

SICILIAN
VIRUS
CHARTS

Oct. 6, 1943

Inoculation of Acute Sandfly Fever Serum
(Sicilian Material)

into 4 patients at Longview Hospital (Cincinnati, O.)

Material: Frozen serum (Datoush and Carr) was thawed out just before inoculation. 2cc was withdrawn from each and mixed in a sterile ampoule. The inoculations were as follows:

0.1cc. intracutaneously flexor surface of forearm

0.75cc subcutaneously in midtriceps region

All inoculations between 3³⁰ and 4 P.M.

Procedures prior to inoculation:

WBC and differential

20cc of blood drawn for serum

Subjects:

1.	[REDACTED]	4 - female
2.	[REDACTED]	8 - "
3.	[REDACTED]	41 - male
4.	[REDACTED]	38 - "

105

104

103

102

~~10~~

101

100

99

98

0 ' 1 ' 2 ' 3 ' 4 ' 5 ' 6 ' 7 ' 8 ' 9 ' 10 '

104

103

102

101

100

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97

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100

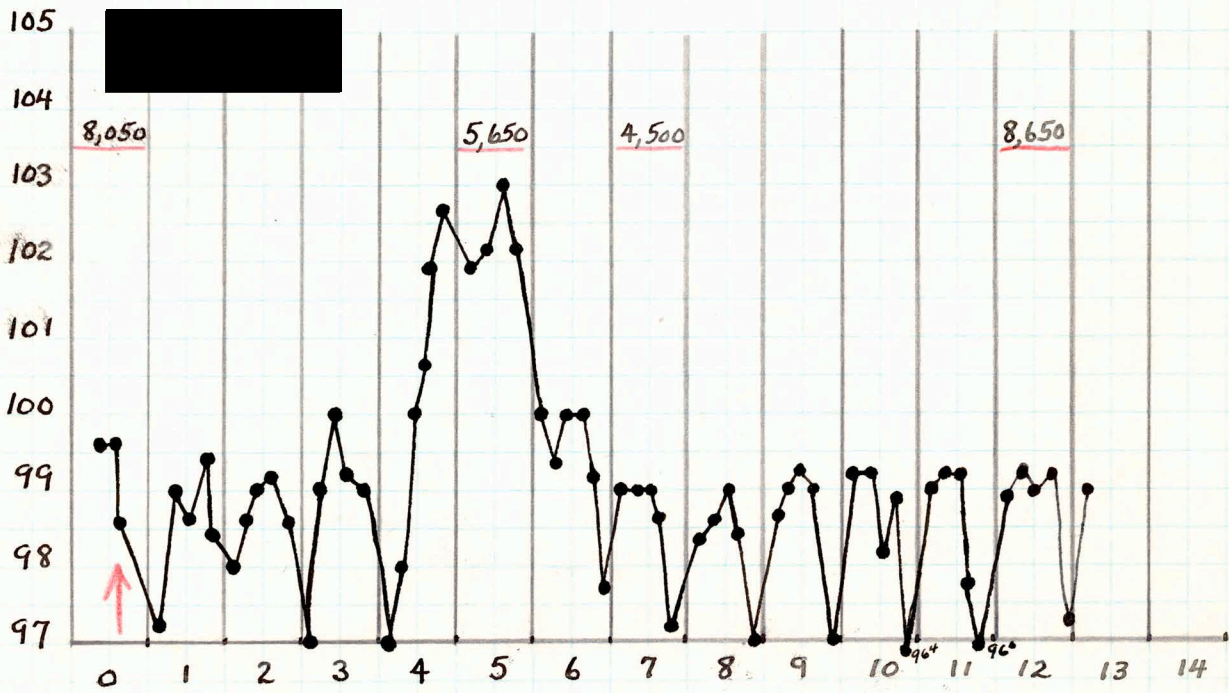
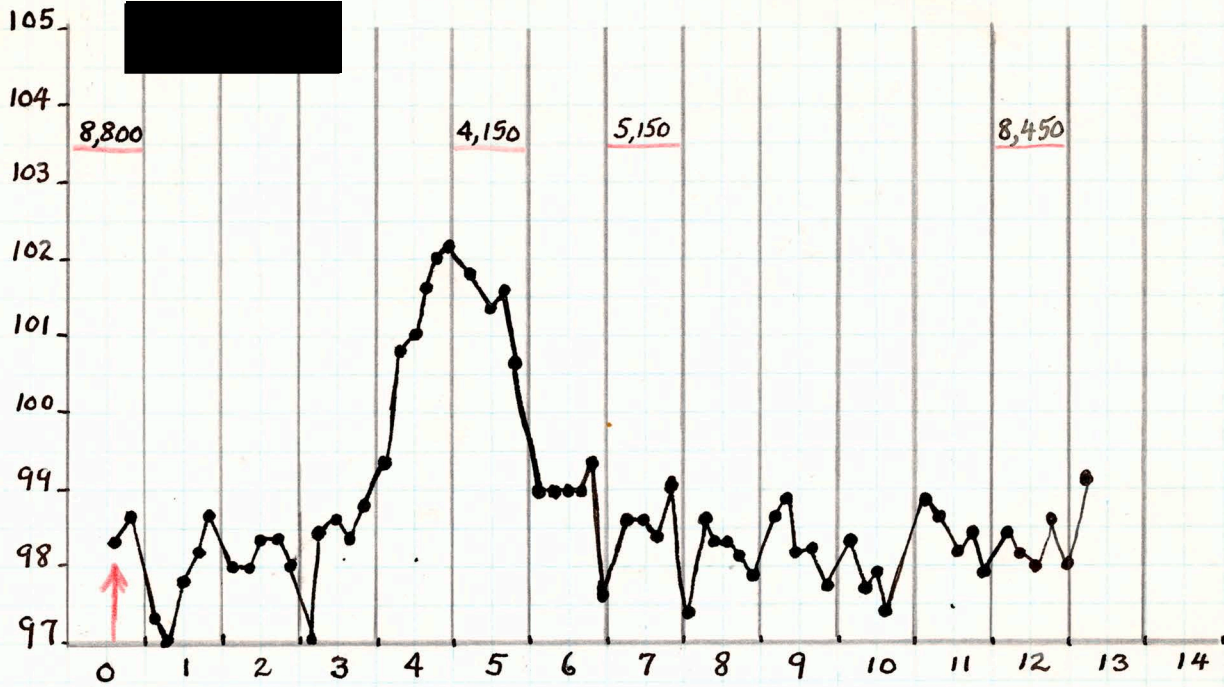
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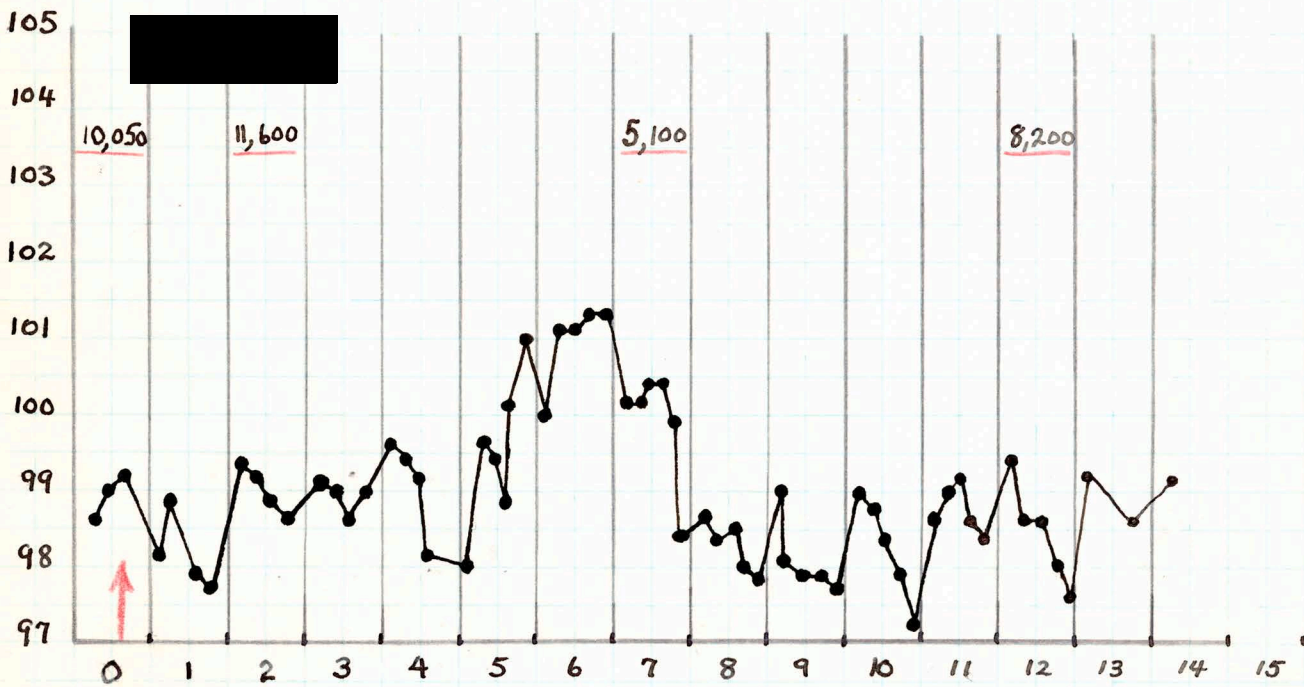
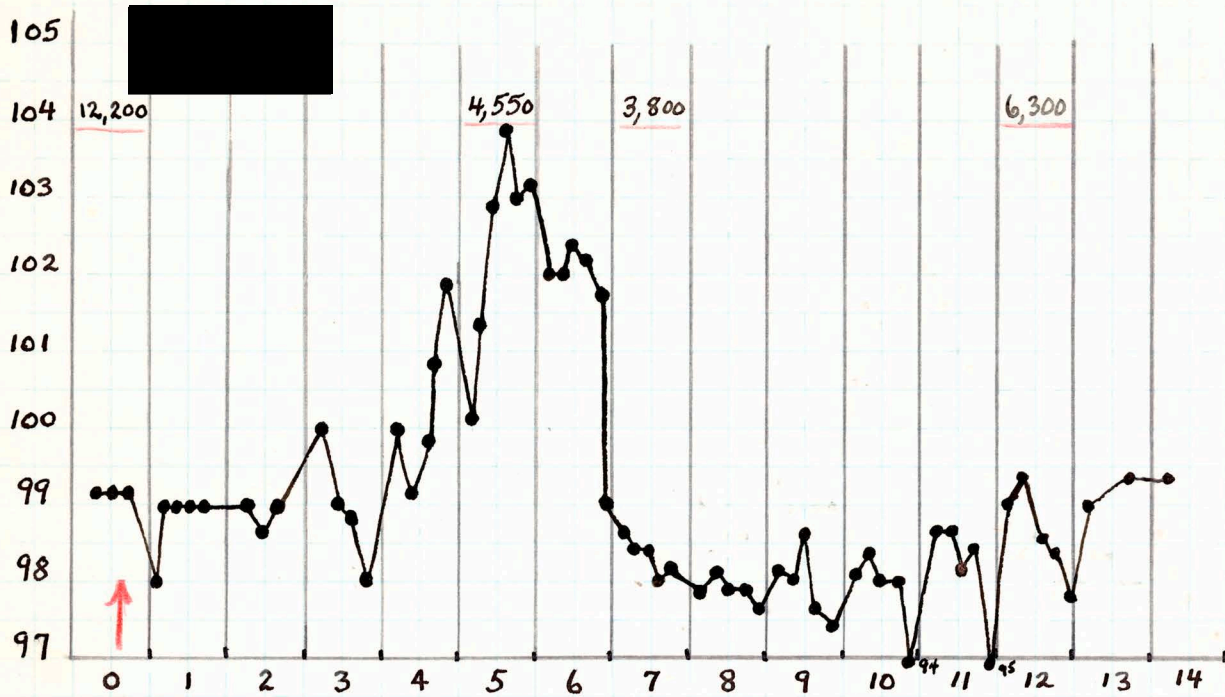
97

ACUTE SICILIAN (PAPPATACI?) FEVER SERUM IN HUMAN SUBJECTS

0.1cc intracutaneously; 0.9cc subcutaneously - Oct. 6, 1943, 3³⁰-4⁰⁰P.M.



Oct. 6 7 8 9 10 11 12 13 14 15 16 17 18 19



Filtration of the Sicilian Virus Through Gradocol

Membranes

— Oct. 21, 1943

Source of Virus — Serum from first experimental passage in human beings, obtained on first day of fever. Frozen in CO₂ box since Oct. 10 or 11.

Pool made up as follows:

■	- 16.5 cc	
■	- 12.0 cc	
■	- 12.5 cc	
■	- 10.0 "	
	<u>51.0 cc</u>	- 24 cc used for this expt. Remainder inoculated in Rhesus monkeys, cotton rats, and mice.

Preparation for Filtration — Serum diluted 1:5 in saline
24cc serum + 96cc saline.

Centrifuged at \approx 1500 rpm for 30 minutes

GRADOCOL MEMBRANES — ELFORD Type

770 μ m membrane — obtained late in 1939 from England

309, 203, 101, 75 and 50 μ m membranes — obtained from

Dr. Johannes Bauer, I.H.D. of the Rockefeller Foundation, about a year old.

Preparation of membranes — Sterilized by steaming for 30 min. Hormone broth which was passed through a Seitz filter the day before and which had a pH of 7.8 to 7.9 was passed through each membrane prior to filtration of the diluted serum.

Procedure — It was planned to filter the centrifuged diluted serum thru the 770 μ m membrane and use this filtrate for subsequent passage thru the 300, 200 and 100 μ m membranes. It was planned to use the 300 μ m filtrate for passage through the 75 and 50 μ m membranes. However

filtration through the 300 μ membrane ~~to~~ proved to be too slow to permit this. For some reason filtration thru the 200 μ membrane was much faster, and part of this filtrate was passed through one 50 μ membrane and 770 μ filtrate thru another 50 μ membrane.

770 μ membrane - 10cc of broth under 10 lbs pressure (Nitrogen) - passed thru in continuous stream.

Total amt filtered 100cc in 4 portions

25cc - 9:32 A.M. - 30 sec

25cc - 9:35 " - "

25cc - 9:38 " - "

25cc - 9:41 " - "

● Part saved - remainder used for filtration thru 300, 200, 100, 75, 50 μ membr.

309 μ membrane - 10cc of broth passed fairly rapidly, under 10 lbs pressure

770 μ filtrate used - 25cc added

10⁰⁰ A.M. - dropwise at 10 lbs pressure; increased to 15 lbs - but filtration still slow

1⁰⁰ P.M. - 10cc filtrate obtained. Filtration stopped

Only single membrane present. New membrane steamed and filtration repeated - but again very slow - much slower than 200 μ .

203 μ membrane - 10cc broth passed rapidly under 15 lbs pressure

770 μ filtrate used.

First 12cc passed thru in 4 minutes

Subsequent 10cc required 60 minutes.

Part inoculated - 11cc used for 50 μ membrane

101 μ membrane - 5cc of broth under 15 lbs pressure - passed rapidly, 770 μ filtrate used

First 5cc passed thru in 5 min.

Subsequent 6cc required 2 hrs.

75 μ membrane - 5cc of broth under 15 lbs. pressure required 30 min.

770 μ filtrate used - Filtration slow

10cc of filtrate obtained in 6 hours.

GRADUOL MEMBRANE FILTRATION - CONTINUED - Oct. 21

50 m μ membrane No. 1 - 5cc of broth under 15 lbs pressure required 3 hrs.

770 m μ filtrate used.
Started 1 P.M.

520 " - 6cc of filtrate present - 5cc removed for inoculation. Filtration continued overnight for another 15 hrs. Flask containing tube & filtrate packed in ice.

Oct. 22 8:30 A.M. - 6.5cc of almost water clear filtrate collected.

50 m μ membrane No. 2 - 5cc of broth under 15 lbs. pressure.
- \approx 4cc passed in 2 hrs.

200 m μ membrane filtrate (10.8cc) used.

Started 6:10 P.M. - filtration slow dropwise. Flask with recipient tube packed in ice.

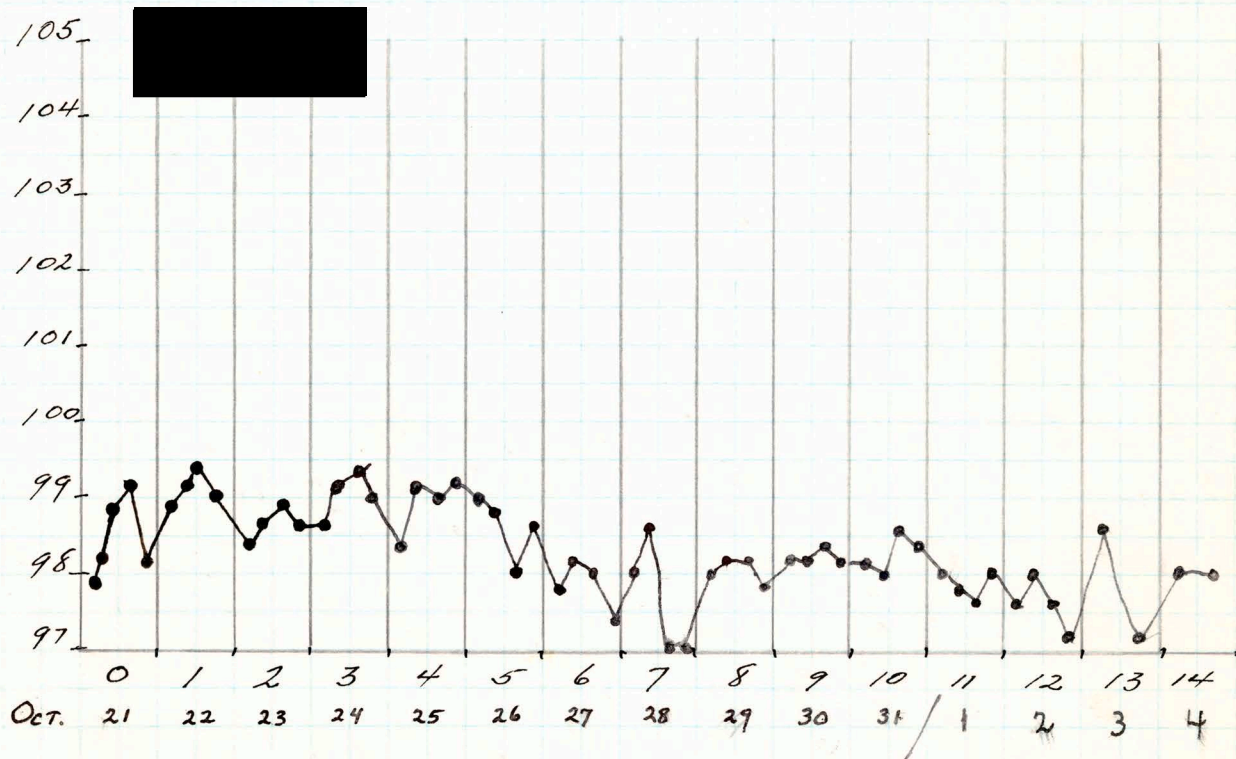
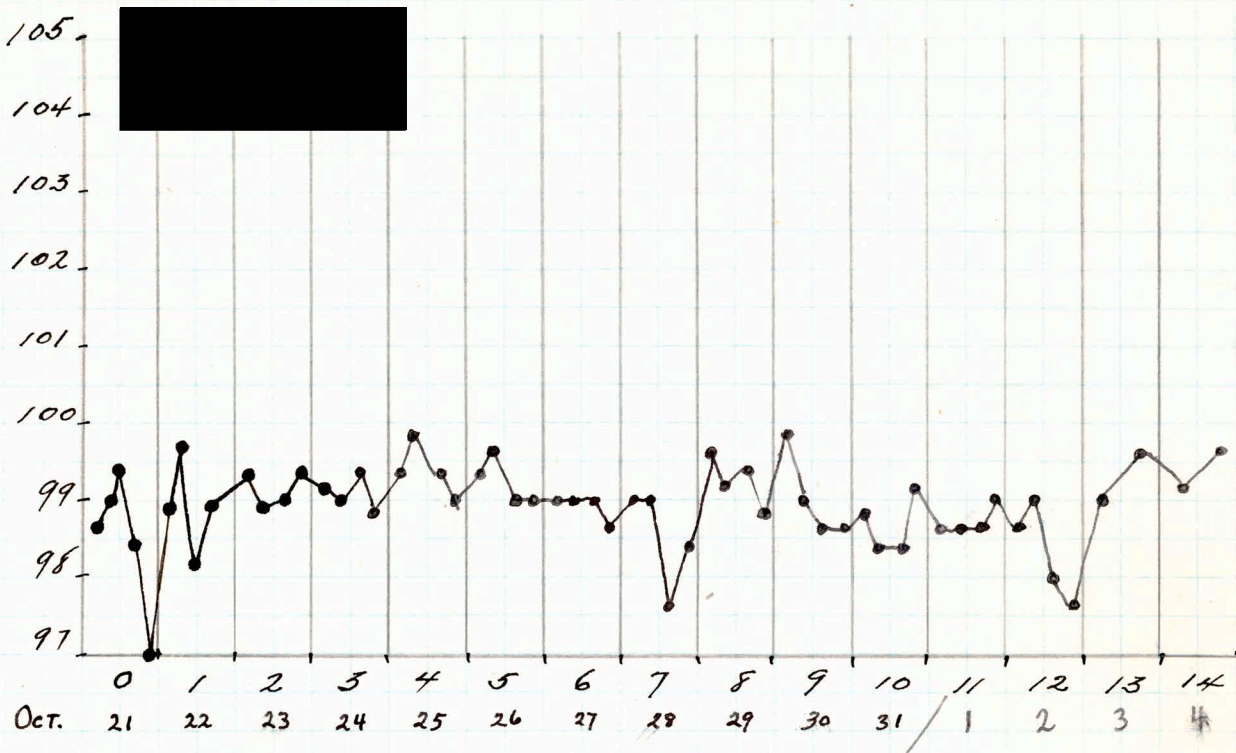
Oct. 22 8:30 AM - 10.4cc of filtrate recovered.

<u>MEMBRANE</u>	<u>Am't of broth passed</u>	<u>Material used for filtration</u>	<u>Pressure lbs.</u>	<u>Total time</u>	<u>Am't. of filtrate collected</u>
770m μ	10cc	Centrifuged serum dil. 1:5 in saline	10	2 min.	100cc
309 "	10cc	770m μ filtrate	15	3 hrs.	10cc
203 "	10 "	" "	"	1 hr.	22 cc
101 101 "	5 "	" "	"	2 "	11 cc
75 "	5 "	" "	"	6 "	10 "
50 " No. 1	5 "	" "	"	19h. 30min	11.5 cc
50 " No. 2	4 "	200m μ "	"	14h 20 "	10.4 cc

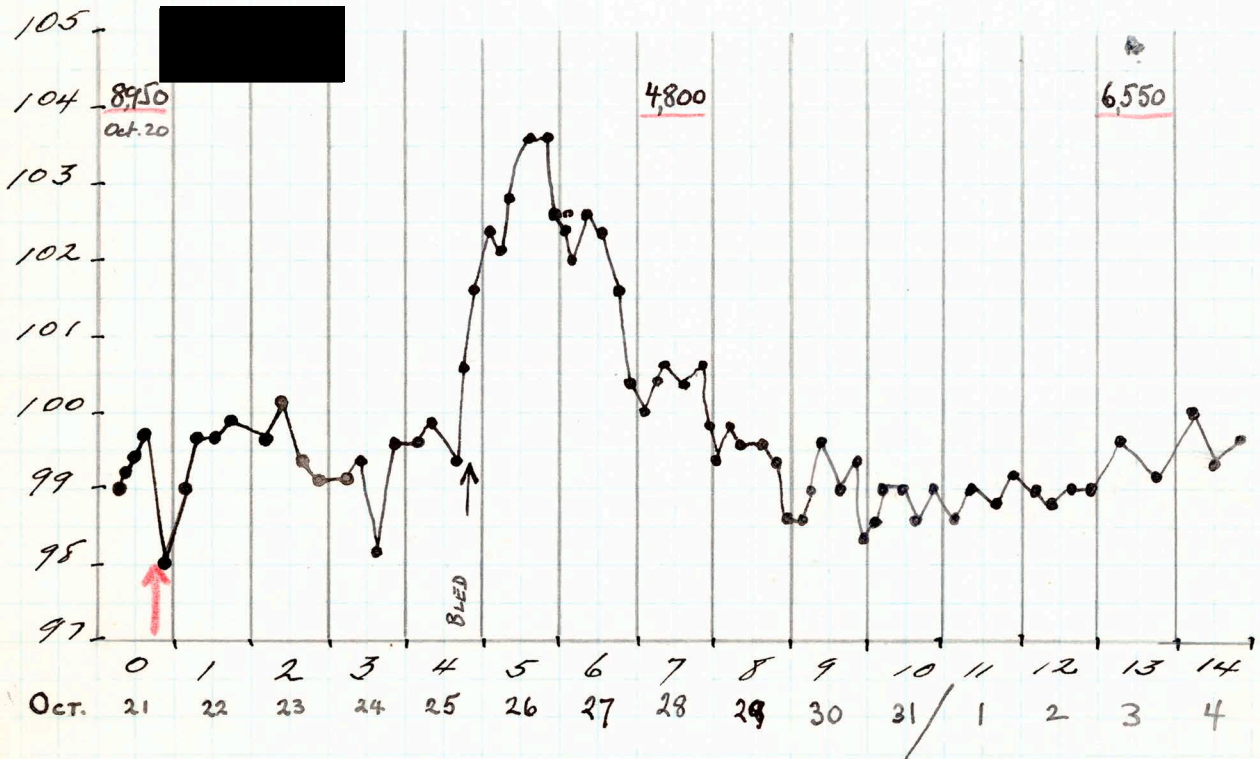
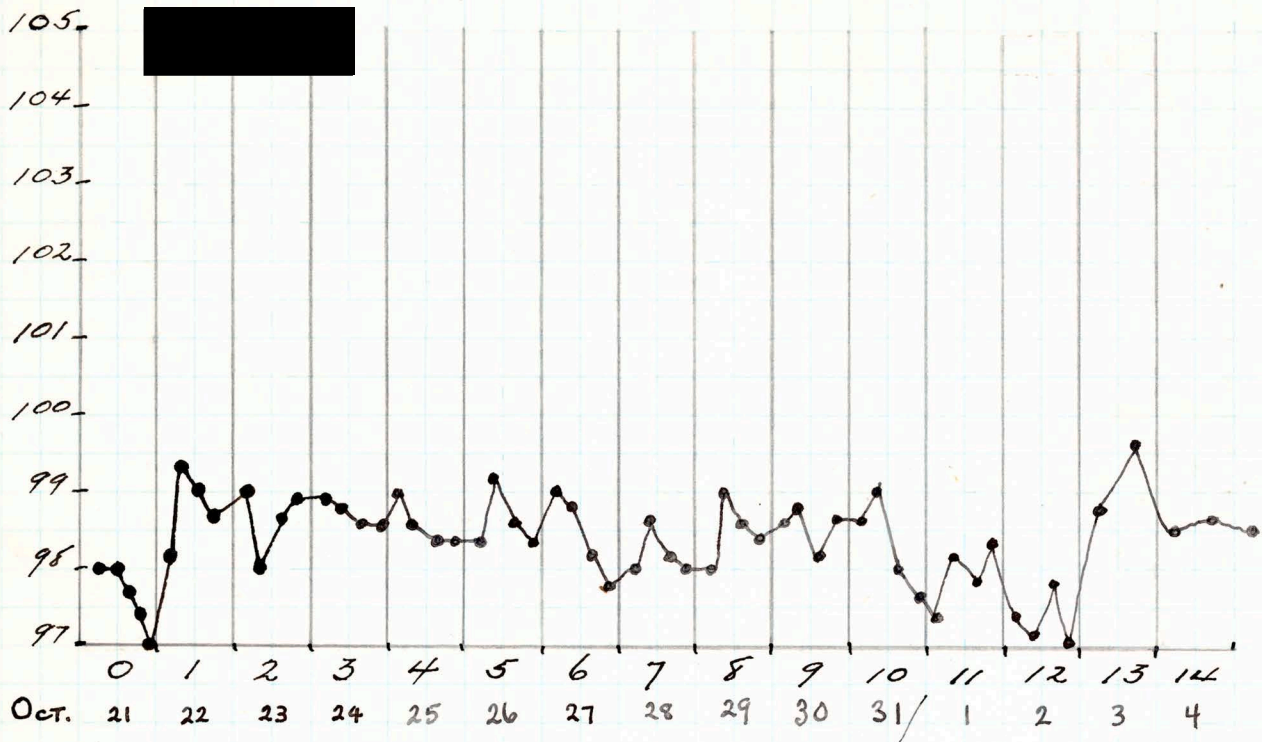
Inoculations - Subcut. and Intramusc.

<u>Material</u>	<u>Dose</u>	<u>Time</u>	<u>Name</u>	<u>Age</u>
Centrifuged unfiltered serum 1:5	5cc	6:30-7 P.M. 10/21	[Redacted]	- white male - 36
	5cc	" " "		- " " - 49
<u>770 mμ</u> filtrate	5cc	" "	[Redacted]	- " " - 41
	"	" "		- " " - 43
<u>309 mμ</u>	5cc	" "	[Redacted]	- " " - 41
	"	" "		- " " - 44
<u>203 mμ</u>	5cc	" "	[Redacted]	- " " - 45
	"	" "		- " female - 40
<u>101 mμ</u>	5cc	" "	[Redacted]	- " " - 46
	"	" "		- " " - 43
<u>75 mμ</u>	5cc	" "	[Redacted]	- Colored " - 45
	"	" "		- " " - 40
<u>50 mμ</u> (770)	5cc	" "	[Redacted]	- white " - 36
" (220)	5cc	10 A.M. 10/22		
<u>50 mμ</u> (770)	6cc	" "	[Redacted]	- colored " - 47
" (220)	5cc	" "		

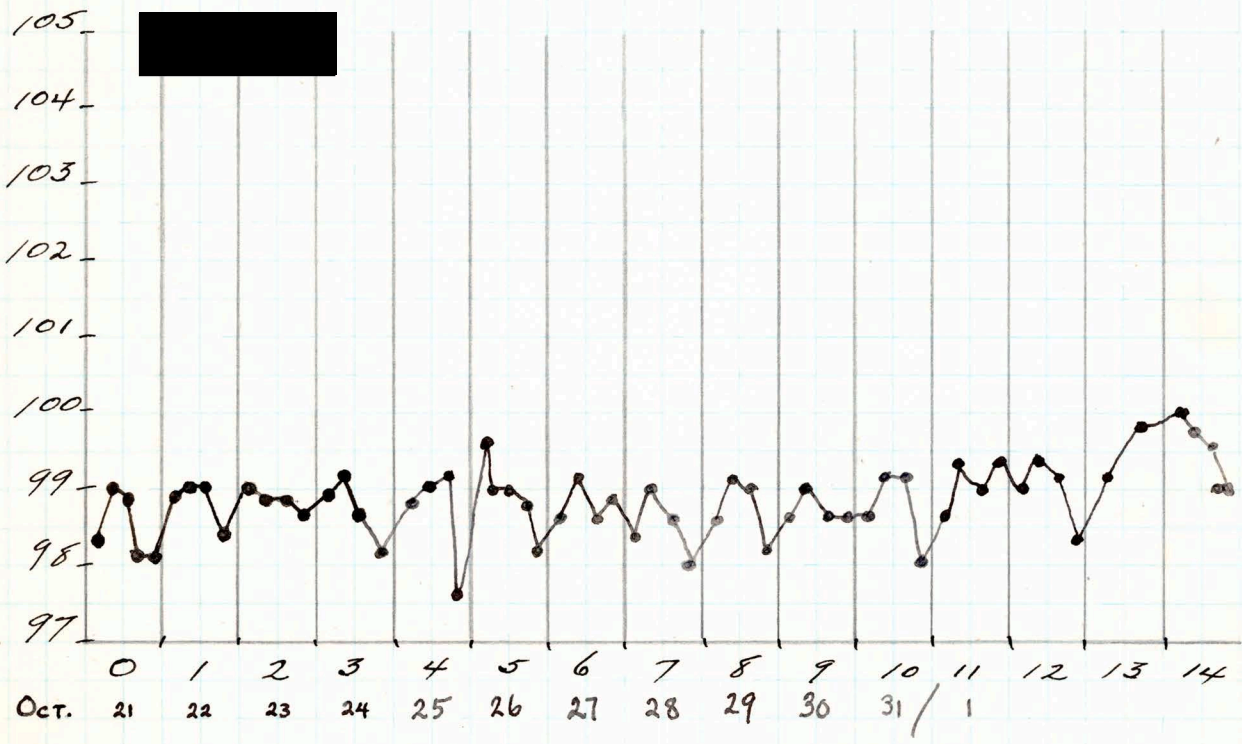
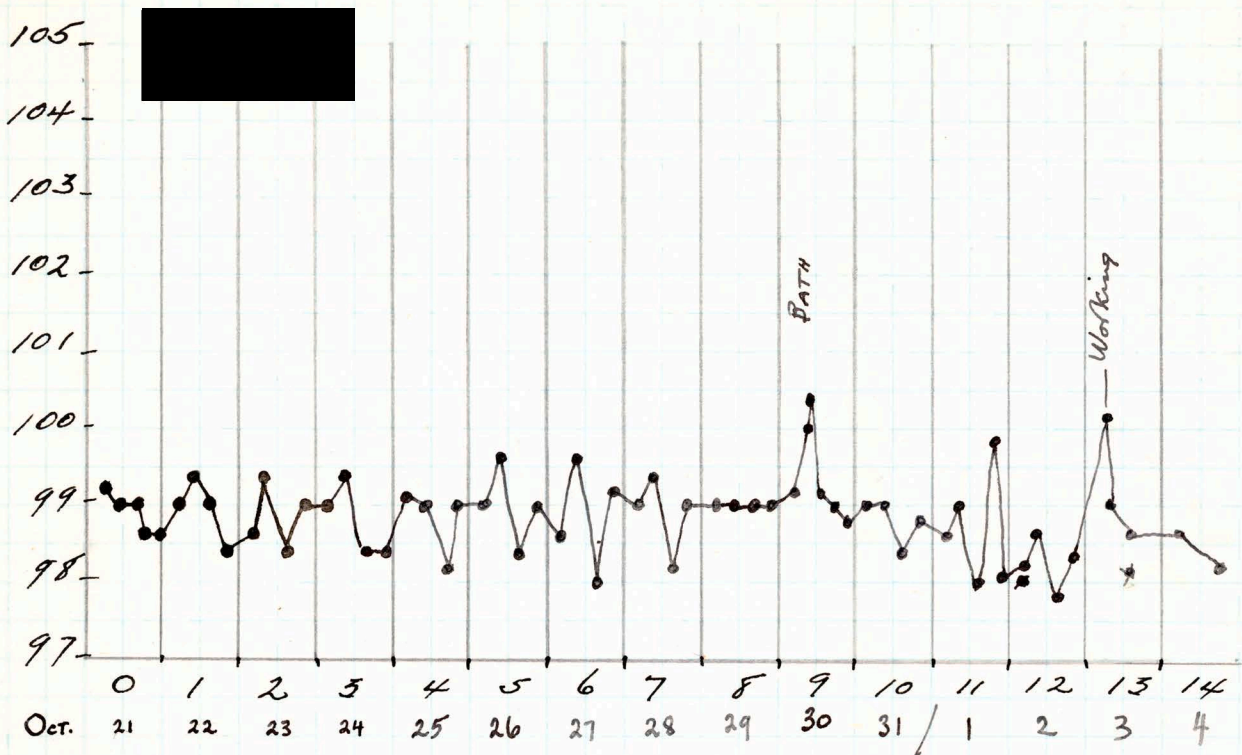
SICILIAN VIRUS



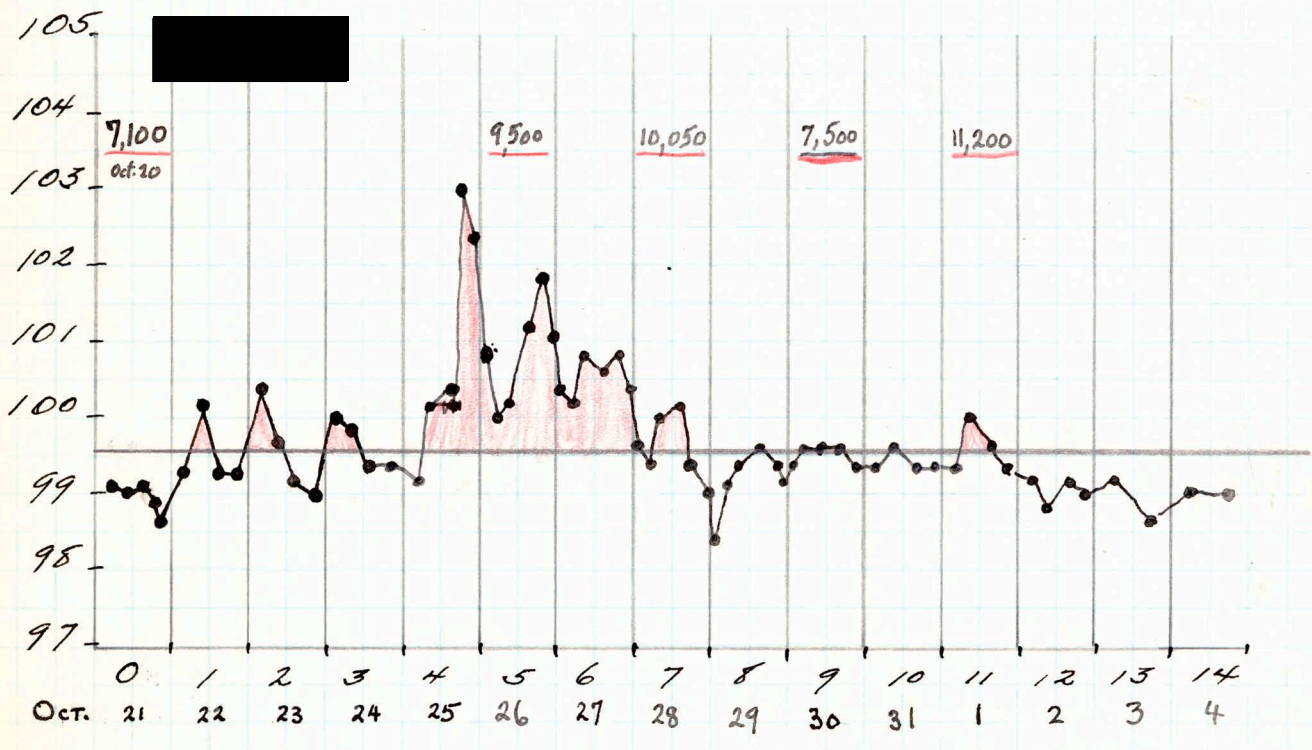
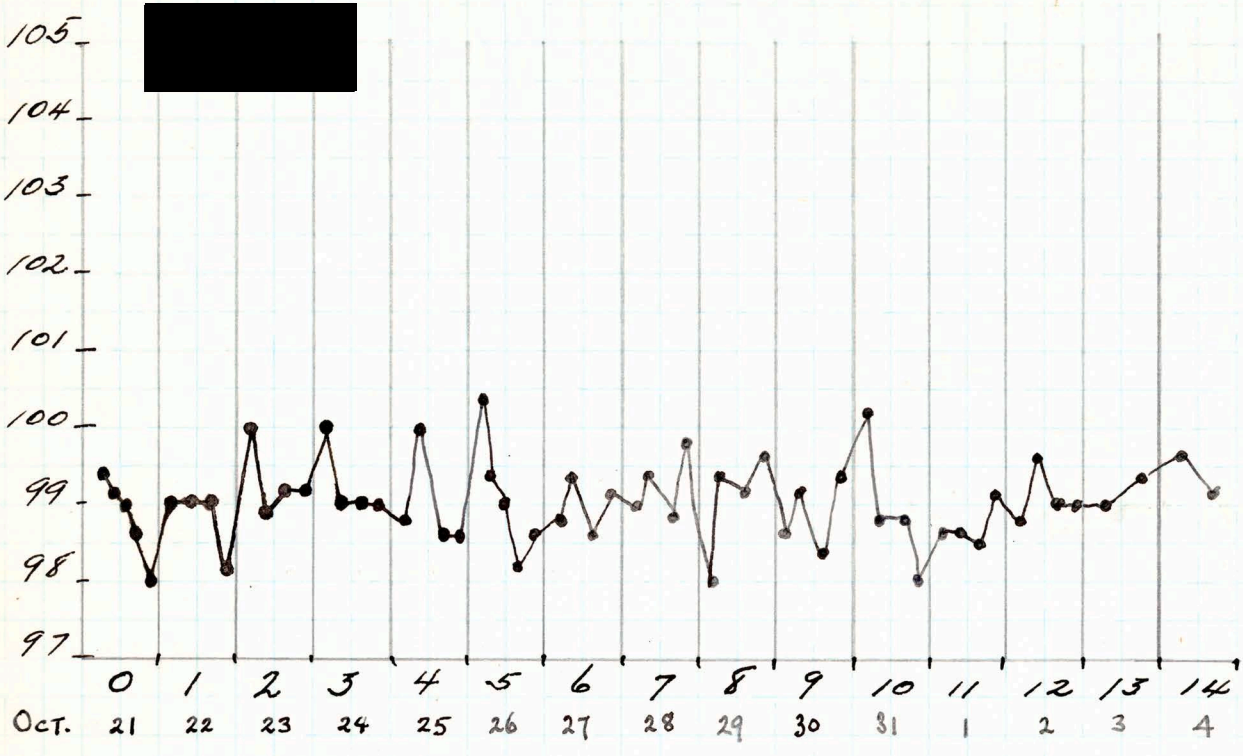
CENTRIFUGED, UNFILTERED SERUM - 5cc



770 mμ SERUM - 5cc.



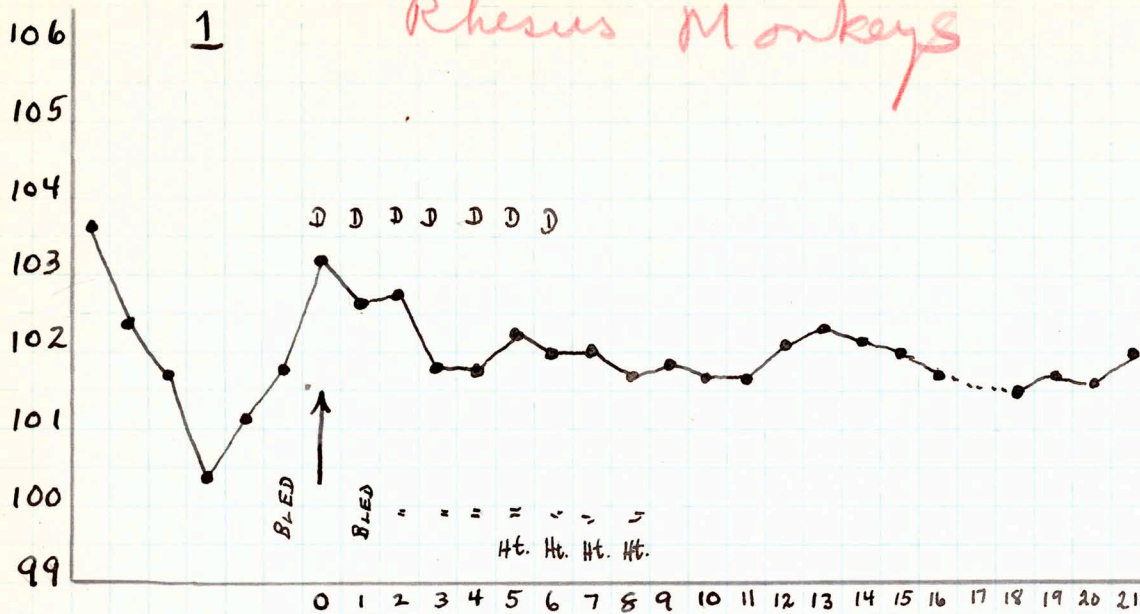
203 μ SERUM - 5cc.



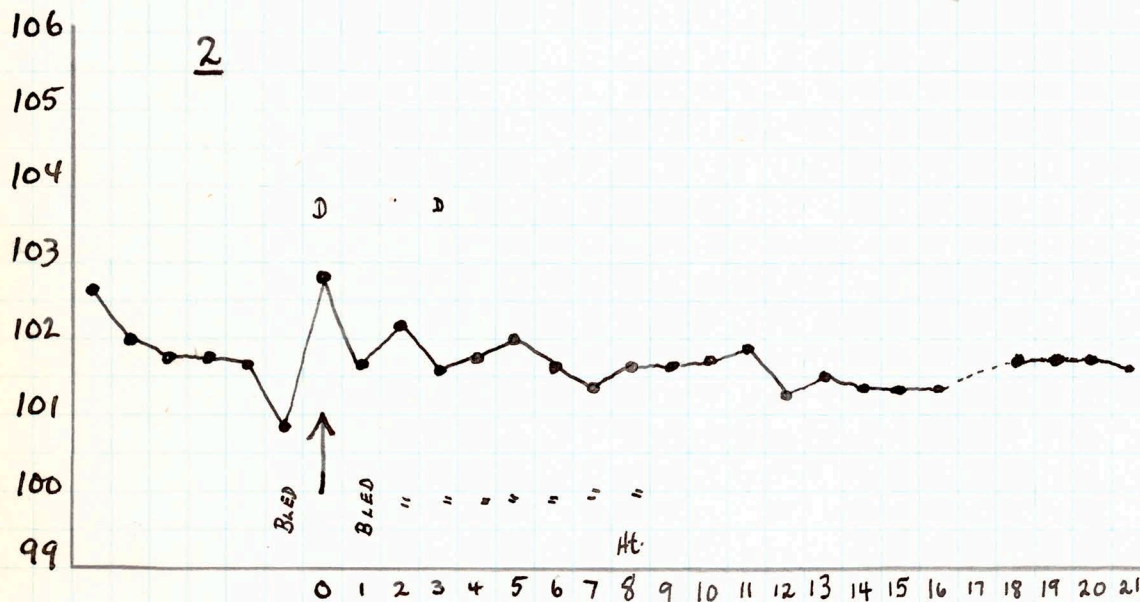
75 μ SERUM - 5cc.

