CLINICAL INQUIRIES

What is the most effective treatment for external genital warts?

EVIDENCE-BASED ANSWER

Podofilox (Condylox), imiguimod (Aldara), cryotherapy, and surgical options all seem reasonable alternatives that are superior to podophyllin. (Grade of recommendation: B, based on systematic review.) No studies of surgical options versus home use preparations have been reported. Trichloroacetic acid and 5fluorouracil (5-FU) have not been sufficiently studied.

EVIDENCE SUMMARY

Nonsurgical treatments that are beneficial in eradicating genital warts are podofilox (Condylox) (8 randomized controlled trials [RCTs] with 1035 participants), imiquimod (Aldara) (2 RCTs with 968 participants), and intralesional interferon (8 RCTs). Cryotherapy is equivalent to trichloroacetic acid1,2 and electrosurgery.3 Although surgical treatments have not been compared with placebo or no treatment, both electrosurgery and surgical excision are superior to podophyllin in clinical trials.⁴⁵ Laser surgery is as effective as surgical excision.⁶ Studies of topical interferon show conflicting results.7 Systemic interferon is not beneficial.7 Topical 5-FU has not been studied with RCTs. Wart clearance rates are summarized in the Table. Treatment duration for nonsurgical options is 4 to 8 weeks. Treatment of genital warts has not been shown to reduce transmission to sex partners.7

Two RCTs^{4,5} showed more frequent recurrence with podophyllin (60% to 65%) than with surgical excision (19% to 20%). Another trial¹ showed recurrence in 22% of participants receiving electrosurgery, in 21% of those receiving cryotherapy, and in 44% of those receiving podophyllin treatment. Data are lacking on recurrence rates with imiquimod, podofilox, and intralesional interferon.

Pain occurs in less than 20% of people with imiquimod, cryotherapy, podophyllin, and electrosurgery; 39% with topical interferon; 44% with electrosurgery; 75% with podofilox; and 100% with surgical excision or laser surgery.7 However, pain has been measured using methods that are unlikely to be comparable across studies. Flulike symptoms, leukopenia, thrombocytopenia, and elevated aspartate transaminase levels are associated with intralesional interferon.7 Topical medications have not been studied in pregnant patients. Cryotherapy is safe in pregnancy based on case series, if only 3 or 4 treatments are given.7

Direct comparisons between home therapies

TABLE -

CLEARANCE RATES R	EPORTED IN CLINICAL TRIALS
Therapy	Clearance Rate (%)
Cryotherapy	63-88
Electrosurgery	61–94
Imiquimod	37–56
Interferon (topical)	6–90
Interferon (intralesional)	17–63
Laser surgery	23–52
Podofilox	45–77
Podophyllin	32-79
Surgical excision	35–72
Trichloroacetic acid	50–81
Placebo or no treatment	0–56

(imiquimod, podofilox) and other treatments are needed. Products for home use are relatively expensive: a 1-month supply of imiquimod costs approximately \$150; a 1-month supply of podofilox, \$110 to \$130. These are average wholesale prices, rounded to the nearest \$10, as of Feb. 15, 2002.

RECOMMENDATIONS FROM OTHERS

The CDC endorses podophyllin, bi- and trichloroacetic acid, podofilox, imiquimod, cryotherapy, intralesional interferon, electrosurgery, laser surgery, and surgical excision.8 A United Kingdom guideline on anogenital warts recommends physical ablative methods such as cryotherapy and surgical options for keratinized lesions and topical medications for soft lesions. The guideline also recommends ablative therapy for persons with a small number of warts regardless of type. Interferon and 5-FU are not recommended.9

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Clinical Commentary by David White, MD, at http://www.fpin.org.

REFERENCES

- 1. Abdullah AN, Walzman M, Wade A. Sex Transm Dis 1993; 20:344-5. 2. Godley MJ, Bradbeer CS, Gellan M, Thin RN. Genitourin Med
- 3. Stone KM, Becker TM, Hadgu A, Kraus SJ. Genitourin Med 1990;
- 4. Khawaja HT. J Reprod Med 1990; 35:1019-22.
- 5. Jensen SL. Lancet 1985; 2:1146-8. 6. Duus BR, Philipsen T, Christensen JD, et al. Genitourin Med 1985; 61:59-61.
- 7. Wiley DJ. Genital warts. Clin Evidence Issue 4, December 2000; 910-8.
- 8. Centers for Disease Control and Prevention. Morbid Mortal Weekly Rep MMWR 1998; 47(RR-1):91-4.
- 9. Clinical Effectiveness Group (Association of Genitourinary Medicine and the Medical Society for the Study of Venereal Diseases). Sex Transm Infect 1999; 75(suppl 1):71-5S.