Combined hormonal contraception

(From FSRH Guideline, 2019)



What is CHC (combined hormonal contraception)?

- Contains an oestrogen and a progestogen. These work synergistically to prevent ovulation, render cervical mucus hostile to sperm and keep endometrium thin and stable. Preparations differ in type/dose of oestrogen/progestogen.
- Includes the combined pill, patch and vaginal ring.
- Oestrogen: most CHC contains synthetic ethinyl oestradiol, but some contains mestranol, estetrol or oestradiol.
- Progestogen: classified according to 'generation' which reflects introduction onto the CHC market (see table below).
 Newer generation progestogens were developed to have fewer androgenic/glucocorticoid effects and so may be better tolerated BUT studies suggest they may incur a higher VTE risk.

Benefits of CHC

- · Light, predictable bleeding patterns.
- Less dysmenorrhea.
- Manages symptoms of:
 - 。 PMS.
 - PCOS.
 - Endometriosis (use continuous regime).
 - Menopause (can use up to 50y).
- Reduces risk of endometrial, ovarian, colorectal cancer.

Progestogens			
1 st generation	Norethisterone		
2 nd generation	Levonorgestrel		
3 rd generation	Desogestrel		
	Gestodene		
	Etonogestrel		
	Norgestimate (sometimes considered 2 nd		
	generation as metabolised to levonorgestrel)		
	Norelgestomin		
4 th generation	Drospirenone		
	Dienogest		
	Nomogestrel		

Efficacy of CHC

- CHC is not as effective as long-acting methods: typical failure rate is 9% in first year of use!
- Highly user-dependent! Remember to follow missed pill rules if vomits within 3h of taking/has diarrhoea >24h.
- No additional precautions needed if taking non-enzyme-inducing antibiotics unless severe vomiting or diarrhoea.

Contraindications and cautions

Common absolute contraindications (UKMEC 4)

- BP ≥160/100.
- Current breast cancer.
- Smoking ≥15 cigarettes/d AND ≥35y.
- Current vascular disease.
- History of VTE or known thrombogenic mutation (e.g. factor 5 Leiden, prothrombin mutation, protein S or C, antithrombin deficiencies).
- Migraine with aura.

Cautions to use

- BMI ≥35 is UKMEC 3 (relative contraindication) (CHC efficacy unaffected by weight, except for patch if >90kg).
- Patients on teratogens should be on a LARC (as CHC not effective enough).
- Avoid CHC if taking enzyme inducers or lamotrigine (can lower seizure threshold).
- Avoid pills in malabsorptive state (e.g. post-bariatric surgery).

Risks (small but greater than with other methods)

Counselling about VTE

- Annual risk of developing VTE $\approx 2/10~000$; this increases around 3x on CHC (but much lower than in pregnancy).
- 1st/2nd generation formulations appear lower risk than 3rd/4th generation.
- Annual risk of death from VTE on CHC $\approx 1-2/100~000$ which is MUCH lower than other lifestyle risks, e.g. driving a car (80/100 000) (Contraception Today 7th Ed, J Guillebaud, Informa healthcare 2012).

Other (small) risks

- Arterial thrombosis (higher risk with higher oestrogen doses).
- Breast and cervical cancer (risk reduces after stopping).

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Always check <u>www.fsrh.org/ukmec</u> if unsure whether a medical condition or lifestyle factor restricts CHC use.

Starting combined hormonal contraception

(From FSRH Guideline, 2019)



Regimes

Traditional (licensed) regimes: involve 21d of pill-taking then a 7d break.

Tailored or extended regimes: recommended by FSRH as involve fewer, shorter or no hormone-free intervals, and therefore potentially less bleeding, fewer side-effects and lower risk of contraceptive failure. Note: the patch and ring may be used in the same way, but phasic pill preparations should NOT.

- Scheduled extended cycles (bi/tricycling): take 2–3 packs consecutively followed by a 4–7d hormone-free interval.
- Unscheduled extended cycle: take pills continuously for >21d, after which, when/if breakthrough bleeding occurs, take a 4–7d break.
- Continuous pill-taking: take a pill daily whether bleeding occurs or not.

Before starting CHC

Before prescribing:

- Check medical eligibility (medical history, lifestyle factors, drug and family history).
- · Measure and record BP and BMI.
- Counsel regarding efficacy, risks, benefits, side-effects and regimes...

Which formulation and how much?

- Prescribe ≤30mcg pill with a 1st or 2nd generation progestogen first line.
- Consider an alternative formulation if likely to be more suitable based on patient experience, tolerance or preference.
- Give up to 12m supply (except vaginal ring: prescribe only 3m due to shelf life).

Starting instructions

If no risk of pregnancy, start CHC immediately:

Up to day 5 of natural menstrual cycle: can be started without using extra precautions.

- After day 5 of menstrual cycle: use extra precautions for 7d.
- If swapping from alternative form of contraception: use extra precautions for 7d if the previous method was not antiovulant (implant, depot, POP).

After oral emergency contraception:

- **Levonorgestrel emergency contraception:** start immediately after giving levonorgestrel, use extra precautions for 7d and do a pregnancy test at 3w.
- **Ulipristal emergency contraception: do NOT start CHC until 5d after giving ulipristal,** then use extra precautions for a further 7d and do a pregnancy test at 3w.
 - EXCEPTION: if an established CHC user misses 2–7d of pills/patch/ring in the first week after the hormone-free interval, we can give ulipristal as emergency contraception and she can restart CHC immediately. She should use extra precautions for 7 days and do a pregnancy test at 3w.

