

The following is a true copy of my laboratory notes, Volume 1, covering the period October 5, 1951 to July 7, 1953, pages 1-218a, with a number of unnumbered pages included.

There are two additional volumes of my notes, one of which is labelled "Summaries".

In this first volume there are a number of unnumbered pages at the front which represent an index, but it should be noted that at page 110 there is a statement "not indexed beyond here".

The chief interest in these notes will be for the specialized transduction produced by the bacteriophage lambda and its discovery. However, many of the notes deal with other matters. Each of us in the Lederberg laboratory had some side assignments and some of these, such as isolation of mutants in E. coli strains believed fertile with K-12, radiation resistance as a function of ploidy and lysogenicity for phage, phage induction by UV, and others were my assignments.

The discovery of the special transduction by lambda was not planned. It came shortly after the discovery of generalized transduction by phage PLT22 in Salmonella by Zinder and Lederberg, and was followed several years later by the generalized transduction in E. coli by phage P1 by Lennox.

As I was producing large amounts of phage lambda in some radiation experiments I was curious as to whether lambda transduced any genetic material. I tried my first transduction (for methionine independence) on March 26, 1952 (page 47) and of course it didn't work. I tried the same experiment again a few days later (pages 48,50).

It is my recollection that Norton Zinder and I were alone in the lab while the Lederbergs were at a meeting at Rutgers University in March or April of 1952, when Norton and discussed the possibility of lambda transducing genetic material. I recall that Norton said he thought that Esther Lederberg had tried some experiments with lambda but he did not know what they were. Since I had a lot of lambda preparations I suggested that we try and so we took all the selective medium plates available, and appropriate recipient strains and mixed lambda and cells and plated them out.

I have always thought that chance played a big part in Norton Zinder's failure to discover special transduction. Norton worked with Salmonella which do not metabolize lactose and therefore he had no EMB lactose plates, on which the first lambda transductions were observed.

It was my good fortune to have had the EMB lactose plates which provided the selective environment for gal⁺ clones.

However, it was still baffling in that the papillae on EMBlac produced by lambda and quantitatively related to the amount of lambda used - proved to be lac⁻ on further examination. I spent much of April 1952 trying to resolve this confusion. It was at this point that Esther Lederberg suggested, on the basis of her prior knowledge and experience with lac-gal interactions and the strains that I was using, that I should look at lambda transduction in terms of selection of gal⁺ clones. So on May 5, 1952 (page 71) I began studying lambda transduction of gal genes on EMB galactose, and it all began to come out. Much more was needed to be done: lambda as vector had to be established; the heterogenetic character of the transductants elucidated; the identification of the alleles involved in the segregants when the donor and the recipient were gal-; quantitative relationships established; the discovery of high frequency (HFT) transducing lysates; that the phage in some of the transductions was a defective phage, and other things.

It was an exciting time and it was an exciting experience and the stimulation by the people in the crowded lab helped - the Lederbergs, Zinder, Alec Bernstein, Tom Nelson, Bob Wright, Luca Cavalli-Sforza, Gaylen Bradley, Tetsuo Iino, Dorothy Gosting, and probably others I overlook.

Prior to writing this introduction I made an audio tape that discusses the rest of my notes in Volume 1, and perhaps when it is transcribed it can added to this copy.

