

Antenatal testing for sexually transmitted infections: Is this the decade of change?

R. Matthew Chico, MPH, PhD

Department of Disease Control
Faculty of Infectious & Tropical Diseases
London School of Hygiene & Tropical Medicine
Keppel Street, London WC1E 7HT, United Kingdom

The first syndrome-based algorithms were developed in the late 1980s to diagnose and treat *Chlamydia trachomatis* and *Neisseria gonorrhoeae* cervicitis in pregnancy (1). These contributed to the first syndromic management guidelines of sexually transmitted infections (STIs) recommended by the World Health Organization (WHO) in 1991 (2). These guidelines have made important contributions to reducing the burden of curable STIs, particularly in low-resource settings where there remains a dearth of laboratory facilities and trained staff. Point of care (POC) tests have since been introduced for syphilis in the antenatal care setting to facilitate rapid treatment (3). Similar advances in POC testing for other curable STIs have been lacking.

Within this context, Peters *et al.* conducted an important non-randomised cohort study of HIV-infected pregnant women in South Africa to evaluate POC screening and treatment for *C. trachomatis*, *N. gonorrhoeae*, and *Trichomonas vaginalis* (n = 427; 51%) compared to syndromic management (n = 414; 49%). Among women in the POC screening group, 40.3% (95% CI: 35.6, 45.1) were positive for at least one STI, 29.5% (95% CI: 25.2, 34.0) for *C. trachomatis*, 5.6% (95% CI: 3.6, 8.3) for *N. gonorrhoeae*, and 20.1% (95% CI: 16.4, 24.3) for *T. vaginalis*. Postpartum, when all women were aetiologically tested, 39% fewer women had a curable STI who had been POC screened and treated at enrolment compared to women who were provided syndromic management (aRR 0.61; 95% CI: 0.35-1.05). Although the upper bound of the 95% confidence

interval crossed the null, the intervention effect seems evident: POC testing and treatment reduced the burden of STIs in pregnancy relative to syndromic management in this setting.

The study was underpowered to detect differences in the incidence of preterm birth (23% vs 23%; aRR 1.2; 95% CI 0.81-1.8) and low birthweight (15% vs 13%; aRR 1.1; 95% CI: 0.66-1.7). Effect modification may have contributed to diluting observable differences. One-fifth of women in the syndromic management group were on antiretroviral therapy at the time of enrolment (n = 80; 19%) compared to less than one percent in the POC screening and treatment group (n = 4; 0.9%). Another effect modifier may have been use of cotrimoxazole prophylaxis against opportunistic infections. Overall use was 4.8% (n = 40), but proportions were not available by treatment group. If antiretroviral use is an indication, women in the syndromic management group may have disproportionately received cotrimoxazole. While cotrimoxazole was unlikely curative of STIs, it is one-part trimethoprim and five-parts sulfamethoxazole, the latter being a sulfanamide. Another sulfanamide combination, sulfadoxine-pyrimethamine, used to prevent the consequences of malaria in pregnancy, has been shown to reduce adverse birth outcomes among pregnant women with curable STIs (4), and a recent mediation analysis has demonstrated strong 'non-malarial' protective effects (5). Regardless of these possible effect modifiers, Peters *et al.* provide key evidence in support of randomised controlled trials that compare POC screening and treatment versus syndromic management powered to improved birth outcomes in a range of antenatal care settings. Is this the decade of change?

References

1. Braddick MR, Ndinya-Achola JO, Mirza NB, Plummer FA, Irungu G, Sinei SK, et al. Towards developing a diagnostic algorithm for Chlamydia trachomatis and Neisseria gonorrhoeae cervicitis in pregnancy. *Genitourin Med.* 1990;66(2):62-5.
2. World Health Organization. Guidelines for the management of sexually transmitted infections. 2001; WHO/HIV_AIDS/2001.01.

3. Mabey DC, Sollis KA, Kelly HA, Benzaken AS, Bitarakwate E, Chagalucha J, et al. Point-of-care tests to strengthen health systems and save newborn lives: the case of syphilis. *PLoS medicine*. 2012;9(6):e1001233.
4. Chico RM, Chaponda EB, Ariti C, Chandramohan D. Sulfadoxine-Pyrimethamine Exhibits Dose-Response Protection Against Adverse Birth Outcomes Related to Malaria and Sexually Transmitted and Reproductive Tract Infections. *Clinical infectious diseases : an official publication of the Infectious Diseases Society of America*. 2017;64(8):1043-51.
5. Roh M, ter Kuile F, Rerolle F, Glymour M, Shiboski S, Gosling R, et al. Overall, anti-malarial, and non-malarial effect of intermittent preventive treatment during pregnancy with sulfadoxine-pyrimethamine on birthweight: a mediation analysis. *The Lancet Global Health*. 2020;8(7):e942-e53.