Review annually

- Check for compliance and any problems.
- Reassess eligibility (including BP, BMI and smoking status).
- Continue up to 50y if acceptable.
- Stopping and starting the pill every few months/years to have a 'pill holiday' is not recommended and may actually increase overall risk of VTE.

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CHC: troubleshooting and missed pills

(From FSRH Guideline, 2019)



Troubleshooting tips

Breakthrough bleeding

- · Common! Usually settles in 3-4m.
- · Higher oestrogen preparations may be associated with less bleeding.
- 2nd generation formulations may be better than 1st generation.
- Extended or continuous regimes may be better.
- Consider other causes (e.g. STIs, pregnancy, compliance, cervical/uterine pathology).

Headache

- No optimal preparation recommended.
- Extended/continuous regimes may be better if occurs in hormone-free interval.

Mood changes

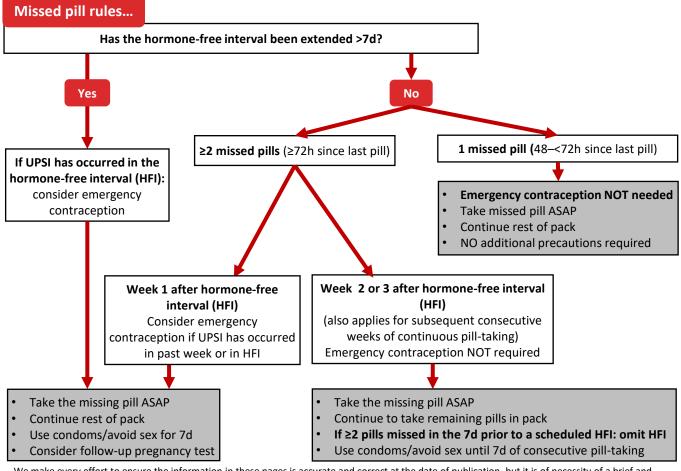
- · Counsel that may be multifactorial.
- There is no clear evidence that CHC causes depression.
- Consider switching to a formulation with a different progestogen.
- Extended/continuous regimes may be better if symptoms occur in hormone-free interval.

Weight gain

- There is no known causal association between CHC and weight gain.
- There is no evidence that different formulations impact weight differently.

Low libido

Evidence contradictory; overall, no clear effect of CHC on libido. Try different formulation or method of contraception.



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Combined hormonal contraception: formulations

(From MIMS 2022) All costs for 3 cycles



20mcg ethinyl oestradiol monophasic pills

Progestogen	Name	Cost		
Third generation				
Desogestrel 150mg	Bimizza	£5.04		
	Gedarel 20	£5.08		
	Mercilon	£8.44		
Gestodene 75mcg	Femodette	£8.85		
	Millinette 20	£5.41		
	Sunya	£6.62		
Fourth generation				
Drospirenone 3mg	Eloine	£14.70		

35mcg equivalent monophasic pills

Vicinity of the second			
Oestrogen	Progestogen	Name	Cost
35mcg ethinyl	First generation		
oestradiol	Norethisterone 500mcg	Brevinor	£1.99
	Norethisterone 1mg	Norimin	£2.28
	Third generation		
	Norgestimate 250mcg	Cilique	£4.65
		Lizinna	£4.64
Mestranol	First generation		
50mcg*	Norethisterone 1mg	Norinyl -1	£2.19
* Equivalent to 35mcg ethinyl oestradiol			

30mcg ethinyl oestradiol monophasic			
Progestogen	pills Name	Cost	
Second generation			
Levonorgestrel	Ambelina	£2.60	
150mcg	Elevin	£29.25	
	Levest	£1.80	
	Maexeni	£1.88	
	Microgynon (ED)	£2.82 (£2.99)	
	Ovranette	£2.20	
	Rigevidon	£1.89	
Third generation			
Desogestrel	Cimizt	£3.80	
150mcg	Gedarel 30	£4.19	
	Marvelon	£7.10	
Gestodene 75mcg	Femodene (ED)	£6.73 (£7.10)	
	Katya	£5.03	
	Millinette 30	£4.12	
Fourth generation		•	
Drospirenone 3mg	Dretine	£8.34	
	Lucette	£9.35	
	Yacella	£8.30	
	Yasmin	£14.70	
	Yiznell	£8.30	

Phasic pills

Oestrogen	Progestogen	Name	Cost
Ethinyl oestradiol 35mcg	First generation		
	Norethisterone 500mcg/1mg	Synphase	£3.60
Ethinyl oestradiol 30/40mcg	g Second generation		
	Levonorgestrel 50/75/125mcg	l 0, , ,	£3.82 (£4.00) £2.43

Oestradiol or estetrol pills

Oestrogen	4th generation progestogen	Name	Cost
Oestradiol valerate 3/2/1mg (phasic)	Dienogest 2/3mg	Qlaira	£25.18
Oestradiol hemihydrate 1.5mg	Nomegestrel 2.5mg	Zoely	£19.80
Estetrol 14.2mg	Drospirenone 3mg	Drovelis	£26.70

Other CHC preparations

Route	Oestrogen content	Third generation progestogen	Name	Cost
Transdermal patch	34mcg/d	Norelgestromin 200mcg/d	Evra	£19.51
Vaginal ring	15mcg/d	Etonogestrel 120mcg/d	NuvaRing	£29.70
			SyreniRing	£23.76

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