Rhesus Monkeys

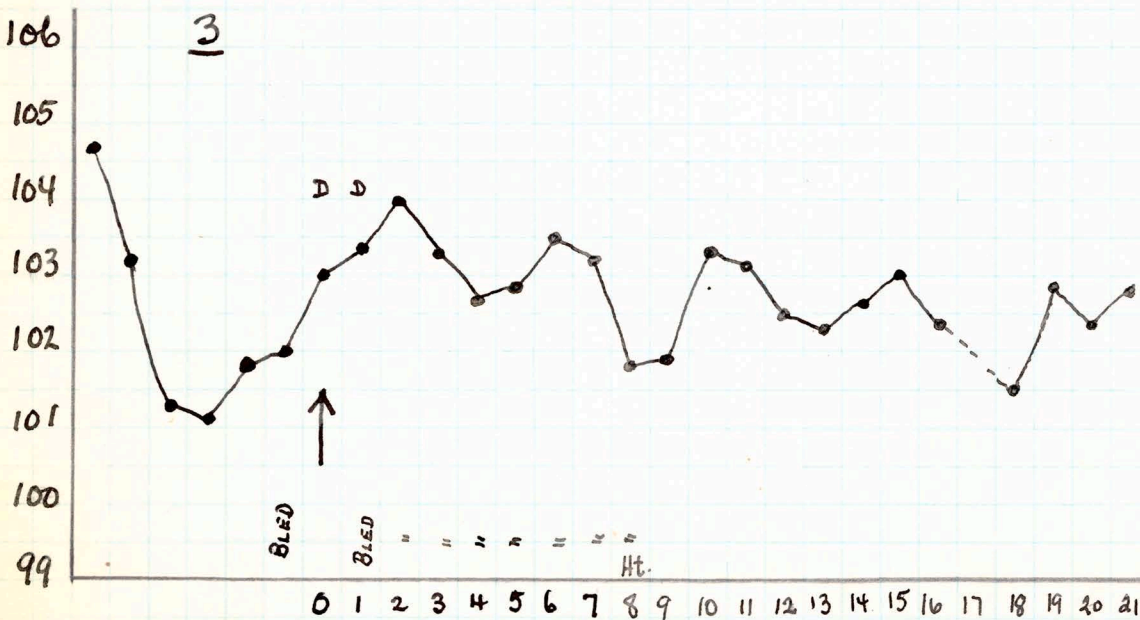
1



2



3



Oct. 14 15 16 18 19 20 21 22 23 24 25 26 27 28 29 30 31 / 1 2 3 4 5 6 7 8 9 10 11

Purpose - to determine whether passage of virus from human to human by means of serum inoculation results in attenuation of virus.

Reason - in the filtration experiment the unfiltered centrifuged virus failed to produce the disease in 2 subjects. The virus ~~was~~ must have been present since at least one of the subjects inoculated with the 770 mu filtrate developed the typical disease. The following questions were raised:

1) Could the previous treatment have modified the response to the virus?

An analysis of the records of these patients did not lend support to this

2) Is there great individual variation in susceptibility?

3) Is the virus becoming attenuated?

The following preparations were inoculated into each of 2 volunteers:

Original - serum from cases occurring in nature (Sicily)
Carr and Datoush sera thawed second time and 1.1 cc removed from each for pool

1st passage - Irwin, Walters, Riggs, Morse - who developed exp. disease after inoculation with "original" serum - sera thawed first time - 0.6 cc from each mixed to form pool

2nd passage - Quer's serum - developed exp. disease after inocul. with 1st passage - 770 mu filtrate 5 cc of 1:5 dilution of serum.

Inoculations - Each subject received

0.2 cc intracut.; on flexor surf. of right forearm
0.8 cc subcut. in right arm.

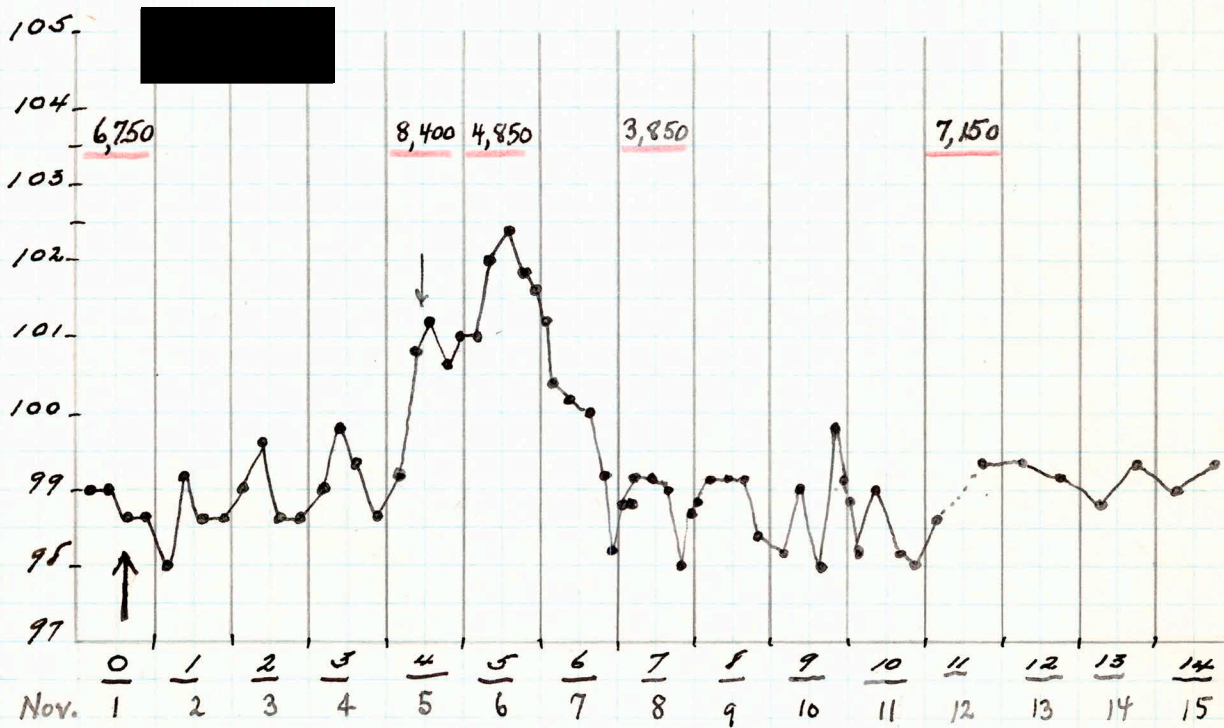
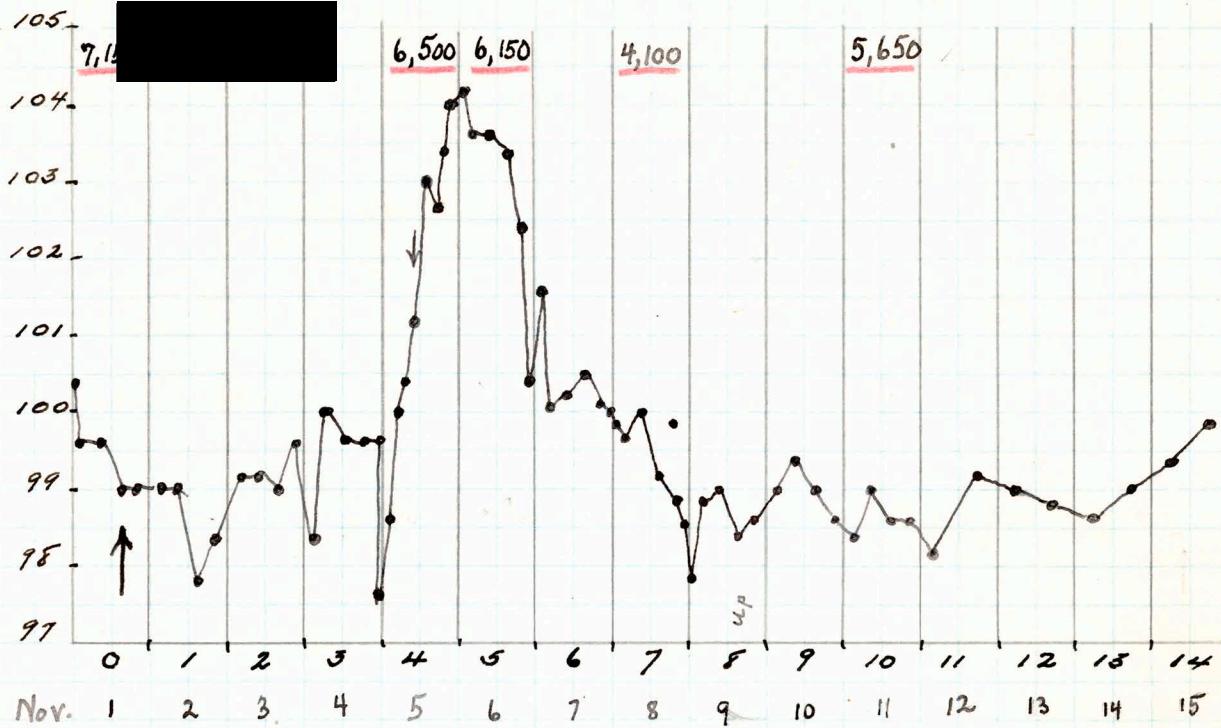
All inoculated between 2⁰⁰ and 2¹⁵ P.M.

Original	—	[redacted]	—	white male	—	52
				white female	—	45
1 st passage	—	[redacted]		white male	—	48
				white female	—	48
2 nd passage	—	[redacted]		white male	—	41
				white female	—	48

Reactions to Intracutaneous Injection to 0.2cc
of Above Sera

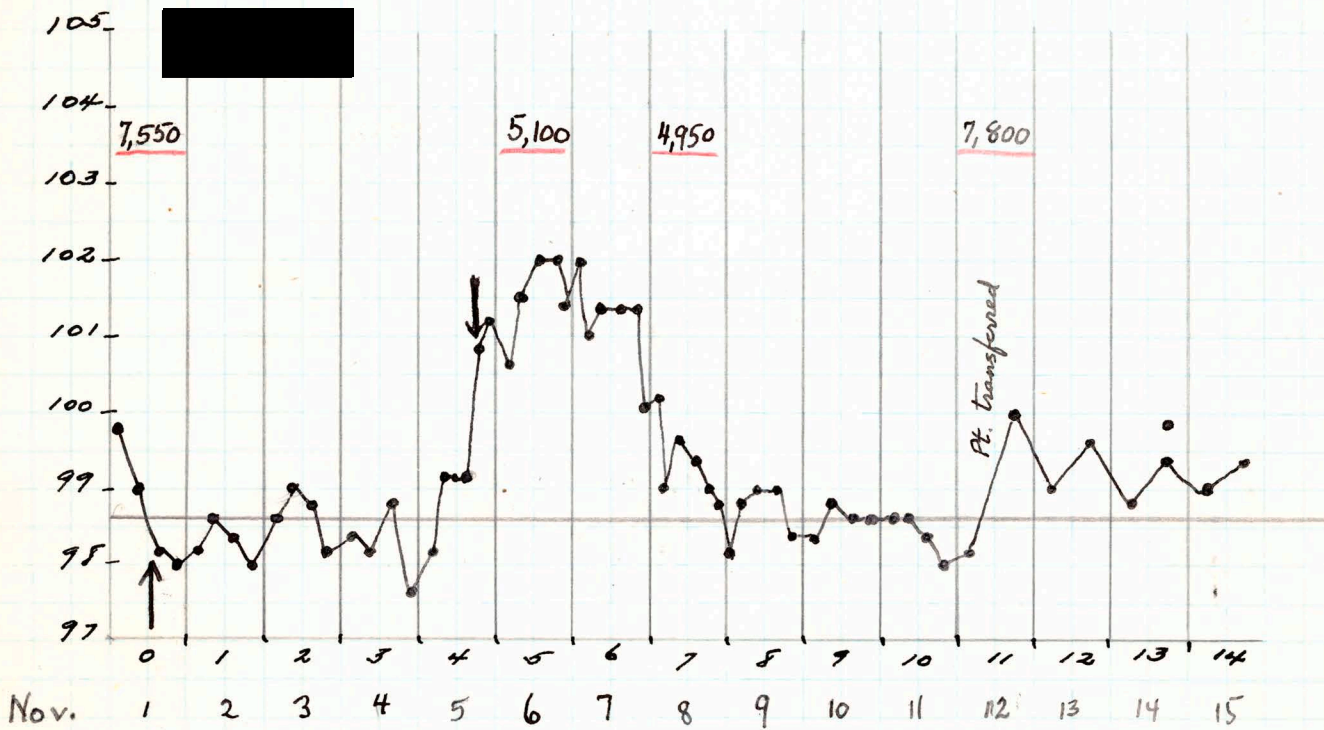
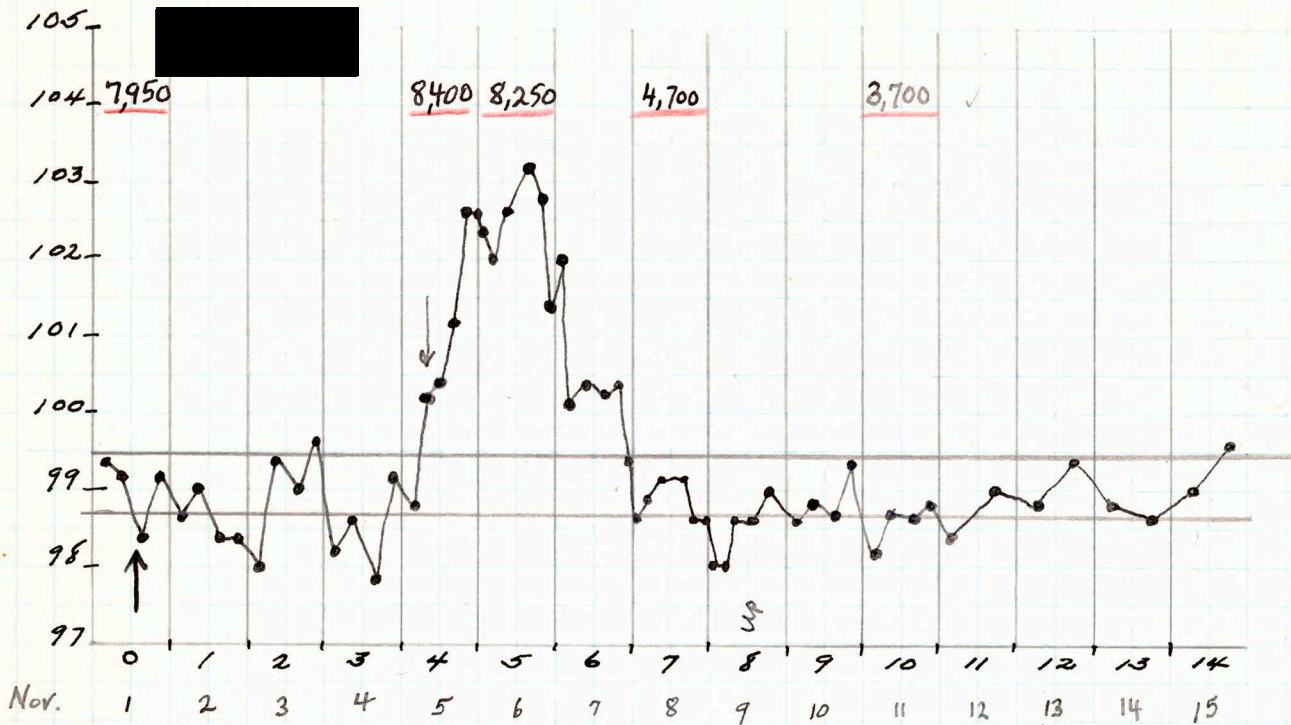
<u>24</u> hours	—	all	completely	negative
<u>48</u> "	—	"	"	"
<u>72</u> "	—	"	"	"

SICILIAN VIRUS



ORIGINAL - 0.2 cc. i.c.,
0.8 cc. subcut.

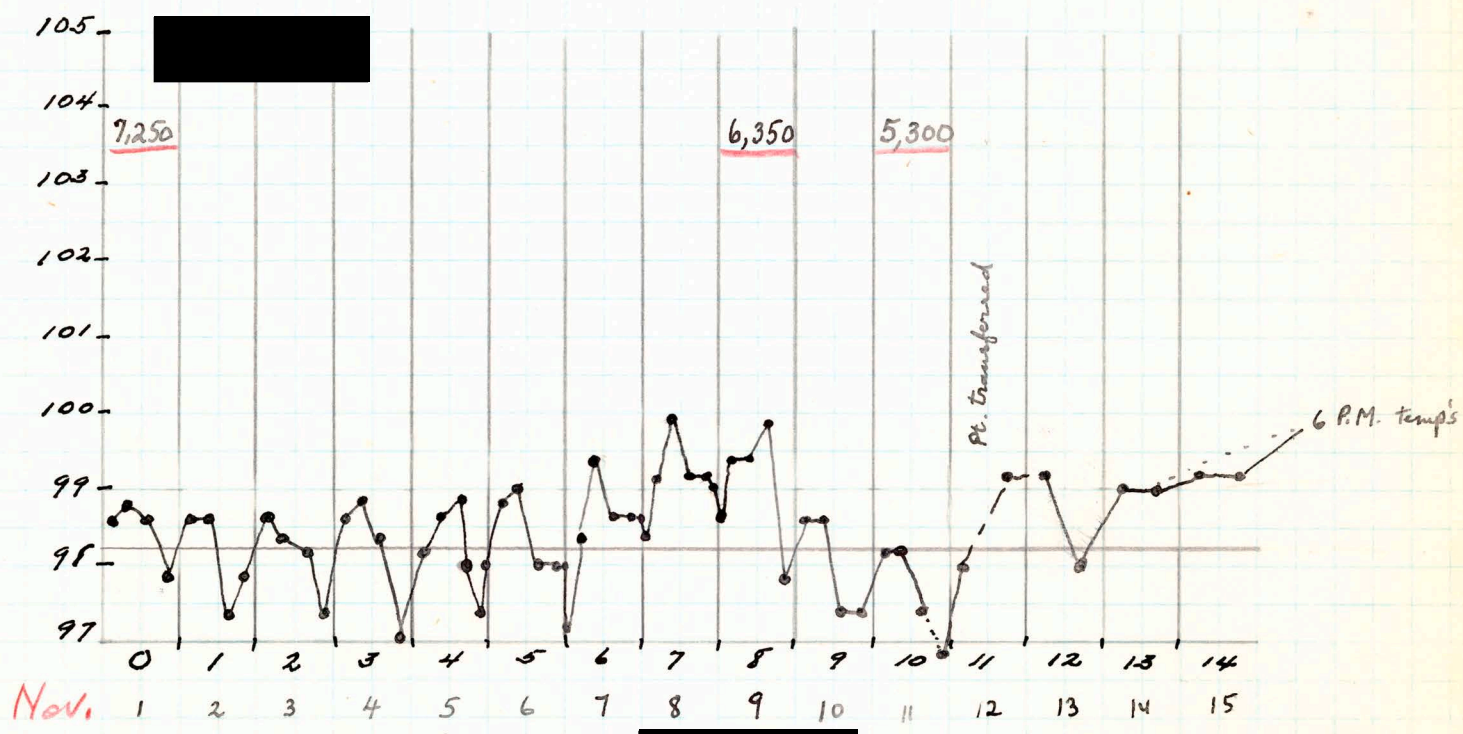
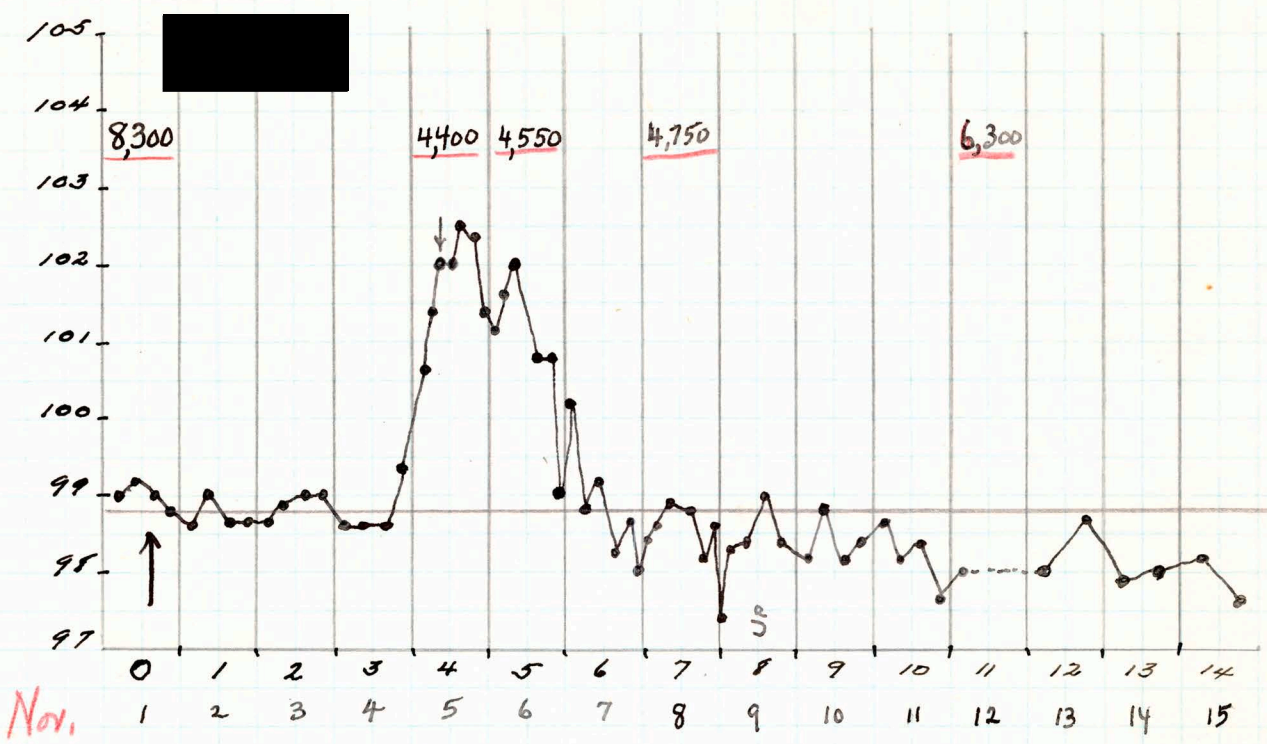
SICILIAN VIRUS



1ST PASSAGE - POOL OF

0.2 cc i.c.t.
0.8 cc subcut.

SICILIAN VIRUS



2ND PASSAGE -

0.2cc i.c. cut
0.8cc s.c. subcut.

SICILIAN VIRUS

Nov. 9, 1943

- A. Immunity to homologous virus 1 month after onset of fever
- B. Influence of route of inoculation on development of disease

VIRUS - First passage had made up of

Routes of inoculation - intracutaneous - 1cc of undiluted serum intracutaneously ~~with~~ in 0.2cc amounts in 5 places on flexor surface of forearm

subcutaneous - Serum diluted 1:5 in saline 5cc of diluted serum subcut. above olecranon in arm.

intramuscular - 5cc of diluted serum into deltoid as follows:
Syringe containing saline used to insert 1" 23G. needle perpendicularly into muscle - syringe was then removed and the one containing serum inserted into needle and inoculation made - syringe again removed and one c saline attached; about 0.5cc of saline injected to wash thru needle

Convalescents

W.M. - 41 -	1cc intracutaneously	left forearm
" " - 38 -	"	" " "
W.F. - 44 -	"	" right "
" " - 48 -	"	" "

Normal controls

W.M. - 38 -	"	"	right	"
W.F. - 43 -	"	"	"	"
C.M. - 43 -	" (5cc 1:5)	intramusc.	left	deltoid
W.M. - 44 -	" " "	"	"	"
W.F. - 42 -	" " "	subcut.	"	arm
" " - 48 -	" " "	"	right	"

Nov. 10 - No reaction at any of the inoculated sites except in Walters who showed area of erythema 10x8mm. at one of the 5 i.cut. sites.

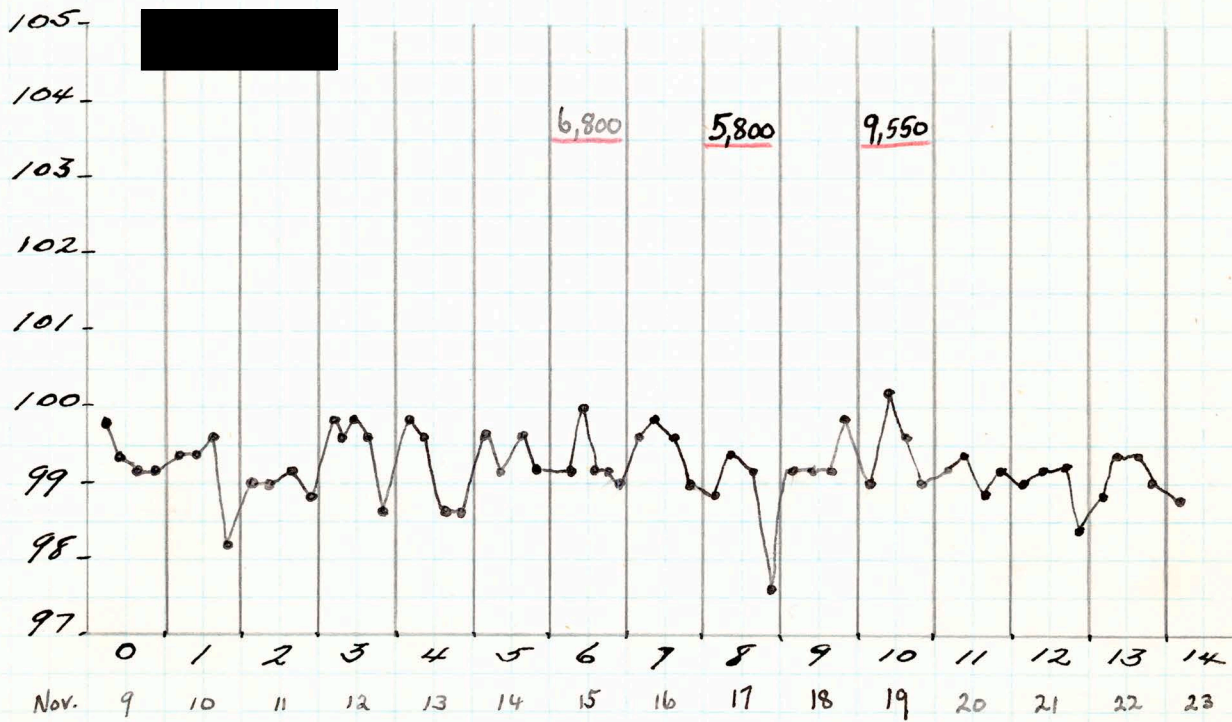
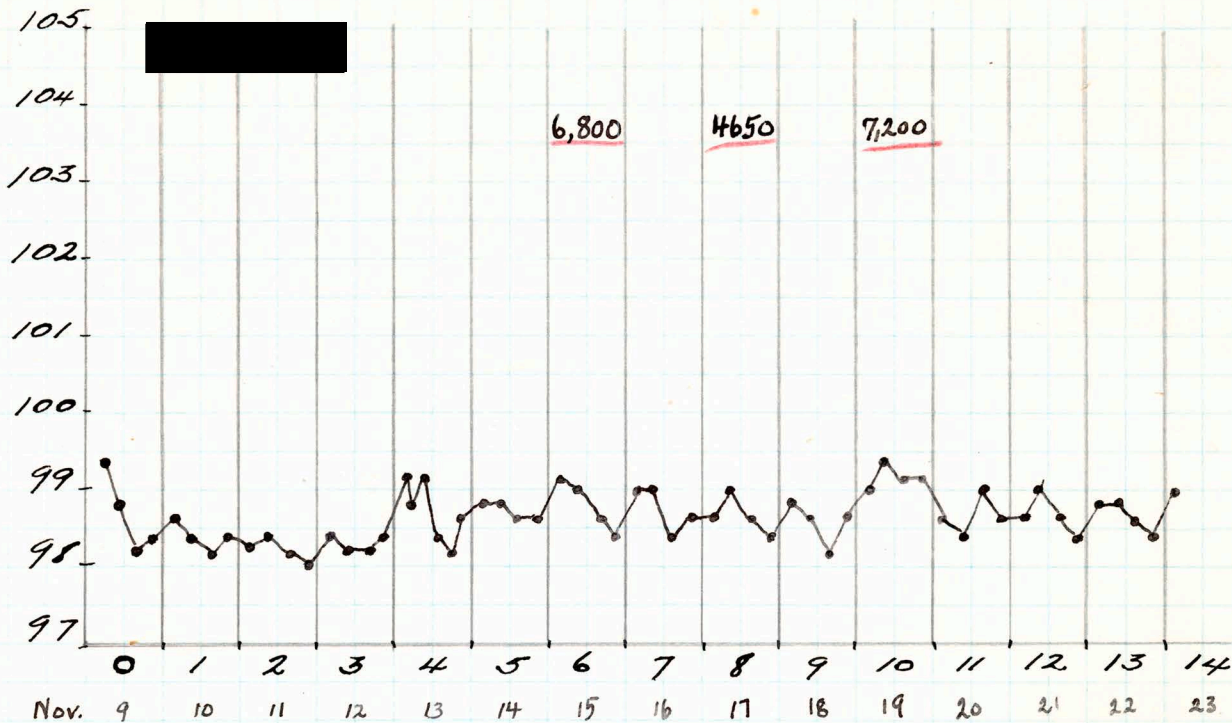
Nov. 11 - all neg.

" 12 - " "

" 13 - " "

SICILIAN VIRUS

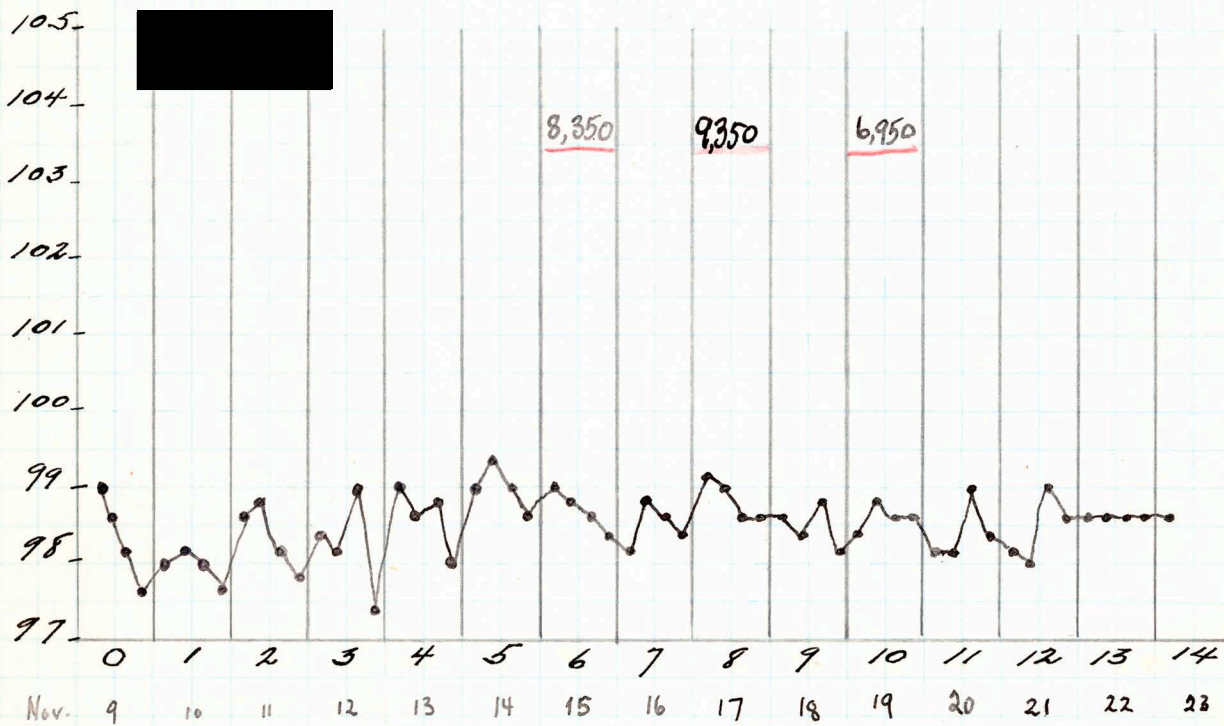
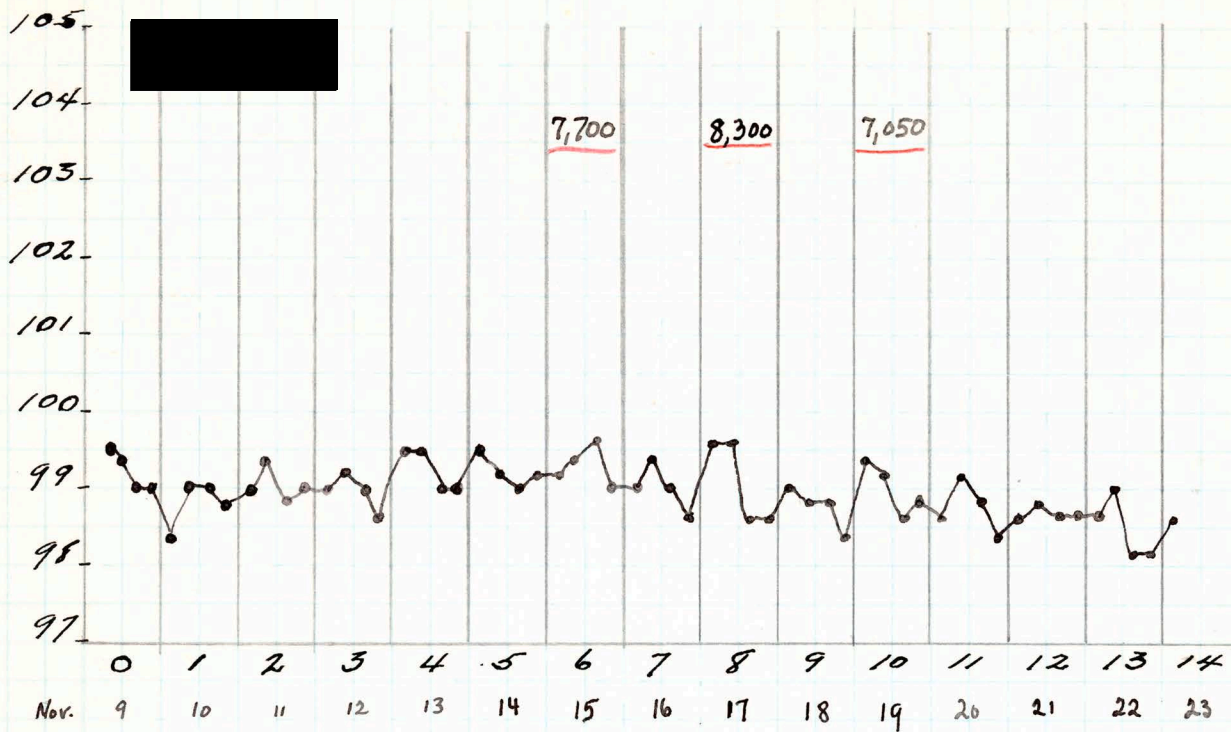
ONE MONTH IMMUNITY TEST - RECOVERED SUBJECTS



1 cc INTRACUTANEOUSLY

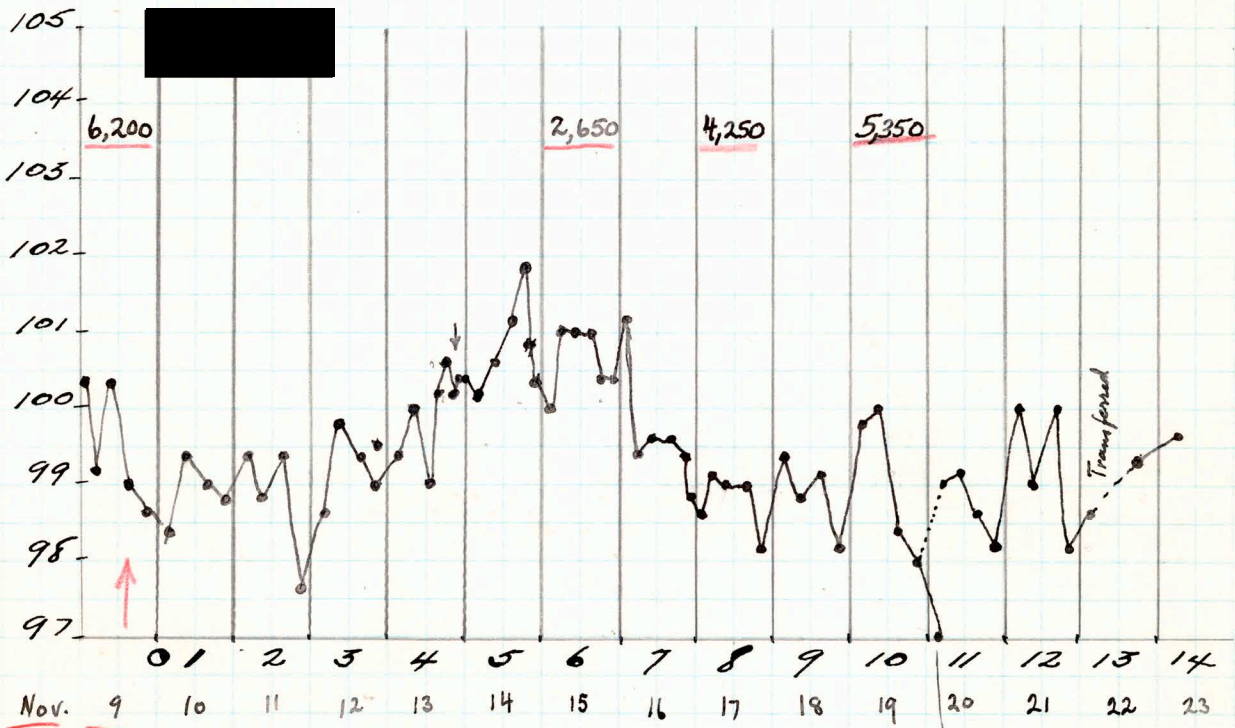
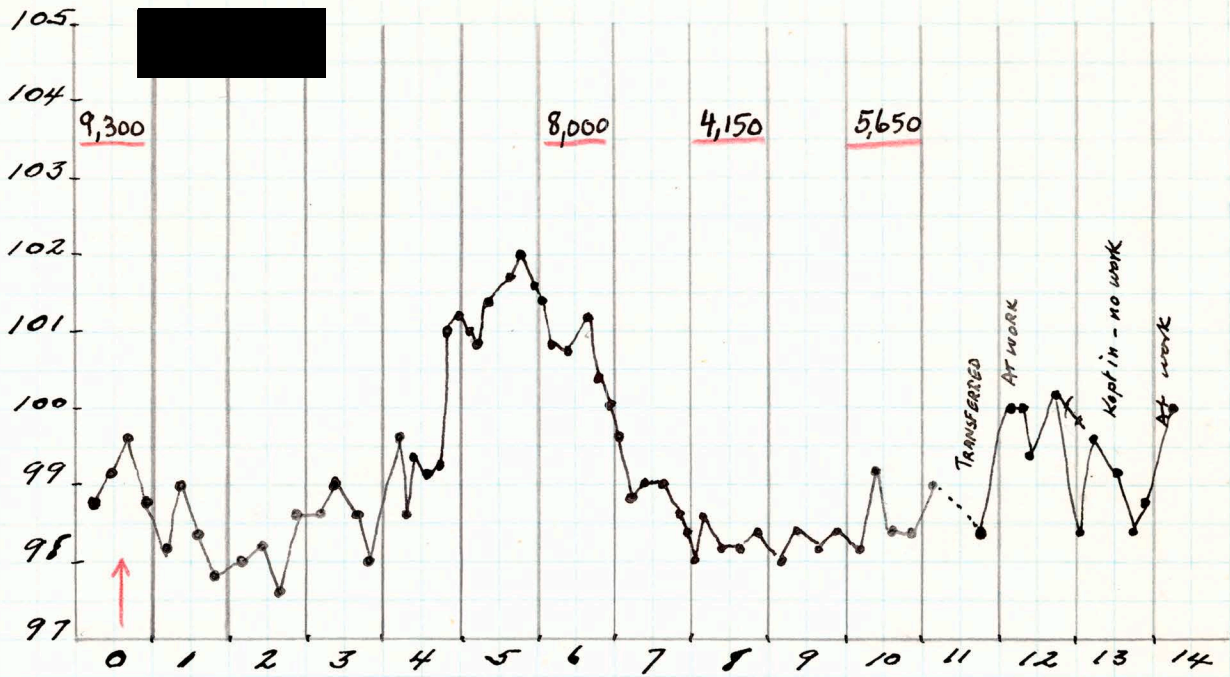
FIRST PASSAGE POOL - [REDACTED]

SICILIAN VIRUS - IMMUNITY TEST - RECOVERED SUBJECTS



1cc INTRACUTANEOUSLY

SICILIAN VIRUS - IMMUNITY TEST - CONTROLS

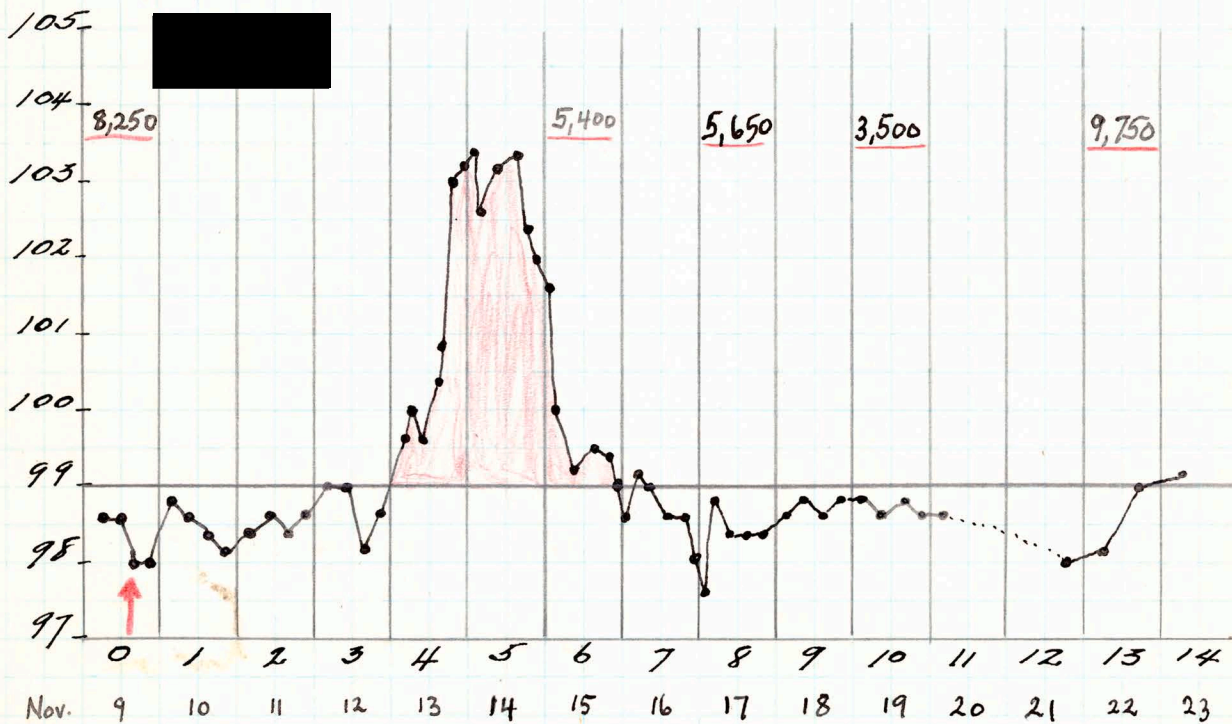
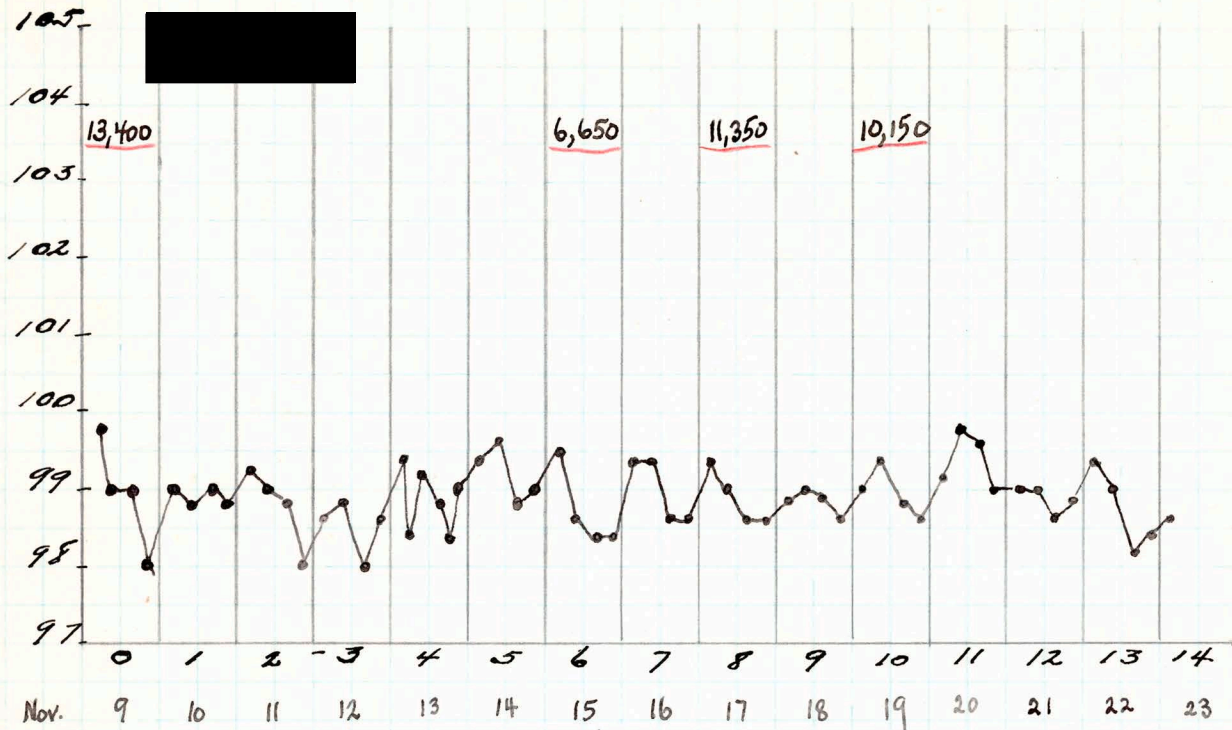


