TOOLS FOR PRACTICE #362 | April 1, 2024



Facing the Evidence in Acne, Part I: Oral contraceptives and spironolactone in females

CLINICAL QUESTION

How effective are combined oral contraceptives (COC) and spironolactone for treating acne of at least mild-moderate severity in females?

BOTTOM LINE

At ~24 weeks, ~80-90% of females report improvement in their acne with COCs, compared to 50-80% placebo, and 30-50% will have clear-almost clear skin versus 10-40% on placebo. Efficacy appears similar between individual COCs. Spironolactone, typically added to topical agents, has similar outcomes. Discontinuations due to adverse events appear comparable to placebo.

EVIDENCE

- COC: Two systematic reviews^{1,2} (19-31 randomized controlled trials [RCTs]; 6199-12,579 patients, 11 different COC combinations) in females aged 14-49 with at least mild to moderate acne. At ~24 weeks:
 - Versus placebo:
 - Patient-assessed improvement: ~80-90% versus 50-80% (placebo).¹ Number needed to treat (NNT)=4-7.
 - Clinician assessed clear-almost clear skin: 30-50% versus 10-40% (placebo), NNT=6-9.
 - Adverse event discontinuations:¹ Usually similar to placebo. When different, number needed to harm (NNH)=25-50
 - o Versus COC:1
 - No consistent statistical differences in 17 comparisons.
 - Adverse event discontinuations: Usually not different.
- Spironolactone: One double-blind RCT (410 females, mean age 29),³ spironolactone 100mg daily versus placebo (~70% using topicals concurrently) for 24 weeks:
 - o Patient-assessed improvement: 82% versus 63% (placebo) (NNT=6).
 - o Patient-assessed clear-almost clear skin: 32% versus 11% (placebo) (NNT=5).
 - Quality of life (30-point scale, higher=better, baseline=13): Increased 8.0 versus 4.5 points (placebo), difference likely clinically meaningful.⁴
 - o Any adverse events: 64% versus 51% (placebo); example: headache 20% versus 12% (placebo).
 - o Adverse event discontinuations: No difference.
 - Other RCT added spironolactone to topical benzoyl peroxide found slightly greater benefit, but benefits possibly exaggerated as smaller, shorter RCT (63 patients, 12 weeks).⁵
- Limitations: Most COC RCTs unblinded, many COC RCTs prohibited concurrent topical agents, no RCTs comparing COCs to topical agents, many industry-funded.

CONTEXT

- Guidelines support adding COC if hormonal contraception desired, or when standard treatments (example: topical benzoyl peroxide or retinoid) inadequate. No clear recommendations for spironolactone (all published prior to recent RCT).⁶⁻⁸
- Two small RCTs (170 patients) found no statistical difference between COC and oral antibiotics.^{1,2}
- Potassium monitoring with spironolactone generally unnecessary unless patient otherwise at risk (example: on angiotensin-converting enzyme inhibitors).⁶

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