

at a clinic or hospital, arrangements are made so that a chemist in the patient's area may supply him with the requisite drugs. Unfortunately, if the chemist is not open on a Sunday the addict receives a double supply on the Saturday and of course if the chemist is closed for Easter Monday or Christmas Day and Boxing Day following a weekend, then the drug addict is given a supply of drugs to carry him over this period.

This system may work well in many cases. As a police surgeon, I meet the cases where the system falls down and very real problems exist. These occur when, because the addict is so dependent on his drugs or for some other reason, he uses up his supply quickly, or when he sells them, and then is without drugs for the next day or so.

Another problem is that if a doctor wishes to confirm that a certain person is a drug addict, then facilities are available to obtain this information from the central register in London. This register is not manned outside normal working hours. Moreover, the information when obtainable is basically the fact that the patient is a drug addict; the name of the drug and the dose in use is not known, though the clinic that the patient is attending can usually be obtained. However, even if the clinic is known the records cannot be obtained outside normal hours.

My object in writing this letter is to stimulate some thought and, I hope, some appropriate action on these problems.—I am, etc.,

A. J. LAIDLAW

Worcester

**Impaired Colour Vision in Diagnosis of Digitalis Intoxication**

SIR,—I was interested in the letter from Dr. Vesa Manninen (14 December, p. 653) with regard to dyschromatopsia in early cases of digitalis intoxication. I have found similar early symptoms in suspected intoxication with digoxin, but I would suggest that the Ishihara plates are not really suitable since the early signs are usually on the blue/yellow axis, for which there are no plates in the Ishihara series. A more suitable test is the Farnsworth Munsell 100-hue test, which tests the whole range of colour hue and is, in addition, much more sensitive than the Ishihara test, which is primarily of value in the detection of the gross defects present in inherited colour blindness. Alternatively, and even better, is the use of the Pickford-Nicholson anomaloscope, but this requires a good deal of experience in a specialist clinic.—I am, etc.,

W. O. G. TAYLOR

Ophthalmic Unit, Heathfield Hospital, Ayr

**Choice of Contraceptives**

SIR,—Having just read the letter from Professor D. B. Jelliffe and E. F. Patrice Jelliffe (14 December, p. 658) I would like to make the following point before breast feeding is advocated instead of contraception.

About 12 years ago I practised in Manitoba and part of my work included 10 colonies of Hutterites. Every married woman in the colonies quite categorically stated that a woman could not become pregnant while

breast feeding. Yet I often saw a 2-year-old, 1-year-old, and 3-month-old child all feeding from the same mother. Certainly such feeding was unrestricted, on demand, and permissive. I have seen a 2-year-old run in from the garden, stand and feed beside its mother, and run out to play again.

I have not had the benefit of a personal communication from Rosa nor have I any knowledge of "curvilinear" compromises, but I would suggest that "linear" technology has a great deal more to offer than "curved" biotraditional contraception.—I am, etc.,

D. L. PRICE

Kingsclere, Newbury, Berks

**Distamine**

SIR,—Some confusion has been caused by the presence on the pharmacists' shelves of three formulations of Distamine (D-penicillamine). The original capsule of penicillamine hydrochloride may be withdrawn in 1975. Its equivalent replacement is a tablet containing 125 mg of penicillamine free base, and this together with the standard 250-mg tablet of base is available now. Will prescribers please note that if a prescription for "Tab. Distamine" or "Tab. penicillamine" is made out without stating the strength required the smaller tablet must be dispensed.—I am, etc.,

W. H. LYLE

Medical Director, Dista Products Ltd.

Liverpool

**Treatment of Genital Herpes**

SIR,—Dr. S. M. Laird and Mr. R. B. Roy (27 July, p. 255) reported the beneficial effect of co-trimoxazole in genital herpes, and Dr. Paula H. Gosling (17 August, p. 473), in support of similar findings, stated that trimethoprim inhibits the growth of the virus in tissue culture, though she did not give the data.