Temp. probably also influenced by activity

checked - pt. observed to pull thermometer part way out - peculiar fever curve may be due to this also

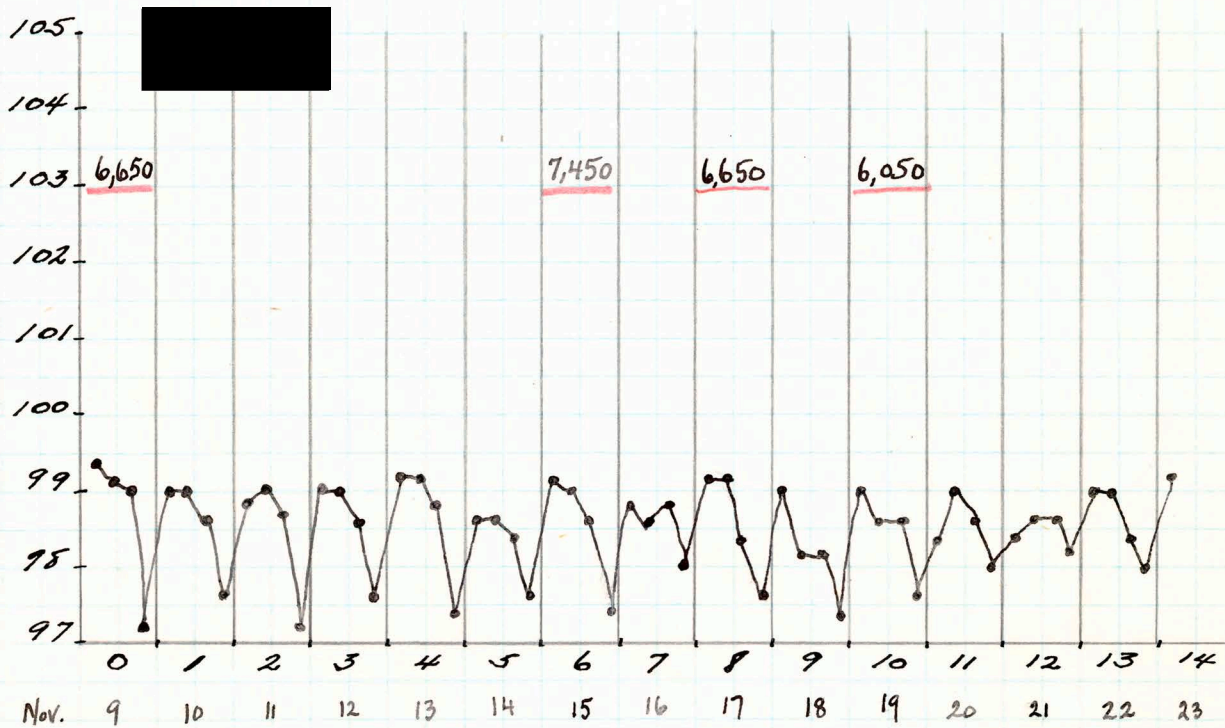
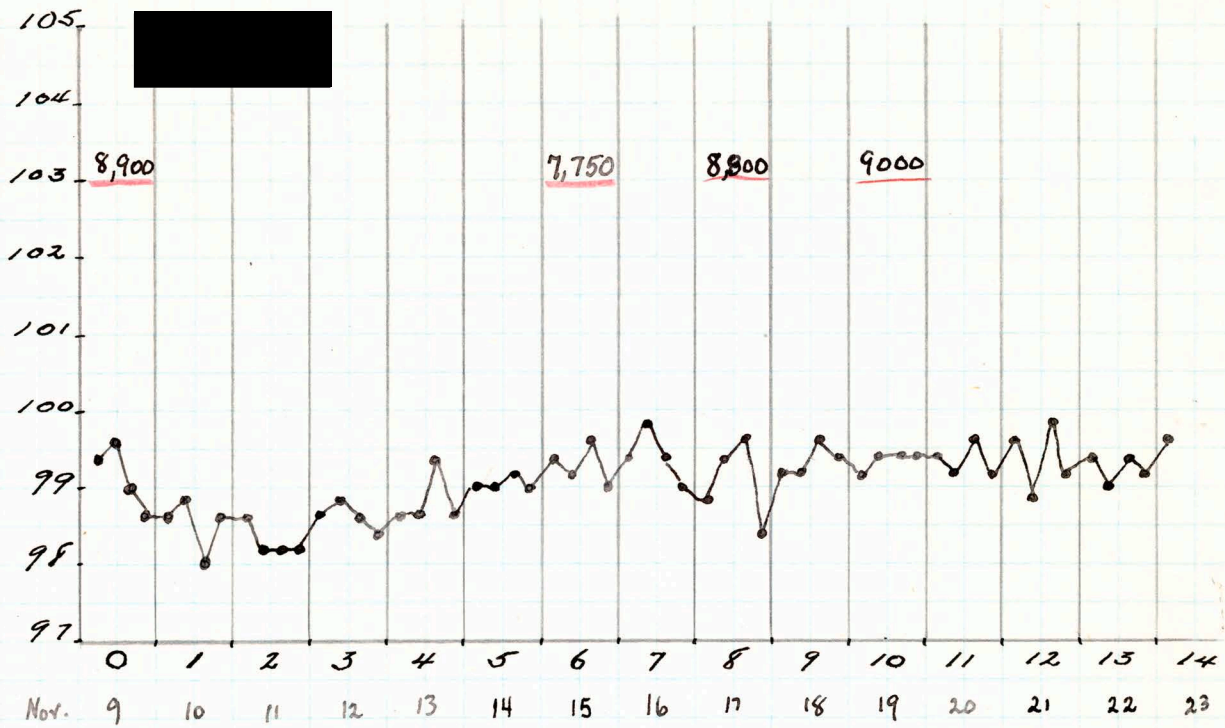
1cc INTRACUTANEOUSLY

SICILIAN VIRUS - INFLUENCE OF ROUTE OF INOCULATION



1cc (5cc of 1:5 dilution) INTRAMUSCULARLY
 FIRST PASSAGE POOL -

SICILIAN VIRUS - INFLUENCE OF ROUTE OF INOCULATION



1cc (5cc of 1:5 DILUTION) SUBCUTANEOUSLY

SICILIAN VIRUS - CROSS IMMUNITY WITH MIDDLE EAST VIRUS

IMMUNITY FOLLOWING INOCULATION OF DILUTED VIRUS BY I.MUSC. OR SUBCUT. ROUTES

1. Resistance of individuals recovered from and proved to be immune to the Middle East Virus to inoculation with the Sicilian Virus [redacted]
2. Does immunity follow the inoculation of an infective amount of virus by the intramuscular [redacted] or subcutaneous [redacted] routes which does not give rise to clinical manifestations of the Disease?

VIRUS USED - Pool of sera representing the 2nd experimental passage of the Sicilian Virus in human subjects.

Kahn - neg	[redacted]	15.0
"	[redacted]	21.0
"	[redacted]	18.0
"	[redacted]	17.0
"	[redacted]	18.2
"	[redacted]	23.5
		<u>112.7</u>

Upon mixing in flask - there was only 105.7 for distribution.
 14cc. used in this experiment
 91.7cc distributed - 5cc each in 18 ampules
 17cc in one amp.
 Frozen in CO₂ box.

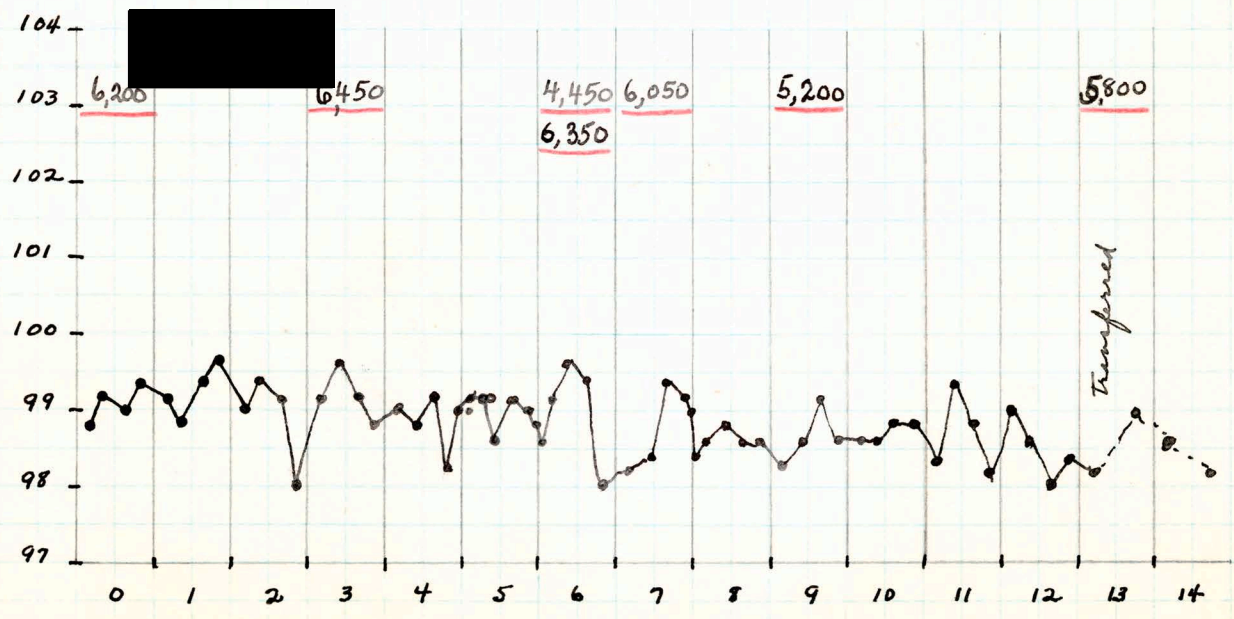
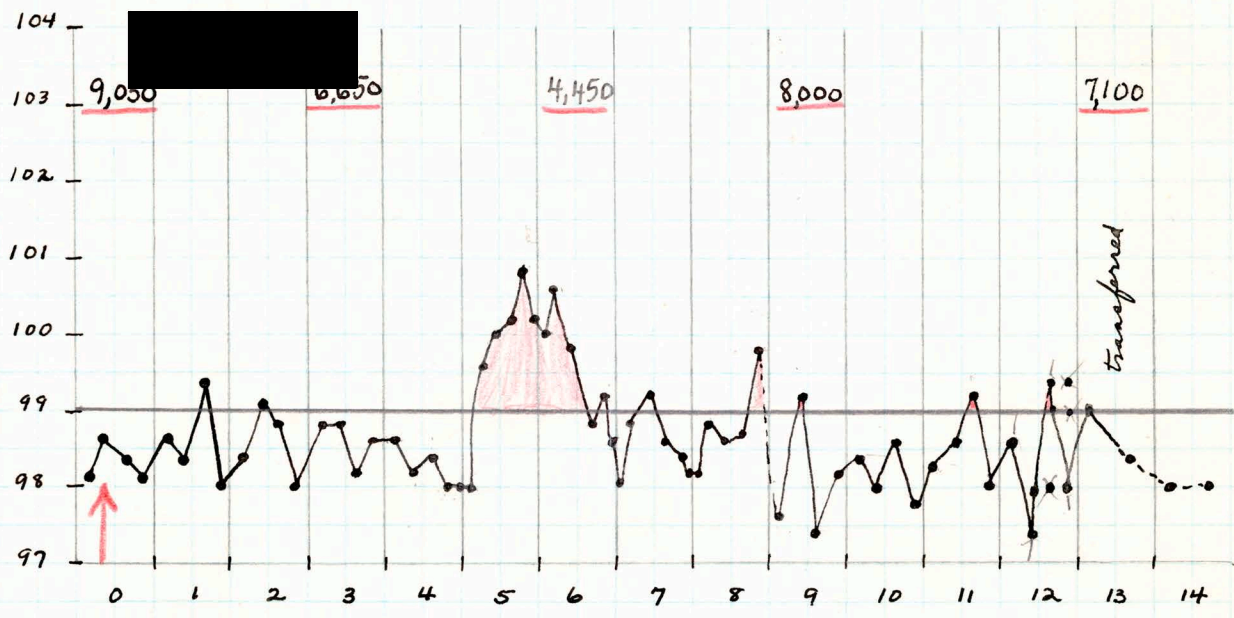
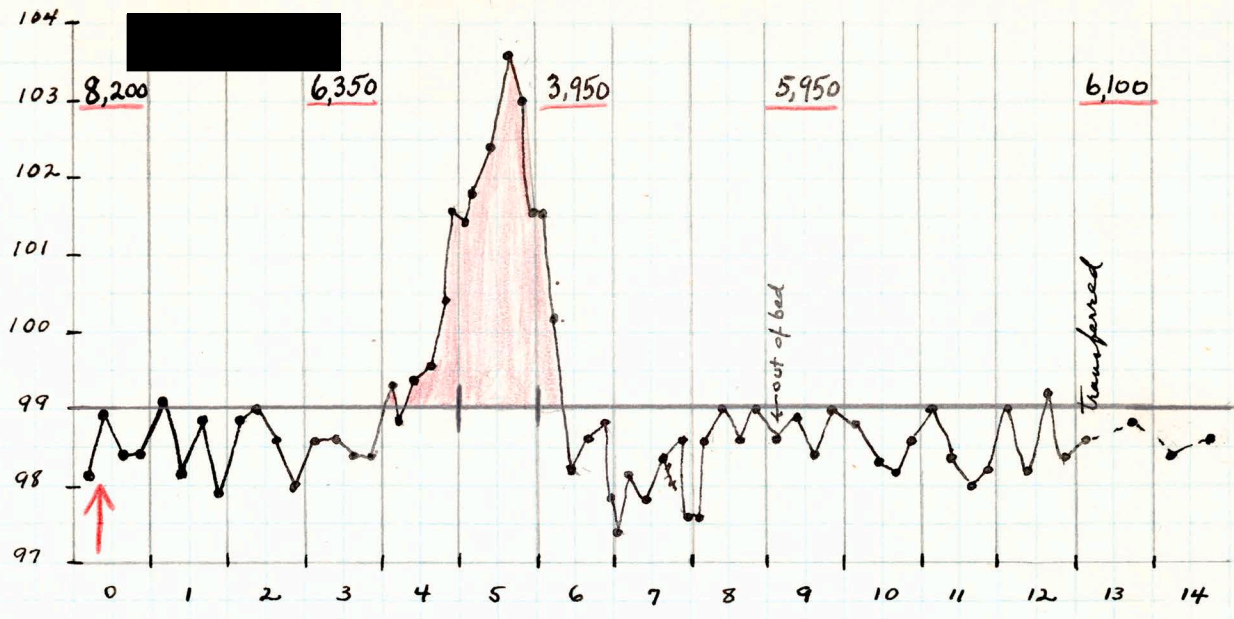
INOCULATIONS - 1cc intracutaneously on right forearm - 0.2cc in 5 places

Controls [redacted] W.M.-47
 [redacted] C.F.-48
 [redacted] C.F.-38

Immune to MIDDLE EAST VIRUS { [redacted]

I. musc. or Subcut. Inoculation of 5cc of 1:5 virus in saline { [redacted] ✓

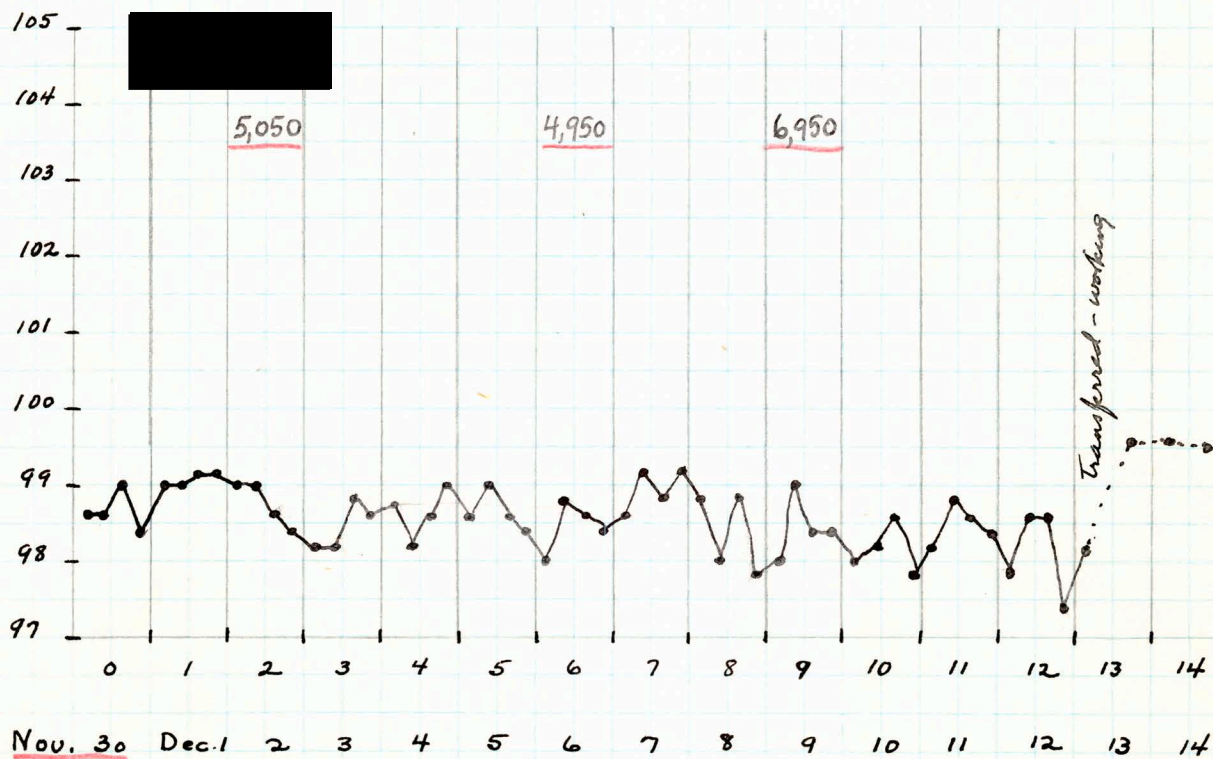
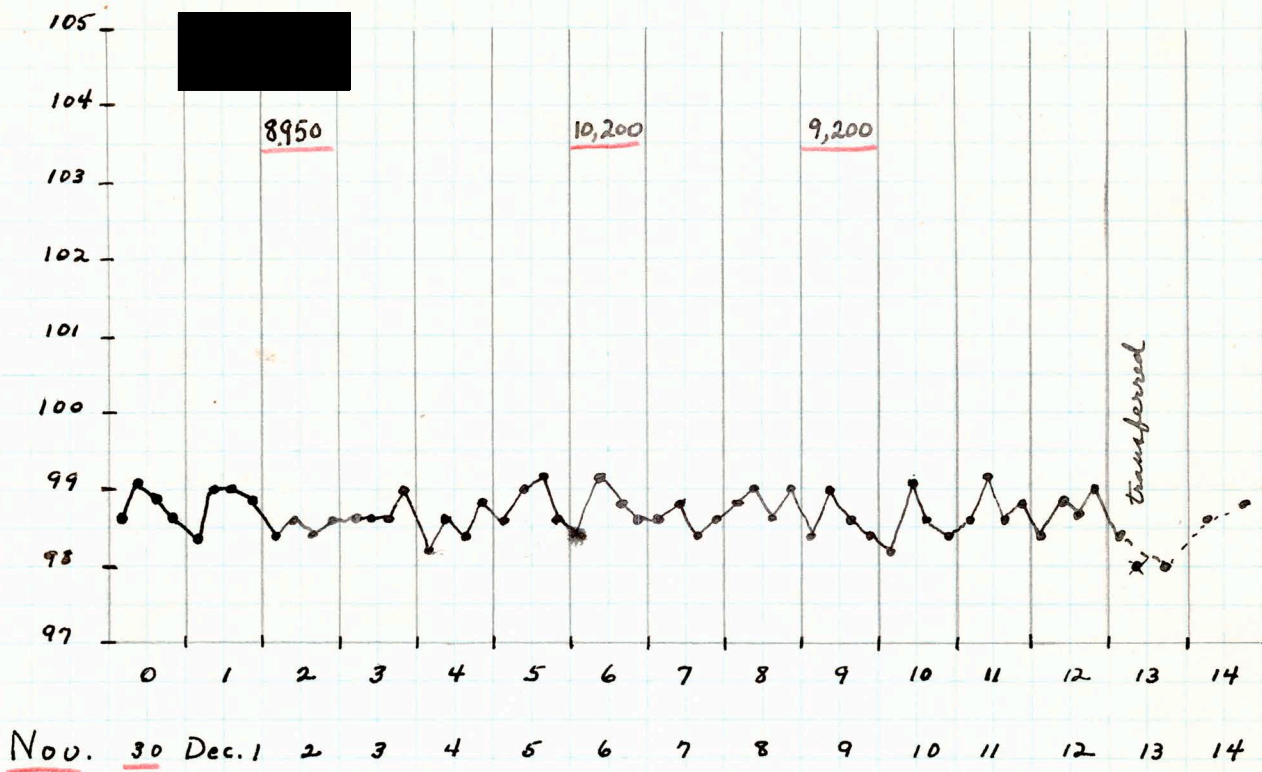
SICILIAN VIRUS - CONTROLS FOR IMMUNITY TESTS
 1cc. 2ND PASSAGE POOL INTRACUTANEOUSLY



Nov. 30 Dec. 1 2 3 4 5 6 7 8 9 10 11 12 13 14

SICILIAN VIRUS - CROSS IMMUNITY TEST

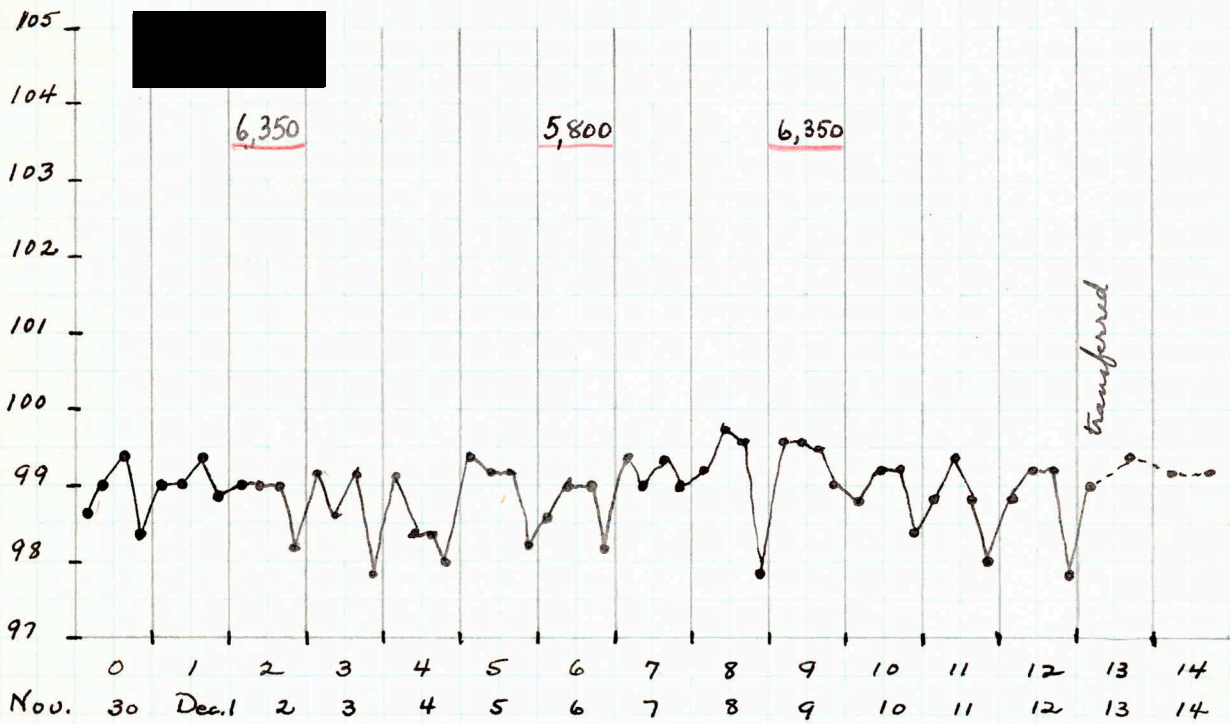
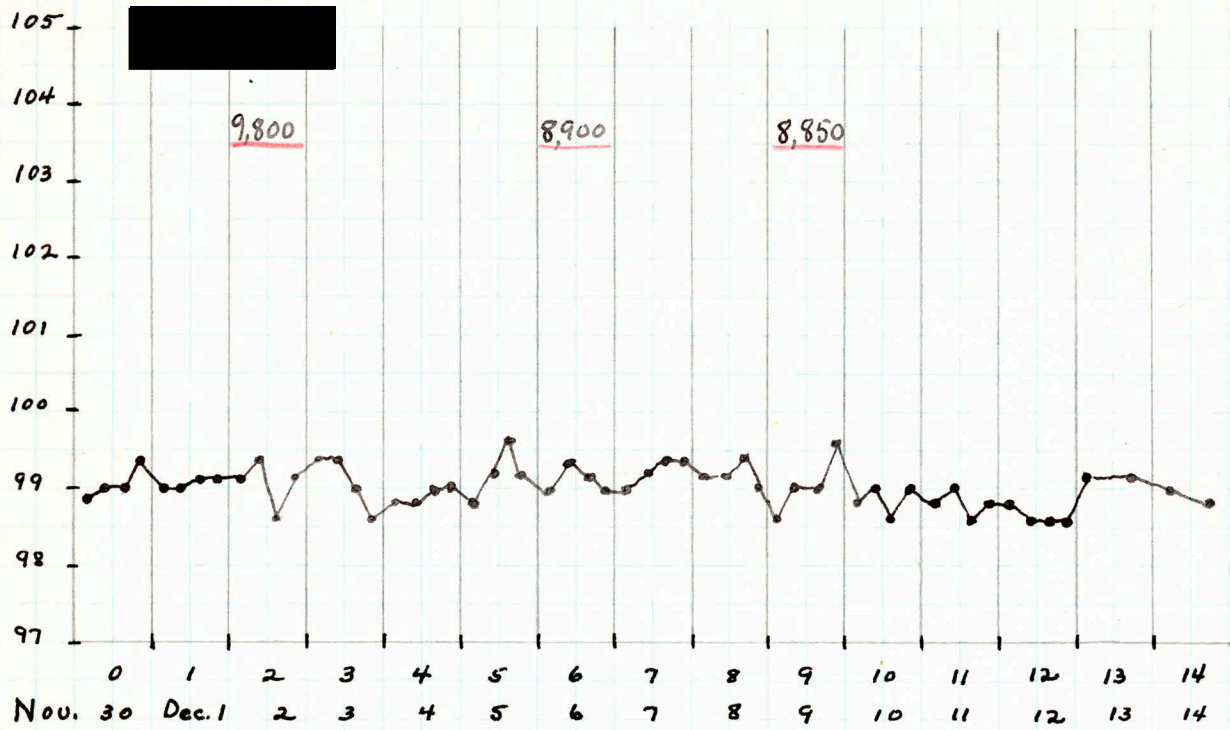
SUBJECTS RECOVERED FROM AND PROVED IMMUNE TO MIDDLE EAST VIRUS



1cc 2nd PASSAGE POOL INTRACUTANEOUSLY

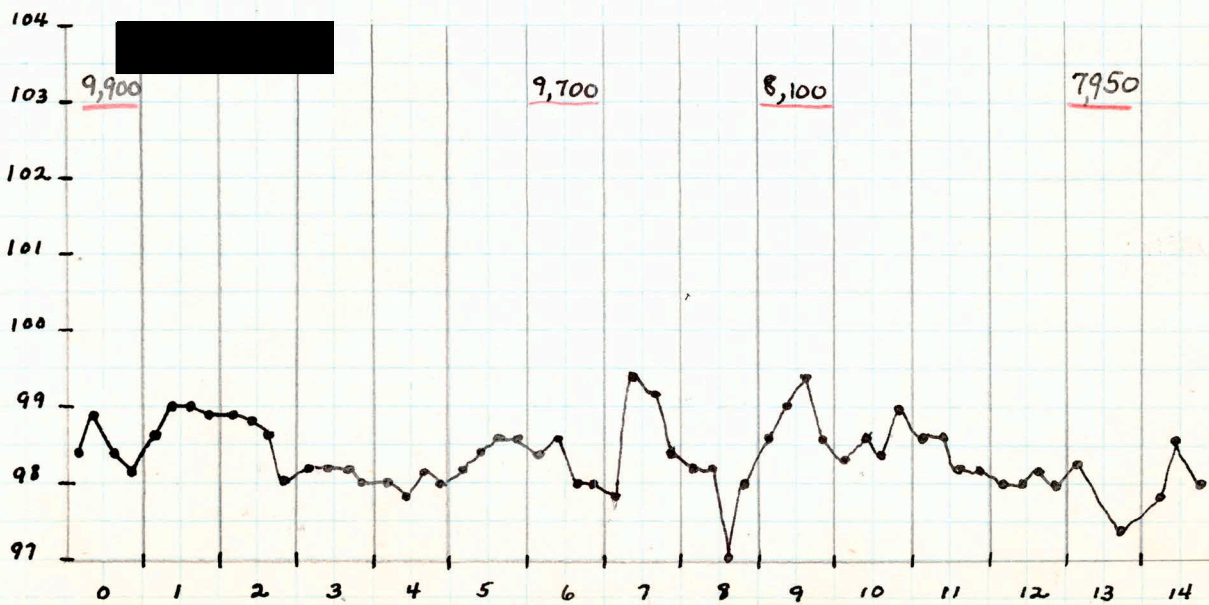
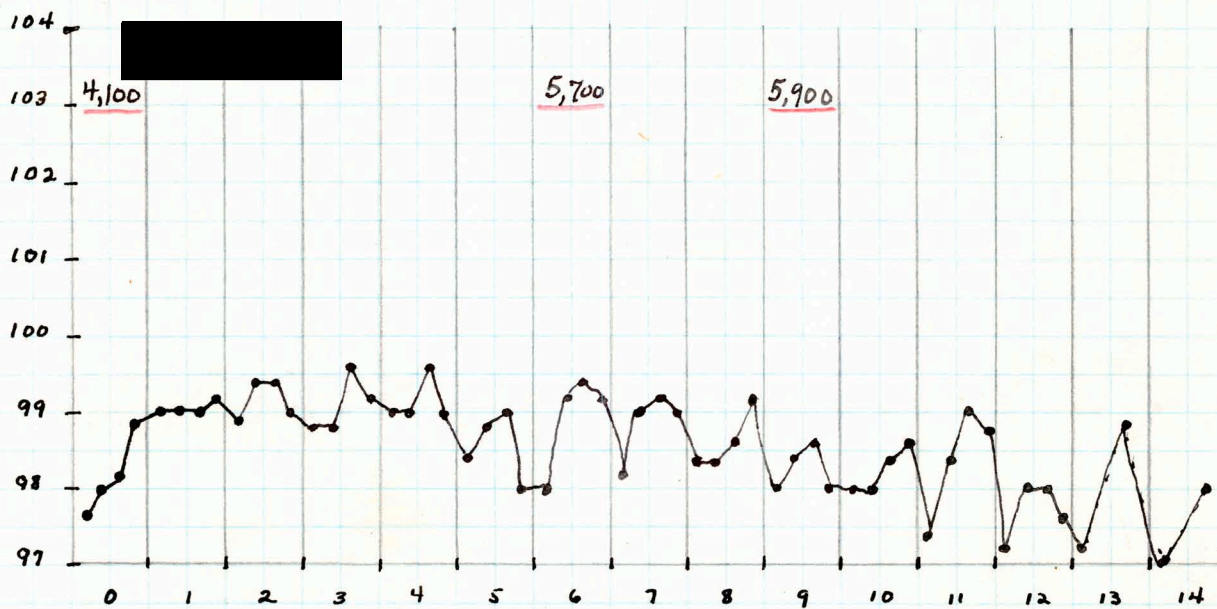
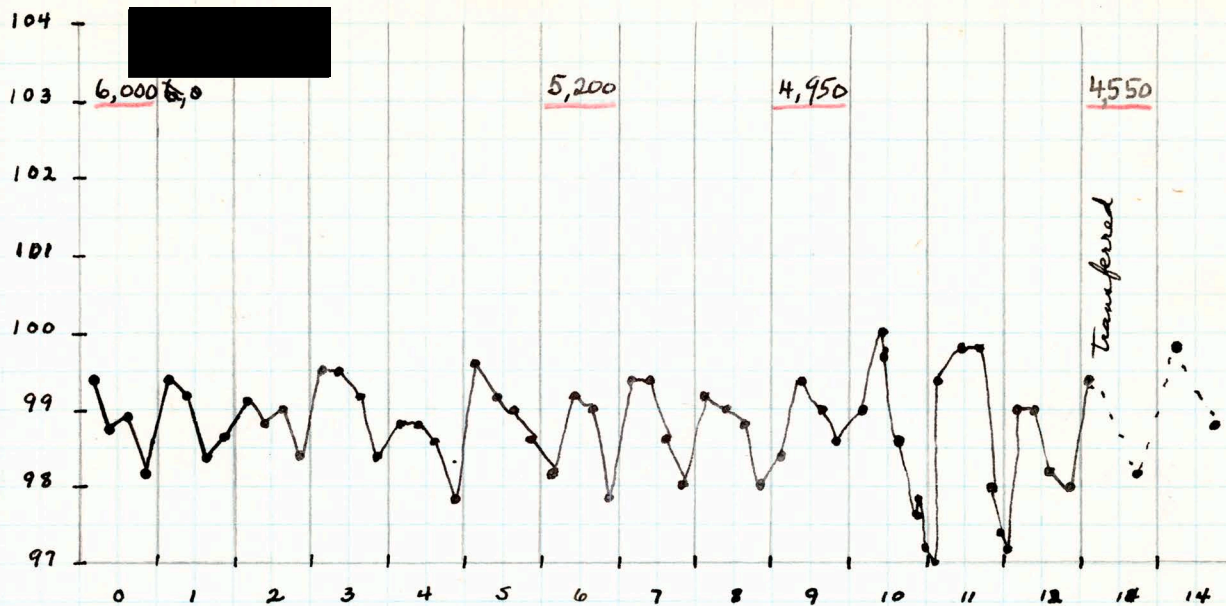
SICILIAN VIRUS - CROSS IMMUNITY TEST

SUBJECTS RECOVERED FROM AND PROVED IMMUNE TO MIDDLE EAST VIRUS

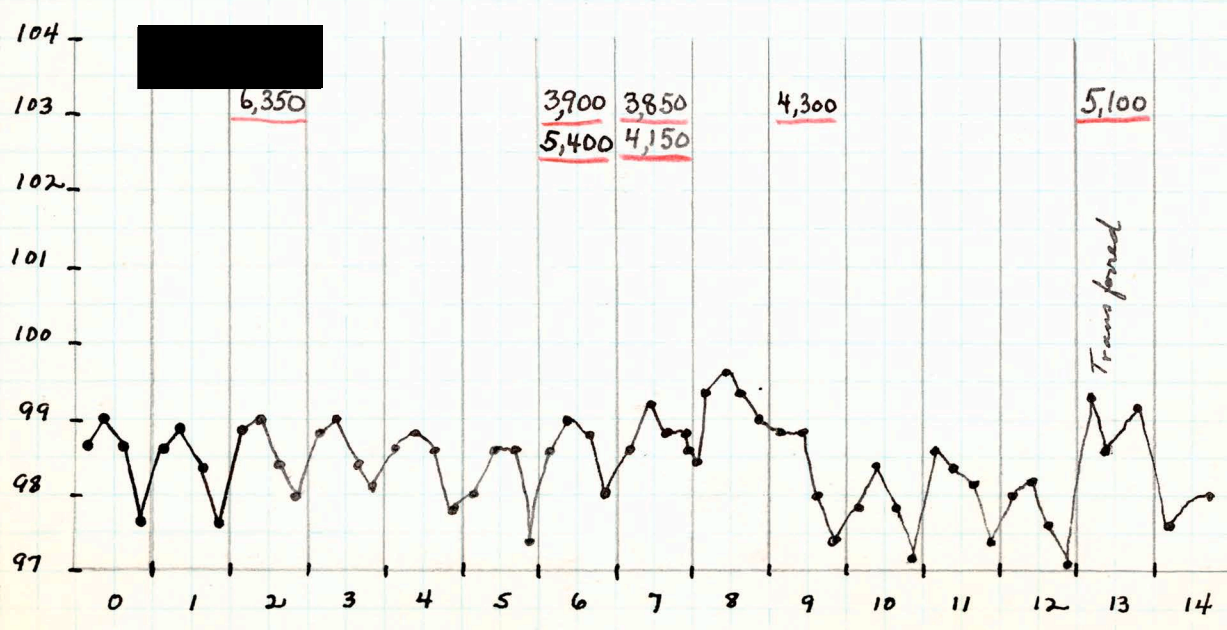
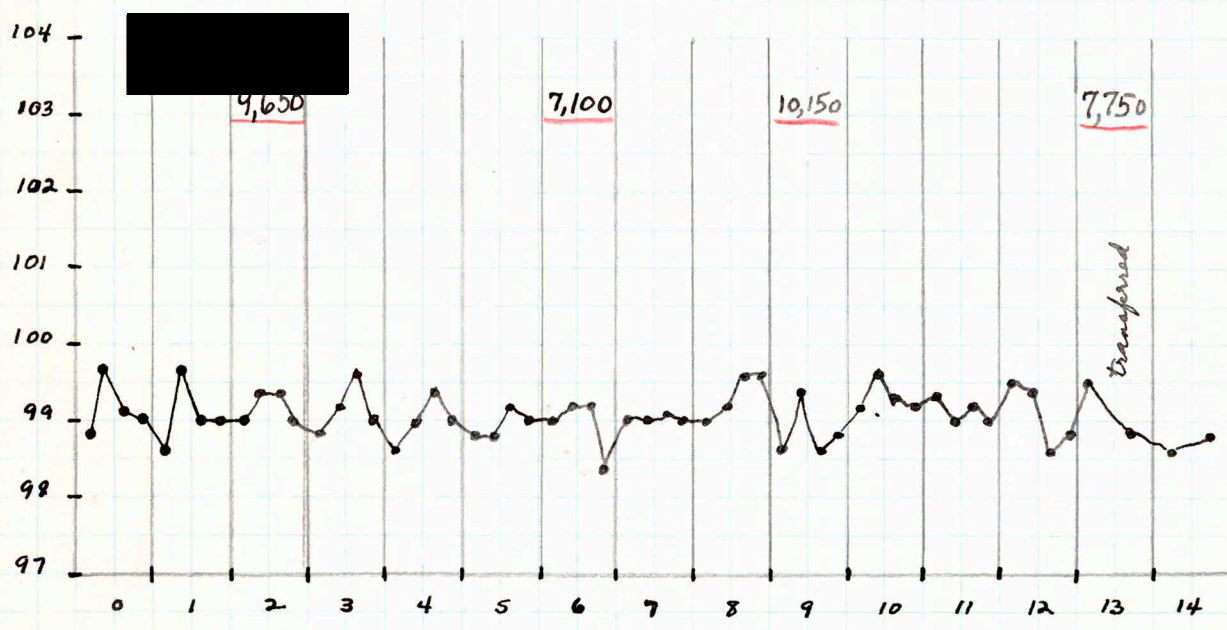
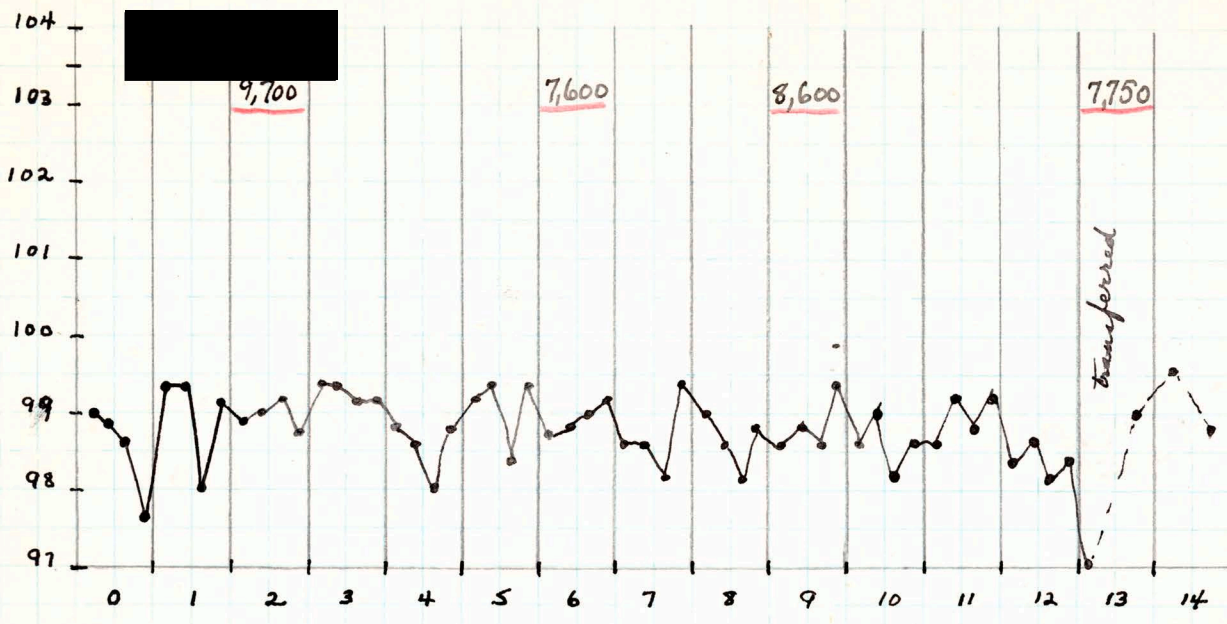


1cc. 2ND PASSAGE POOL INTRACUTANEOUSLY

SICILIAN VIRUS - IMMUNITY FOLLOWING INOCULATION OF DILUTED VIRUS
INTRAMUSCULARLY AND/OR SUBCUTANEOUSLY



Nov. 30 Dec. 1 2 3 4 5 6 7 8 9 10 11 12 13 14



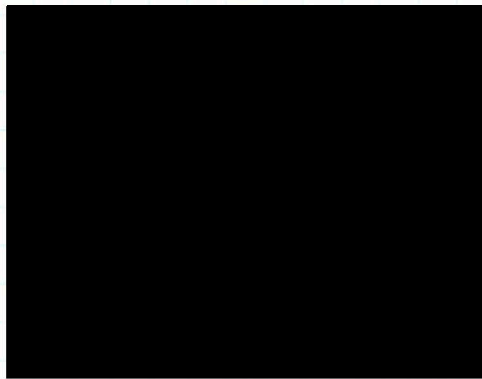
Nov. 30 Dec. 1 2 3 4 5 6 7 8 9 10 11 12 13 14

DEC. 14, 1943

SICILIAN VIRUS - 1. IMMUNITY OF INDIVIDUALS INOCULATED
WITH VARIOUS GRADOCOL MEMBRANE FILTRATES

2. To PROVIDE SUBJECTS FOR Aedes Aegypti TRANSMISSION EXPERIMENT

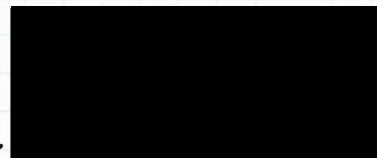
1. ON OCT. 21, 1943 the following were inoculated subcutaneously and/or intramuscularly with GRADOCOL membrane filtrates and failed to develop fever.



- 309 μ filtrate
- 203 " "
- 101 " "
- 75 " "
- 50 " "

[redacted] had a 3 day fever but it was associated with tonsillitis and she failed to develop a leukopenia

2. Controls for above - 1.



NEGRO MALE - AGE 35

2.

