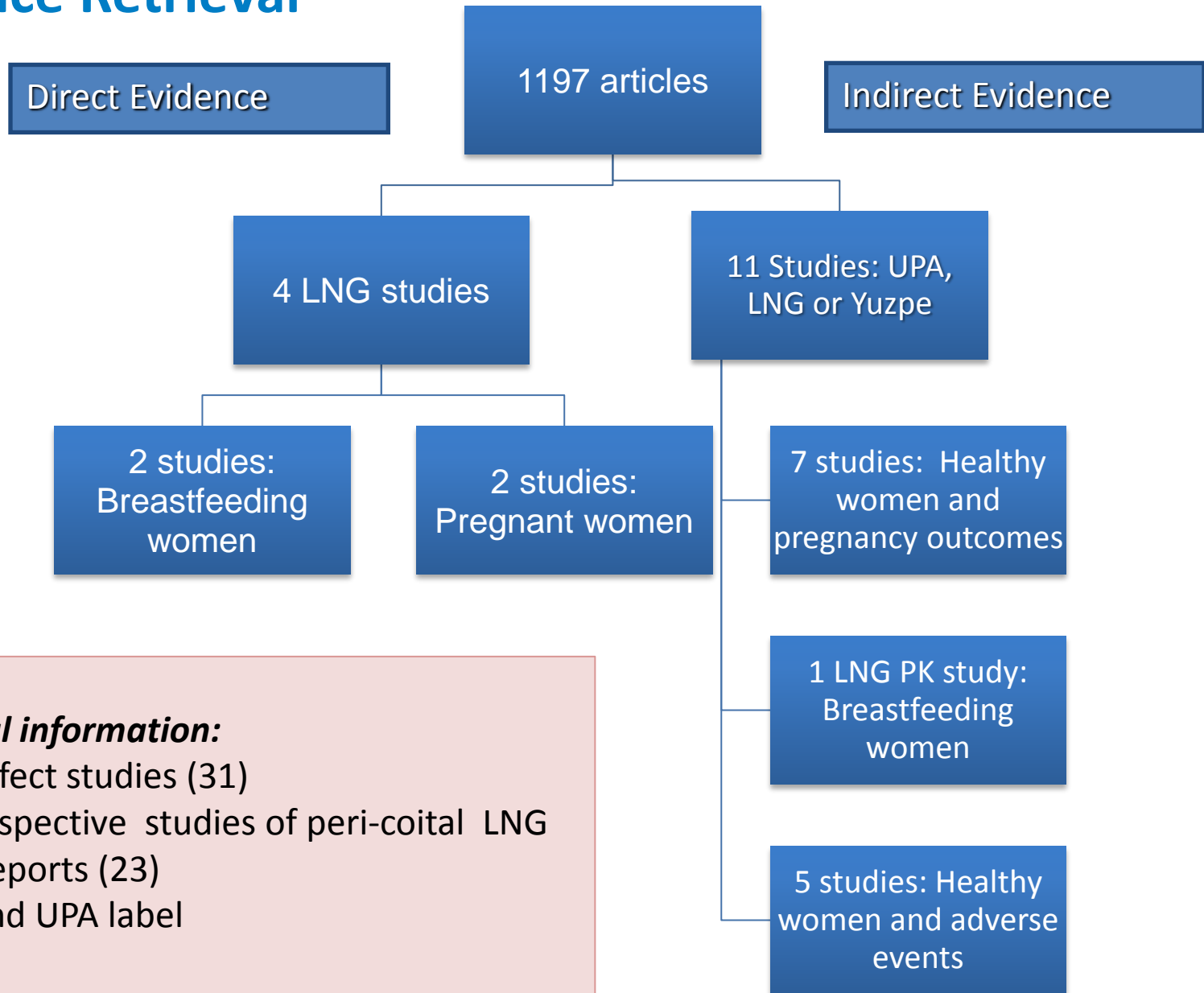
SEVERE LIVER DISEASE (INCLUDING JAUNDICE)

The duration of use of ECPs is less than that of regular use of COCs or POPs and thus would be expected to have less clinical impact.