M. L. Morse
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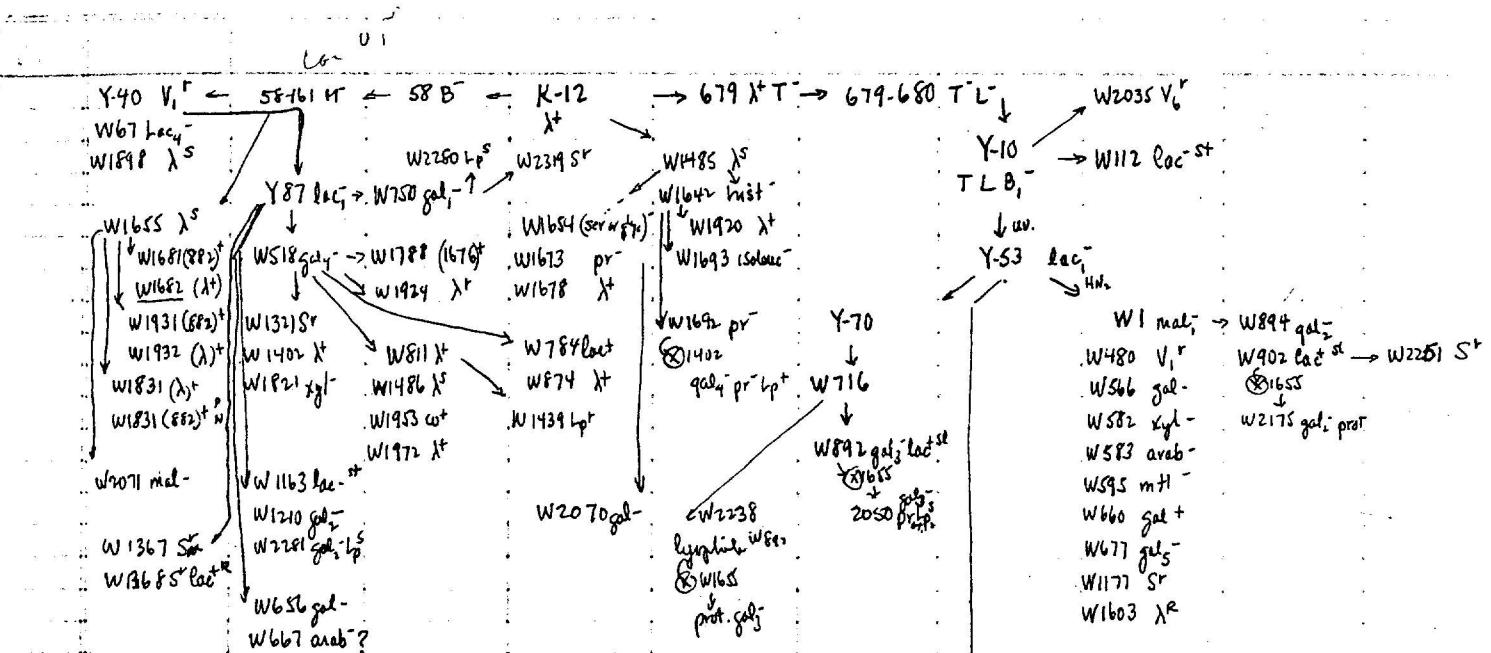
July 23, 1986.

Research Notes Vol. I

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Room 200



58-161

W416 V_i^r $V_{i_h}^r$ \otimes Y64 V_i^r

W1296 λ^r ← W588 lac^+ \otimes W518
W1503 λ^r (λ^r lac^+)

H217
↓ segregation

W1267 (lac^-) lac^- gal^- λ^r
W1436 S^r
W1661 (882)⁺
W2196 mal^-
W1662 λ^+
W1736 (1670)⁺
W2193 mal^-
W2202 V_i^r
W2234 F^r

Cultures in Stand 8/20/53

✓ K-12									
✓ 58-161									
✓ W67									
✓ W578									
✓ W588									
✓ W687									
✓ W750									
✓ W811									
✓ W874									
✓ W892									
✓ W902									
✓ W1177									
✓ W1210									
✓ W1368									
✓ W1402									
✓ W1436									
✓ W1439									
✓ W1485									
✓ W1503									
✓ W1655									
✓ W1673									
✓ W1678									
✓ W1692									
✓ W180P ^(u) ₍₃₁₎									
✓ W1821									
✓ W1924									
✓ W2035									
✓ W2050									
✓ W2062									
✓ W2070									
✓ W2071									
✓ W2175									
✓ W2196									
✓ W2202									
✓ W2251									
✓ W2280									
✓ W2281									
✓ W2297									
✓ W2319									