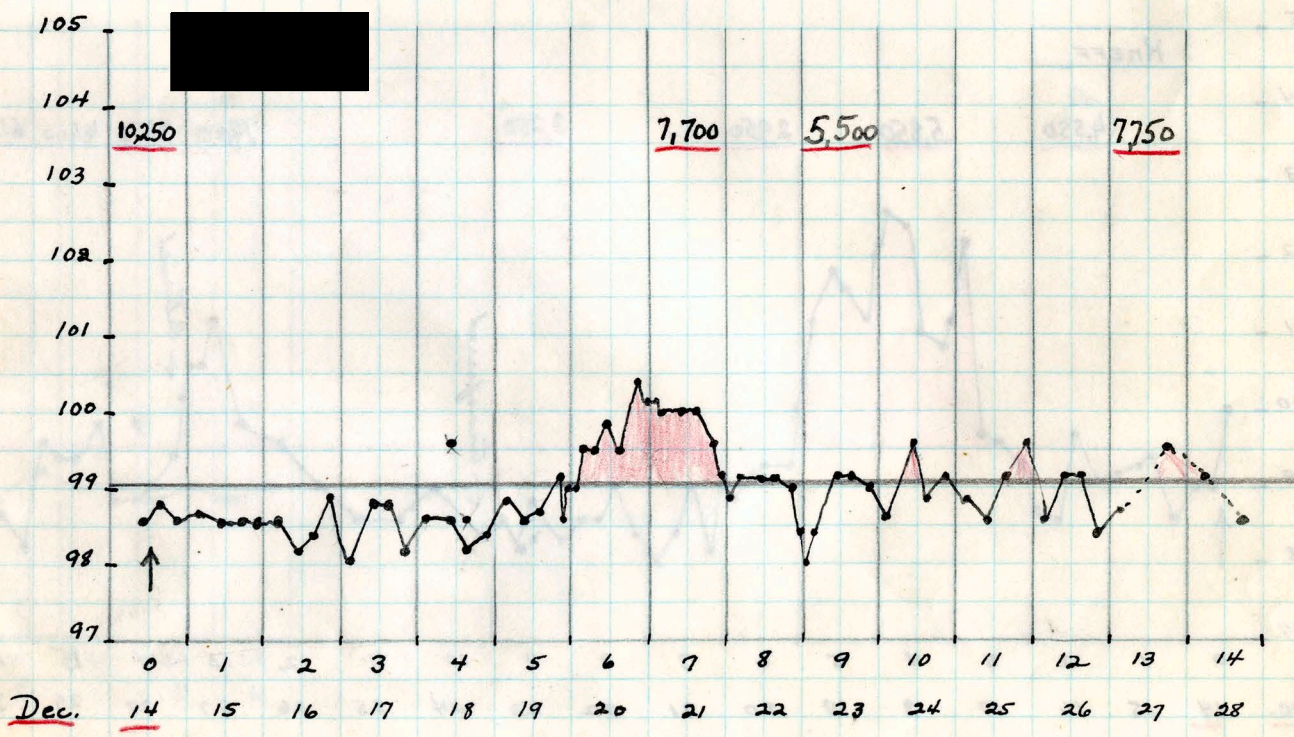
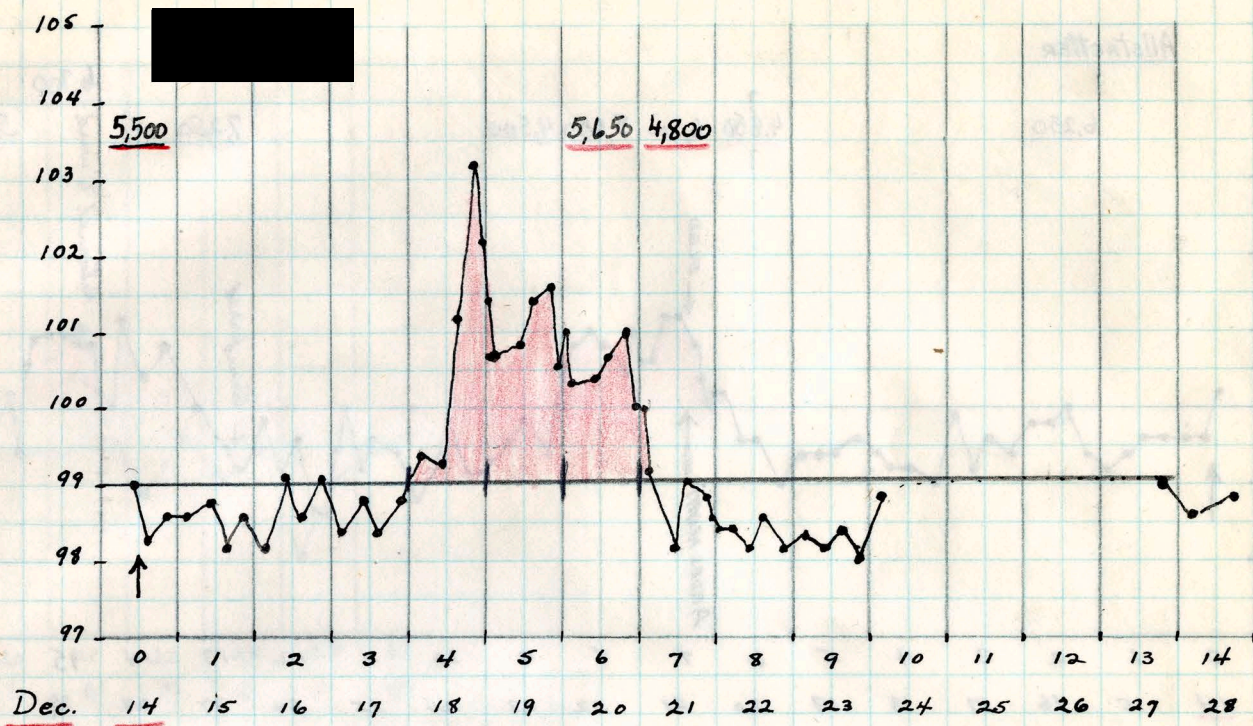
" " - " 45

VIRUS - 2ND PASSAGE Pool prepared Nov. 30 (used on Orben, Calmeise, and Bonner)

Dose and route - 1cc intracutaneously (men on rt. forearm, women on left) - 0.2cc in 5 sites.
Inoculations @ 2 P.M.

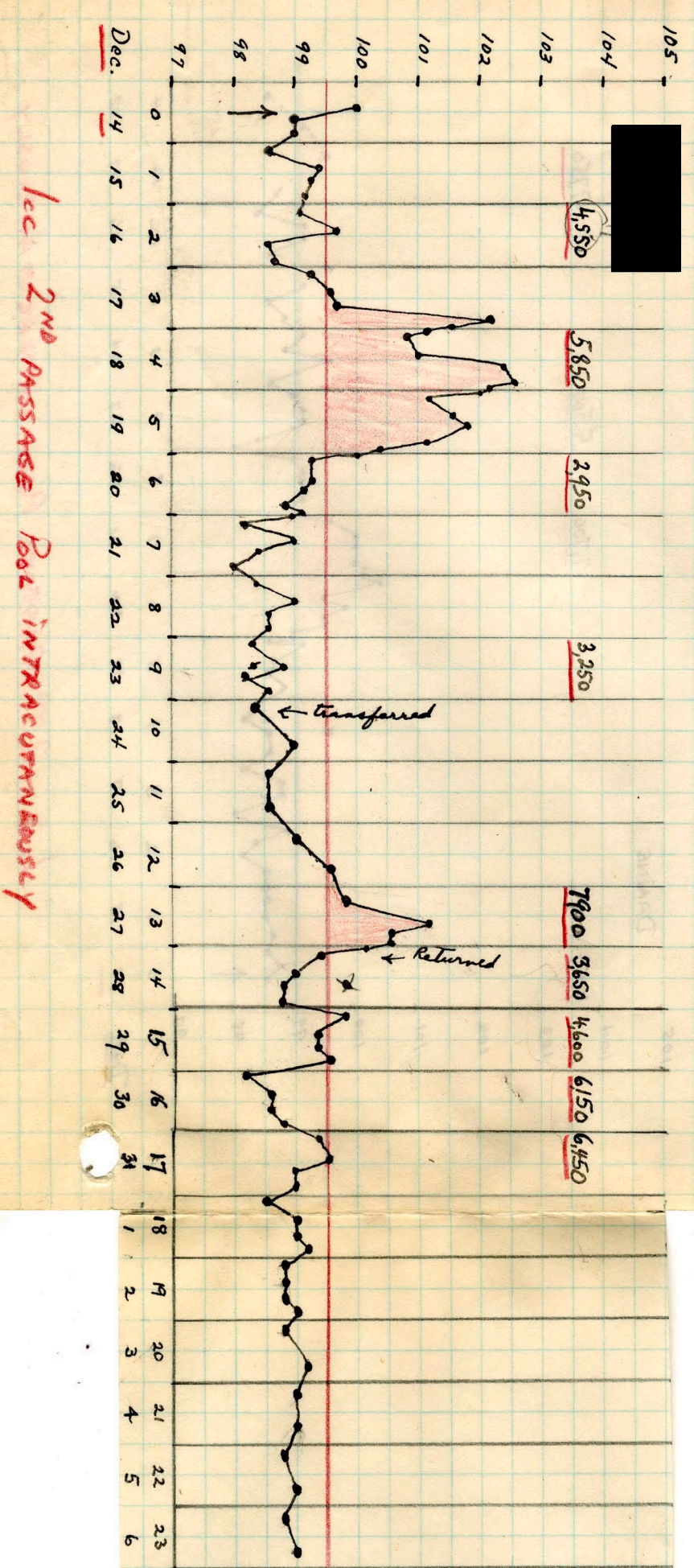
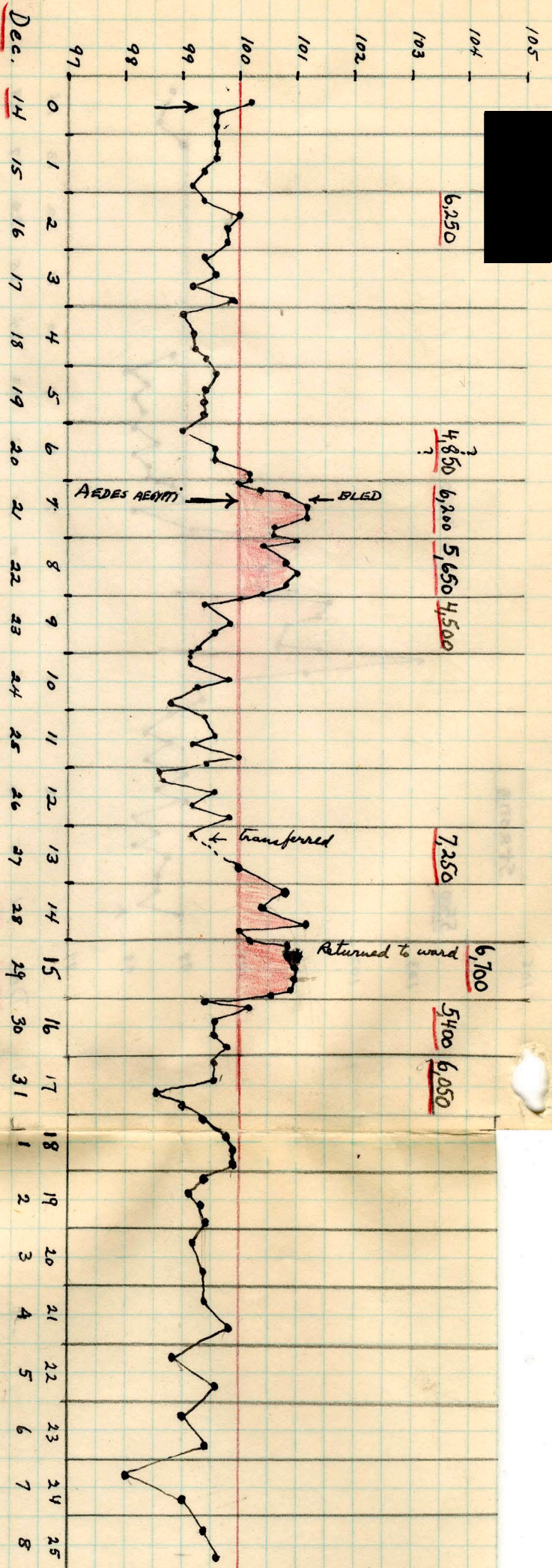
Dec. 15 - all show papules (\approx 5x5mm) and varying degrees of erythema at sites of inoculation (except Divido)

SICILIAN VIRUS - CONTROLS FOR GRADUOL MEMBRANE IMMUNITY TEST
 AND DONORS FOR Aedes Aegypti



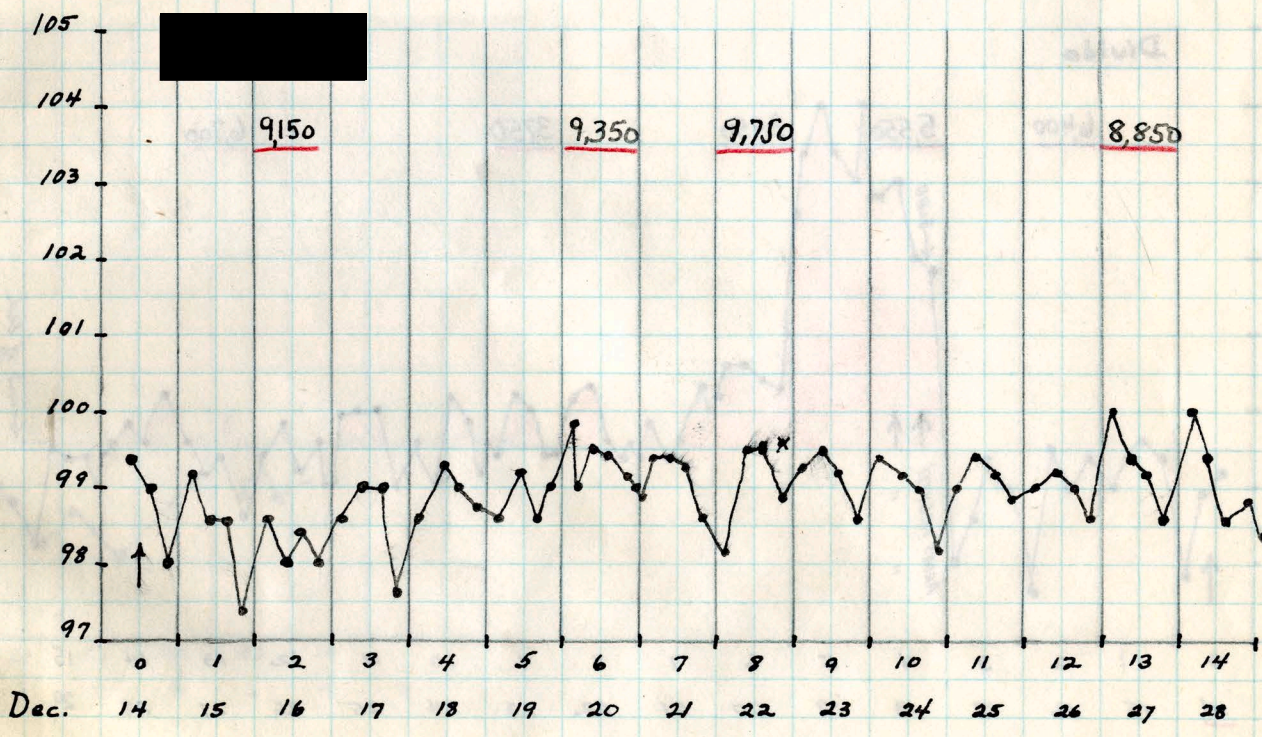
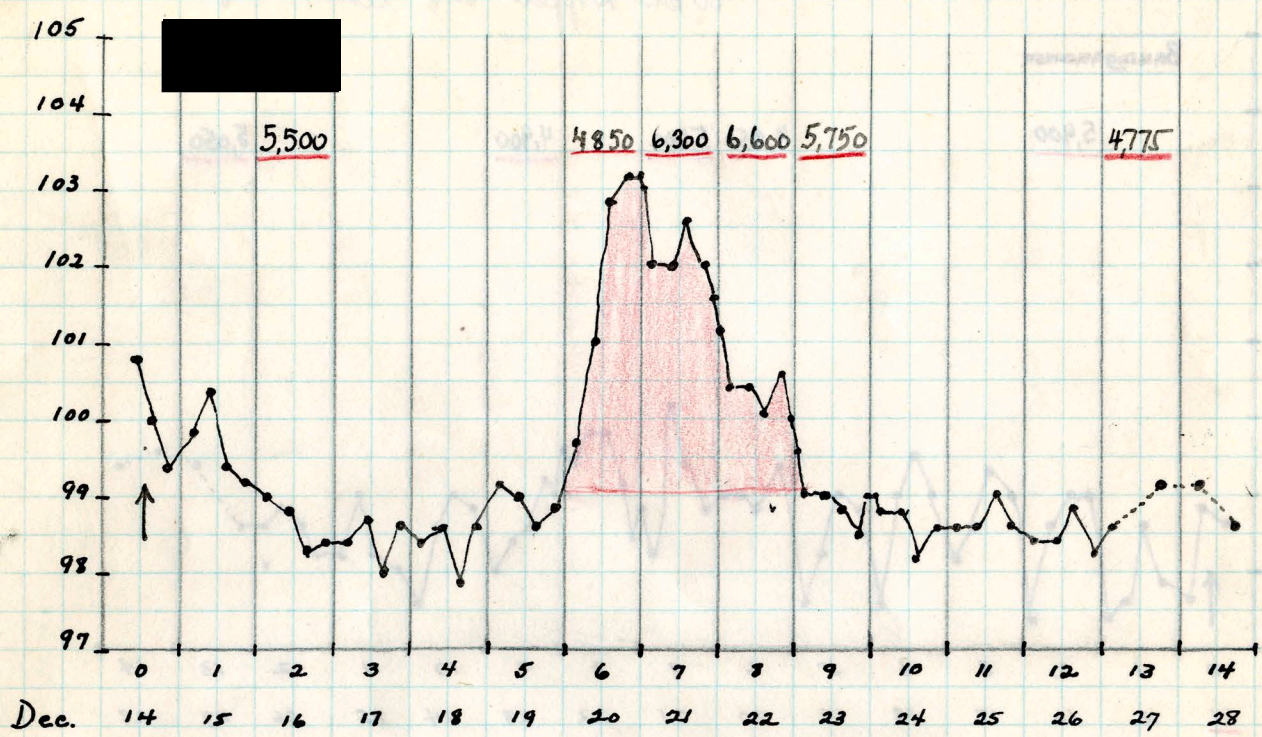
1cc 2ND PASSAGE POOL INTRACUTANEOUSLY

Sicilian Virus - 309 m μ filtrate on Oct. 21



1cc 2ND PASSAGE POL INTRACUTANEOUSLY

SICILIAN VIRUS - 203 ml FILTRATE ON Oct. 21



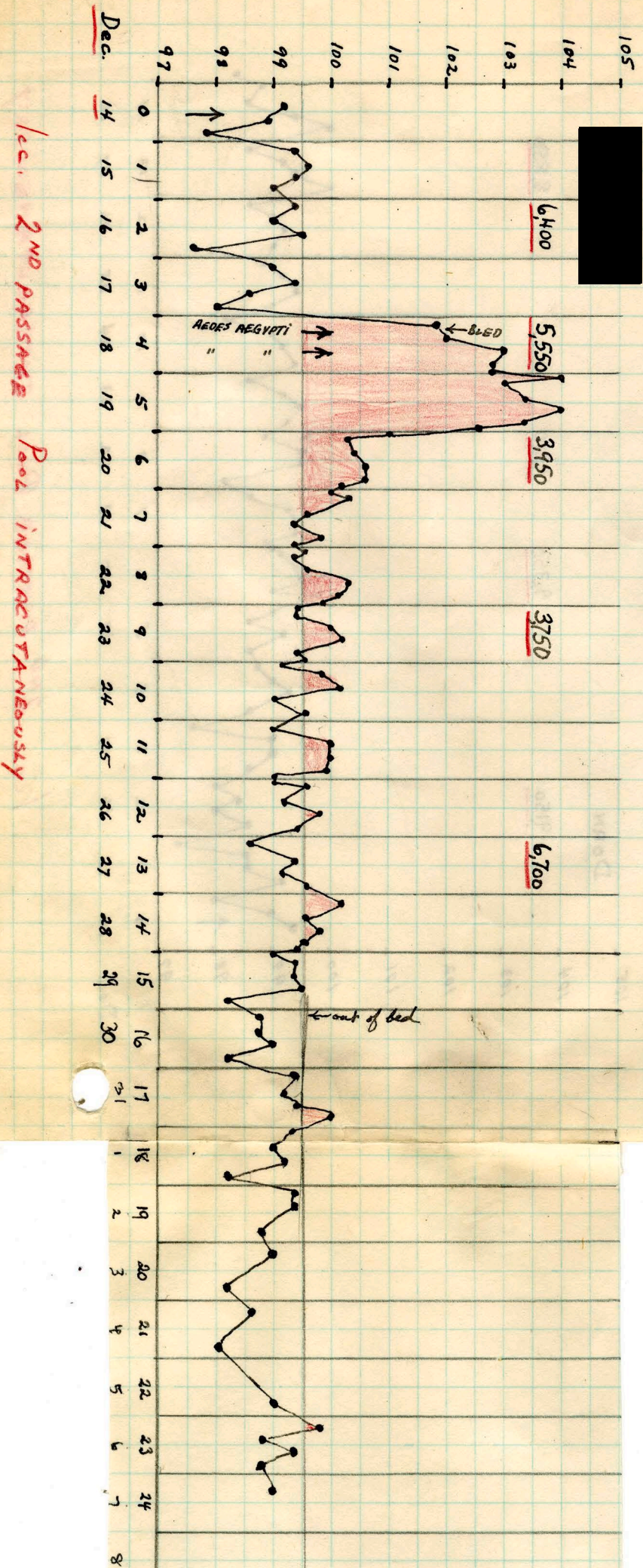
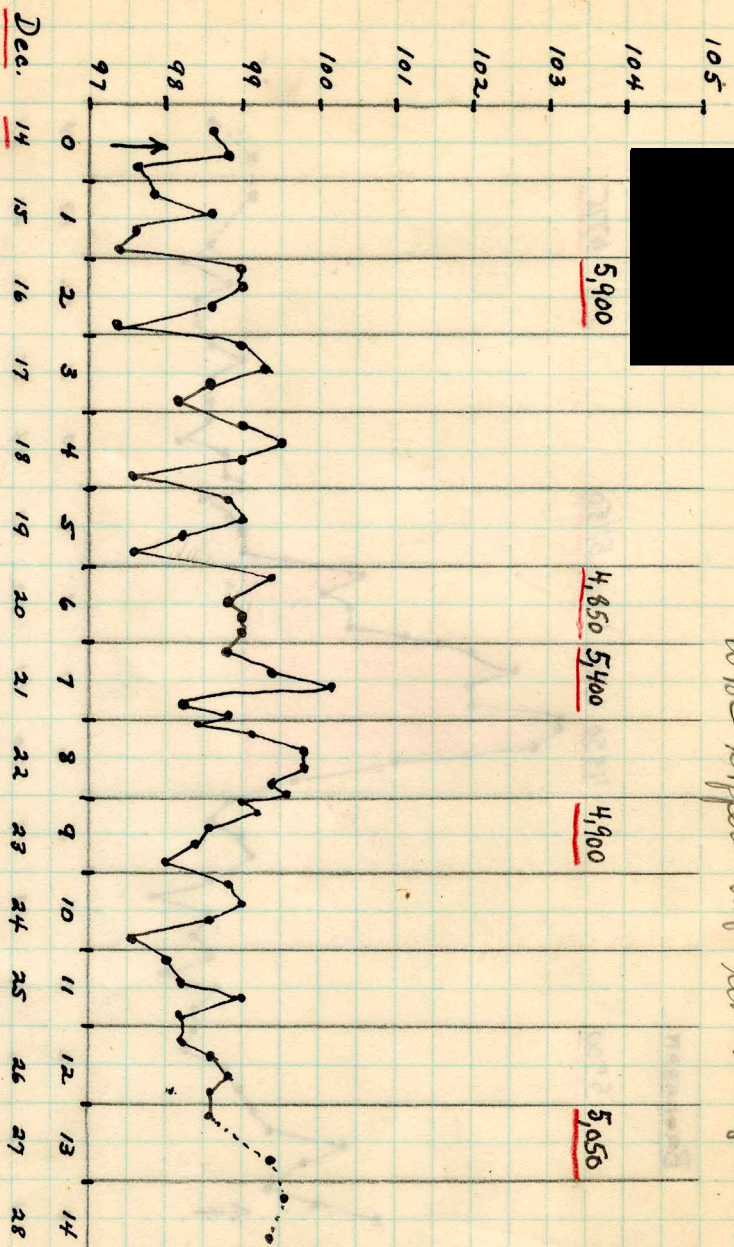
1cc. 2ND PASSAGE POOL INTRACUTANEOUSLY

SICILIAN

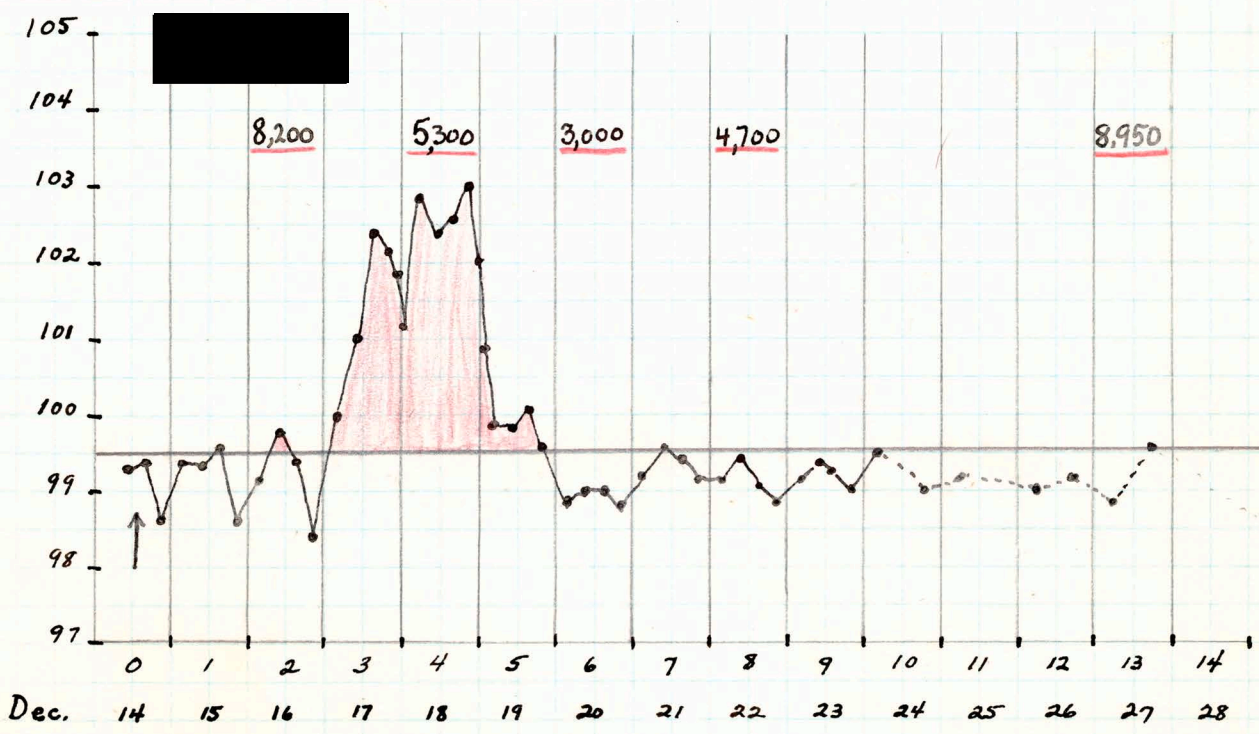
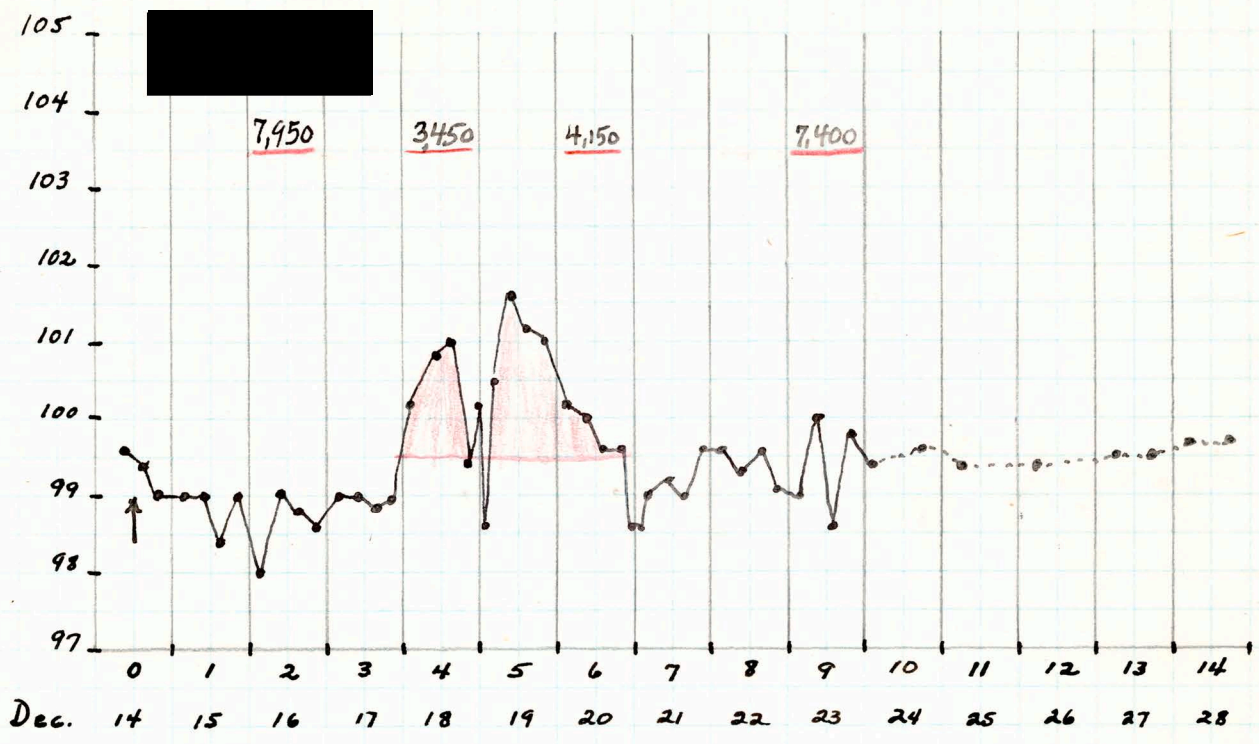
VIRUS

101 m μ FILTRATE ON OCT. 21

wbc Differential unchanged

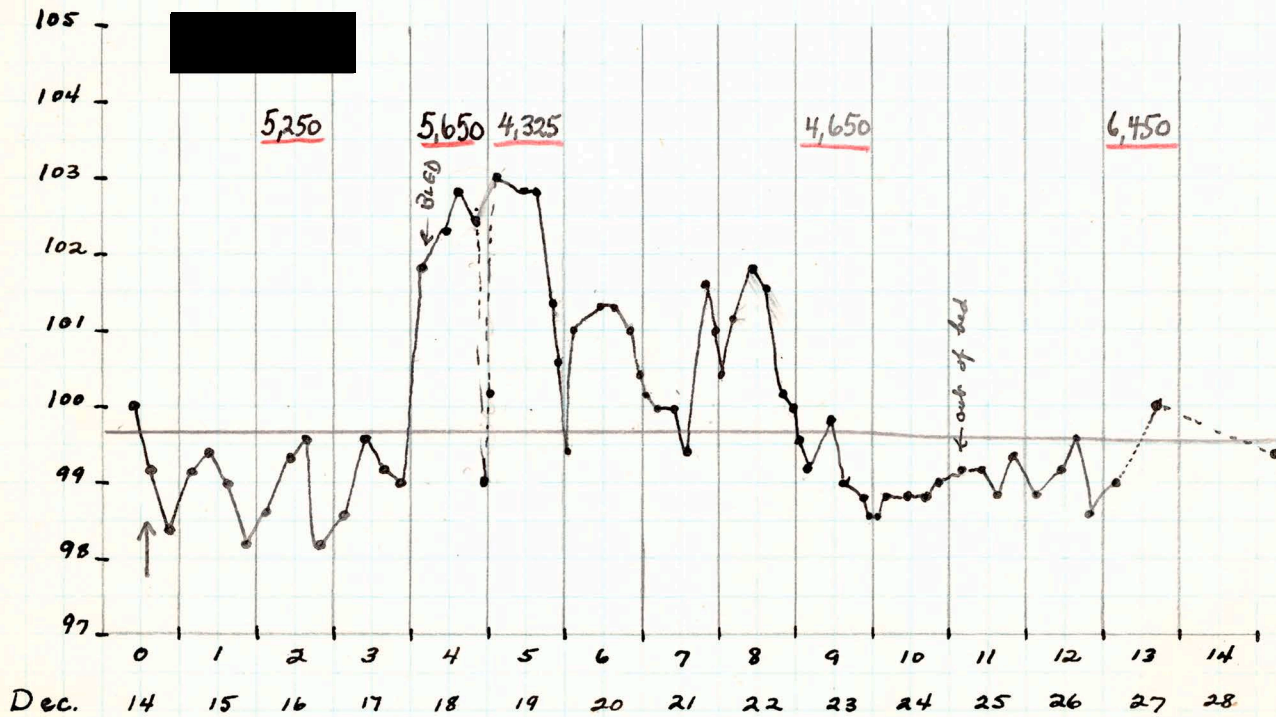
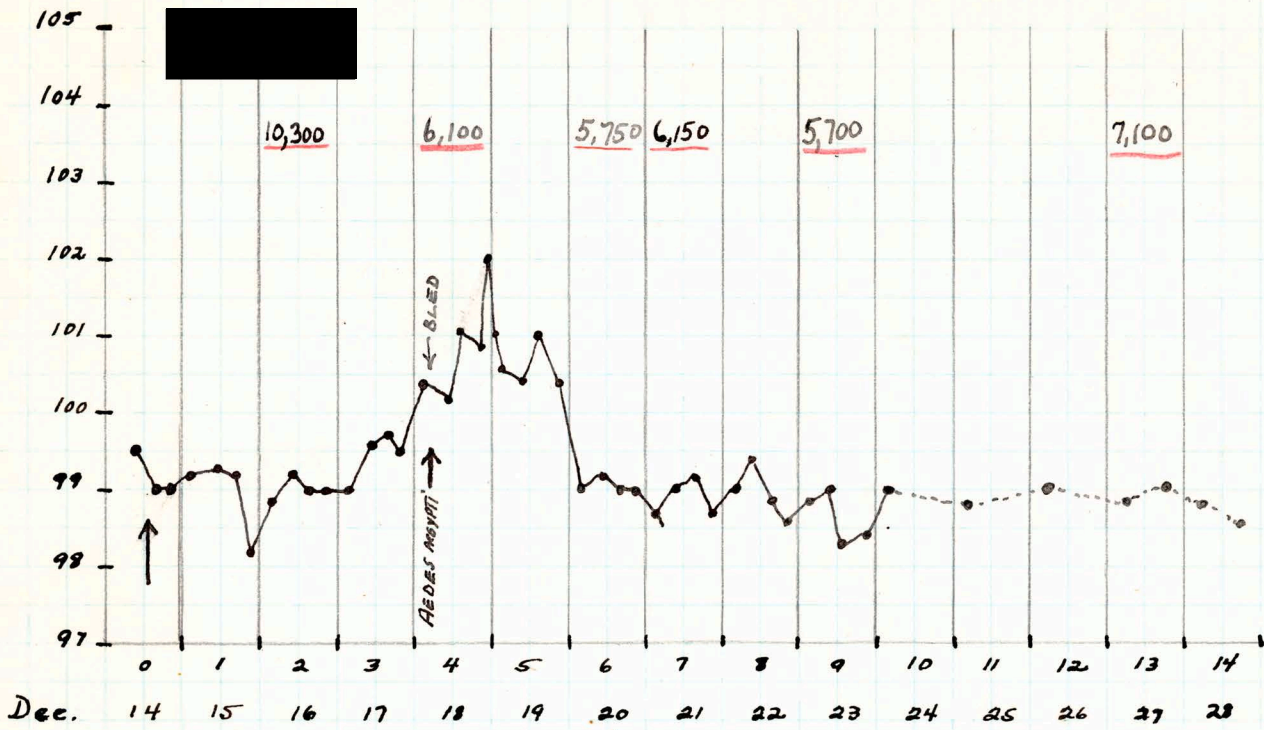


SICILIAN VIRUS - 75 μ m FILTRATE ON OCT. 21



1cc. 2ND PASSAGE POOL INTRACUTANEOUSLY

SICILIAN VIRUS — 50 μ l FILTRATE ON OCT. 21



1cc. OF 2ND PASSAGE POOL INTRACUTANEOUSLY

SICILIAN VIRUS

DEC. 24, 1943

1. Capacity of Aedes aegypti which fed on [REDACTED] (Dec. 18) and [REDACTED] (Dec. 21) to transmit infection

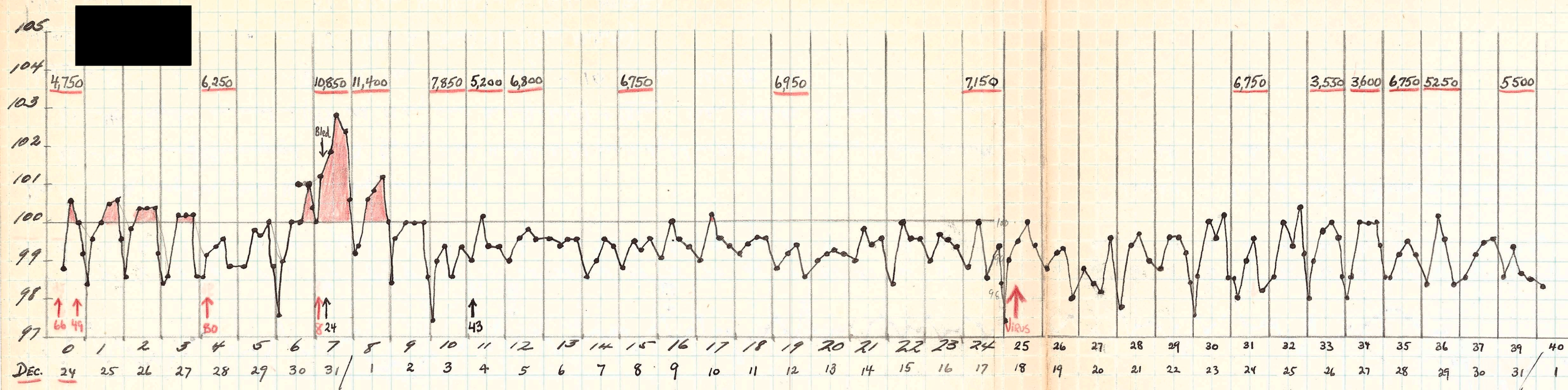
SUBJECTS - [REDACTED] - white male - age 33
- " " - " 39

2. To establish that the blood of the donors [REDACTED] was infective at the time the mosquitoes engorged - and also to provide donors for feeding of additional normal mosquitoes for another test.

1cc of pool of acute [REDACTED] sera injected intracutaneously (11 A.M.) into:

SUBJECTS - [REDACTED] - white female - age 44
- " male - "

SICILIAN VIRUS - ATTEMPTED TRANSMISSION BY Aedes Aegypti



DEC. 24 - 115 mosquitoes engorged [redacted] lot - (6 day incub) - 66 engorged at 11-11⁴⁵A.M.; 49 at 7 P.M.

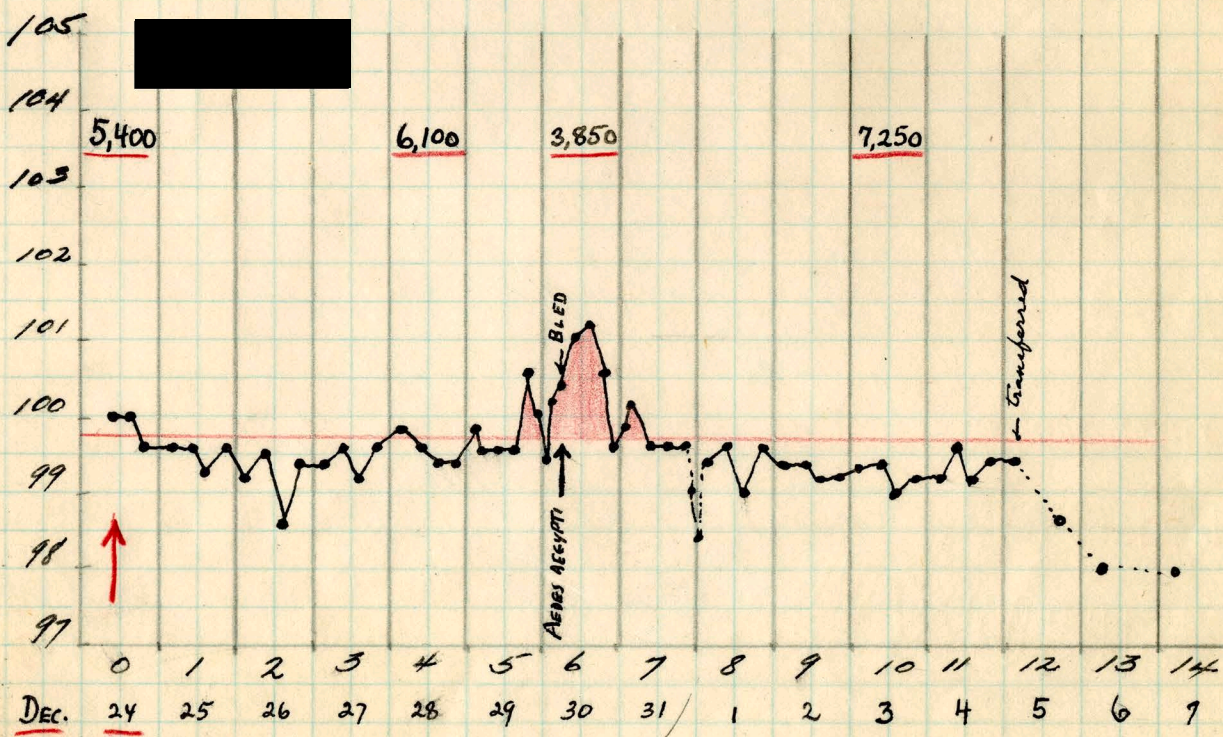
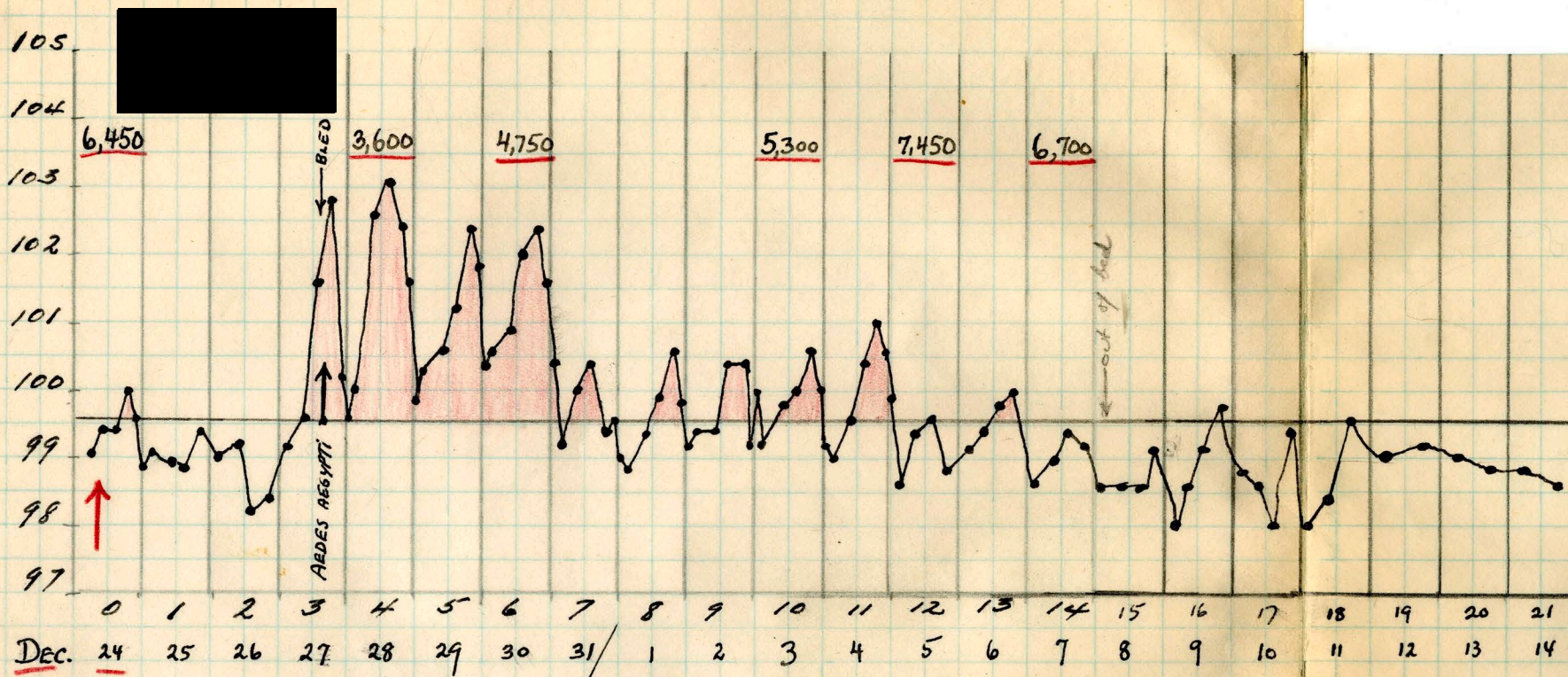
" 28 - 80 new " " (70 ten day incub. [redacted] prev. fed on [redacted] or unfed) + (10 seven day [redacted])

" 31 - 32 mosquitoes engorged (24 & 13 day [redacted] lot) + (8 new - 10 day [redacted] lot prev. fed on [redacted])

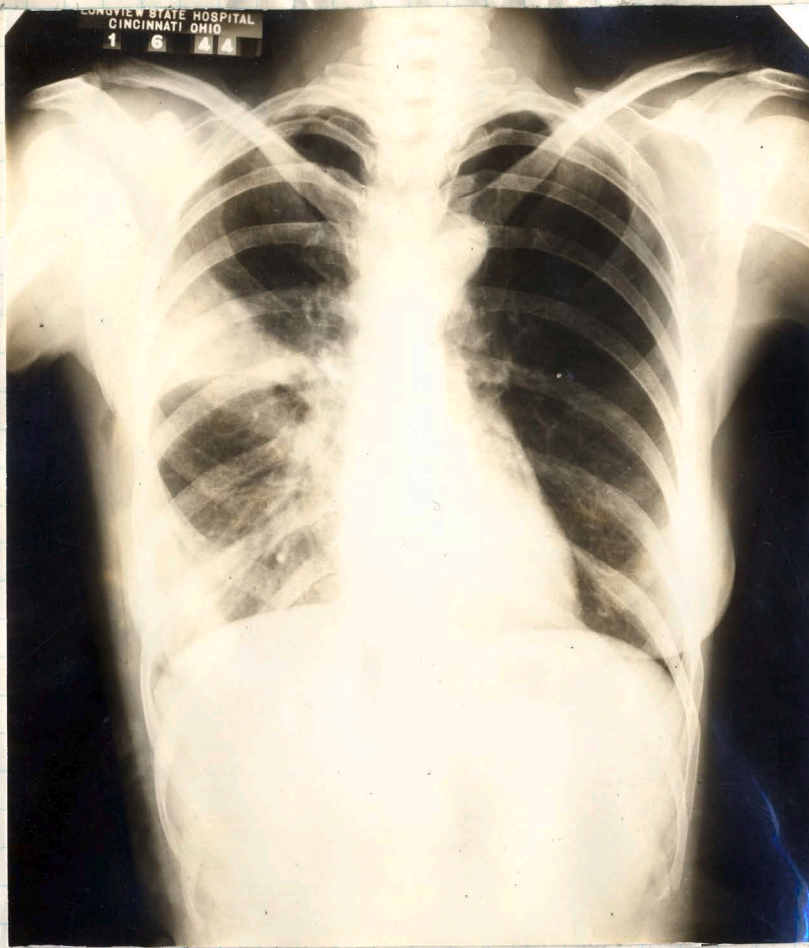
JAN. 4 - 43 " " (40 - 17 day [redacted] lot) + (3 - 14 day [redacted] lot)

JAN. 10 - 1cc of 3rd passage pool Sicilian Virus i.cult. on right forearm

SICILIAN VIRUS - INFECTIVITY OF BLOOD OF [REDACTED]
AT TIME MOSQUITOES FED ON THEM



1cc. of pool of [REDACTED] serum intracutaneously,



SICILIAN VIRUS

JAN. 5, 1944

CAPACITY of Aedes Aegypti mosquitoes which engorged on [REDACTED] (12/27) and [REDACTED] (12/30) to transmit infection.

