## LETTERS TO THE EDITORS

## **GENETICS**

## Salmonella-Escherichia Hybrids

Until recently, genetical studies with Salmonella typhimurium have been carried out mainly by the transduction method. In an effort to facilitate such studies, we attempted to develop a means of applying conjugation techniques also, using S. typhimurium and Escherichia coli, since Baron and his collaborators had been able to produce hybrids between these two bacteria. We have been successful in experiments with a Salmonella culture carrying a mutator gene (mut) and a culture of E. coli strain K-12 carrying the factor for high frequency of recombination (Hfr). The mutator strain was isolated by Miyake² from strain LT-7. Its main characteristic is a 10- to 100-fold increase in spontaneous mutability.

Our procedure for the detection of recombinants is as follows. Equal volumes of over-night broth cultures of the S. typhimurium mut strain and E. coli Hfr are mixed; 0·1-ml. samples of the mixture are plated immediately on appropriate medium; and the plates are incubated for 24–36 hr. at 37°C. Under these conditions the frequency of recombinants is about 1 per 10° plated bacteria. Thus it is considerably lower than the frequency observed in recombination between two strains of E. coli.

Two E. coli strains have been used so far: HfrCS101 and HfrC. With the first we obtained hybrids in which the following S. typhimurium mutant genes were replaced by their wild-type alleles from the E. coli genome: ability to ferment lactose (lac); requirements for leucine (leu), arginine (arg), threonine (thr), aspartic acid (asa), thiamine (thi), proline (pro), glutamic acid (glt) and pantothenate (pan). With HfrC we have obtained hybrids incorporating the lac, pro, leu and thr wild-type alleles.

Thus it appears likely that the hybrids may carry any region of chromosome of the donor bacterium.

The results so far can be summarized as follows:
(1) Hybrids have been obtained which combine certain genetic markers carried separately by the two parent bacterial strains.

- (2) Among these hybrids three types have been recognized: (a) sensitive to virulent Salmonella phage H5, but resistant to E. coli phage T6; agglutinated by Salmonella serum; (b) resistant to phages H5 and T6; very slight or no agglutination by Salmonella serum; (c) resistant to phage H5, but sensitive to T6; not agglutinated by Salmonella serum.
- (3) Temperate Salmonella phage (PLT-22 H1) grown on a type-(a) hybrid will transduce markers carried in the Salmonella part of the chromosomal complex, provided the recipient Salmonella bacteria have the mut (mutator) gene.
- (4) This same phage forms about  $10^1-10^2$  times more plaques on hybrid-(a) bacteria than on LT-7 bacteria carrying the mut gene, and  $10^2-10^4$  times more plaques than on LT-7 bacteria without mut. Adsorption tests indicate no difference between hybrid LT-7 mut and LT-7 mut bacteria with regard to phage adsorption.

(5) The approximate order of certain loci in the Salmonella genome as determined by these recombination tests is: mut—thr—leu—pan—pro—lac. There is evidence that pro and lac are close together and close also to asa, glt and arg.

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 Baron, L. S., Carey, W. F., and Spilman, W. M., Abst. Comm. at Paper Sessions, Seventh Int. Congr. Microbiol. (1958).
 Demerec, M., Lahr, E. L., Miyake, T., Goldman, I., Balbinder, E., Banic, S., Hashimoto, K., Glanville, E. V., and Gross, J. D., Carnegie Inst. Wash. Year Book, 57, 390 (1958).

## A Further Example of the Kell Blood Group Phenotype K-,k-,Kp(a-b-)

In an earlier communication<sup>1</sup>, we reported two sisters having the Kell phenotype K-,k-,Kp(a-b-). The parents were second cousins, and it was assumed that the sisters were homozygous for a Kell gene the product of which does not react with any of the known Kell antisera. Allen  $et\ al.^2$  later gave this gene the abbreviation  $K^c$ . Alternatively, there may be a deletion or suppression. We have sought for other examples of this phenotype by using the serum of the propositus of the above family by the capillary-papain method<sup>3</sup>; by this method the serum reacts with all K+ or k+ bloods. The 3,122nd blood so tested did not react, indicating that it was of the phenotype we sought. This was confirmed by testing with anti-K, anti-K, anti-K, anti-Kpa and anti-Kpb; it reacted with none of these.

The parents of the propositus, Mrs. Kan, are first cousins. Her husband, two children, daughter-in-law and four grandchildren are all K-,k+,Kp(a-b+). While the two children are presumably of genotype  $kKp^b.K^o$ , titration with anti-k and with anti- $kp^b$  failed to differentiate between assumed genotypes  $kKp^b.kKp^b$  and  $kKp^b.K^o$ . The family is of Finnish descent. The mother and six brothers and sisters of the propositus live in Finland.

It may or may not be significant that two members of the original family in which K<sup>0</sup> occurred, and two of the nine members examined in the present sibship, had suffered abnormal bleeding for which they were transfused; further, the new-born baby of the propositus in the first family suffered from a form of erythroblastosis feetalis in which petechial hæmorrhages of the skin were a prominent sign.

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<sup>1</sup> Chown, B., Lewis, Marion, and Kaita, Hiroko, Nature, 180, 711 (1957).

<sup>2</sup> Allen, jun., Fred H., Lewis, Sheila J., and Fudenberg, H., Vox Sang., 3, 1 (1958).

<sup>3</sup> Lewis, Marion, Kaito, Hiroko, and Chown, B., J. Lab. Clin. Med., 52, 163 (1958).