<u>Subject</u>	<u>Rays</u>					
Gel directions						
SR-161	47, 82					
W518 gel+	54, 92(sup), 93, 95(2), 97(stable) (S18C16-12) ¹⁶ 99(stable)					
W1655	48, 49, 50, 51					
W1736 gel+	59, 61, 62, 65(gas), 71(gas) 72 ^(gas) , 74 ^(gas) , 75 ^(gas) , 76 ^(gas) , 77, 78 ^(gas) , 80(gas), 81(lab), 82, 83(acetate), 86, 87, 88 ^(acetate) , 95(stable) 96(stable), 99(4,4)(stable),					
W112	71, 83(princip.), 85, 94(derived)					
W1678	74, 76					
W1662	80, 82, 84					
W811 gel-	81, 82, 83(BM), 84, 86, 87, 88(5 ¹), 89 ^(S18C16-12) , 92(80%), 93, 94, 96 ^(S18C16-12) 99(4,4)(stable), 103(stable)					
W1439 gel- ¹⁶ ₁₈	82					
W1821 gel-	83 ^(lab) , 85(gel, x9, BM), 87					
W902 gel-	88, 100(derived) 110(derived) 110(gel, stable)					
W750 gel-	88, 91, 93 ^(second) 94(75 ¹⁰), 96(stable)(200C90), 98, 99(4,4)(stable), 105(stable), 107(stable)(stable), 109(stable)					
W1692	96					
W1920	96					
W2057 gel-	99 ^(K-12) , 106, 107(stable)					
W1578 gel+, F-	99					
W2062	100, 101, 104, 105, 106 102, 103, 104(lab), 104 ^(lab) , 106, 107(stable), 109(stable)					
W1924						
Adsorption Exp.	89					

SubjectTransductinsPages

86, 87
85a (creatin)
87
81 (lac⁻)
80 (gal⁻)
89

47, 82
54
48, 49, 50, 51
55, 61, 62, 63, 64 (gal), 71 (gal), 72 (lac⁻), 74 (gal), 75 (leucine), 76 (lac⁻), 77, 78 (lac⁻)
71, 83 (arginine), 85 (lac⁻)
74, 76
80, 82, 84
81, 82, 83 (B6), 89, 90, 97
82
83 (gal, lac⁻ B6), 87

Reconstruction eq.

82.

Crosses

58-161 X Wq 14 pr- trypt⁻
W112 X W1655 [A^r T18, lac⁻ + B6, lac⁻]
W902 X W1655
811 tK-r X 1436
518 tK-r X 1436
780 X 1503
(177 X 1635)

50, 52
85a, 88, 97
95, 96, 98, 100 (lac⁻)
100, 101, 102 (+), 103 (-), 104 (os (repeat)), 106, 107 (shuttle), 108, 109
100, 101, 102 (+), 103 (-), 105 (repeat), 106, 107

SubjectPages

Lwoff effect with λ (for other phages see particular strain)

determined

by gross examination

K-12
SB-161
W67
W1177
W1661
W1662
W1736
W1682 (w^{eff+})
H267
90r
W1821
W811
750
W1934

26, 30, 32, 33, 36, 38, 43, 51, 61, 90
36, 38, 39, 43, 92
57,
57,
59,
59,
59, 67, 90 (gal⁺)
63,
58
91
90
90, 90 (gal⁺)
94
98

by plaque count (see also λ prep.)

K-12
K-12A, K-12B
W1177
W1678
W1932
H267
W1682 (w^{eff+})
W1934 (w^{gal})
W1972 (gal⁺)
W1998 (gal⁺)

22*, 23*, 53* (gal⁺), 68 (gal⁺, cult) 69*
34, 35
13
58
62*
60*, 68*
63
69*
69*, 78*
78*

effected by post incubation:

K-12
W1177
W1603
W1736 gal⁺
W811 gal⁺
S194L-12
S111L-12

29, 25, 26
13
13
88
88
102
102

SubjectPages

λ Preparations

51

K-12 vs λ

-

K-12 L1

PK

32

L2

PK

32

L3

PK

32

L4 (PK + Spn)

PK

33

L5

PK

43, 47

L6 (Pen)

Pen

51, 61, 62

L7 (Pen)

Pen

37,

W67

L1

58-161

L1

L2

L3 (Pen)

93, 93

W1177

L1

59, 71, 80

W1485

39, 35

W1662

59

W1655

31, 44 (aggr) 48 (agg)

W1736

59, 71, 74, 80, 80 (agg)

W811

84, 84, 85, 86, 90, 90 (agg)

H-267

58, 59

882 prep

69, 74, 77

1821

85, 90, 93

Jrrad. of λ 90%

80, 80

93

99

16

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Subject

Ultraviolet effects -

Survival

K-12

S8-161

W 518

W-811

W-1177

W-1985

W-1603

W-1655

W-1655 (882) = 1921

W-1673

W-1678

W-1681

W-1682

W-1831

W-1831 (882) (W) (P)

W-1931

D [S8-161 (882)]

W-1932

H267

W1736

W1959 (W_g 31)

W1972 (X_g 2)

W1898 (A_g)

W1503 (A_g)

W1998 (W_g 28 X_g)

W1436 (X_g)

λ

W1953 (wt)

W1661

Survival curves

K-12

H267

Mice W1832 for J.L.