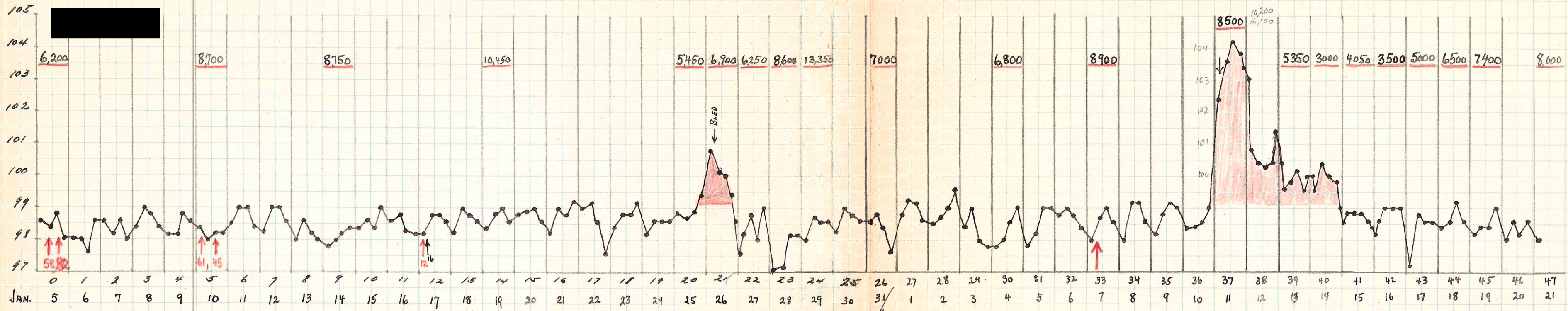
SUBJECTS

[REDACTED]

white male - age 44

" " - " 48

SICILIAN VIRUS - ATTEMPTED TRANSMISSION BY MEDES AEGYPTII

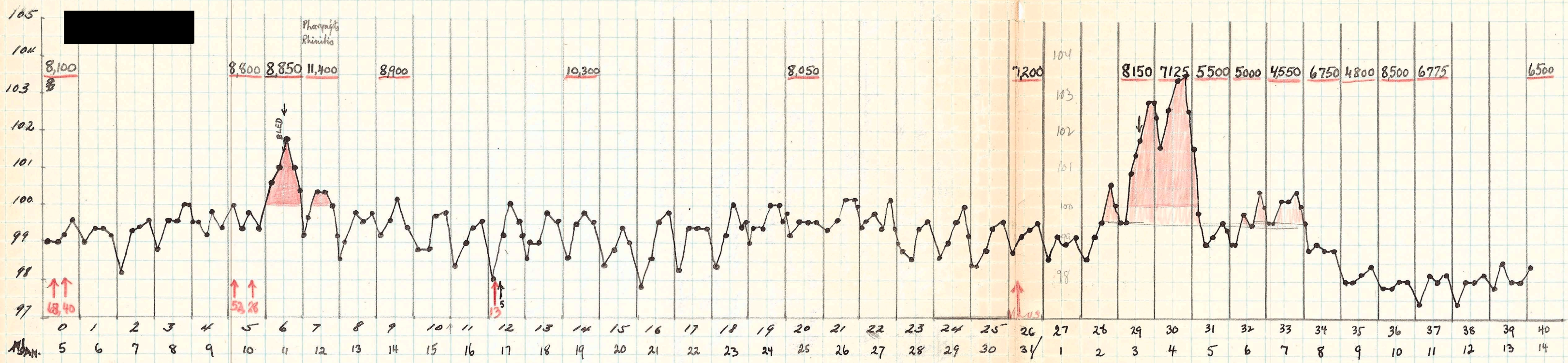


JAN. 5 - 140 mosquitoes engorged; 10 A.M. - 31 [redacted] + 9 day + 27 [redacted] - 6 day; 5:40 P.M. - 53 [redacted] 9 day + 25 [redacted] - 6 day
JAN. 10 - 106 new " " ; 10 A.M. - 36 (" " 14 ") + 25 (" - 11 "); 4:20 P.M. - 23 (" - 14 ") + 22 (" - 11 ")
JAN. 17 - 28 mosquitoes engorged (12 new + 16 old); 13 [redacted] lot 21 day + 15 [redacted] lot - 18 day

JAN. 31 - NOTE - PATIENT was not inoculated with virus at this time because of his bout of fever ^{only} 15 days before. The possibility of nonspecific interference was considered

FEB 7 - TEST FOR SUSCEPTIBILITY OR IMMUNITY - 1cc of [redacted] Pool (used Jan. 31 on Wilhite, [redacted] injected intracut. in 5 sites

SICILIAN VIRUS - ATTEMPTED TRANSMISSION BY TIGRES AEGYPTI



JAN. 5 - 108 mosquitoes engorged; 10 A.M. - 49 (19 day) + 20 (6 day); 5:40 P.M. - 23 (6 day) + 17 (6 day)

JAN. 10 - 80 new " " ; 10 A.M. - 35 (" - 14 ") + 17 (" - 11 "); 4:20 P.M. - 19 (" - 14 ") + 9 (" - 11 ")

JAN. 17 - 18 mosquitoes engorged (13 new + 5 old); 15 (10 day) + 3 (18 day)

JAN. 31 - TEST FOR SUSCEPTIBILITY OR IMMUNITY - 1cc of [redacted] Pool (SERA of JAN. 22, 1944) injected intracutaneously - 0.2cc in 5 places

JAN. 11 - 10 A.M. - patient was up and about and cooperative as usual

1 P.M. - Temp. 101 - patient still maintains there is nothing wrong with him. Put to bed. General physical examination revealed nothing significant.

Skin - hot - presents severe reactions to mosquito bites

Eyes - slight lemon tinge to sclerae - impression was that it was natural. No conjunctival injection, eyes not tender.

Mouth - slight injection of fauces and post. pharyngeal wall.

Chest and abdomen - negative; liver + spleen not palpable; no tenderness

Neck - not stiff or painful

Lymph nodes - not enlarged or painful


WBC + blood drawn at 1:30 P.M.

JAN. 12 - DEFINITE PHARYNGITIS AND RHINITIS PRESENT

INFECTIVITY of blood of rhesus monkeys 3 and 4
days after inoculation with Sicilian virus by various
routes.

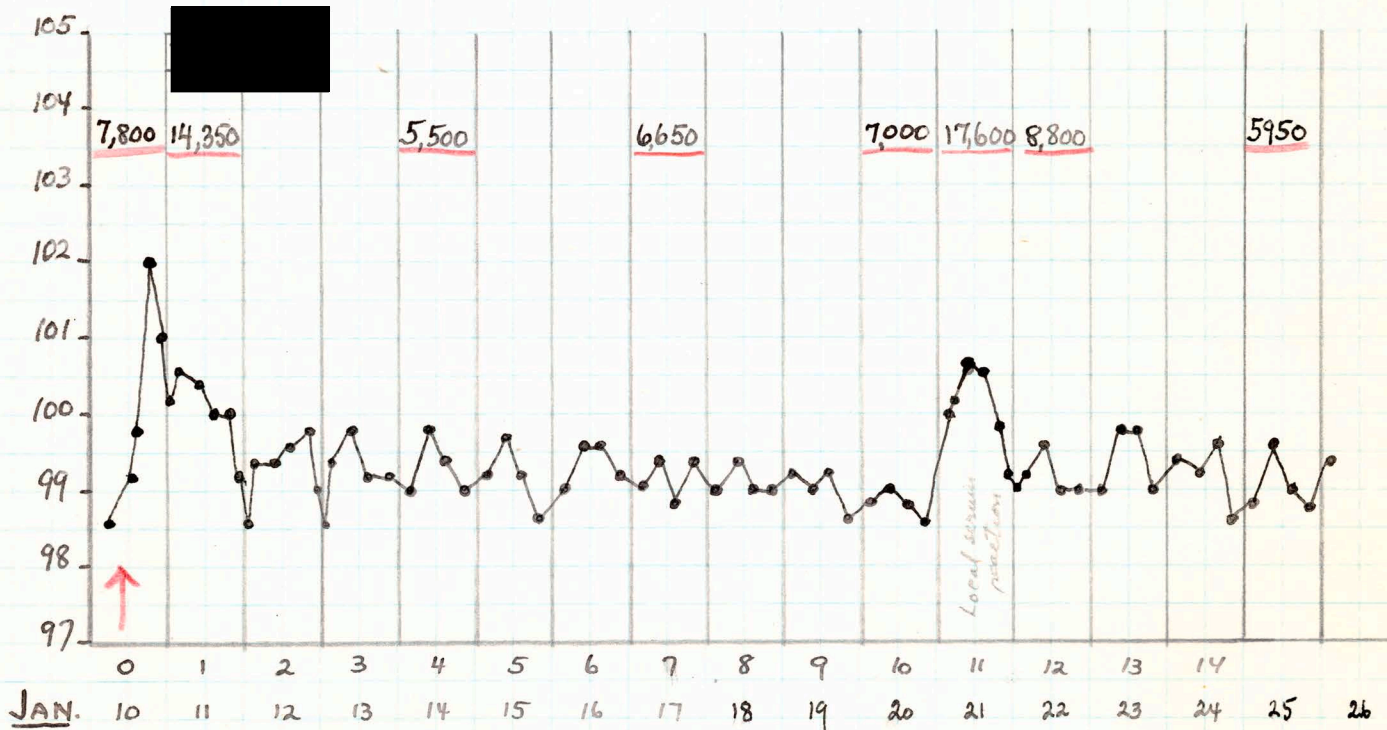
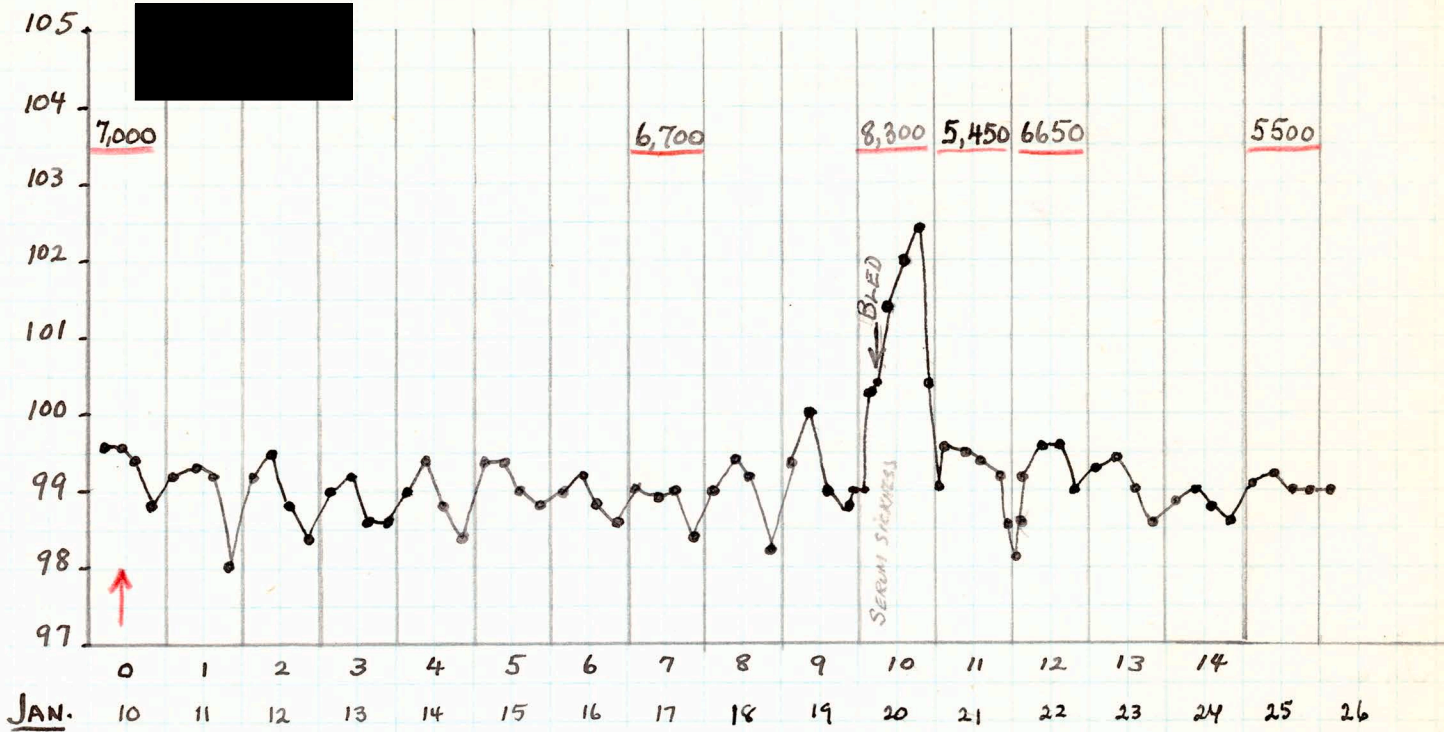
The serum of Rh. monkeys 1, 2, and 3 which
were bled on Oct. 24 and Oct. 25 and which had
been frozen since then, was thawed and pooled.

2cc was injected intracutaneously (0.25cc
in 8 sites) into each of the following 2 subjects:

-  - white male - age 46
- " " - " 51

Inoculation - 11 A.M.

JAN. 11 - No reaction at sites of i.cut. inoculations



NOTE: [redacted] had an acute pharyngitis associated with his fever which developed on the day he was admitted - JAN. 10.

2 cc. Rhesus serum intracutaneously

JAN. 20 - ^{10:30 AM} [REDACTED] - flexor surface of left forearm - site of inoculation of virus serum, presents an inflamed area about 10_{cm} x 8 cm. suggestive of a receding urticarial reaction. Patient states that earlier this morning this area itched but not now. No other complaints. Physical examination negative, except for some injection of the throat which almost everyone seems to present at this time.

JAN. 21 - [REDACTED] - a generalized urticarial reaction has appeared at the entire flexor surface of the left forearm presents a confluent inflammatory reaction with local raised areas suggestive of receding wheals. Itching present. No other complaints.

[REDACTED] whose inoculated arm was negative yesterday presents a diffuse inflammatory reaction similar to, but less marked than that of Wallace. Itching present - no urticaria anywhere else.

JAN. 22 - [REDACTED] - local reaction still present. Still no generalized eruption or other signs - nothing beyond local reaction which is mild and shows signs of scratching.

JAN. 23 - [REDACTED] - only large area of erythema left - multiple wheals over entire area more marked than before

JAN. 24 - All clear in both Wallace and Bockhurst

REPETITION OF WORK WITH ULTRAVIOLET LIGHT ~~IRRADIATED~~

IRRADIATED VIRUS

In view of the fact that 5 subjects who had been inoculated with virus (MIDDLE EAST) inactivated by ultraviolet light proved to be immune to reinoculation with active virus, it was desirable to extend the data

VIRUS USED - A new pool of virus consisting of 3rd passage sera obtained on Dec. 18, 1943 - all Kahn negative - was prepared.

3 RD PASSAGE POOL	{	[REDACTED]	- 23.5cc
		[REDACTED]	- 20.0 "
		[REDACTED]	- 15.0 "
		[REDACTED]	- 4.0 "
			<u>62.5cc</u> - slightly opalescent - lipids

19cc used for irradiation

3.5cc " untreated for inoculating controls

Remainder put in vials in 5cc amounts and refrozen

IRRADIATION WITH ULTRAVIOLET LIGHT - 15 MINUTES

SAME PROCEDURE AS USED ON Nov. 24

7cc removed for inoculation; remainder frozen

1cc intracutaneously
(left forearm)

1cc subcutaneously
(above left olecranon)

- white male - age 31
- " " - " 42
- colored female - " 29
- white male - " 33
- " female - " 47
- colored " - " 37

All inoculations 10-11 AM

CONTROL - UNTREATED VIRUS

1cc intracutaneously

- white male - age 51
- " female - " 49
- " male - " 39

(over)

JAN. 19 - No reaction at any of the inoculated sites in any of the subjects

JAN. 31

REINOCULATION OF [REDACTED]

to determine a) whether failure to develop disease was due to inoculation of subinfective amounts of virus the first time in which case they should be susceptible to reinoculation

b) ^{or} to completely inapparent infection or resistance the first time in which case they should be immune or again resistant to inoculation.

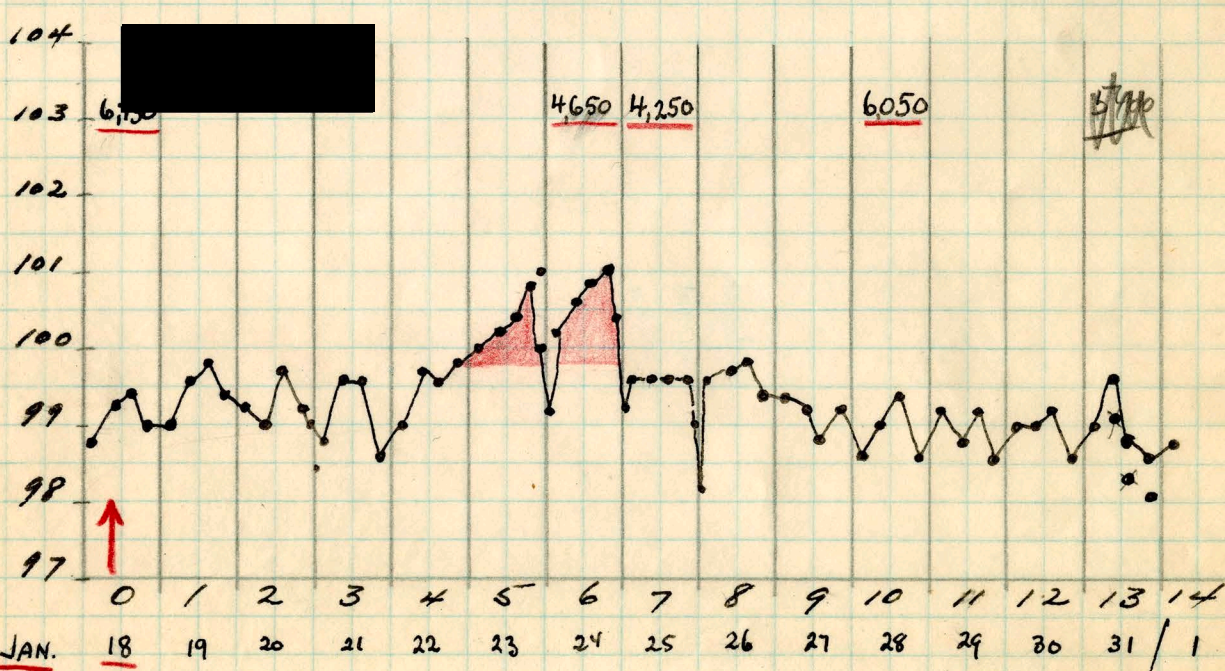
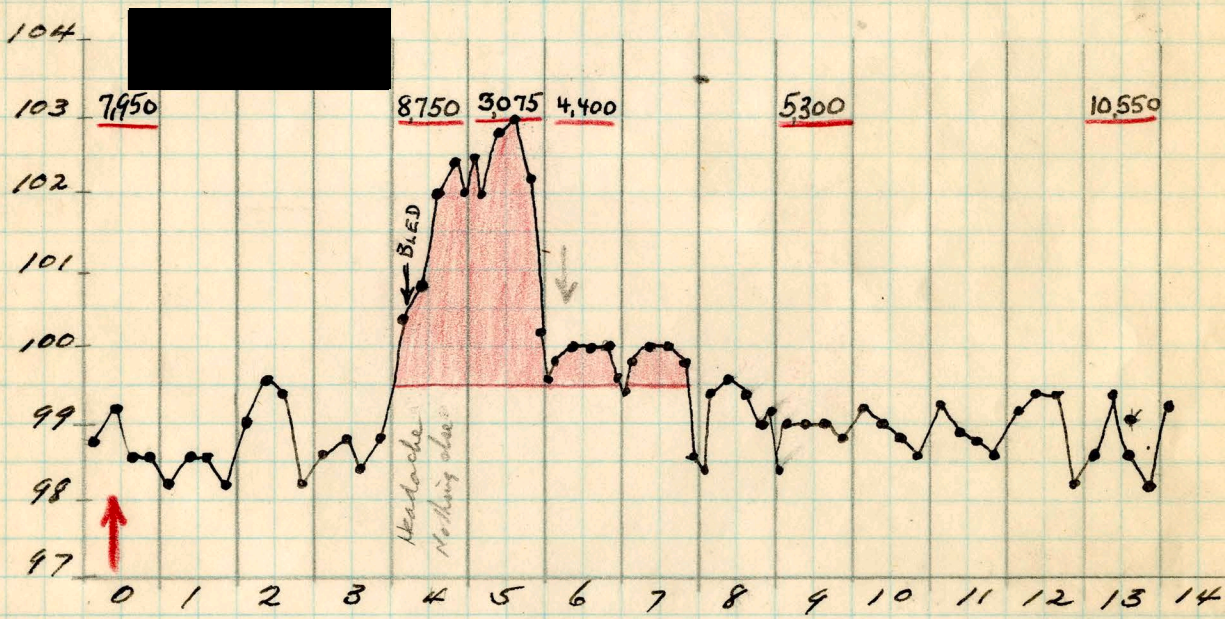
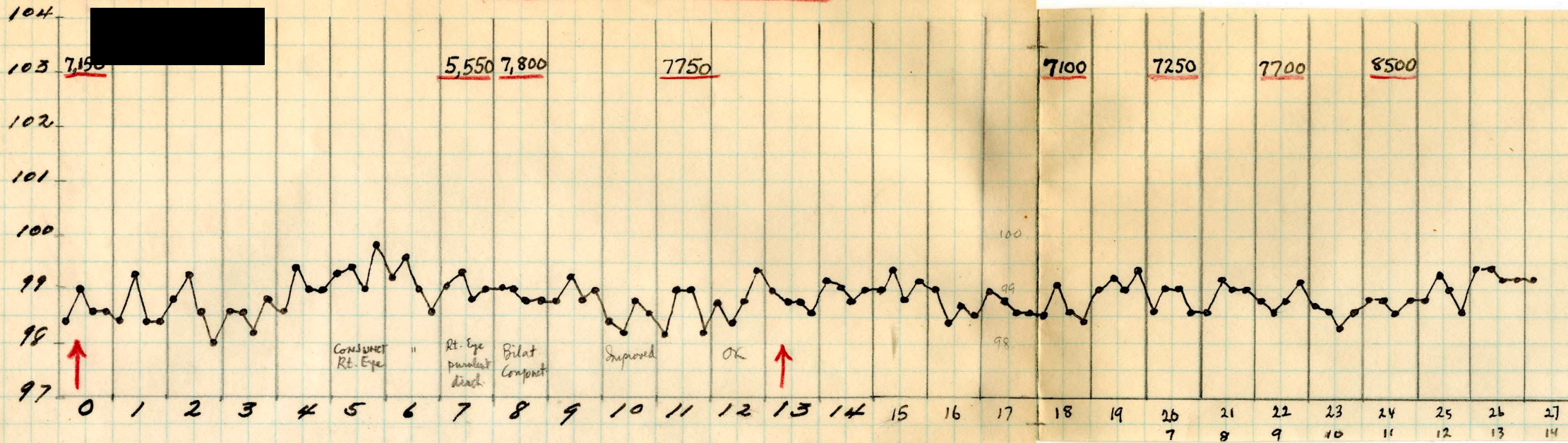
VIRUS USED - Pool of frozen serum of Lakeman and [REDACTED] - obtained Jan. 22, 1944

1cc intracutaneously - 0.2cc in ~~two~~ five sites

INOCULATION TIME - 11 A.M.

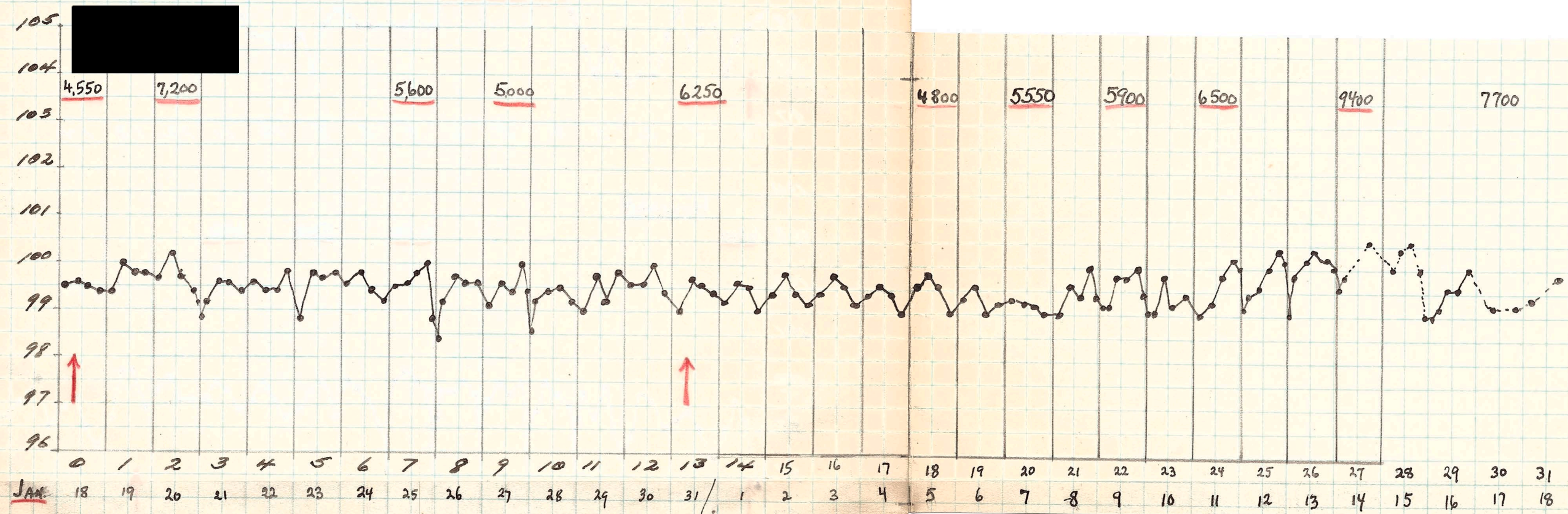
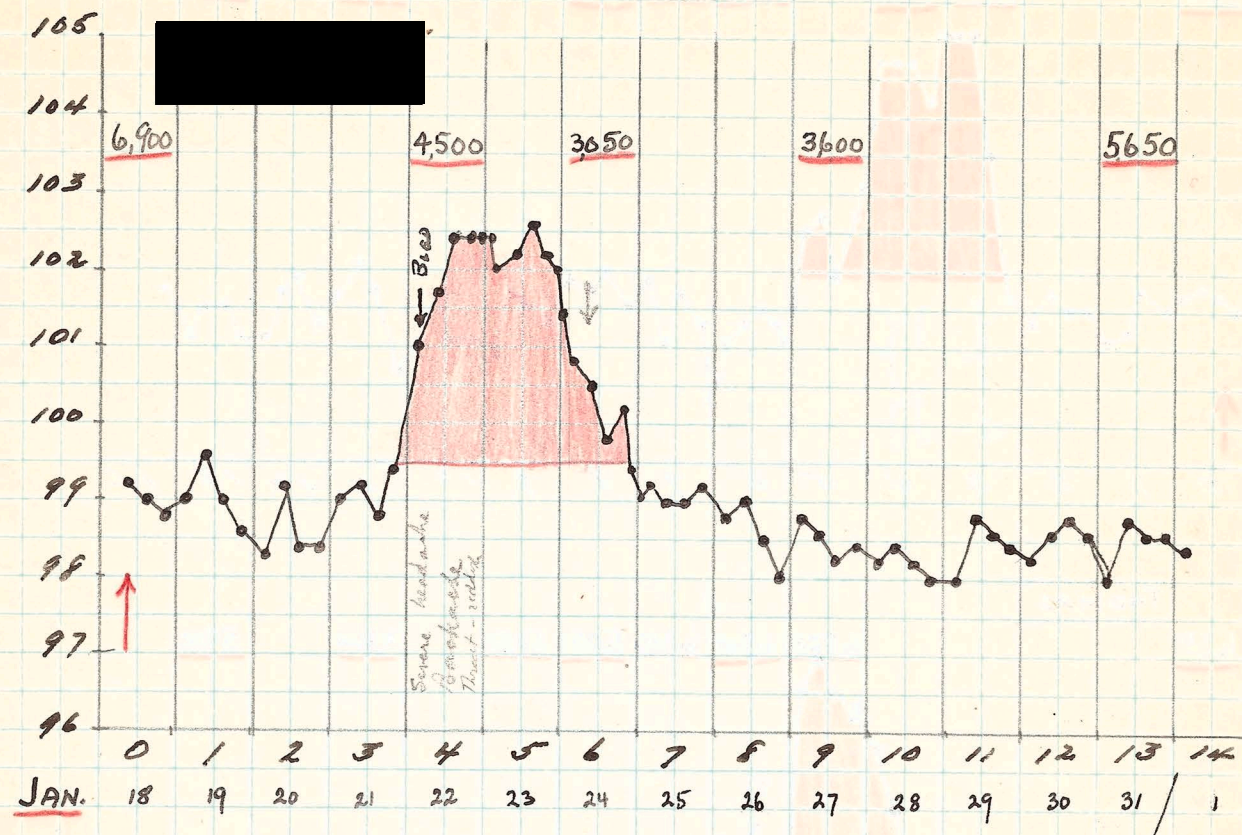
OTHERS INOCULATED SIMULTANEOUSLY - WITH SAME DOSE
[REDACTED]

SICILIAN VIRUS - 3RD PASSAGE POOL - ULTRAVIOLET 15 MINUTES



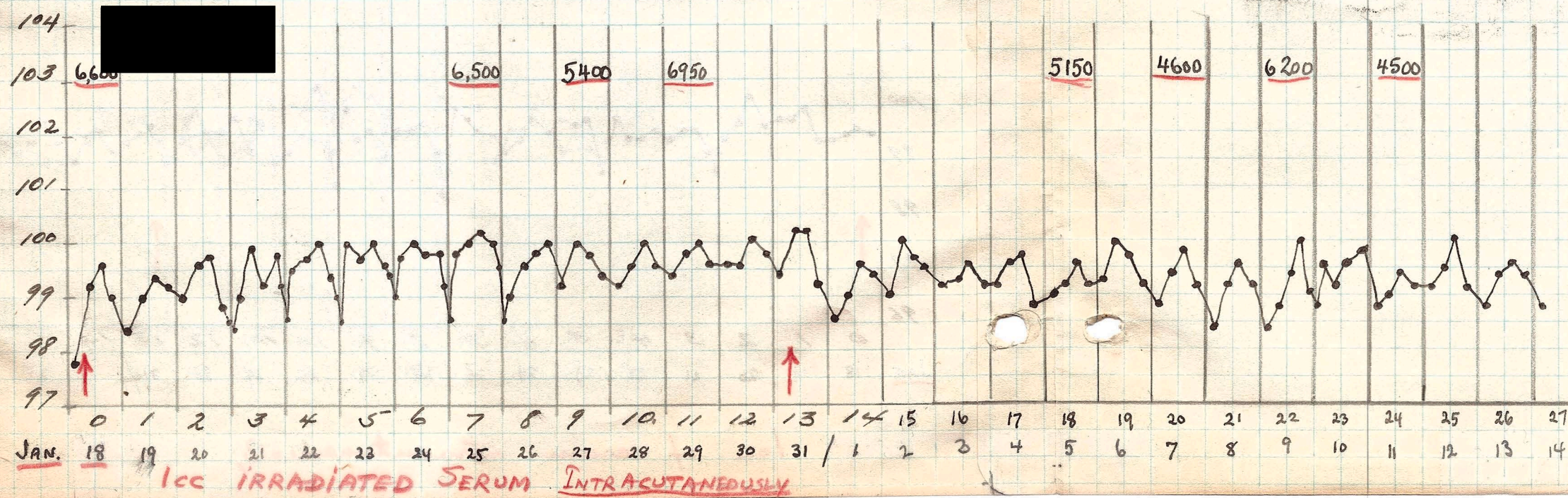
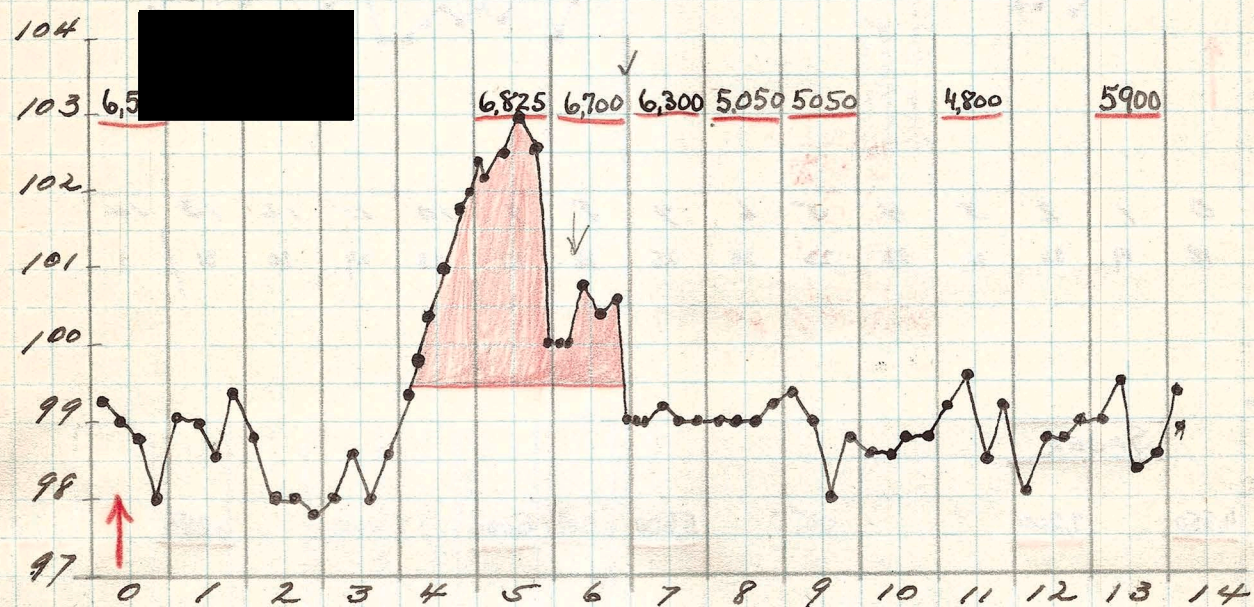
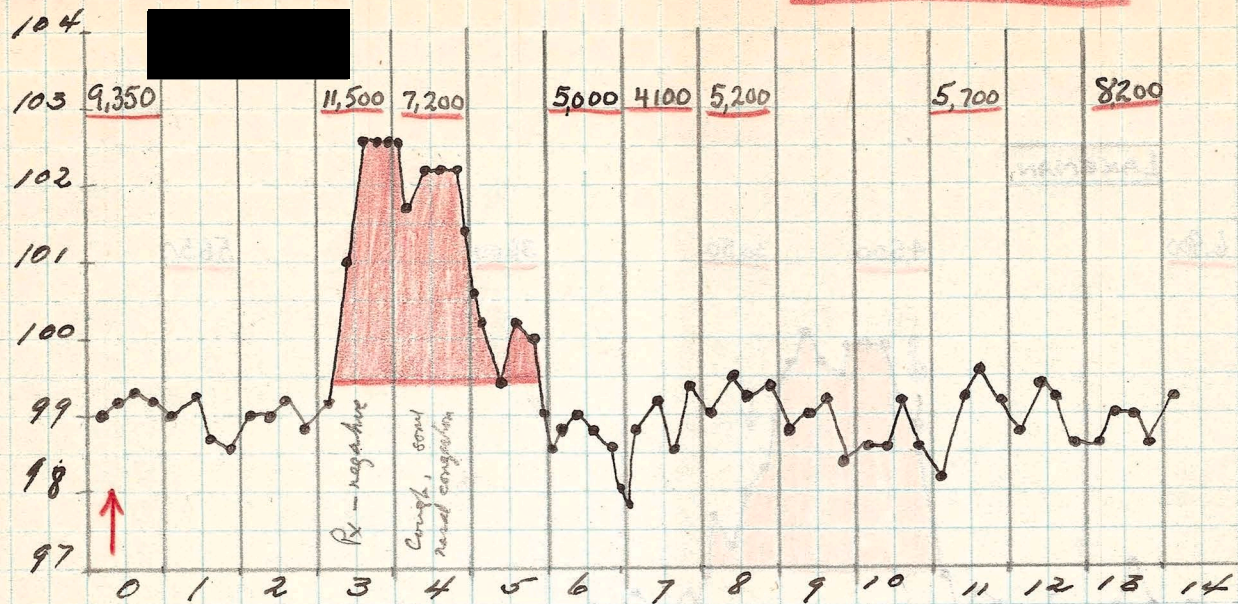
1cc. IRRADIATED SERUM SUBCUTANEOUSLY

SICILIAN VIRUS - 3RD PASSAGE POOL - CONTROLS FOR ULTRAVIOLET IRRADIATED MATERIAL



1cc of serum intracutaneously

SICILIAN VIRUS - 3RD PASSAGE POOL - IRRADIATED WITH ULTRAVIOLET LIGHT FOR 15 MIN.



ICC IRRADIATED SERUM INTRACUTANEOUSLY

REPETITION OF ULTRAVIOLET IRRADIATION

Since it became evident that the virus irradiated for 15 minutes on JAN. 18 was not inactivated, it was ~~to~~ deemed desirable further to irradiate the same preparation (which was frozen on JAN. 18)

11cc of serum which had already been irradiated for 15 minutes was exposed for another 20 minutes.

3.5cc was removed for inoculation and the rest was frozen

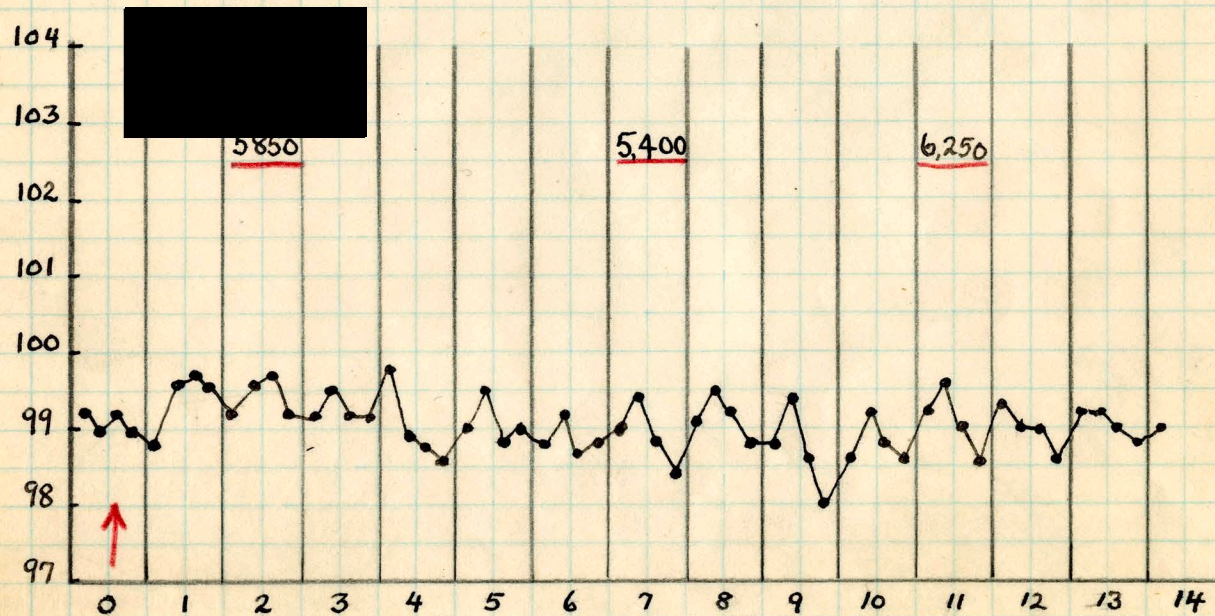
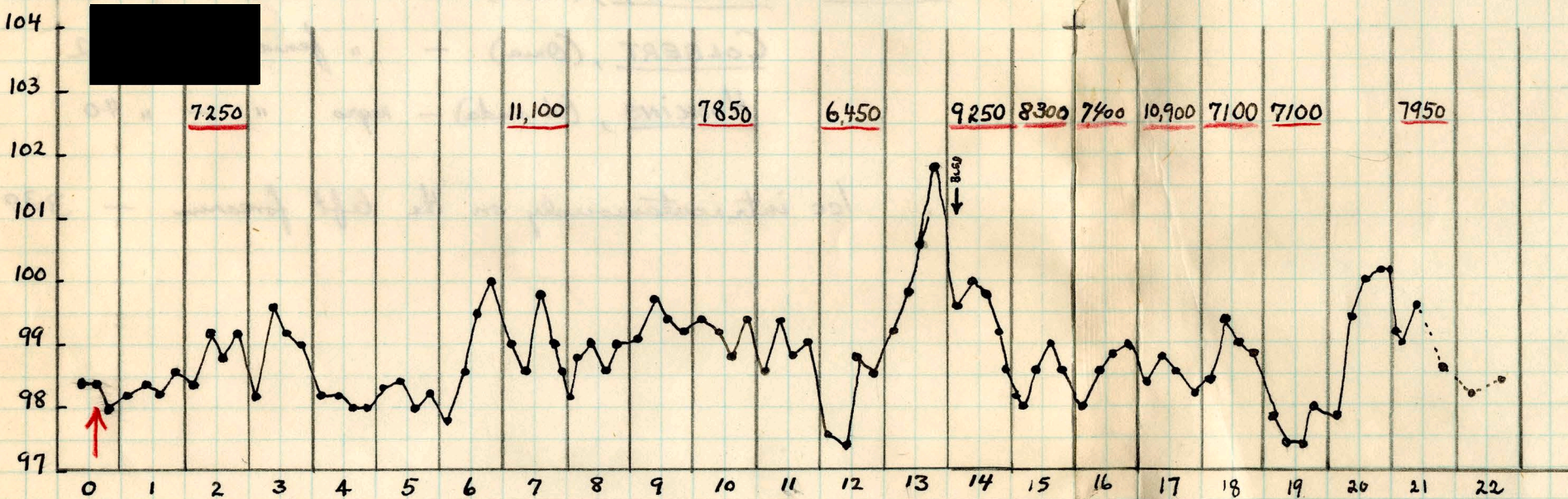
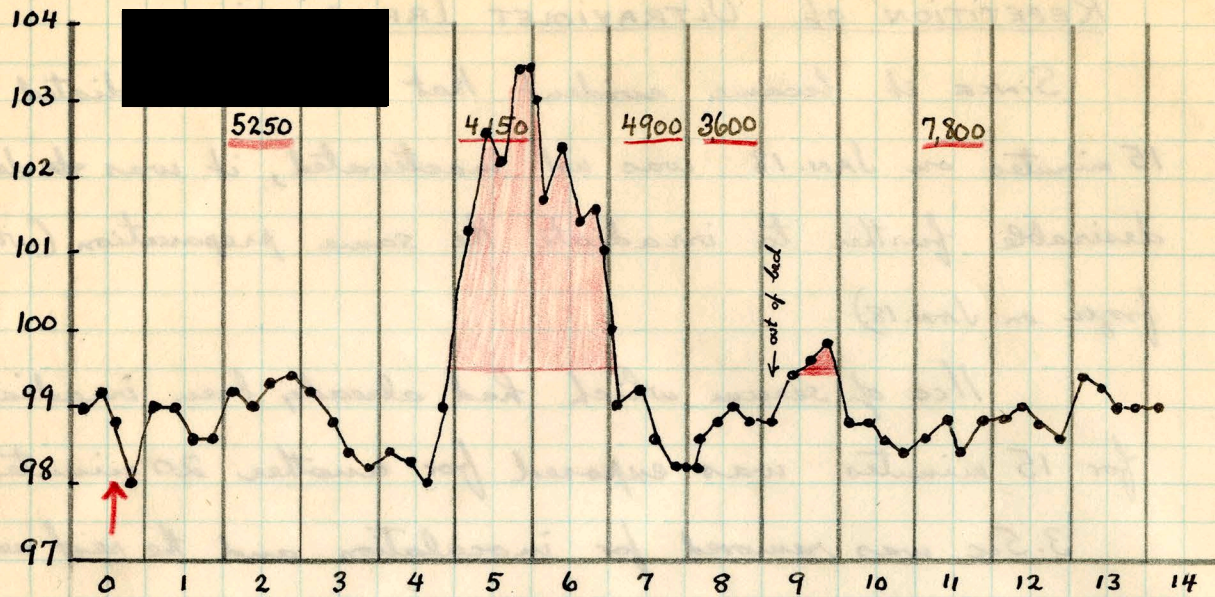
SUBJECTS:

[REDACTED] white male - age 29

" female - " 42

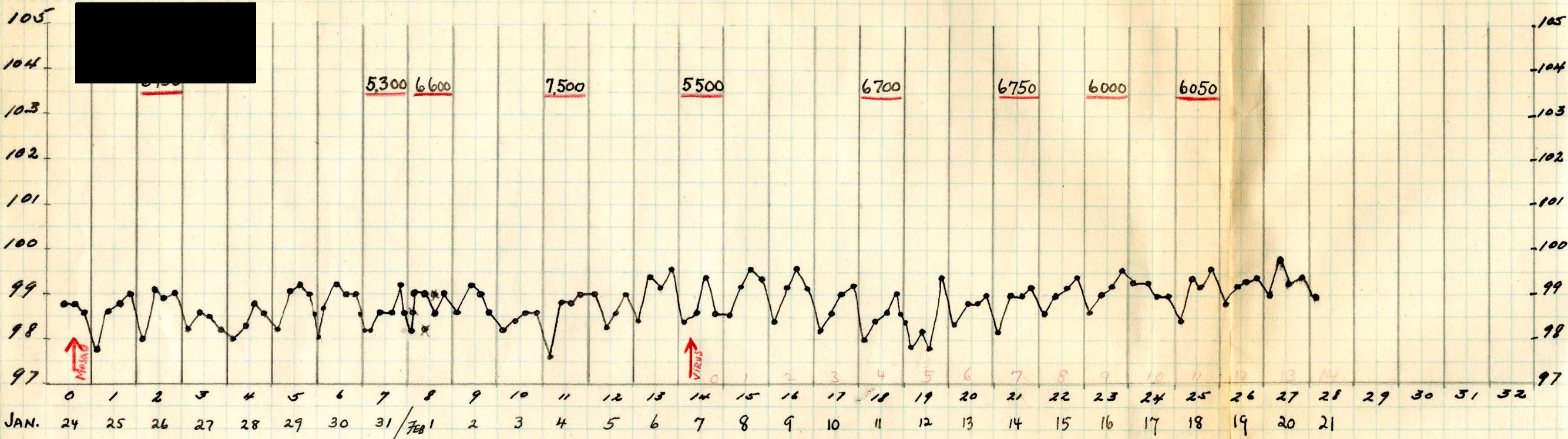
negro " - " 40

1cc intracutaneously on the left forearm - 3³⁰ P.M.