Co-trimoxazole depends for its antibacterial action on inhibition of bacterial purine synthesis, the sulphonamide component inhibiting dihydrofolate synthetase and trimethoprim dihydrofolate reductase, the bacterial dihydrofolate reductase being at least 10 000 times more depressed than the mammalian enzyme.<sup>1</sup> Suppression of herpes simplex growth would not be expected since these enzymes are not induced by herpes virus infections<sup>2</sup> and are probably not necessary for virus replication, the virus utilizing the pre-existing cellular pool of purines in the synthesis of its D.N.A.

We have studied the effect of trimethoprim on the growth of herpes simplex types

TABLE I—Plaque Assay of Herpes Simplex Virus Types 1 and 2

Drugs Added*	Virus	
	Type 1	Type 2
None .. .. .	37	82
TMP 1 µg/ml .. .	42	94
TMP 10 µg/ml .. .	46	110
TMP 100 µg/ml .. .	36	106
P and S .. .. .	41	74
TMP 1 µg/ml + P and S .. .	43	100
TMP 10 µg/ml + P and S .. .	35	82
TMP 100 µg/ml + P and S .. .	39	72

\*TMP = Trimethoprim. P and S = Penicillin and streptomycin (100 U/ml and 100 µg/ml respectively).

1 and 2 in B.H.K.21 cells using plaque assays (see table I) and one step growth curves (tables II and III) as indices of virus growth. The growth curves were carried out in the absence of penicillin and streptomycin and both dividing and resting cells were studied, the growth in resting cells being more closely analogous to infection in the intact host. Levels of trimethoprim up to 100 µg/ml, alone or in combination with sulphamethoxazole, produced no more than 50% reduction in virus growth, a value insignificant in comparison with other antiviral agents. This level of trimethoprim is some 30 to 50 times that achieved in serum after a 240-mg oral dose,<sup>3</sup> reduces mammalian dihydrofolate reductase activity by 50%<sup>1</sup> and is toxic to B.H.K. cells (personal observations). Patients receiving trimethoprim in a daily dose of 1 g have been shown to have significant bone marrow depression.<sup>4</sup>

TABLE II—Virus Growth at 18 Hours after Infection in Absence of and in Presence of Trimethoprim

Virus	Plaque-forming Units per Cell		
	No Tri-methoprim	30 µg/ml Tri-methoprim	100 µg/ml Tri-methoprim
Type 1 ..	10		10.1
Type 2 ..	15	13.3	12.6

TABLE III—Virus Growth at 18 Hours after Infection in Absence of and in Presence of Co-trimoxazole

Virus	Plaque-forming Units per Cell		
	No Co-trimoxazole	Co-trimoxazole (= 30 µg/ml TMP*)	Co-trimoxazole (= 100 µg/ml TMP)
Type 1 in dividing cells	{ 40 54 38	27	23 28 28
Type 2 in resting cells	141		67

\*TMP = Trimethoprim.

Our studies failed to show significant inhibition of the growth of herpes simplex in tissue culture. The reported effect of co-trimoxazole in the clinical situation is unexplained.—We are, etc.,

DAVID H. WATSON  
DAVID HAIGH

Department of Microbiology, School of Medicine, Leeds

- Hitchings, G. H., and Burchall, J. J., *Advances in Enzymology*, 1965, 27, 417.
- Freearson, P. M., Kit, S., and Dubbs, D. R., *Cancer Research*, 1966, 26, 1653.
- Brumfit, W., et al., *Postgraduate Medical Journal* (Suppl. Nov.), 1969, 45, 56.
- Kahn, S. B., Fein, S. A., and Brodsky, I., *Clinical Pharmacology and Therapeutics*, 1968, 9, 550.

**Screening for Gynaecological Abnormalities**

SIR,—In her interesting survey of gynaecological abnormalities found at a cytology screening clinic Miss Diana Edwards (26 October, p. 218) makes the important point that abdominal and pelvic examinations should always be carried out at such clinics. However, I would contest her statement that this assessment should be done only by doctors with gynaecological experience. This is an ideal arrangement in countries with a high doctor-patient ratio but not in the under-doctored areas of the world where "well-women" screening services are at present inadequate or totally absent.