W1931 for E.L.

Regions from λ in W112

W1736 goalt

Lysates streaked out

Pages

12, 17*, 19*, 20*, 22*, 23*, 25*, 26*, 53 (age+) 68 (sym w/)
6, 8, 9*, 10*, 11*, 15, 18*, 69*, 71 (age+)
2*, 30*, 13, 90, 73 (sym)

13, 17*, 19*, 20*, 22*, 23*, 64

13, 6, 7, 8, 11*, 15, 18*, 20*, 70*

146*

12*

58*

6, 8, 15, 53

6, 8, 15, 63*

18*, 55, 64

29*, 45*, 55*

54*

15

62*

57*, 59*, 60*, 64*, 66*, 68*, 73* f5 (sym goalt)

66

69*

69*, 78*

70*

69*

78*

81

80, 81

81

80

76

78

77

80

104, 106

Phage Stocks - lysates

#	Source	titer						
1.	750 (gal ₁ -)	$>2.4 \times 10^{10}$						
2.	902 (gal ₁ -)	$4.9 \times 10^{10} \leftarrow ?$						
3.	K-12	1.4×10^{10}	↓ exhausted					
4.	58-161	1.8×10^9						
5.	811 (sp. gal ⁺)	4.0×10^{10}	(contam?)					
6.	1821 (gal ₉ -)	1.0×10^{10}	(contam?)					
7.	K-12	2.3×10^{10}	wasted					
8.	1485 (fulcate)	-						
9.	750+1821	$6.5 \times 10^9 \leftarrow ?$	(contam?)	0.1 ml / EMB gel gave no col	10/9			
10.	1439 (gal ₄ -)	1.1×10^{10}						
11.	811 (gal ₉ -)	1.7×10^{10}						
12.	811 (gal ₉ -)	?						
13.	1736 (sp. gal ⁺)	?						
14.	892 (gal ₃ -)	$3.8 \times 10^9 \leftarrow ?$						
15.	K-12 (HEATED Δ)	-						
16.	1954	$>140 \times 10^9$						
17.	2096	2.1×10^{10}						

SubjectPages

Wg-14

pr- verified
pr+
pr- trypt- crosses
lac- verified
lac^t(+₁, +₂, +₃, +₄)
pr- trypt+ lacst

27, 28, 32, 34, 35, 36, 38, 39, 40, 41, 42, 56, 57, 59, 60, 61, 66, 67, 68, 85a
35
36
35, 44, 46, 78
50, 52
45
60, 61
75

Wg-16

pr-
pr- x⁻(1)
pr- x⁻(2)

27, 28, 36, 37, 38, 39, 40, 41, 42, 55, 64, 67, 68, 75
42, 47, 48, 49, 52
53
56 - not su on testing

Phase sensitivity

$$\frac{1682}{1681} = \lambda$$
$$882/1503$$

W9 14

W1584

$\text{pr}^- \text{ F}^+ \text{loc}^{-8} \text{Malt} + \text{Suc}^{+8} \text{Gal} + \text{celo}^- \text{phenam}^+ \text{Acrit}^R \text{Juice} + V_1^R V_2^L V_{3,7}^L \lambda^L \rho^R \times 370R$
stock

(pr⁻loc⁺)

#1, 2, 3, 4

pr-loc

(pr-tryp)

p⁺

W1716

W1716

$\text{F}^+ \text{loc}^{-8} \text{gal} + \text{celo}^- \text{phenam}^+ \text{Acrit}^S \text{milk} + V_{1,7}^R \lambda^R \rho^R \times 370R$ hypo for K-12

Stock

pr-

pr-x₁

pr-x₂