JAN. 24 25 26 27 28 29 30 31/ 1 2 3 4 5 6 7

TEST FOR VIRUS IN FILTERED SUSPENSION OF MOSQUITOES USED IN TRANSMISSION TESTS



JAN. 24 - 2cc of filtrate injected intracutaneously (0.25cc in 8 sites on left forearm) at 3:30 P.M.

" 25 - area somewhat "dusky" and warm - patient complains of local itching

" 26 - negative

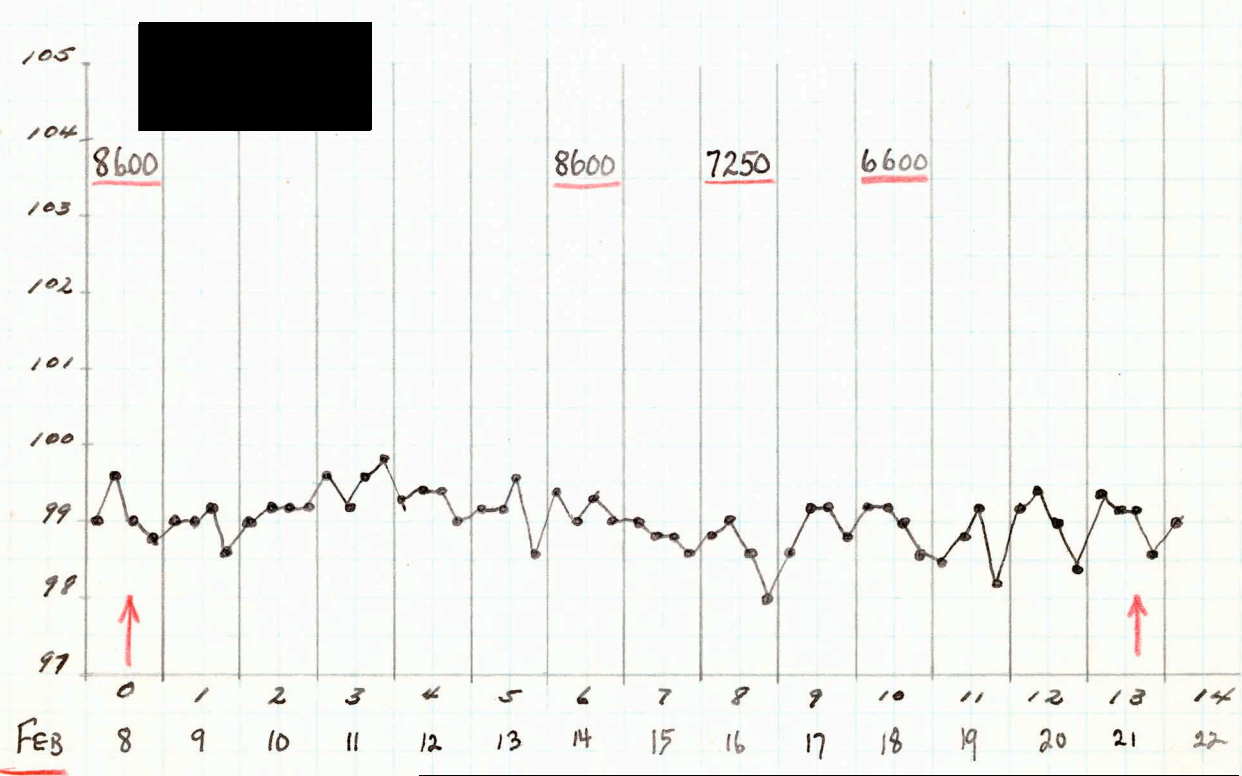
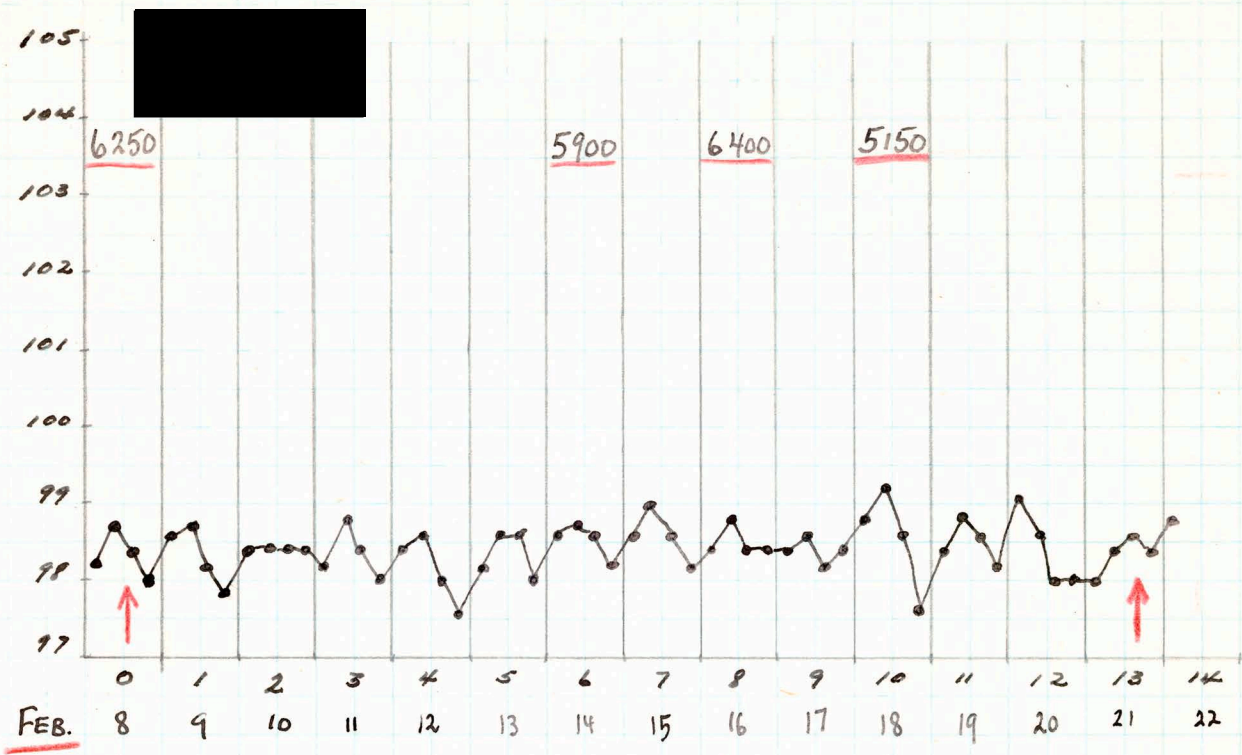
PREPARATION OF MOSQUITO FILTRATE: JAN. 22 - 80 mosquitoes of [redacted] lots frozen on Jan. 14

24-27 days after engorging on S.F. patients were mixed with 69 mosquitoes of the Barbours-Gonia lots 23 to 26 days after original feeding. Total weight - 0.46 gm. Ground with sand in 6cc of a 10% (human serum (carral) - saline diluent, centrifuged at @ 2000 rpm. and sn. liquid put over 770 mμ membrane (saturated with broth). After first 1cc - no more would pass. The fluid in the filter was then diluted further - 4cc + 8cc saline - centrifuged for 40 min. and put over a new 770 mμ membrane - Again filtration was very slow despite 15 lbs pressure (nitrogen). Total amount filtered - 4cc. 0.1cc amounts cultured immediately - remainder frozen

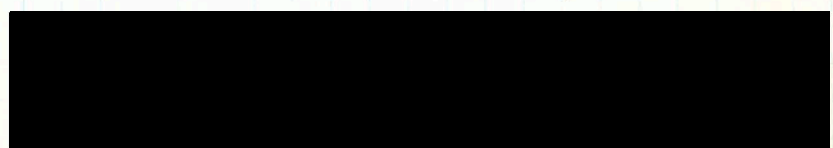
- CULTURES:
- BAP I - 0000
 - " II - 0000
 - Broth I - 0000
 - " II - 0000
 - Anaerobic I - 0000
 - medium II - 0000

SUBJECT - CLARENCE CARTER - NEGRO MALE - AGE 51

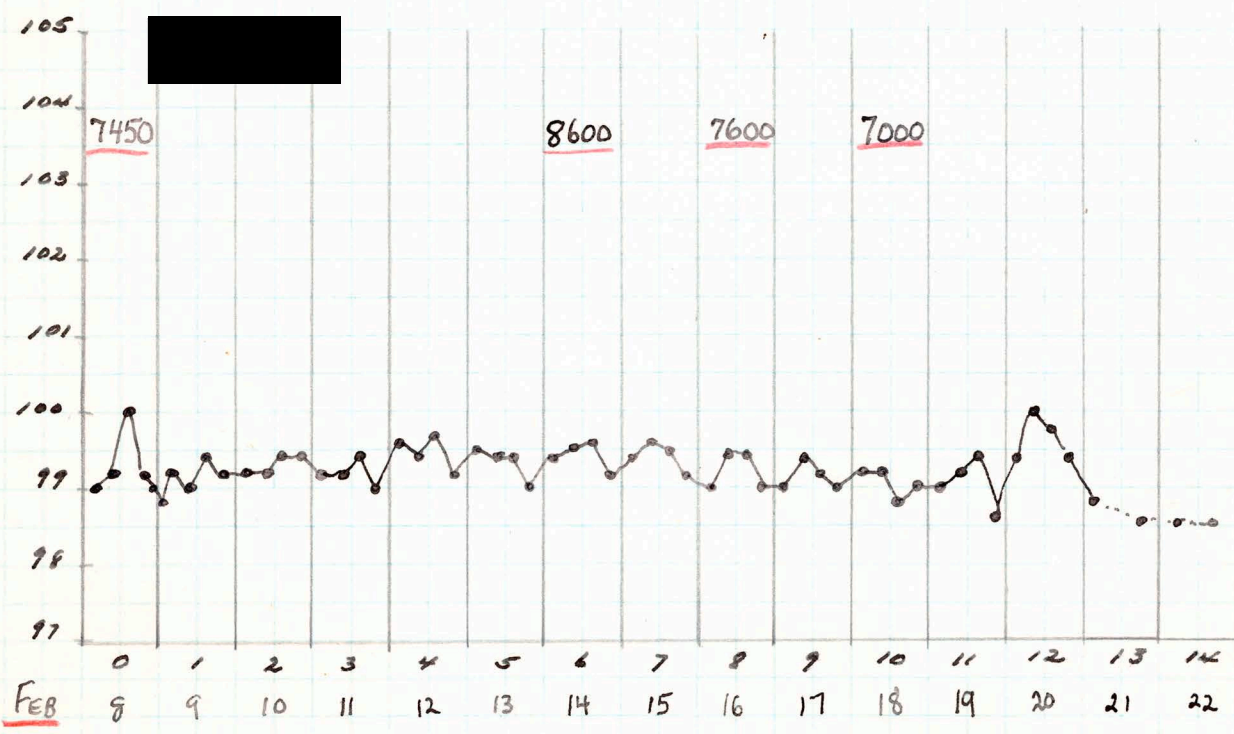
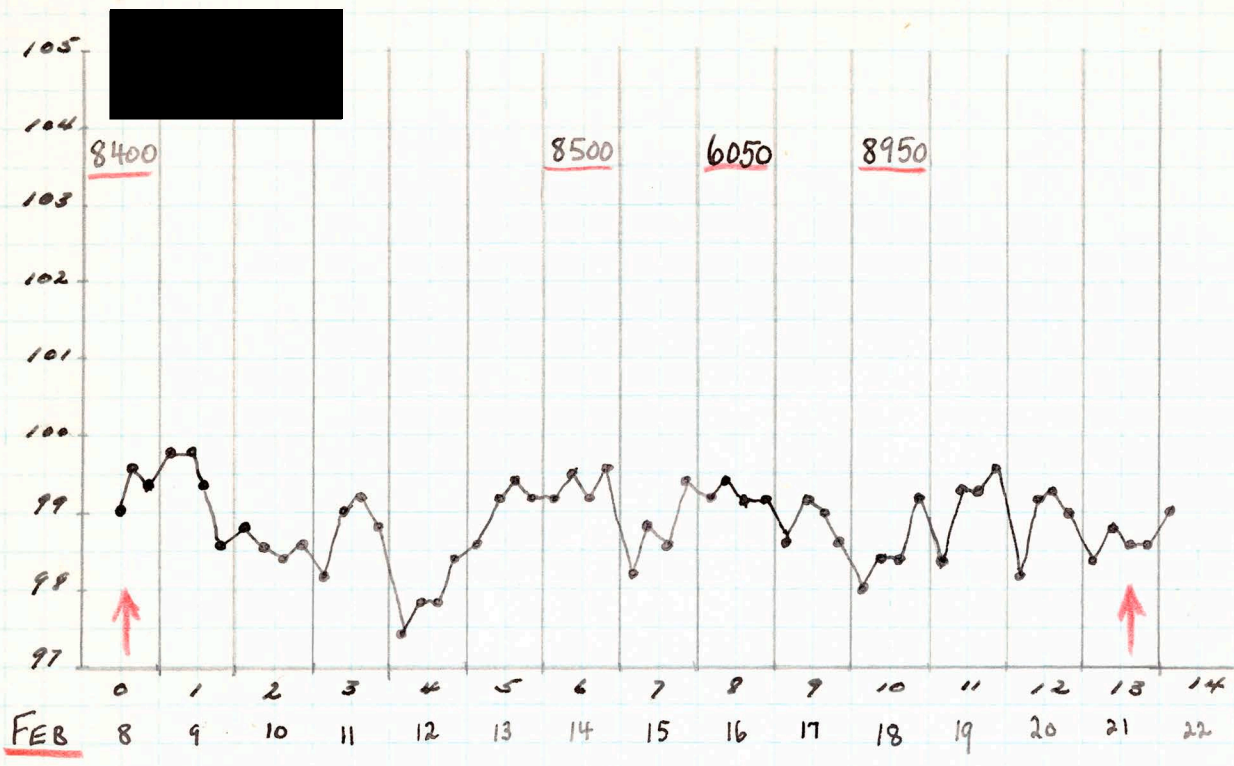
SICILIAN VIRUS - TEST FOR PRESENCE OF VIRUS 48-54 HOURS
 AFTER ONSET OF FEVER



1.5cc OF



TEST FOR PRESENCE OF VIRUS IN SERUM OF PATIENTS
 WITH "INTERCURRENT FEVERS" DURING COURSE
 OF Aedes Aegypti TESTS



1.5 cc of

POOL

Inocul

Inoculations FEB. 15, 1944

1045 - 11¹⁵ A.M.

[REDACTED] SERUM
UNTREATED
1cc INTRACUT.

[REDACTED]

Pass IV

[REDACTED] SERUM
IRRADIATED WITH
ULTRAVIOLET LIGHT
FOR ONE HOUR
1cc. INTRACUT.

[REDACTED]

- C.M. - 3 AGE 31
- W.M. - " 44
- C.F. - " 39
- W.F. - " 47

[REDACTED] SERUM
24 HOURS BEFORE ONSET
OF FEVER
1cc INTRACUT.

[REDACTED]

W.F. - AGE 50

[REDACTED] SERUM
1.0.0. 3 days
after irrad. virus
1cc INTRACUT

[REDACTED]

C.M. - AGE 50

3RD PASSAGE CHICK EMBRYO
20% SUSPENSION
2cc. INTRACUT.

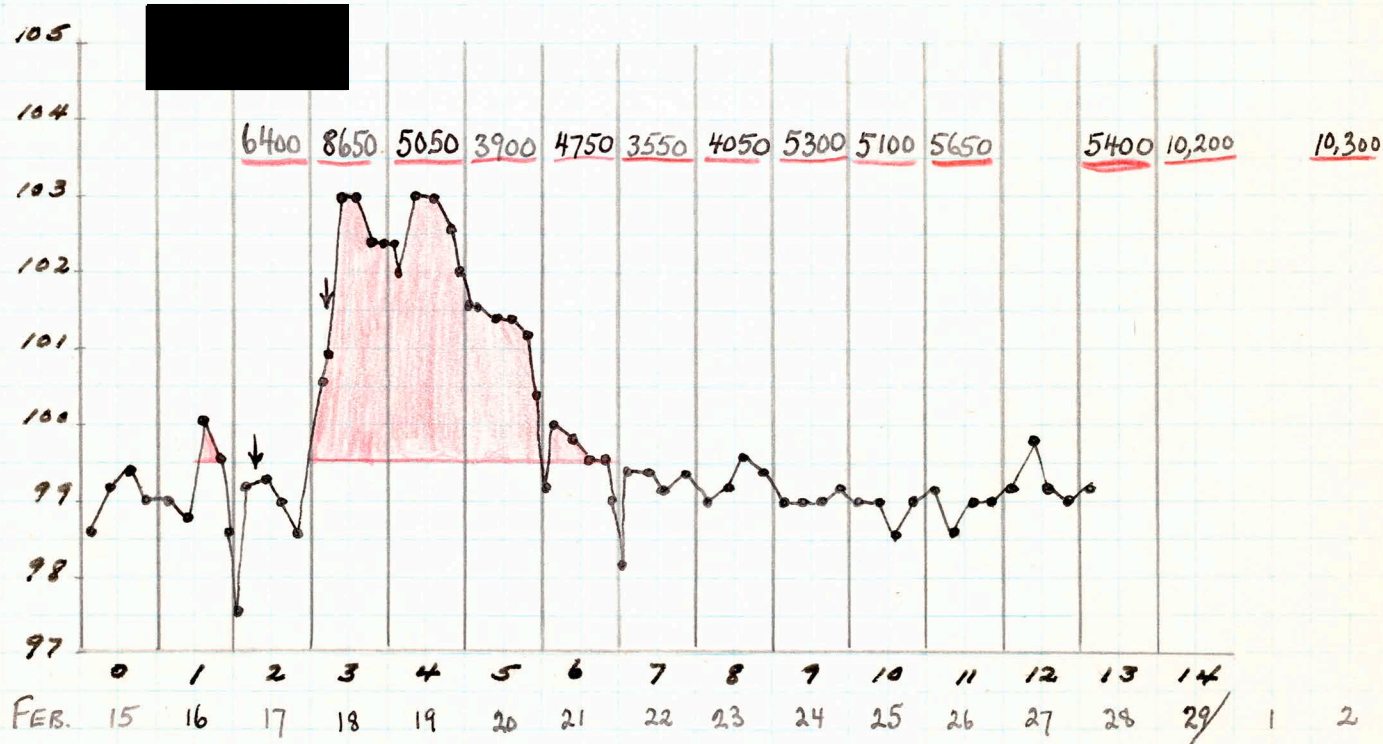
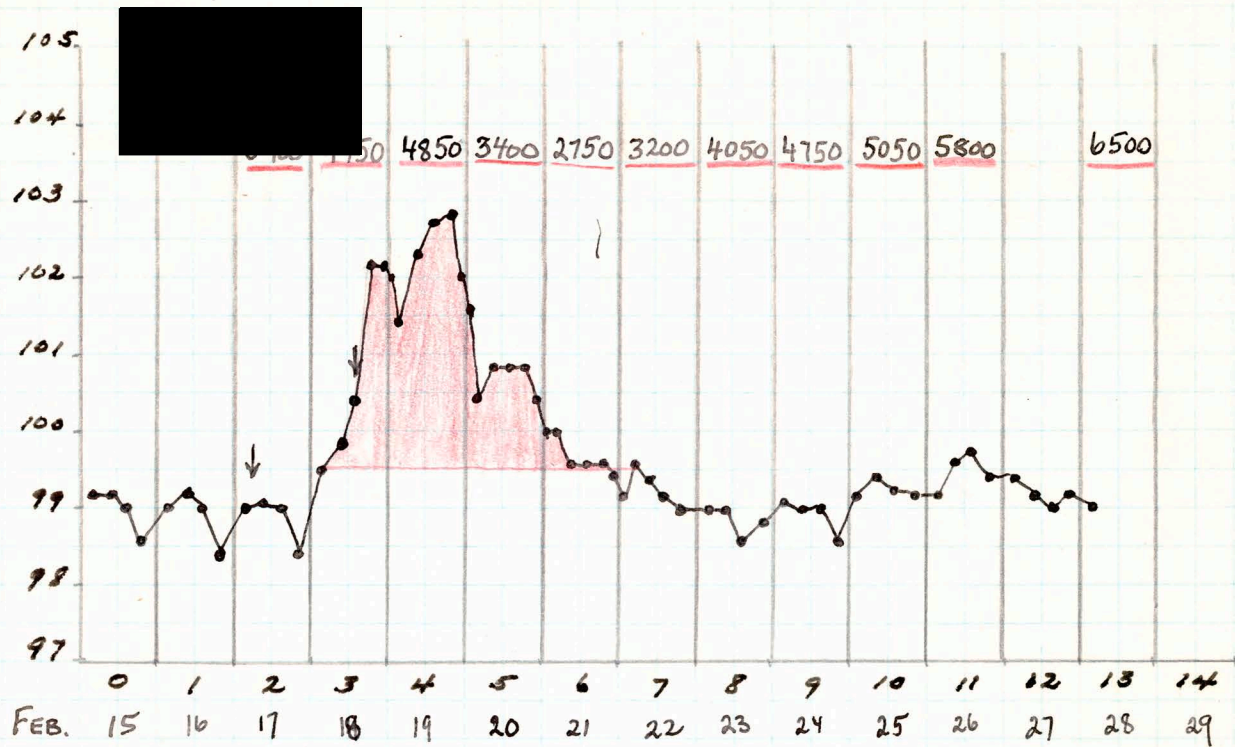
[REDACTED]

- W.M. - AGE 52
- C.F. - " 43

SICILIAN VIRUS

- CHECK ON SUSCEPTIBILITY

CONTROLS FOR ULTRAVIOLET IRRADIATION

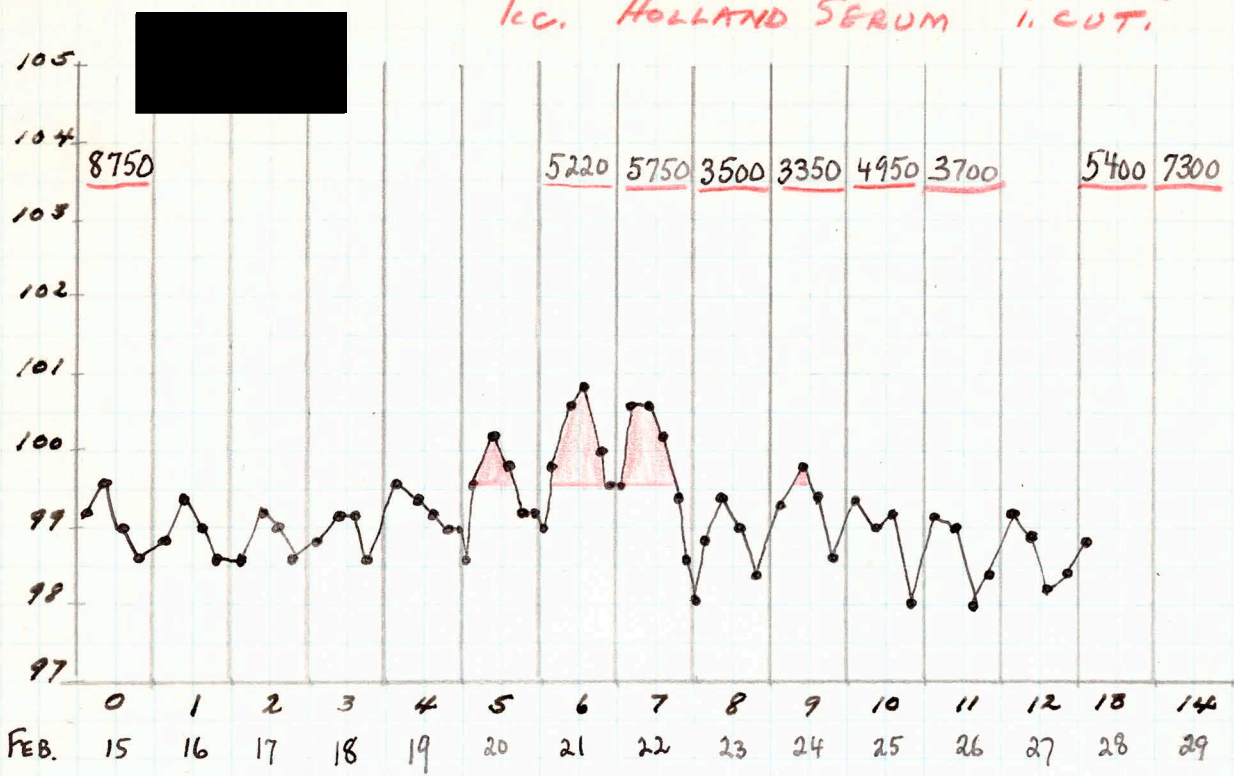


1cc

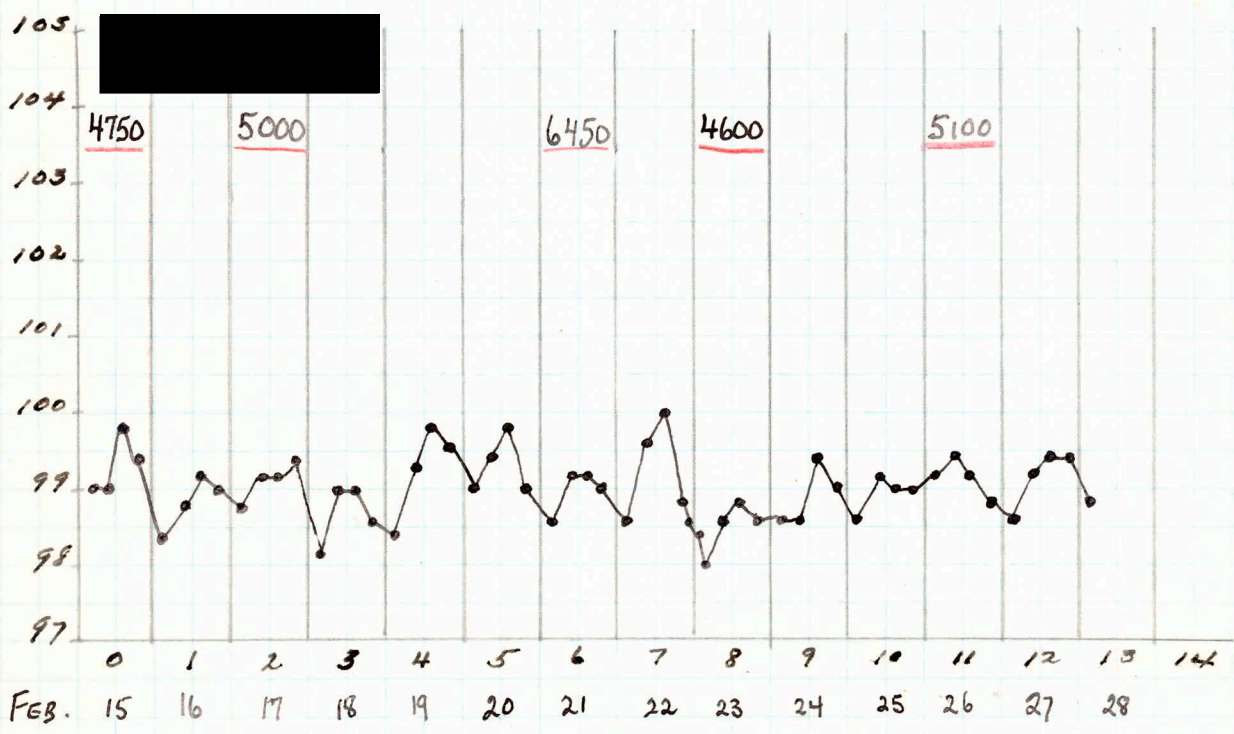
(4TH PASSAGE) SERUM i.c.u.t.

MIDDLE EAST VIRUS -

TEST FOR PRESENCE OF VIRUS IN
SERUM OBTAINED 24-28 HOURS
BEFORE ONSET OF FEVER
I.C.C. HOLLAND SERUM 1. CUT.



TEST FOR PRESENCE OF VIRUS IN COLBERT SERUM
(FEVER 13 DAYS AFTER INOCULATION OF IRRADIATED VIRUS)

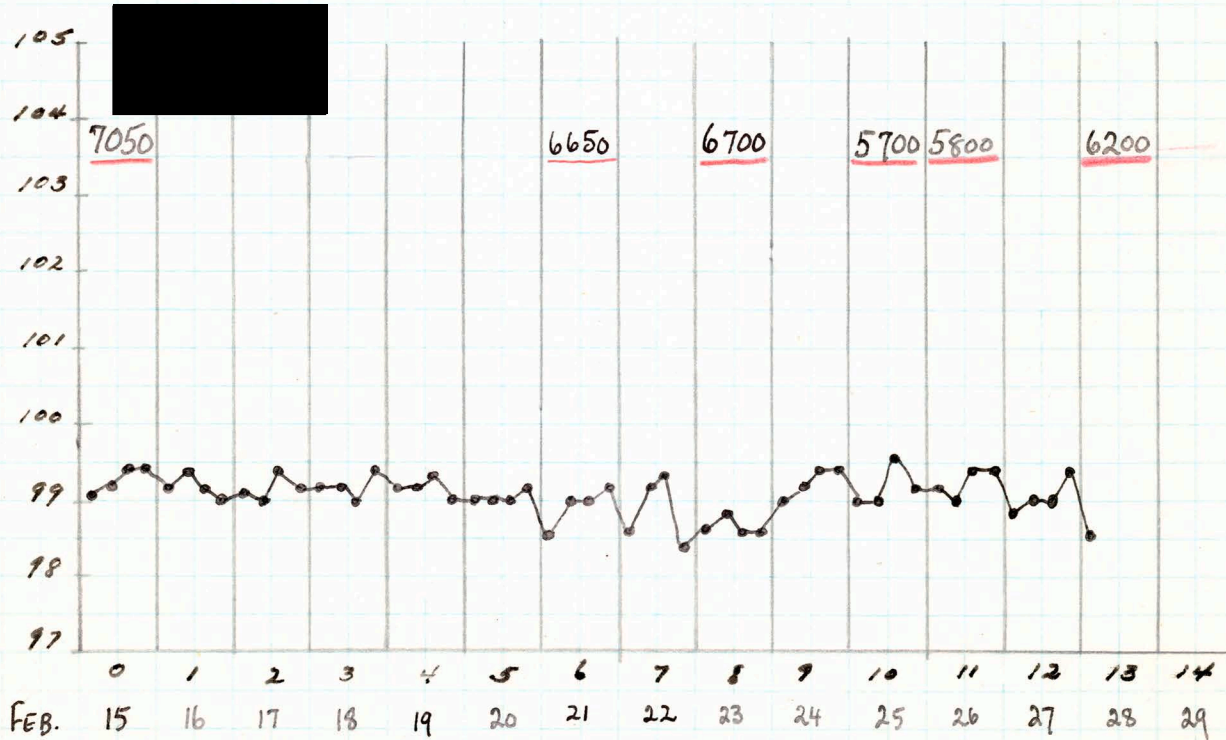
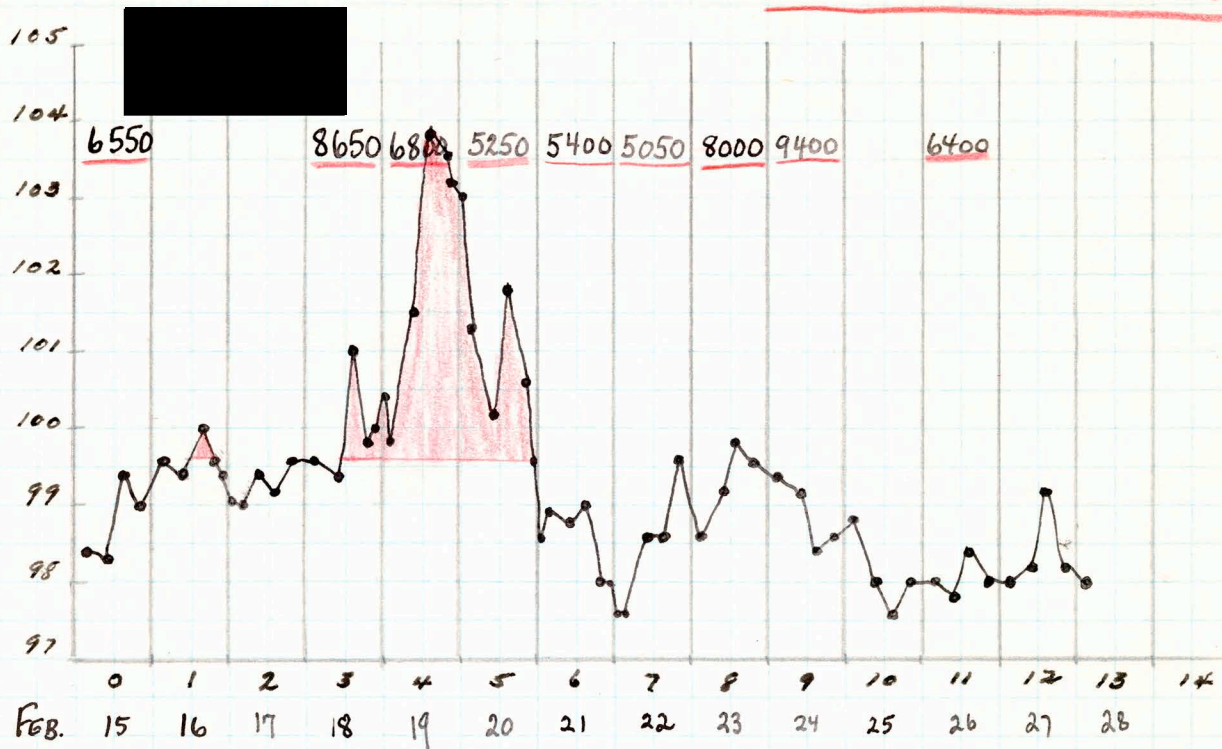


I.C.C. COLBERT SERUM 1. CUT.

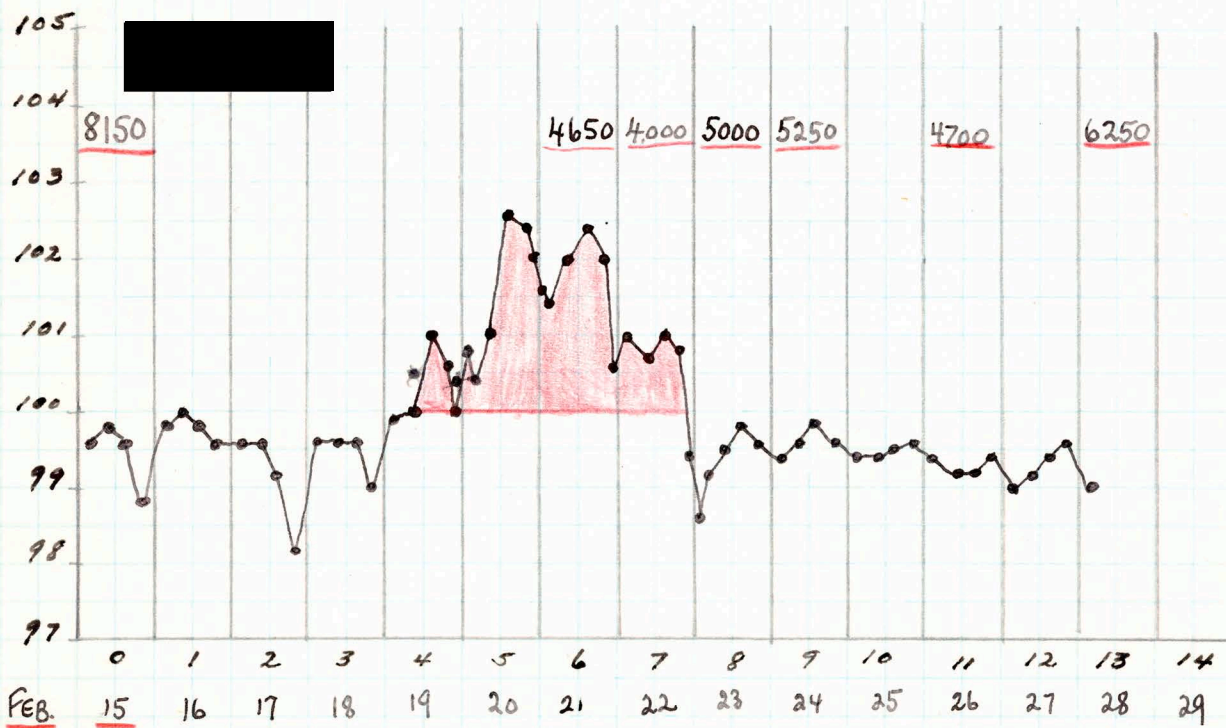
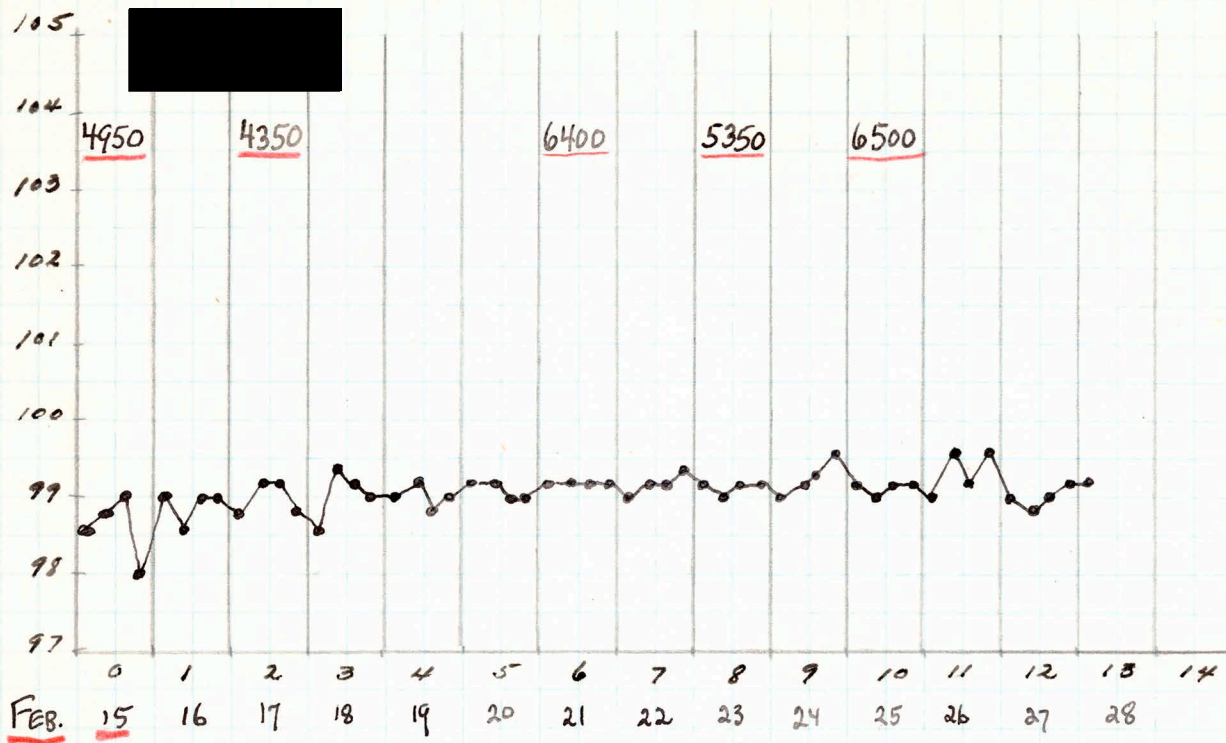
SICILIAN VIRUS - 4TH PASSAGE LAKEMAN - BOSART SERUM

IRRADIATED WITH ULTRAVIOLET LIGHT

FOR ONE HOUR



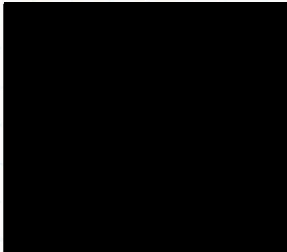
1cc of IRRADIATED SERUM INTRACUTANEOUSLY



1cc OF IRRADIATED SERUM INTRACUTANEOUSLY

ESTIMATION OF SIZE OF *PHLEBOTOMUS* FEVER VIRUS BY FILTRATION THROUGH
PRELIMINARY NOTES ON GRADOCOL MEMBRANE FILTRATION
FEB. 21, 1944

VIRUS - USE SICILIAN STRAIN - 5TH PASSAGE SERA

1.  since FEB. 3 - 10cc
" " " " 7cc
" " " 11 - 10cc
" " " 17 - 10cc
" " " 18 - 6.9cc

36.9cc

2. DILUTE 1:4 WITH SALINE, I.E. ADD 3 VOLUMES OF SALINE TO 1 OF SERUM
3. CENTRIFUGE DILUTED SERUM ON HORIZONTAL CENTRIFUGE FOR 15 min. at about 2000 rpm.
SAVE 20cc for control inoculation
4. SET UP 600 μ , 206.9 μ , and 100 μ filters
5. Pass 10cc of broth through each of these
6. Filter 110-120cc through the 600 μ membrane
(NOTE - if the filtrate does not pass rapidly - change to a 770 μ membrane for the stock filtrate.)
7. Reserve about Set aside 20cc of the 600 μ filtrate
8. Pass about 22cc of the 600 (or 770 μ) filtrate through the 100, 200, 300, and 400 μ membranes
9. Each of two subjects is to receive - 2cc intracutaneously
6cc intravenously
or the equivalent of 0.25cc undil. ser. i.cut.
= 1.75cc " " i.VEN.

NOTES ON ACTUAL FILTRATION ON FEB. 21, 1944

600 mu membrane - 10cc of broth passed rapidly

25cc of centrifuged diluted serum passed in 3 minutes, but second portion of 25cc was going thru dropwise slowly at 14-15 lbs. pressure

It was therefore decided to pass the remainder of the serum thru a 770 mu membrane ~~for~~ as a preliminary to other filtrations

770 mu - 10cc of broth passed thru rapidly under pressure

Remaining 74cc of diluted serum passed thru in about 4-5 min. of filtration time at 14-15 lbs.

400 mu - 10cc of broth passed rapidly under pressure.

