

Misima. March 20, 1958

TRANSDUCTIONAL ANALYSIS OF MONOPHASIC TYPES OF SALMONELLA

(Report by Tetsuo Iino and Joshua Lederberg\*)

The specificity of flagellar (H) antigen in Salmonella is controlled by two distinct ~~genes~~ loci, phase-1 by  $H_1$  and phase-2 by  $H_2$ . Which one is manifested in a given clone depends on the phase determinant at the  $H_2$  locus. That is, the alternation of  $H_2$  state leads to the alternative expression, which has been known as phase variation of H-antigen (the Annual Report, 1956).

Some Salmonella strains do not express two phases but only one. Those strains are called monophasic-1 or -2 strains depending on their fixed phase, either phase-1 or phase-2 respectively. Three additional groups of the genes which are involved in the production of H antigen were disclosed by transductional analysis of the monophasic strains.

S. abortus-equi CDC-26 is stable in phase-2, enx-type.

A rare alternative phase can occasionally be obtained by anti-serum selection, resulting in <sup>an</sup> equally stable phase-1 (a). Transductions were carried out from enx-phase of CDC-26 to i-phase of a diphasic strain of S. typhimurium, TM2 i:1.2 (such transduction is designated CDC-26 (a):enx  $\rightarrow$  TM2 i:1.2). Among 65 transductional clones which had been selected on semisolid nutrient gelatine-agar media (NGA), 4 expressed diphasic <sup>h</sup> a:1.2 type, 42 diphasic i:enx and 19 monophasic-2 enx which carry a hidden  $H_1^i$ . Thus, when a is transduced ~~a fraction of the transductional clones become monophasic.~~

\* Department of Genetics, University of Wisconsin, Madison, Wis., U. S. A.

the resulting transductions remain diphasic, whereas when *enx* is transduced a fraction of the transductional clones become monophasic. By anti-*enx* selection, i-phase cultures were obtained from the monophasic *enx*-transductional clones. The i-phase cultures obtained are also monophasic. The stabilization of the  $\underline{H}_2$  state in *S. abortus-equus* is therefore caused by a factor which is linked to  $\underline{H}_2$ . The factor will be given a symbol  $\underline{Vh}_2^-$ .

*S. typhimurium* SW1061 is a monophasic-2 mutant of a diphasic strain TM2 i:1.2. The culture reacts to anti-1.2 serum but not to anti-i serum. However, the strain frequently produces nonmotile H-negative (non-flagellar) subclones, which in turn revert to motile cells with 1.2 antigen in successive cultures. From the transduction, diphasic *S. abony* CDC-103 b:*enx* ---x SW1061, monophasic<sup>h</sup>-2 *enx*, diphasic b:1.2 and a small number of i:1.2 types were obtained. The change from the monophasic type to diphasic types was always coupled with the loss of the ability to oscillate between motile and non-motile types. These results are consistent with the following explanation. In SW1061,  $\underline{H}_1^i$  is inactive; on the other hand  $\underline{H}_2^{1.2}$  changes its state as in usual diphasic strains. Consequently, when  $\underline{H}_2$  is active, phase-2 antigen, 1.2, is produced, and when  $\underline{H}_2$  changes to inactive, that is both  $\underline{H}_1$  and  $\underline{H}_2$  are inactive, the cell

cannot  
~~mutants~~ produce H antigen and become non-motile. The production of diphasic  $\times$  1:1.2 type suggests that the inactivation of  $H_1$  is not caused by an intrinsic change of  $H_1$  itself but by an inhibition of its function by a gene linked to  $H_1$ . The linkage and the recombination between  $H_1$  and  $H_1$ -~~inhibits~~activity controller were confirmed on  $Fla_1^+$  (linked to  $H_1$ ) transductions from SW1061  $\rightarrow$  S. heidelberg  $Fla_1^-$  r:1.2, from which monophasic-2 -(1):1.2 and diphasic 1:1.2 as well as diphasic r:1.2 were obtained.

The  $H_1$ -activity controller is designated  $Ah_1^-$ .  $Ah_1^-$  was discovered in two other monophasic-2 strains of S. typhimurium, SW629 and SW547. All of three  $Ah_1^-$  are nonallelic and linked to  $H_1$  and  $Fla_1$ . The ~~parent~~ type of gene,  $Ah_2^-$ , which inhibites the function of  $H_2$  was disclosed on four monophasic-1 strains of S. typhimurium. ~~XXXXXXXXXXXX~~ All of them are linked to  $H_2$ . Both  $Ah_1$  and  $Ah_2$  are phase specific but are not concerned with the specificity of antigen types, which are determined exclusively by  $H_1$  and  $H_2$ . (the detail will be reported in )