RAPE

There are no restrictions for the use of ECPs in cases of rape.

Evidence Retrieval



Additional information:

- Side effect studies (31)
- Six prospective studies of peri-coital LNG
- Case reports (23)
- LNG and UPA label

NEW ECP Recommendations

Condition	COC	LNG	UPA
Pregnancy	NA	NA	NA
Breastfeeding	1	1	2
Past ectopic pregnancy	1	1	1
Obesity	1	1	1
History of severe cardiovascular disease	2	2	2
Migraine	2	2	2
Severe liver disease (including jaundice)	2	2	2
CYP3A4 inducers	1	1	1
Repeated emergency contraceptive pill use	1	1	1
Rape	1	1	1

COC= combined oral contraceptives; LNG = levonorgestrel; UPA = ulipristal acetate;
 NA = not applicable

ECPs and Breastfeeding

Condition	COC	LNG	UPA
Breastfeeding	1	1	2

Clarification: Breastfeeding is not recommended for 1 week after taking UPA since it is excreted in breast-milk. Breast-milk should be expressed and discarded during that time.

ECPs and Obesity

Condition	COC	LNG	UPA
Obesity	1	1	1

Clarification: ECPs may be less effective among women with BMI ≥ 30 kg/m² than among women with BMI < 25 kg/m². Despite this, there are no safety concerns.

Evidence: There is limited evidence from one study that suggests obese women with BMI ≥ 30 kg/m² experience an increased risk of pregnancy after use of LNG compared with women with BMI < 25 kg/m². Two studies suggest obese women may also experience an increased risk of pregnancy after use of UPA compared with non-obese women, though this increase was not significant in one study.

ECPs and CYP3A4 INDUCERS

Condition	COC	LNG	UPA
CYP3A4 INDUCERS (E.g. rifampicin, phenytoin, phenobarbital, carbamazepine, efavirenz, fosphenytoin, nevirapine, oxcarbazepine, primidone, rifabutin, st john's wort/hypericum perforatum)	1	1	1

Clarification: Strong CYP3A4 inducers may reduce the effectiveness of ECPs.

Evidence: According to labelling information, rifampicin markedly decreases UPA levels by 90% or more which may decrease its efficacy. Theoretical concerns therefore extend to use of other CYP3A4 inducers as well as to COC and LNG ECPs, which have similar metabolic pathways to UPA. A small pharmacokinetic study found that concomitant efavirenz use decreased LNG levels in women taking LNG ECP (0.75 mg) by 56% compared with LNG ECP alone.

PROGESTERONE-RELEASING VAGINAL RING (PVR)

Progesterone- Releasing Vaginal Ring (PVR)



- Matrix design
- Diameter 58 mm/ Cross-section 8.4 mm
- Delivers 10mg/day over 3 months
- Serum levels of 10-20 nmol/L [\sim 7ng/ml]
- Specifically designed for breastfeeding

- Registered in 9 Latin American countries -
Acceptability studies underway in Africa

NEW Recommendations for PVR

Condition	Category
Pregnancy	NA

Clarification: Use of PVRs is not required. There is no known harm to the woman, the course of her pregnancy, or the fetus if PVRs are accidentally used during pregnancy.

NEW Recommendations for PVR

Condition	Category
Breastfeeding \geq 4 weeks	1

Clarification: The woman must be actively breastfeeding (i.e. at least 4 breastfeeding episodes per day) during PVR use to maintain efficacy.

Evidence: No differences were observed between various measures of breastfeeding performance among PVR users compared with users of non-hormonal or progestogen-only (synthetic progesterone) contraceptives during 12 months of observation. No statistically significant differences in infant weight gain were observed among PVR users compared with women using a non-hormonal or progestogen-only contraceptives, and similar patterns of infant weight gain were observed in another study that compared PVR and IUD users. One study reported no significant difference in infant health.

Thank you!

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