Permethrin (PER) plus trimethoprim/sulfamethoxazole (TMP/SMX) was better than PER but not TMP/SMX for head lice

Hipolito RB, Mallorca FG, Zuniga-Macaraig ZO, et al. Head lice infestation: single drug versus combination therapy with one percent permethrin and trimethoprim/sulfamethoxazole. Pediatrics. 2001 Mar;107:E30.

QUESTION

Is a combination of topical permethrin (PER) and oral trimethoprim/sulfamethoxazole (TMP/SMX) better than either PER or TMP/SMX alone for treating head lice infestation (HLI)?

DESIGN

Randomized (unclear allocation concealment*), blinded (unclear),* controlled trial with 4-week follow-up.

SETTING

3 private pediatric and family practices in San Joaquin County, California, United States.

PATIENTS

115 children who were 2 to 13 years of age (70% girls) and were diagnosed with HLI (clinical suspicion of lice or eggs). Children with hypersensitivity to TMP/SMX or 1% PER or with a history of parental noncompliance or neglect were excluded from the study. 97% of children completed follow-up.

INTERVENTION

In all treatment groups, parents were instructed to first wash their child's hair with regular shampoo and then rinse. 39 children were allocated to 1% PER, for which parents were instructed to apply the 1% PER creme

rinse to the scalp and hair for 10 minutes, rinse, towel dry, and comb the hair thoroughly with the comb provided. The procedure could be repeated after 1 week if necessary. 36 children were allocated to oral TMP/SMX at a dose of 10 mg/kg per day in 2 divided doses for 10 days. 40 children were allocated to the combined 1% PER and TMP/SMX treatment.

MAIN OUTCOME MEASURE

Cure of HLI (absence of adult lice and nymphal stage or eggs) at 2 and 4 weeks.

MAIN RESULTS

The combined treatment was better than PER alone (Table) but not TMP/SMX alone for treating HLI at 2 weeks (95.0% vs 83.3%, $\{P=0.10\}^{\dagger}$) and 4 weeks (92.5% vs 77.8%, $\{P=0.07\}^{\dagger}$). PER alone was not better than TMP/SMX alone at 2 weeks (79.5% vs 83.3%, $\{P=0.67\}^{\dagger}$) and 4 weeks (71.8% vs 77.8%, $\{P=0.55\}^{\dagger}$). 3 children with an allergic-appearing rash from TMP/

SMX were excluded from the analysis. 3 children with mild scalp irritation from 1% PER use and 9 children with transient pruritis and 3 children with nausea and vomiting from TMP/SMX use were not excluded.

CONCLUSIONS

The combination of topical permethrin (PER) and oral trimethoprim/sulfamethoxazole (TMP/SMX) was a better second-line treatment for head lice infestation than was PER alone but not TMP/SMX alone. TMP/SMX use was often associated with adverse reactions.

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*See Glossary.

 $\dagger P$ values calculated from data in article.

Combined 1% permethrin creme rinse (PER) and trimethoprim/sulfamethoxazole vs PER alone for cure of head lice infestation in children‡

Outcomes	Combined	PER	RBI (95% CI)	NNT (CI)
Cure at 2 wk	95.0%	79.5%	19.5% (1.1 to 48.4)	7 (4 to 117)
Cure at 4 wk	92.5%	71.8%	28.8% (5.3 to 66.2)	5 (3 to 25)

‡Abbreviations defined in Glossary; RBI, NNT, and CI calculated from data in article.

COMMENTARY

Recurrences after the treatment of HLI are common. Most of these reinfestations are treatment failures, although some are new infestations, but evidence exists that some head lice are resistant to the drugs usually used in treating HLI. Because the turnover of different generations of lice is much less than in bacteria, the ability of lice to become drug resistant is rare.

Hipolito and colleagues confirmed in their randomized controlled trial that a combination of 1% PER creme rinse with oral TMP/SMX was more effective than 1% PER alone, with an NNT of 5 at 4 weeks. TMP/SMX is an antimicrobial drug with adverse effects caused mainly by the sulfa component. It is cheap, relatively safe, and widely used for acute otitis media and urinary tract infections. TMP/SMX works

by killing the intestinal flora of the head lice via the blood they suck from the host. The effect of PER creme rinse is additive because it paralyzes the respiratory muscles of the lice.

Some of the patients who participated in this study had recurrent HLI. The authors recommended that the use of TMP/SMX be reserved for patients with suspected drug resistance. Even though the primary outcome was confirmed by an office visit rather than by telephone in only half of the patients among the treatment groups, it is easy to recommend at least offering this combination treatment in patients who have reinfestation.

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