## UNUSUAL PATTERNS OF ANTIBIOTIC RESISTANCE FROM A CLINICALLY ISOLATED R-FACTOR

W.D. Willis and J.T. Smith, Microbiology Section, Department of Pharmaceutics, The School of Pharmacy, Brunswick Square, London, WClN 1AX

It is common for clinical strains of antibiotic resistant bacteria to harbour R-factors. Consider an organism with R-factor mediated resistance to four drugs. If these resistances are borne on a single R-factor then on mating all four resistances will transfer as a unit. If there are two R-factors the resistances will transfer as two independent groups. Let us suppose a strain with two R-factors transfers one of them at a rate of  $10^{-5}$  per input donor and the other one at a rate of  $10^{-6}$  per input donor, then only very rarely would an individual recipient cell receive both R-factors.

An Escherichia coli harbouring an R-factor, which we term  $R_{\rm SS}^{28}$ , was isolated from a patient suffering from an urinary tract infection which had failed to respond to co-trimoxazole therapy.  $R_{\rm SS}^{28}$  was transferred to a laboratory strain of E.coli J62-1 using trimethoprim (Tm) selection, and resistance to Tm, Spectinomycin (Sp), Sulphonamide (Su) and Streptomycin (Sm) was found in the recipient strain. At this stage  $R_{\rm SS}^{28}$  seemed to be a single R-factor. However, when J62-1 (RSS28) was mated with another laboratory recipient, strain J53-2, the following results were obtained.

Selective drug	Туре	Resistance patterns			transfer rate per input donor	
		Tm	Sp	Su	Sm	
Tm	1	+	+	+	+	$1.65 \times 10^{-6}$
	2	+	+	-	-	$0.15 \times 10^{-6}$
Sp	3	_	+	+	-	$2.26 \times 10^{-4}$
Su	4	_	+	+	+	$0.61 \times 10^{-4}$
	5	-	+	+	-	$2.63 \times 10^{-4}$
	6	-	-	+	+	$3.1 \times 10^{-4}$
Sm	7	_	+	+	+	$0.2 \times 10^{-4}$
	8	_	_	+	+	$2.1 \times 10^{-4}$

+ = resistance found in recipient, - = resistance absent from recipient

As before the Tm result (type 1) would seem to suggest that we are dealing with a single R-factor conferring resistance to all four drugs. However, the other results would seem to suggest that there are two R-factors, one conferring resistance to Tm Sp (type 2) and another conferring resistance to Su Sm (types 6 and 8). However, types 3 and 5 which received Sp Su suggest a third R-factor. This last R-factor would seem to occur together with an Su Sm R-factor to give types 4 and 7. These results are puzzling because (i) the patterns of resistance do not fit into clear cut linkage groups and (ii) the frequencies of the cotransfer of three or four resistances seem too high.

Types 4 and 7 were analysed further. It was found as forecast that when they were mated with other recipients these received either Sp Su (types 3 and 5) or Su Sm (types 6 and 8) or all three resistances together.

As  $R_{\rm SS}28$  seems to be comprised of three R-factors, i.e. Tm Sp, Sp Su and Su Sm it is possible that there were two different Sp genes and two different Su genes in the original strain.