About 26cc of 770 filtrate passed rapidly at 14 lbs. requiring only 2 to 3 minutes

17.5cc saved for inoculation - 8cc used for 101 mu membrane

310.6
~~300~~ mu

- Same as 400 mu membrane - 770 filtrate used
About 22cc filtered in 3 to 4 min.

17.5cc for inoculation - 3cc saved for 101 mu

206.9 mu - 10cc of broth passed rapidly

About 25cc of 770 mu filtrate filtered at 15 lbs.

15cc passed in first 3 minutes

20cc " " " 5 minutes

25cc " " " 8 "

17.5cc saved for inoculation - 7cc for 101 mu

101 mu - MEMBRANE "A" - 10cc Broth passed rapidly within 3 minutes

25cc of 600 mu filtrate used at 15 lbs.

Filtration dropwise - slow - 6cc in first hour
10cc " " 3 hours

MEMBRANE "B" - Broth as above. Fluid previously passed thru 400 mu (8cc), 300 mu (3cc), 207 mu (7cc) was pooled (total 18cc) and filtered thru at 14 lbs.

Filtration relatively rapid though dropwise

10cc in first 5 minutes

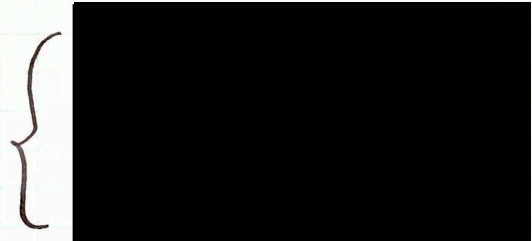
15cc in " 15 "

SICILIAN VIRUS - GRADUOL MEMBRANE FILTRATION TEST

INOCULATIONS - FEB. 21, 1944

UNFILTERED VIRUS
SERUM 1:4-CENTRIFUGED

2cc INTRACUT.
6cc INTRAVEN.



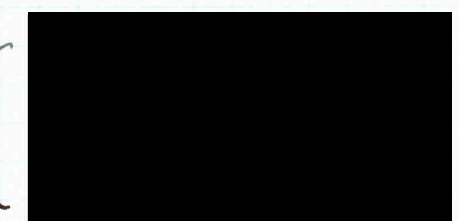
600 mμ FILTRATE

2cc INTRACUT.
6cc INTRAVEN.



400 mμ FILTRATE

2cc INTRACUT.
6cc INTRAVEN.



- W.M. - AGE 56

- C.F. - " 32

300.6 mμ FILTRATE

2cc INTRACUT
6cc INTRACUT



- W.M. - AGE 48

- C.F. - " 35

206.9 mμ FILTRATE

2cc INTRACUT
6cc INTRAVEN



- C.M. - " 37

- C.F. - " 46

101 mμ FILTRATE

2cc "B" INTRACUT
10cc (A+B) INTRAVEN.



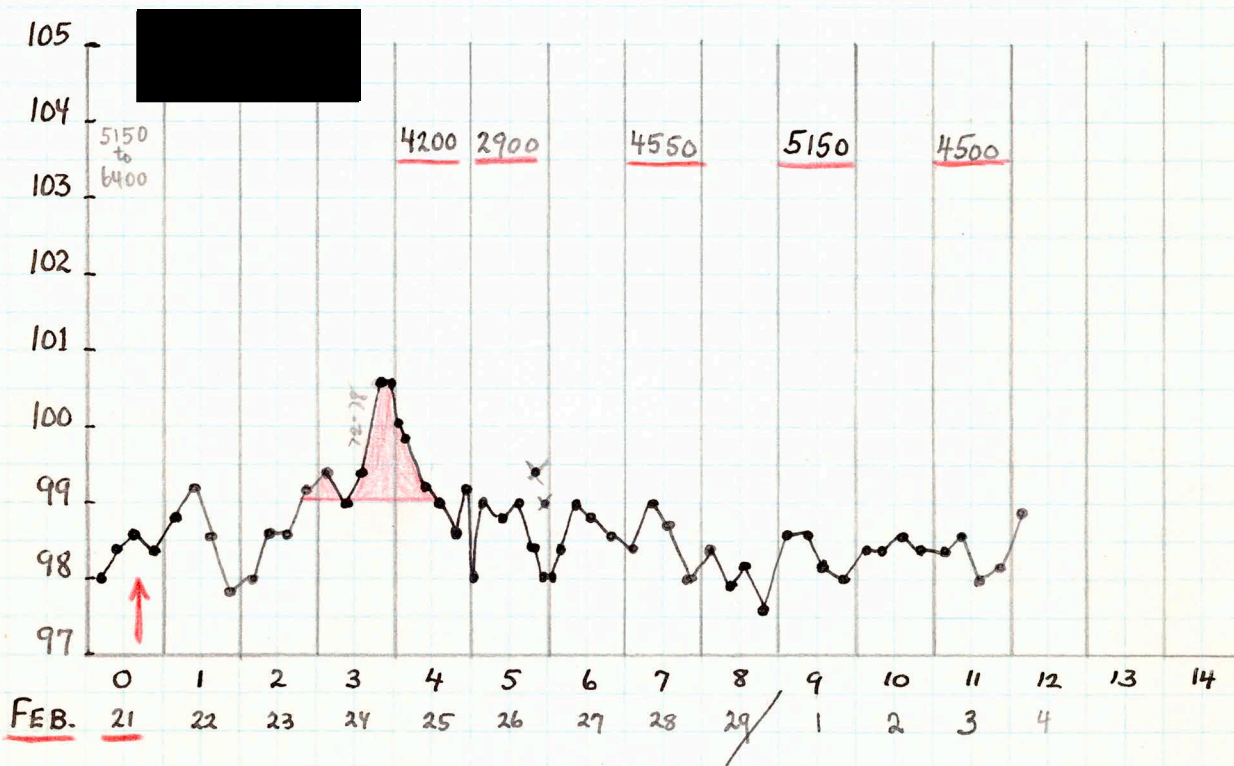
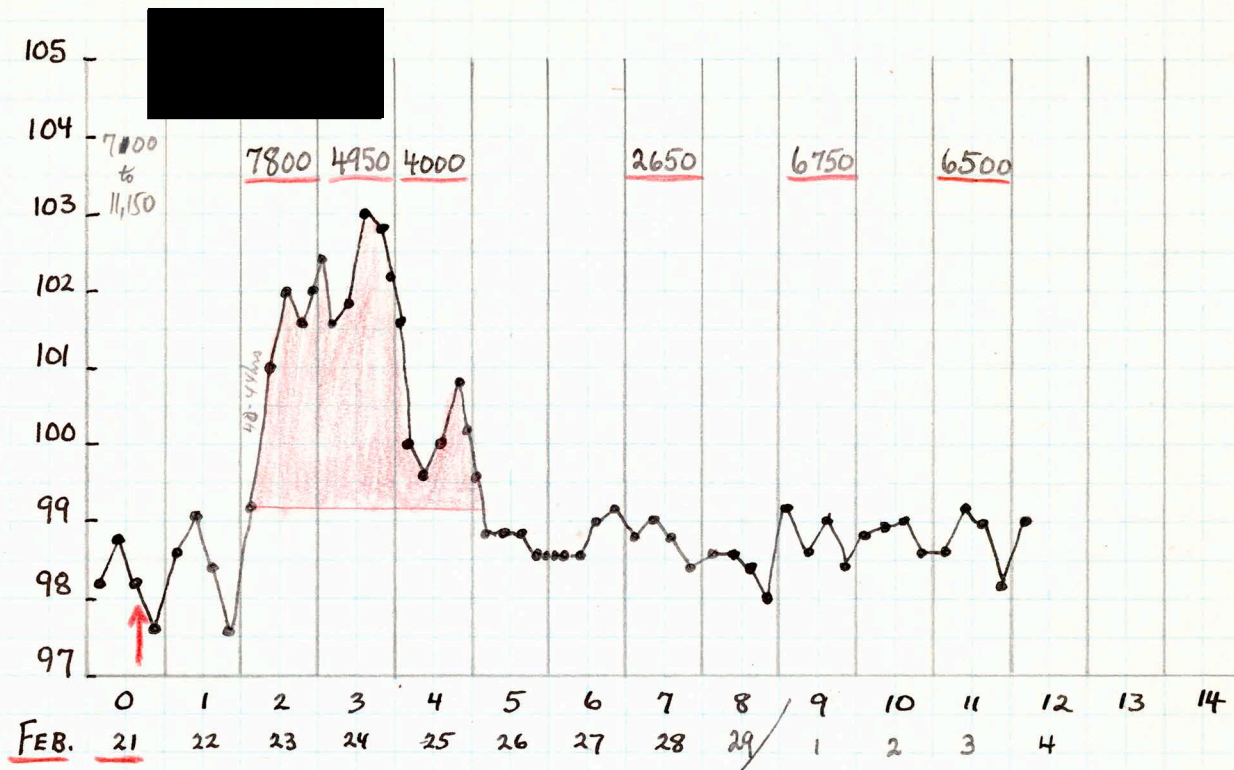
- C.M. - " 45

- W.F. - " 15

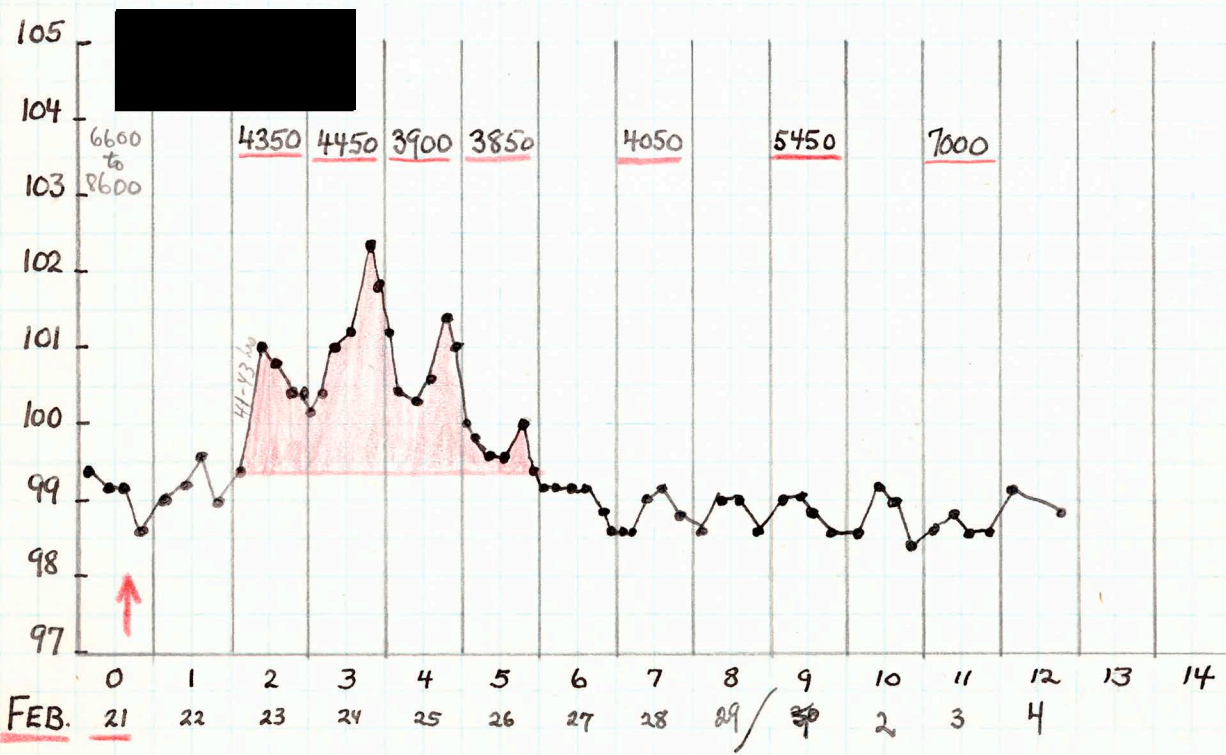
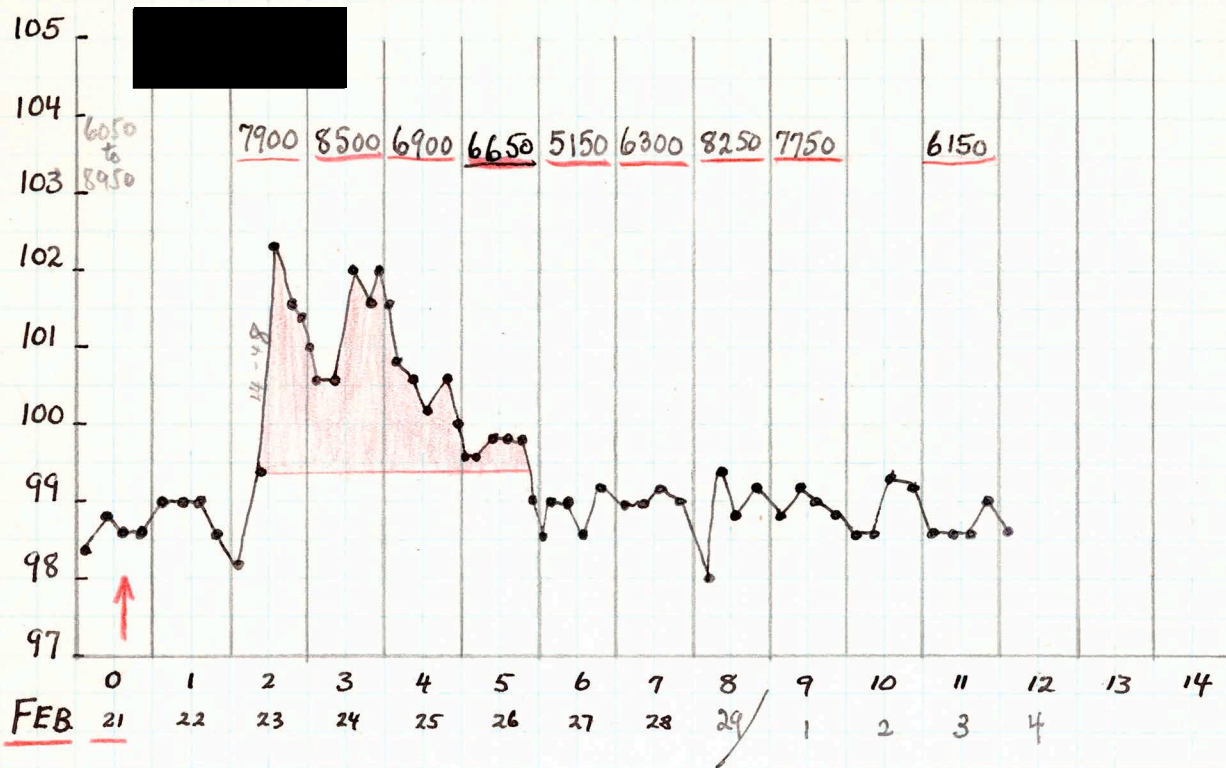
MEN 345-445 P.M. ^{8:10 cc} WOMEN 450-545

SICILIAN VIRUS -

UNFILTERED, DILUTED, CENTRIFUGED VIRUS



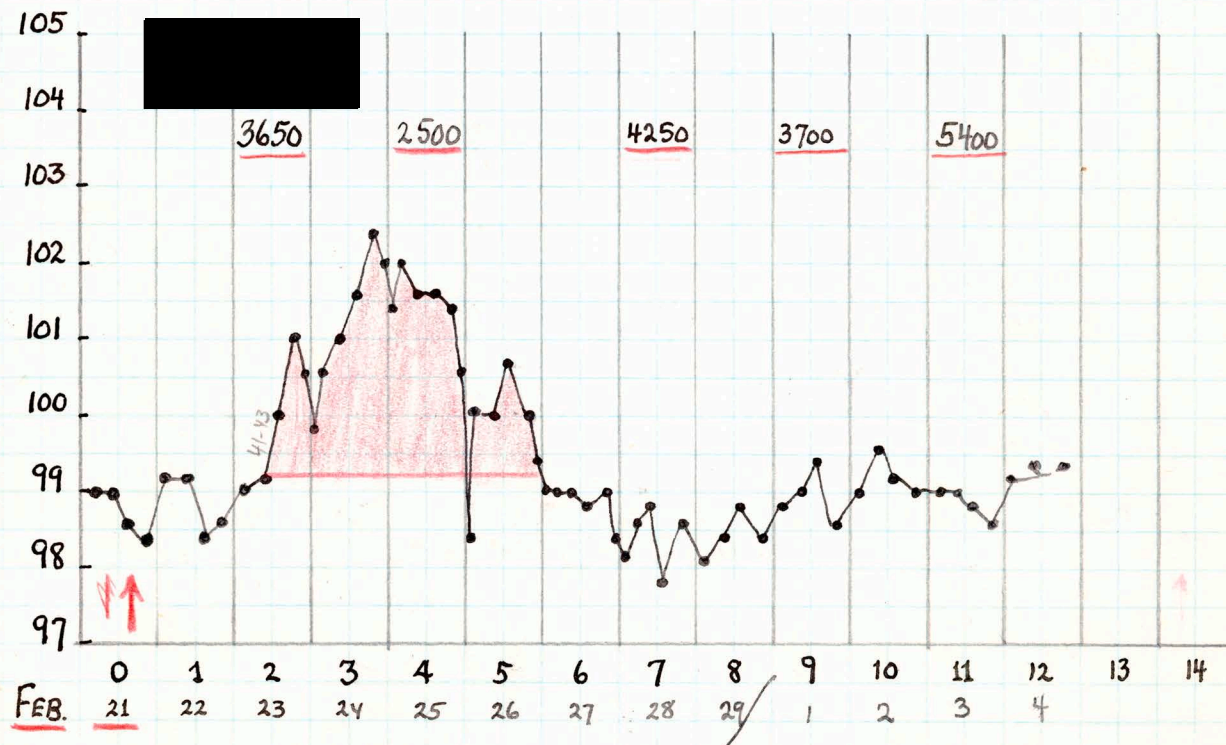
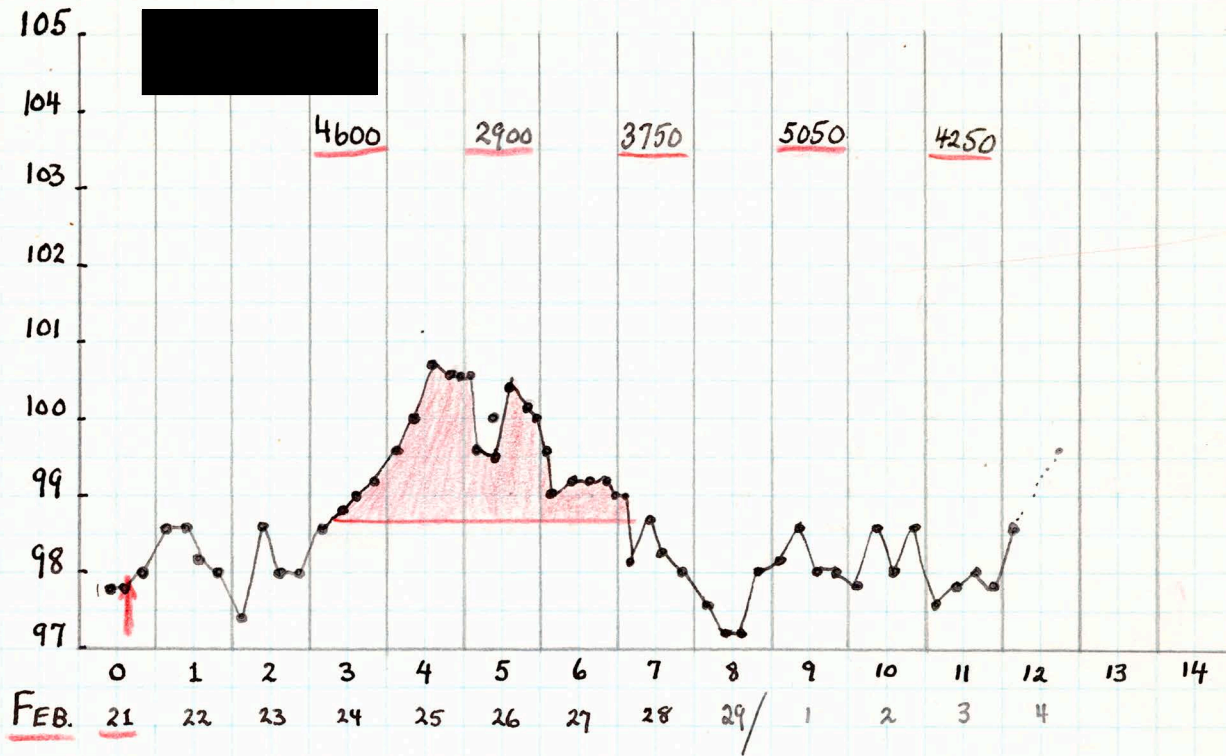
2cc INTRACUT
6cc INTRAVEN



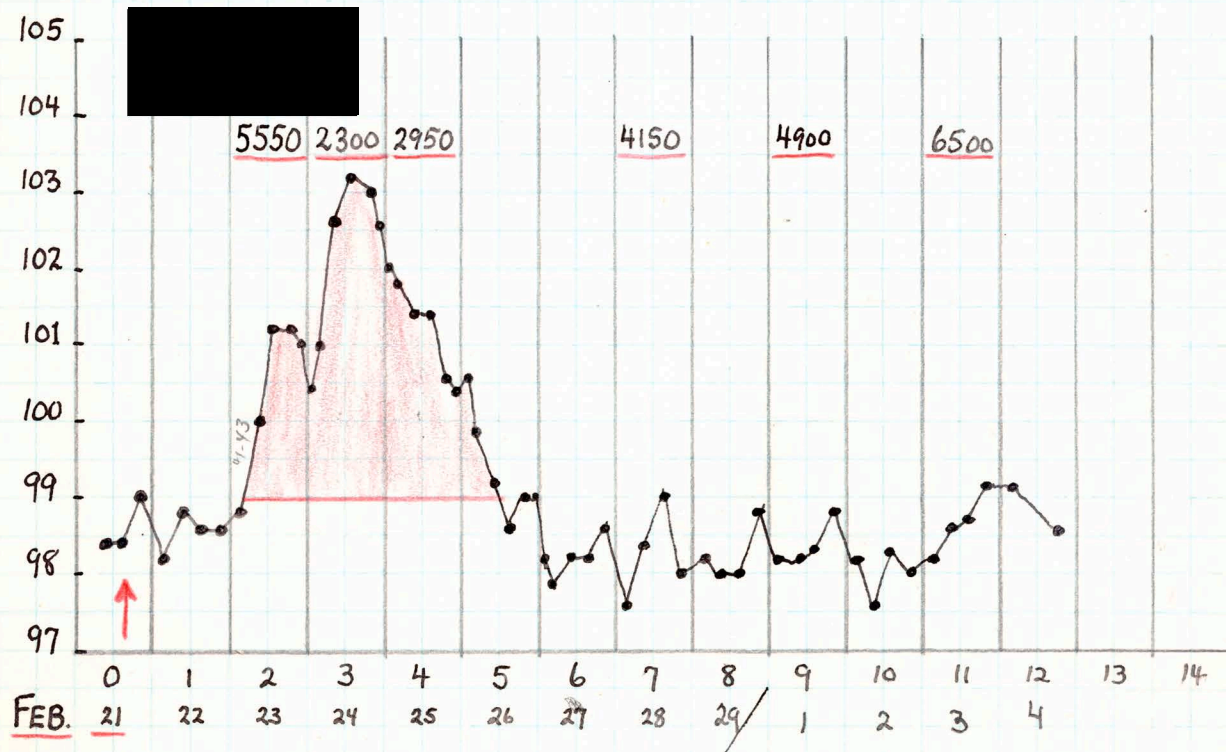
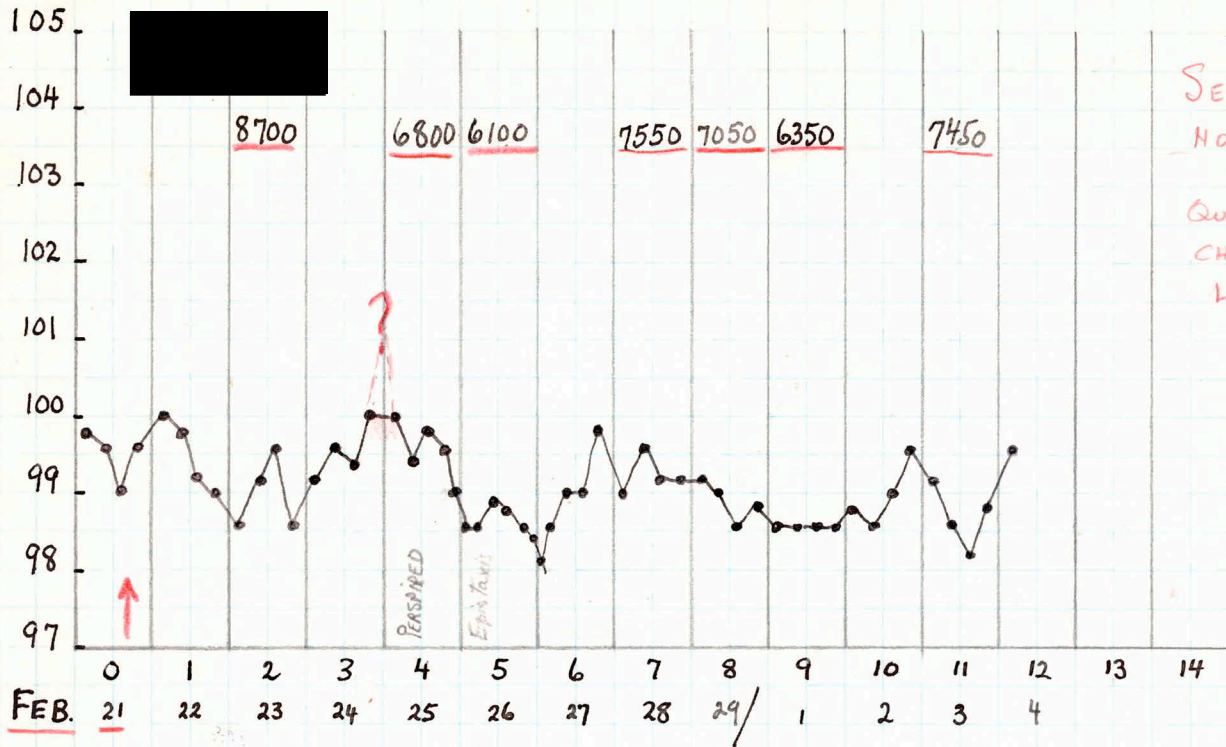
2 cc INTRACUT.
6 cc INTRAVEN.

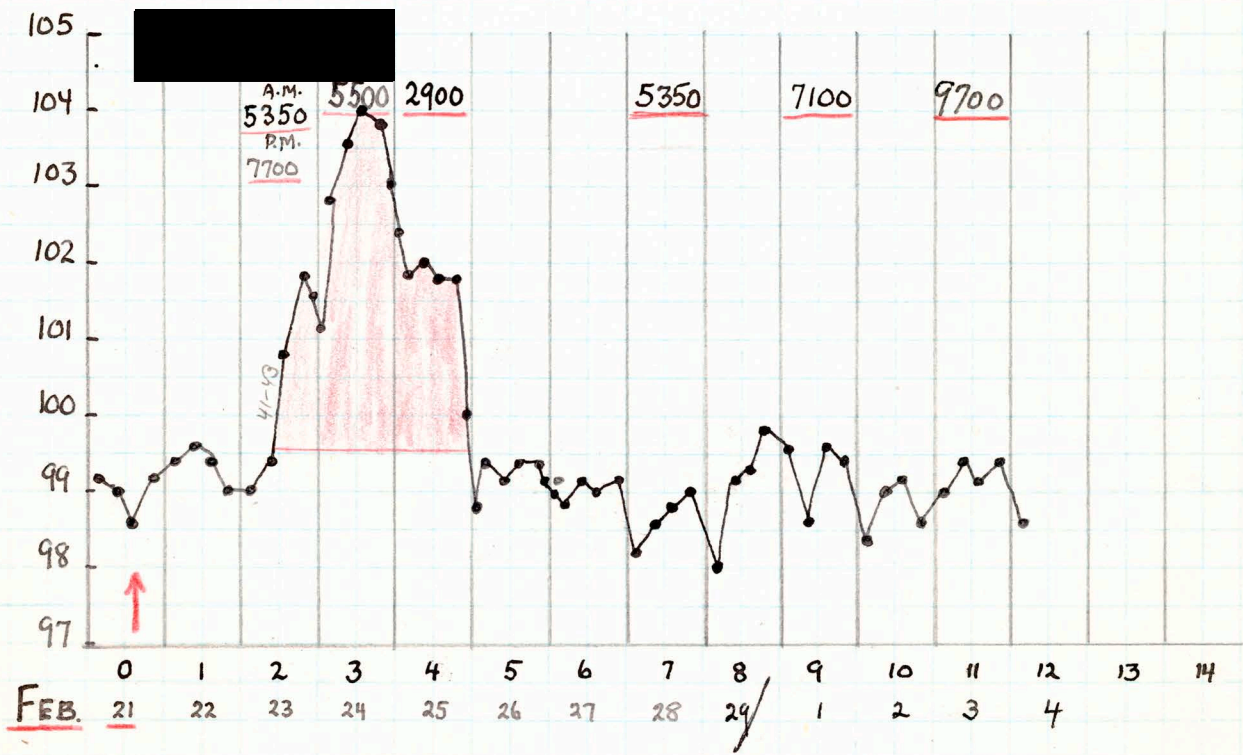
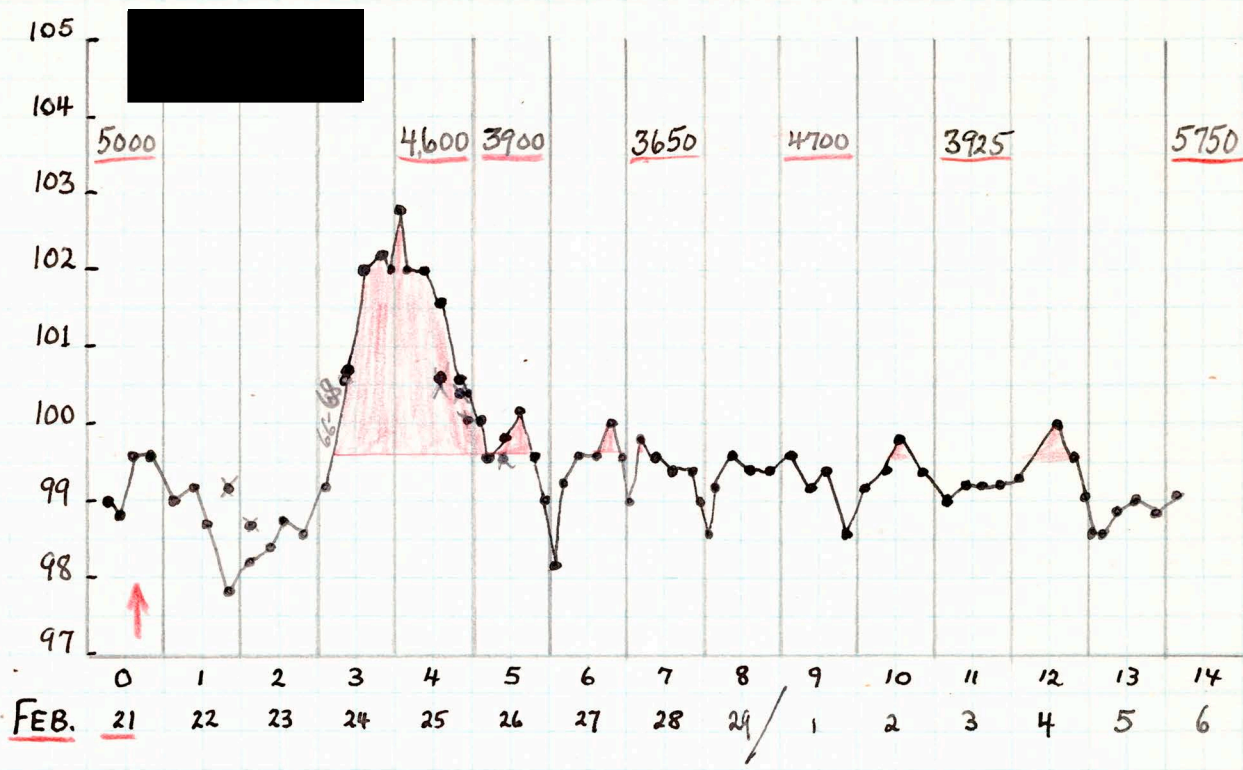
SICILIAN VIRUS

400 mju FILTRATE

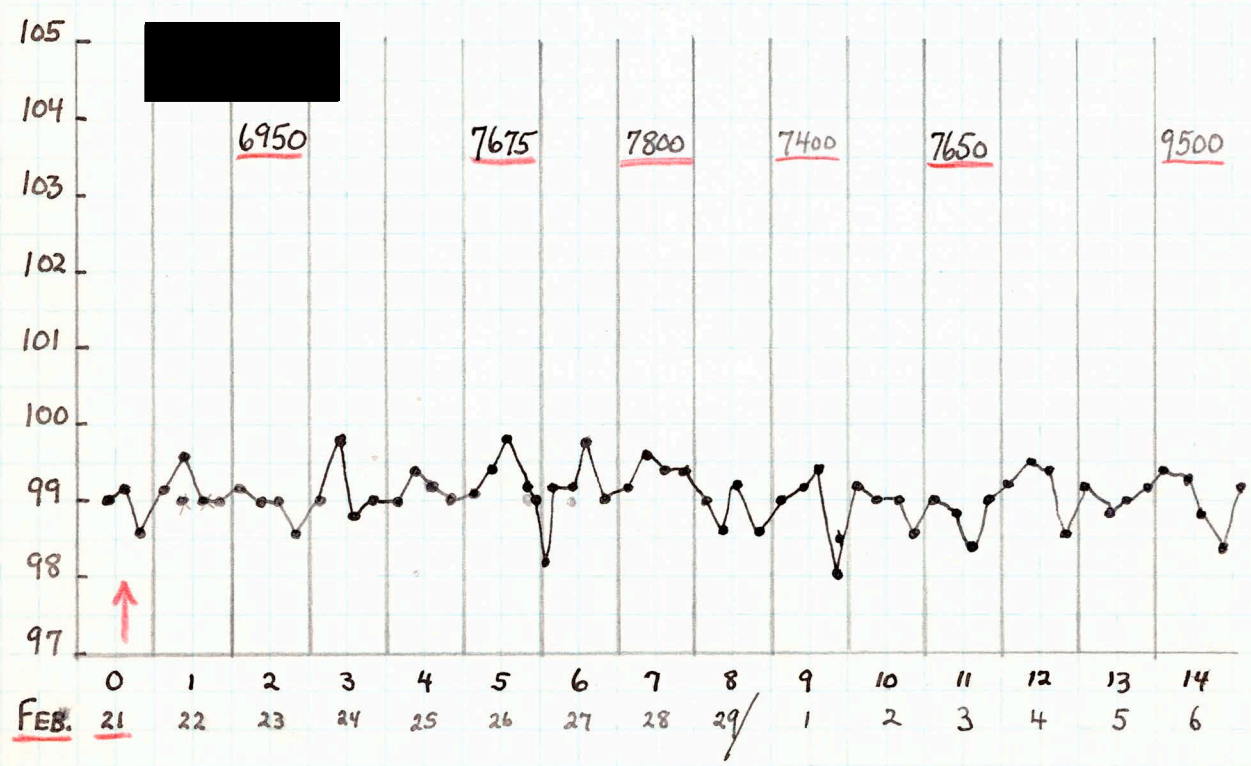
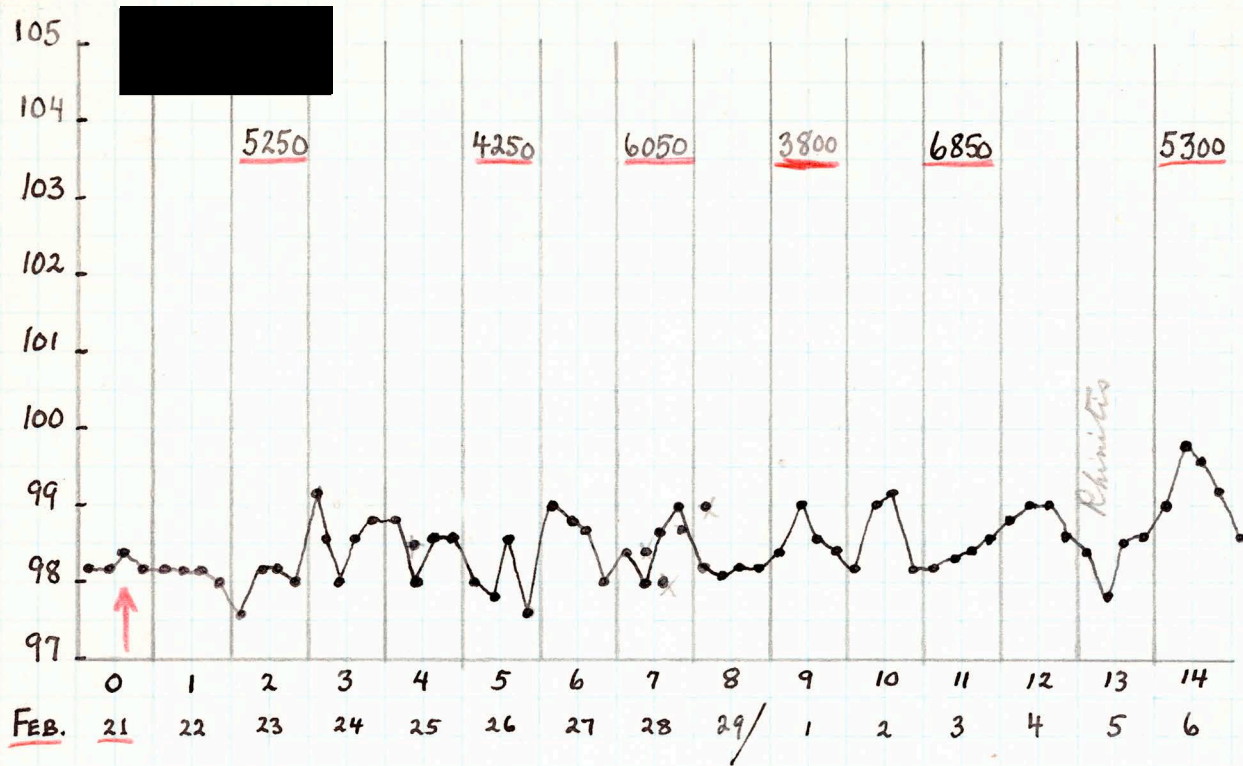


2 cc INTRACUT
6 cc INTRAVEN.





2 cc INTRACUT.
6cc INTRAVEN.



2cc INTRACUT.
10cc INTRAVEN.

1. CONCENTRATION OF VIRUS IN POOL USED FOR GRADOCOL
MEMBRANE FILTRATION TEST

VIRUS - POOL OF SNAPP, GILBERT, BOCKHORST, WALLACE AA
1:10,000 and 1:100,000 dilutions prepared in
saline shortly before inoculation

1cc, 1:10,000
intraven.

1cc, 1:100,000
i. VEN.

white male - age 49

colored female - " 37

" male - " 34

" female - " 35

2. TEST FOR NEUTRALIZING ANTIBODY IN SERA OF RECOVERED,
IMMUNE PERSONS

"IMMUNE" SERUM - SAME AS USED IN PREVIOUS TEST, I.E.
POOL OF IRVIN, WALTERS, MORSE, RIGGS
(frozen ampules - not previously thawed)

VIRUS - POOL USED ABOVE

2.2cc serum + 1.1cc virus - mixed at 12:30 P.M.
left at room temp. until
inoculation 2:50 P.M.

1.5cc of
mixture
intracutan.

WHITE MALE - AGE 39

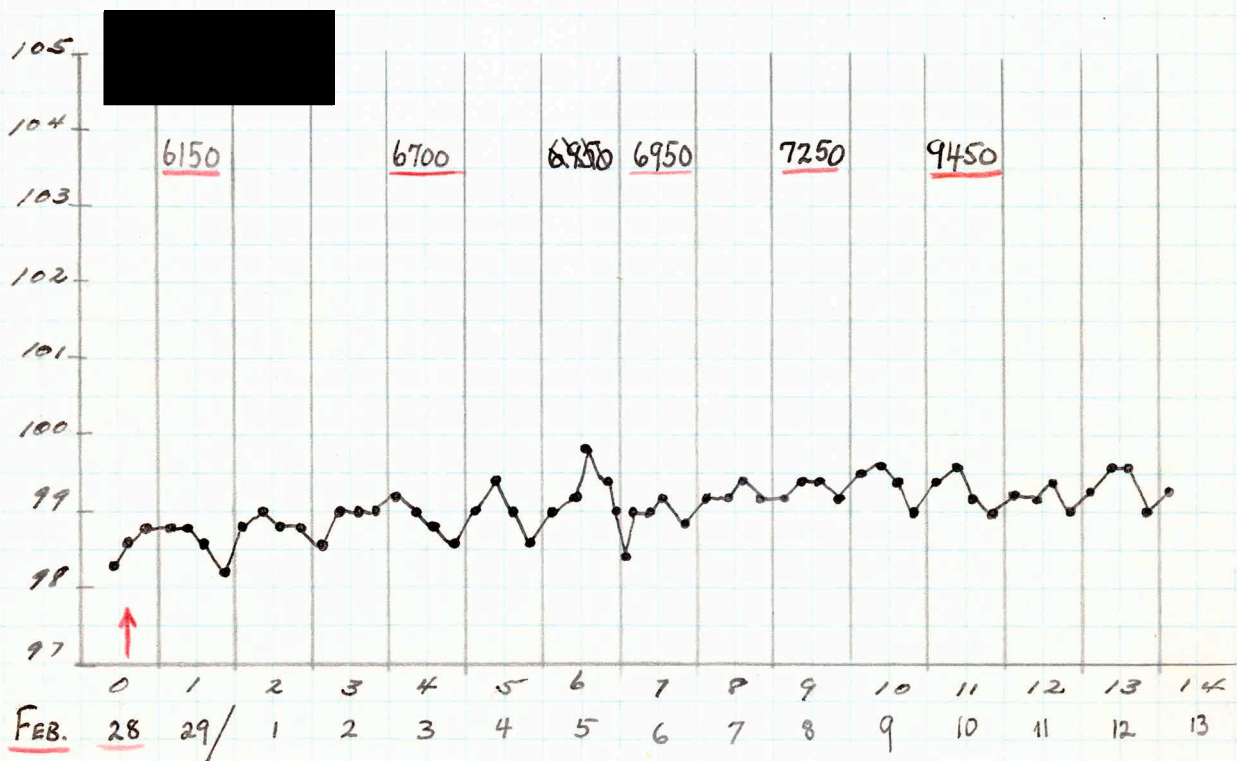
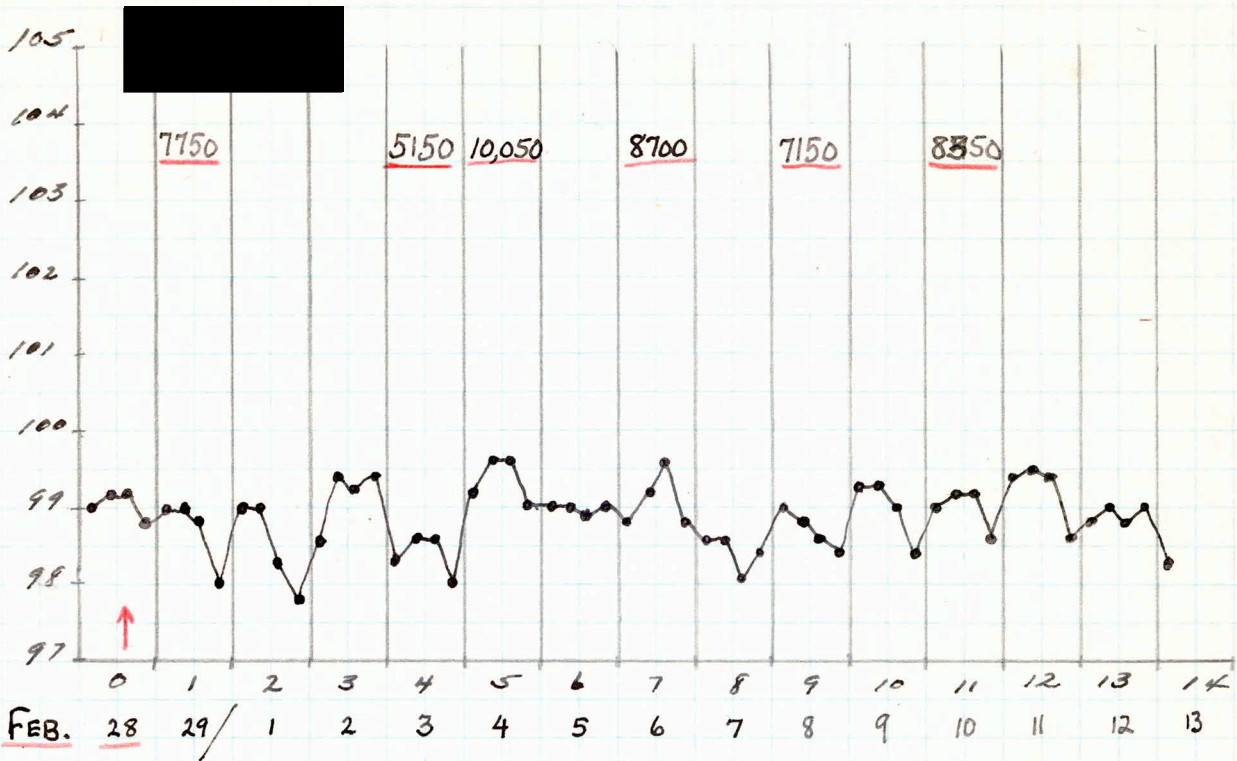
COLOR " - " 40

3. TEST FOR SUSCEPTIBILITY AND CONTROL FOR ABOVE

0.5cc of above -
virus i. CUTAN.

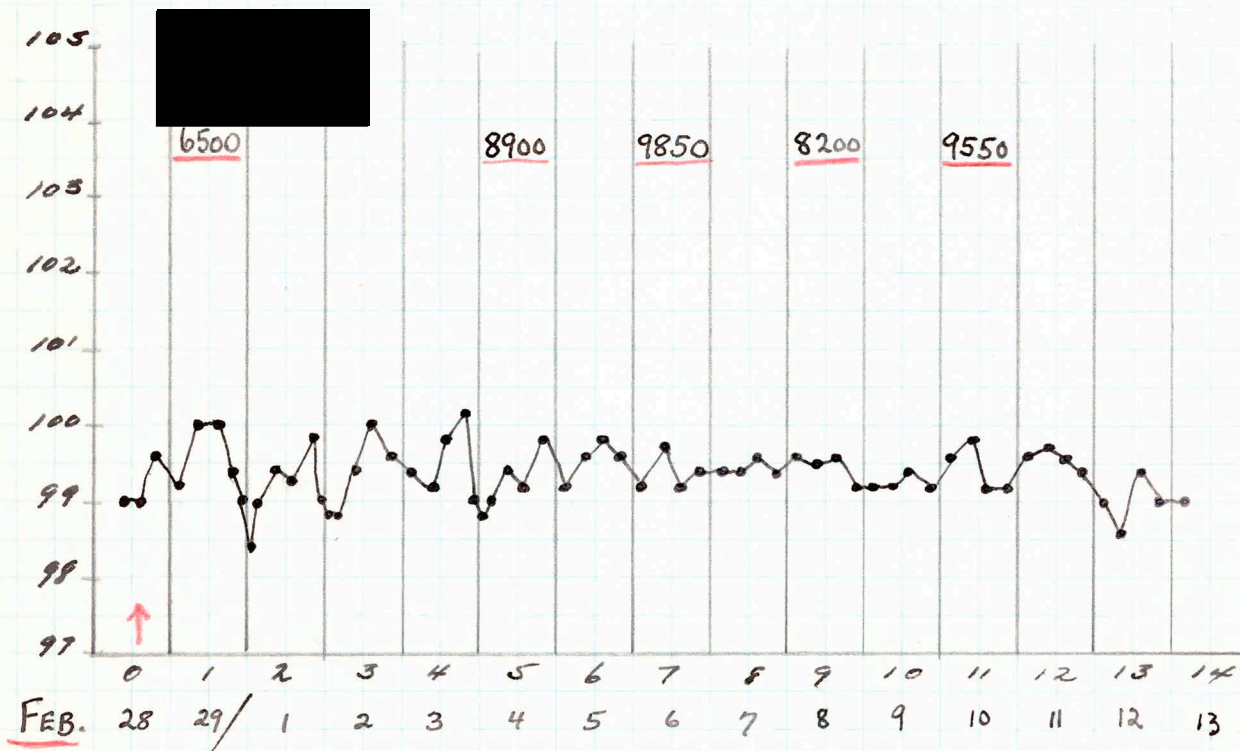
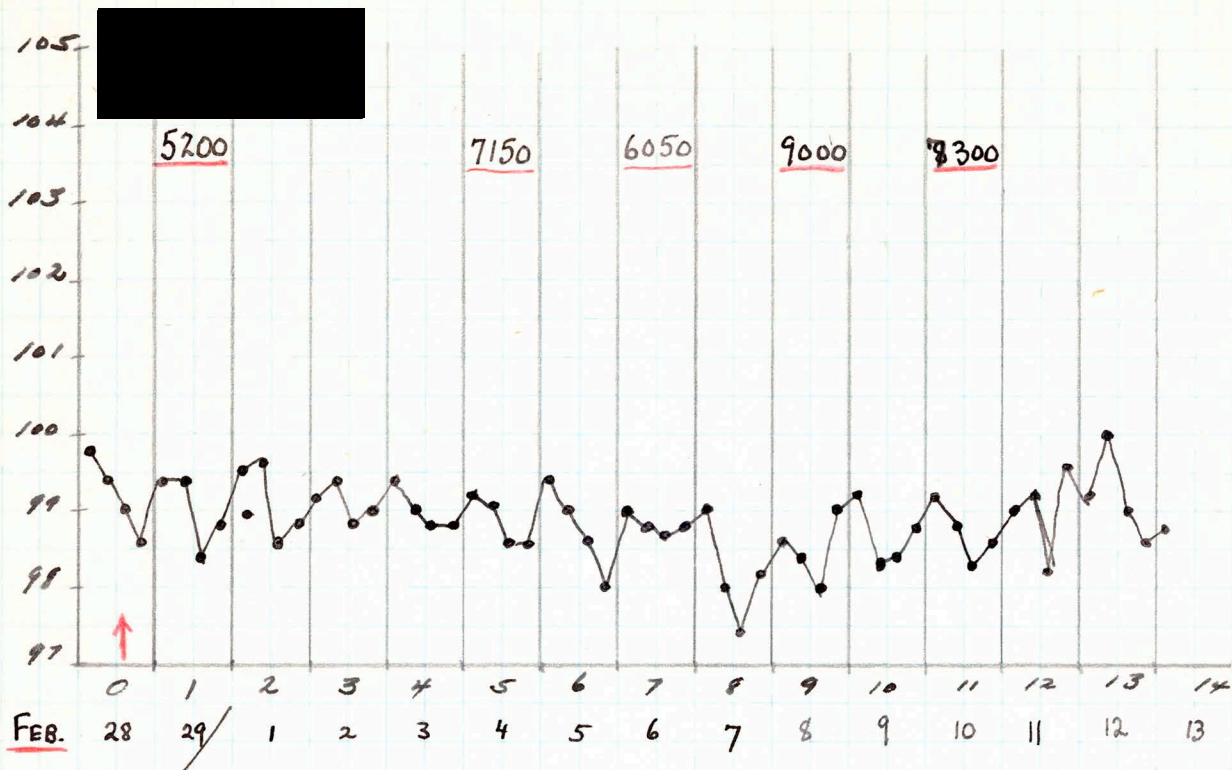
WHITE FEMALE - AGE 50

SICILIAN VIRUS - 5TH PASSAGE POOL - CONCENTRATION OF VIRUS



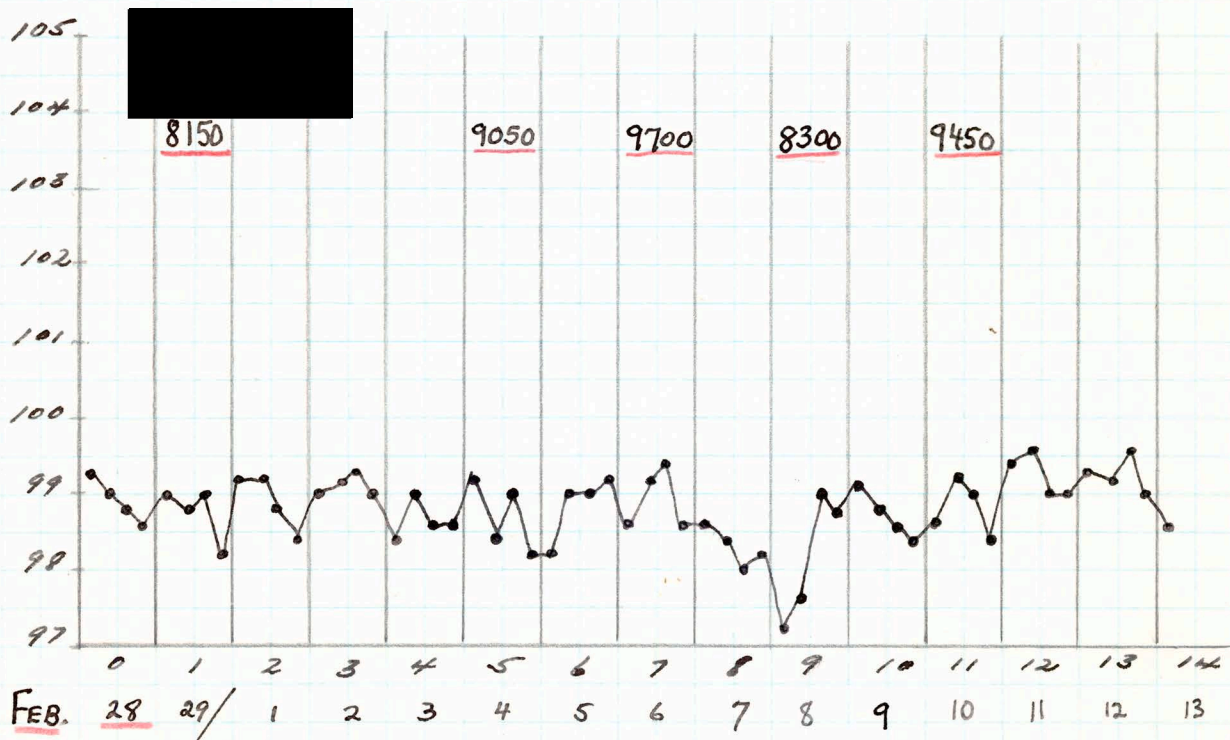
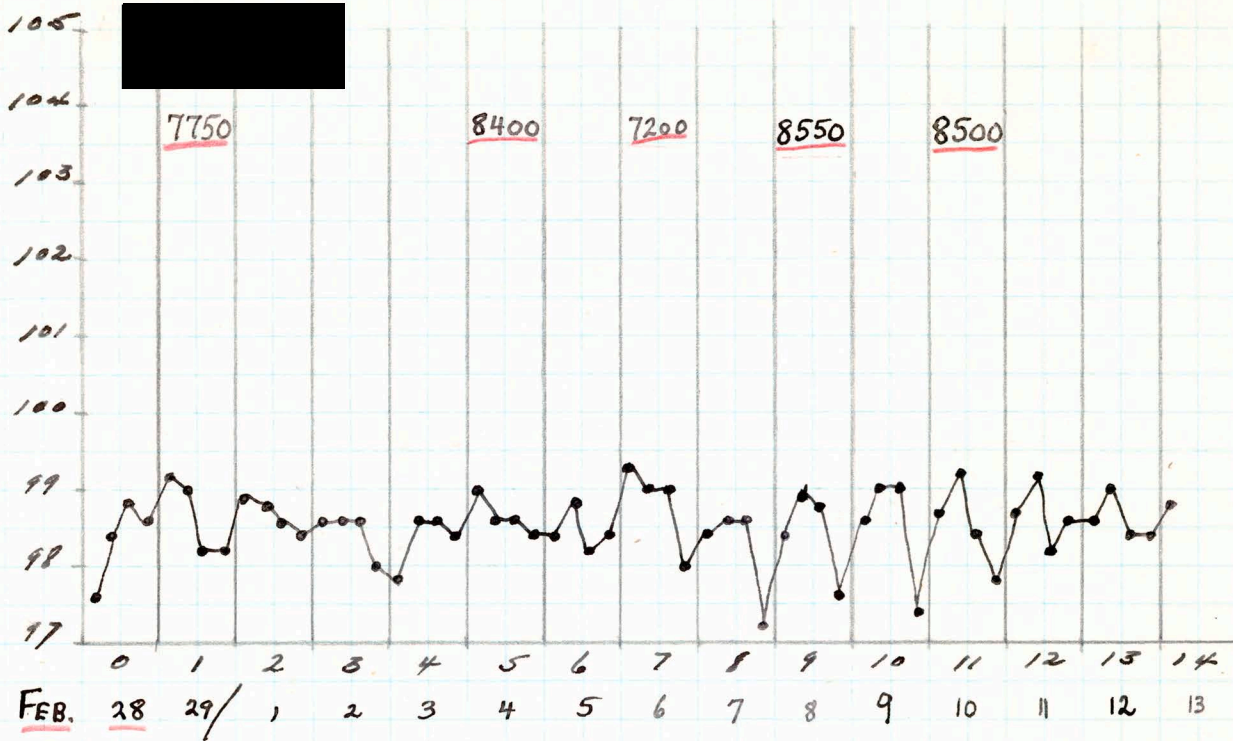
1cc of 1:10,000 INTRAVENOUSLY

SICILIAN VIRUS - 5TH PASSAGE POOL - CONCENTRATION OF VIRUS



1cc OF 1:100,000 INTRAVENOUSLY

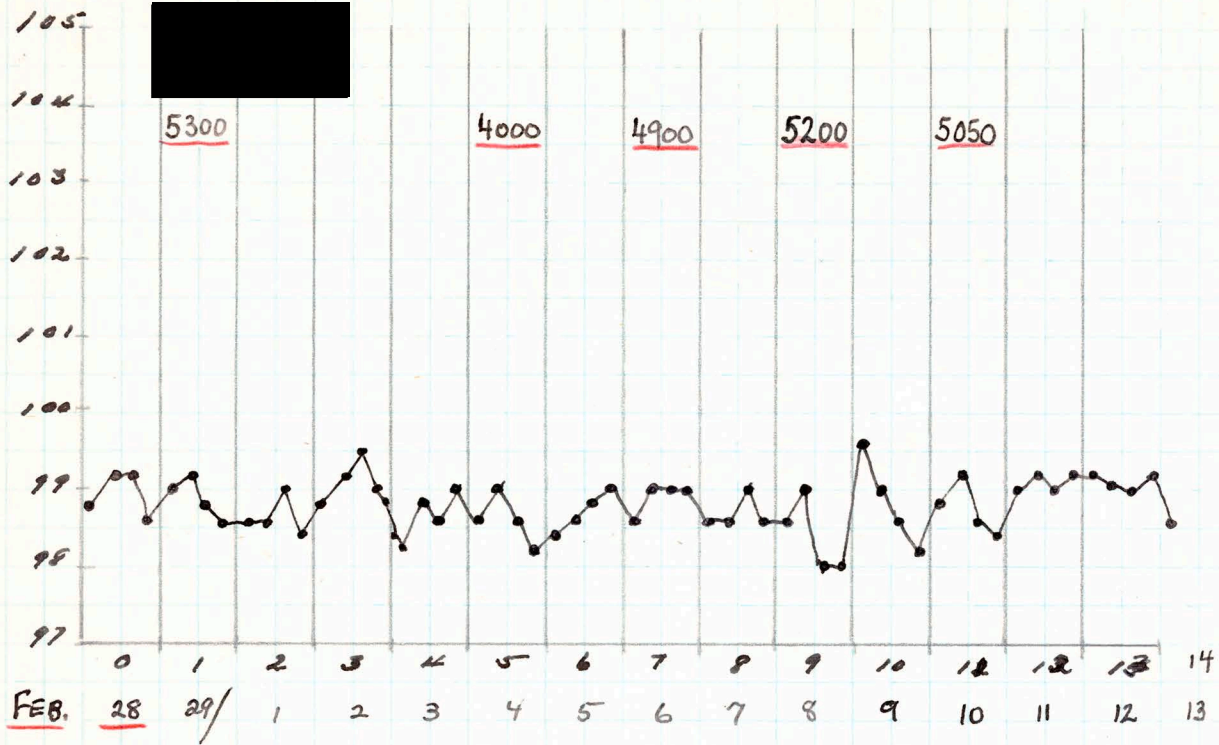
SICILIAN VIRUS - TEST FOR NEUTRALIZING ANTIBODY



1.5 cc OF CONVALESCENT SERUM - VIRUS MIXTURE
 INTRACUTANEOUSLY
 (1cc of IMMUNE SERUM + 0.5cc VIRUS)

SICILIAN VIRUS - TEST FOR SUSCEPTIBILITY

0.5cc 5th PASSAGE POOL INTRACUTAN.



SICILIAN VIRUS

MARCH 7, 1944

1. FOUR MONTH IMMUNITY TEST
2. TEST FOR SUSCEPTIBILITY OF SUBJECTS INOCULATED WITH 100 m μ FILTRATE

VIRUS USED - POOL OF 5TH PASSAGE SERA USED IN FILTRATION AND NEUTRALIZING ANTIBODY TESTS

4.0 cc
4.5 cc
4.7 cc
2.7 cc
15.9 cc

<u>PREVIOUS INOCULUM</u>	<u>1ST FEVER DATE</u>	<u>NAME</u>	<u>TEST DOSE</u>
SICILIAN	OCT. 25/43	[REDACTED] W.M. - AGE 43	1 cc
"	NOV. 5 "	[REDACTED] " " - " 41	"
"	" "	[REDACTED] " " - " 48	"
"	" "	[REDACTED] " " - " 52	"
"	" "	[REDACTED] W.F. - " 45	"
"	NOV. 13	[REDACTED] " " - " 43	"
MIDDLE EAST	NOV. 7	[REDACTED] W.M. - " 42	"
" "	" "	[REDACTED] C.M. - " 55	"
" "	" "	[REDACTED] W.F. - " 39	"
" "	NOV. 6	[REDACTED] " " - " 46	"
0.01 cc	No FEVER OCT. 11	[REDACTED] W.M. - " 46	<u>0.5 cc</u>
0.001 cc	OCT. 11 - NO FEVER	[REDACTED] W.F. - " 50	"
101 m μ FILTRATE		[REDACTED] C.M. - " 45	1 cc
" "		[REDACTED] W.F. - " 15	"

All inoculated intracutaneously 10 to 11 A.M.

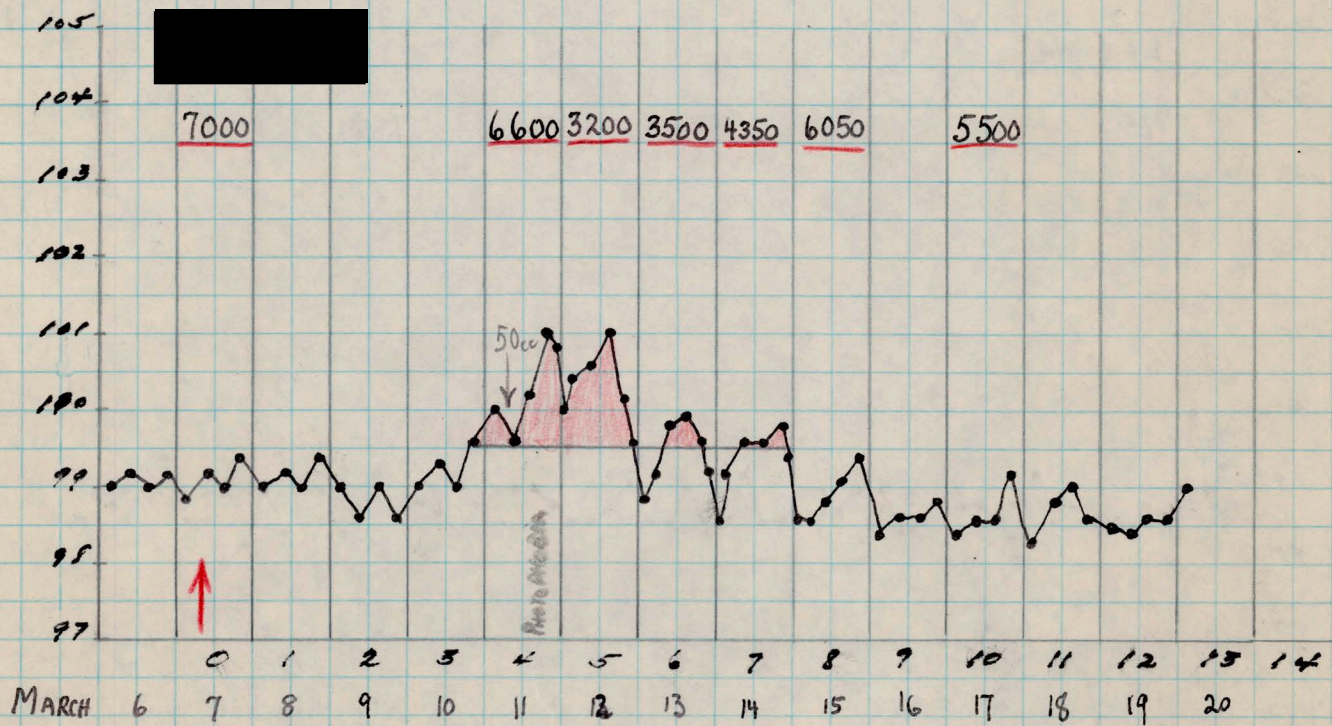
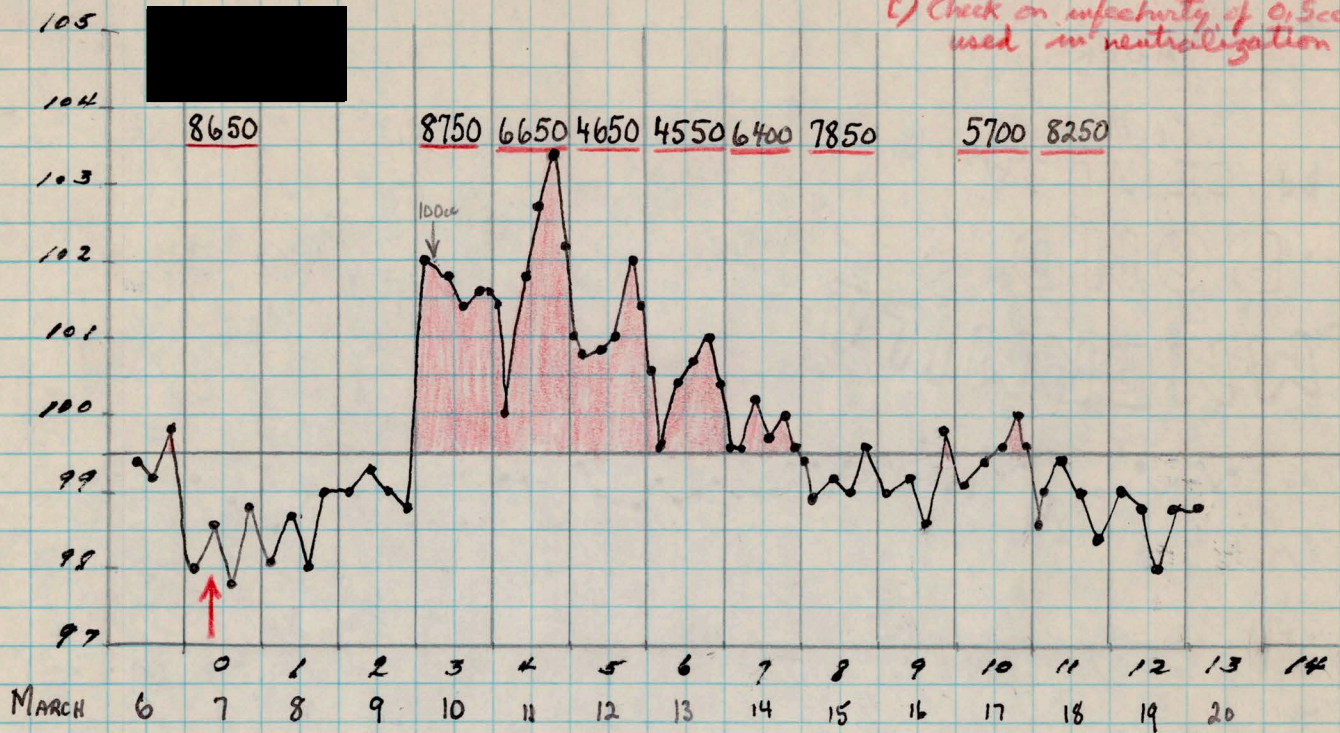
MARCH 8, 1944 - No reaction at any of the inoculated sites

SICILIAN VIRUS

- CONTROLS FOR a) 4 MONTH IMMUNITY TEST

b) 100mu subjects' IMMUNITY TEST

c) Check on infectivity of 0.5cc dose, used in neutralization test 3/23/44

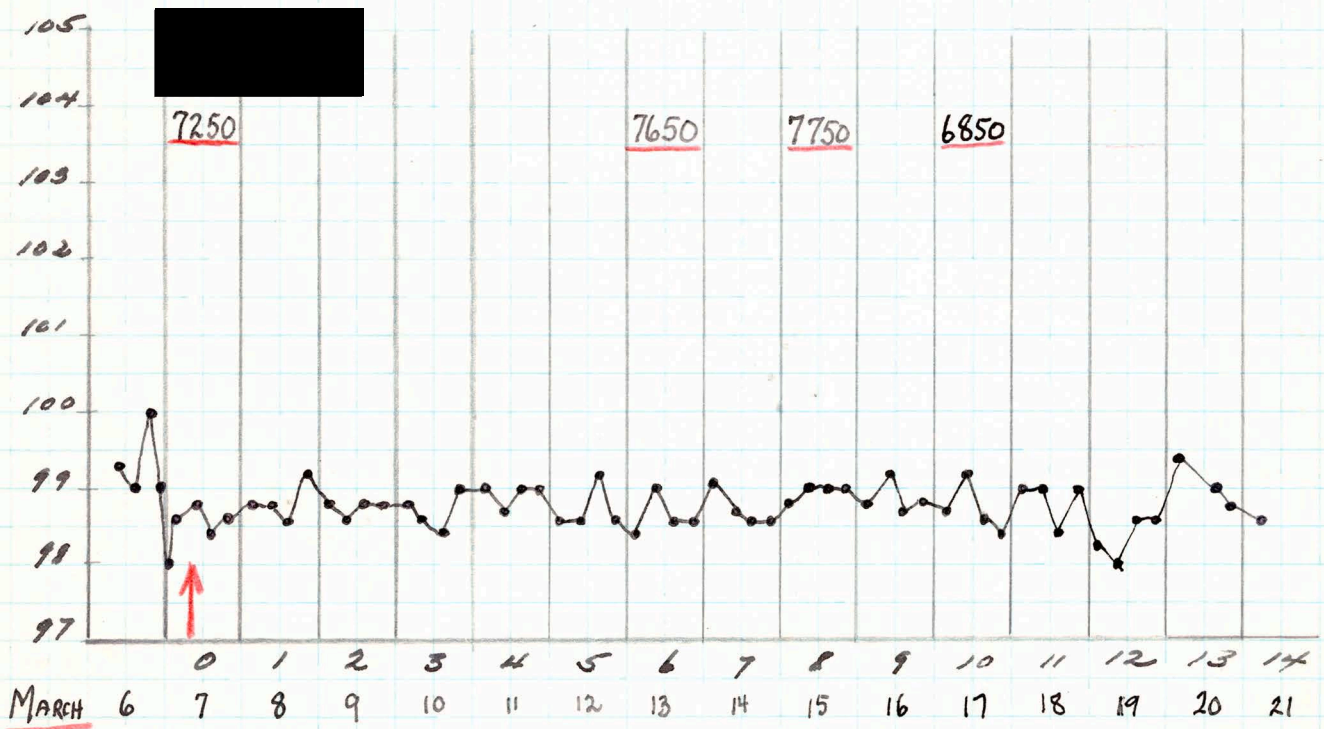
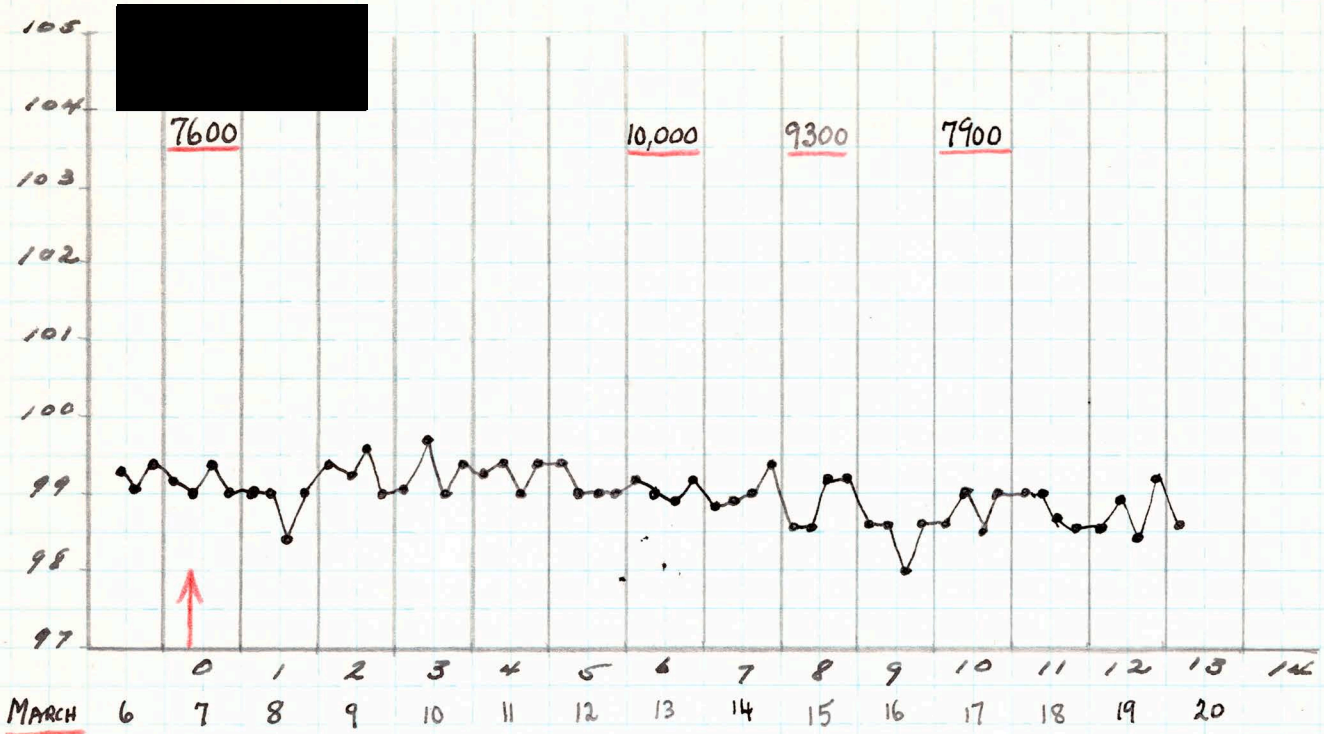


0.5cc 5th PASSAGE VIRUS INTRACUTANEOUSLY

SICILIAN VIRUS

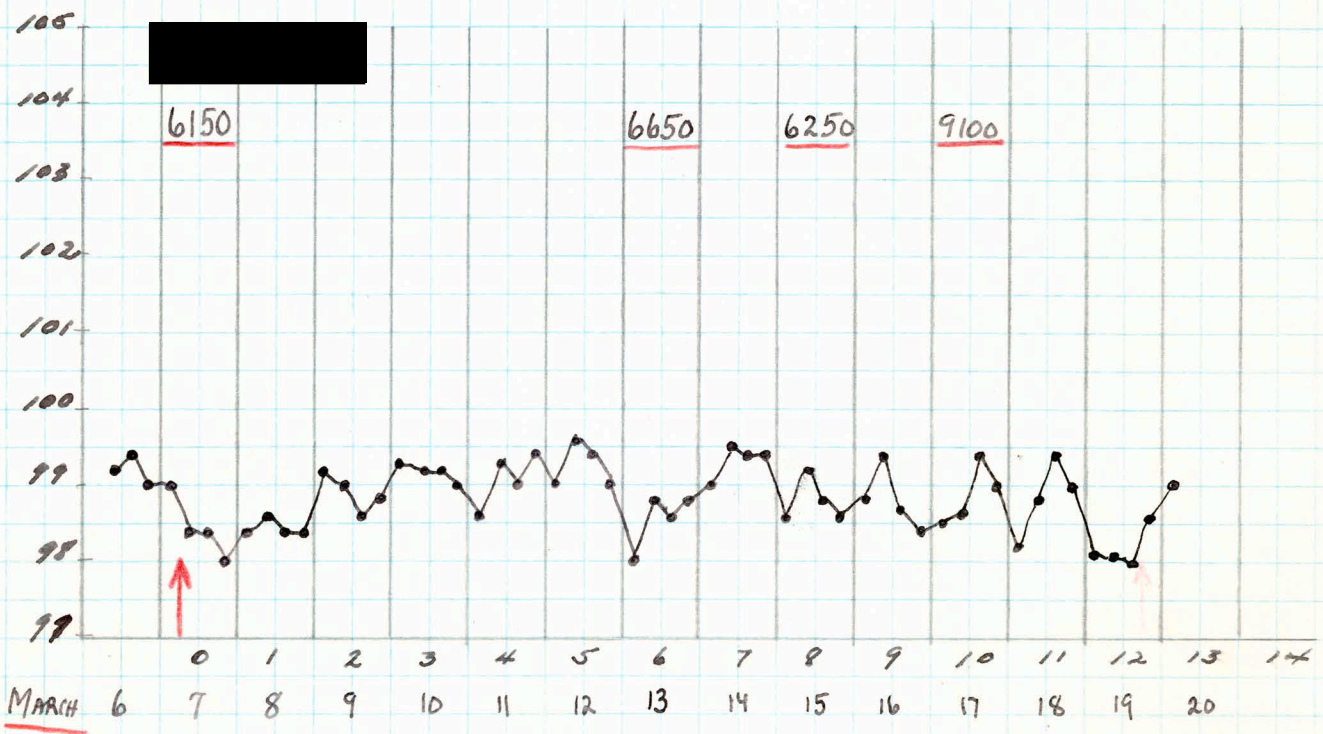
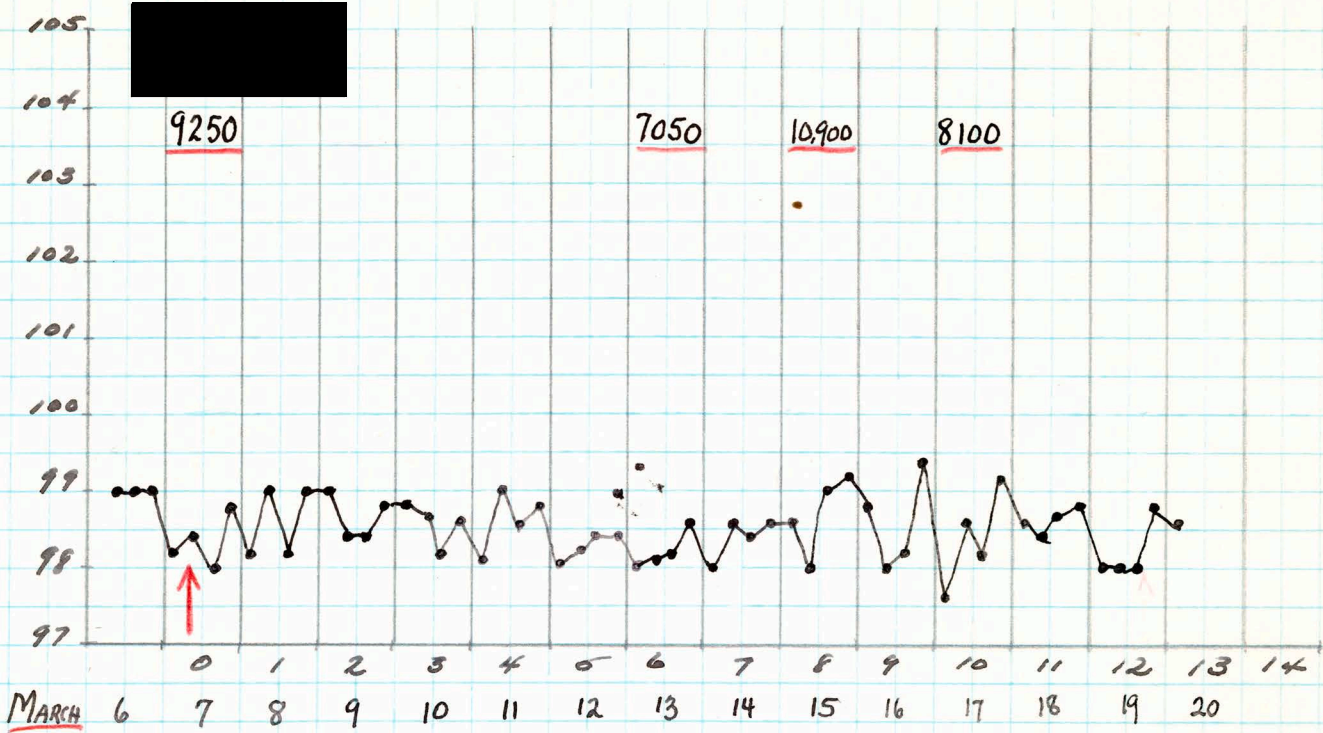
Sicilian Virus Attack 4 months ago

1944



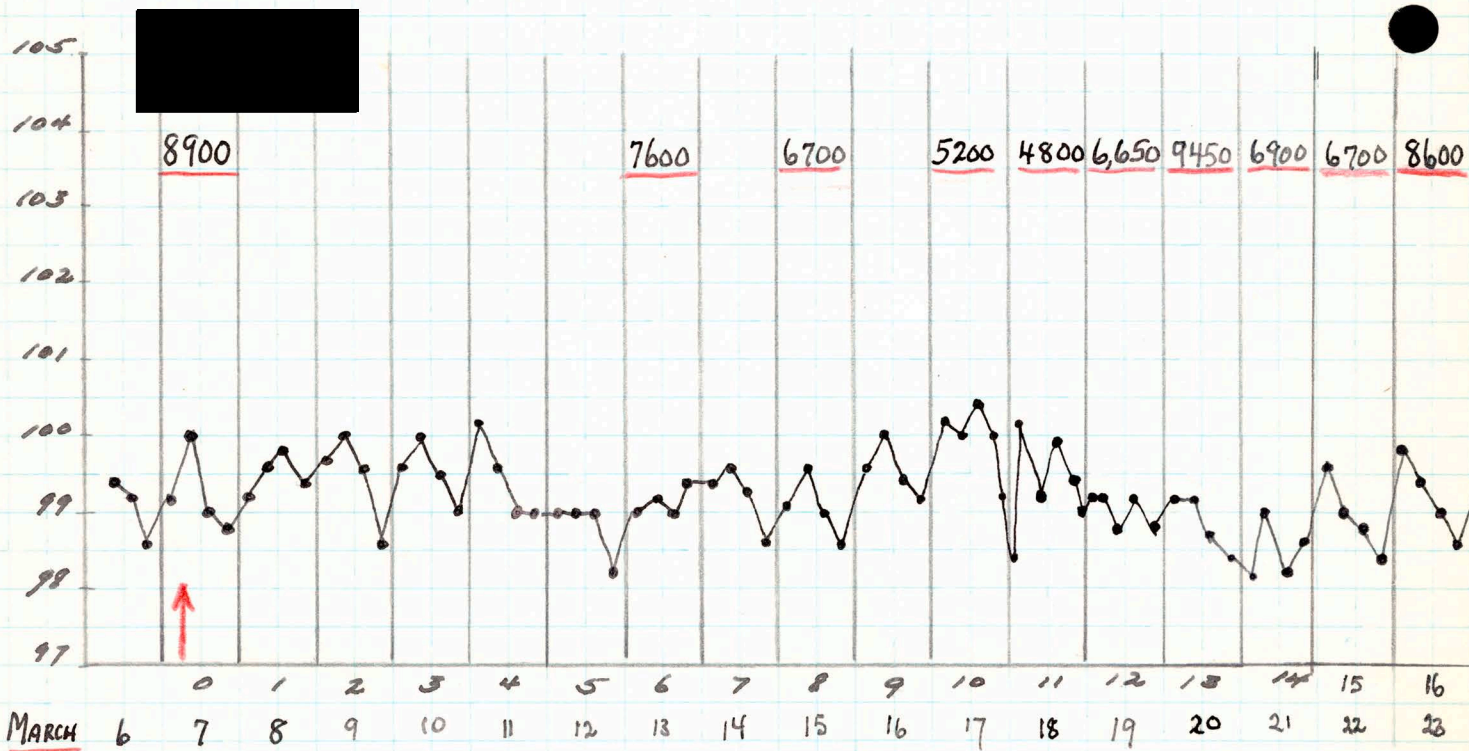
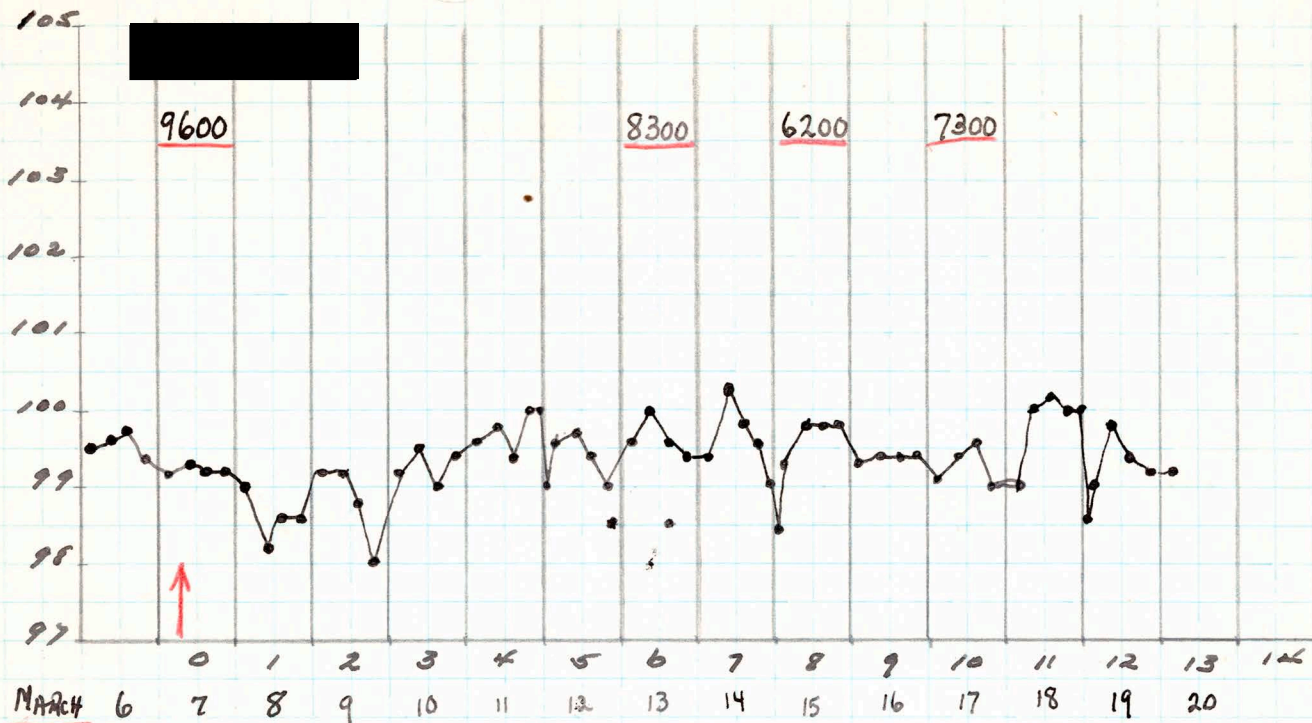
1cc 5TH PASSAGE VIRUS INTRACUTANEOUSLY

SICILIAN VIRUS - SICILIAN VIRUS ATTACK 4 months ago



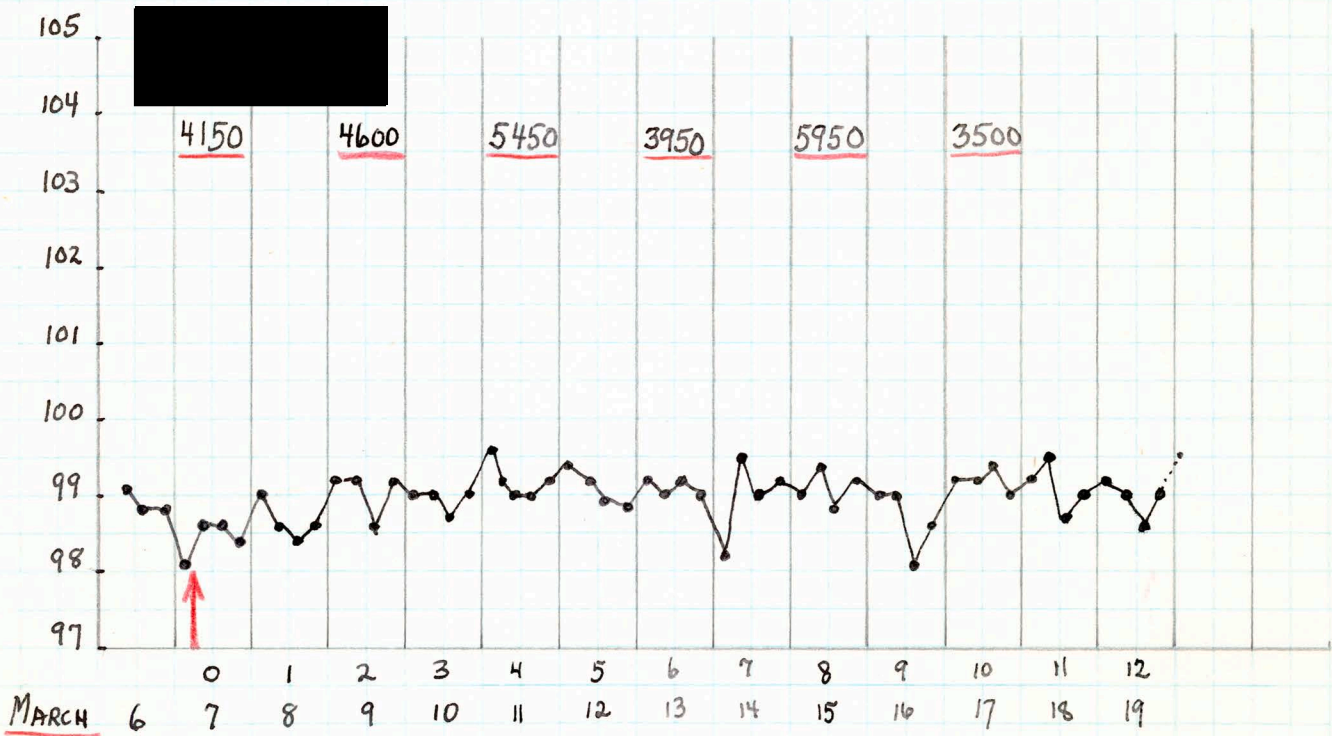
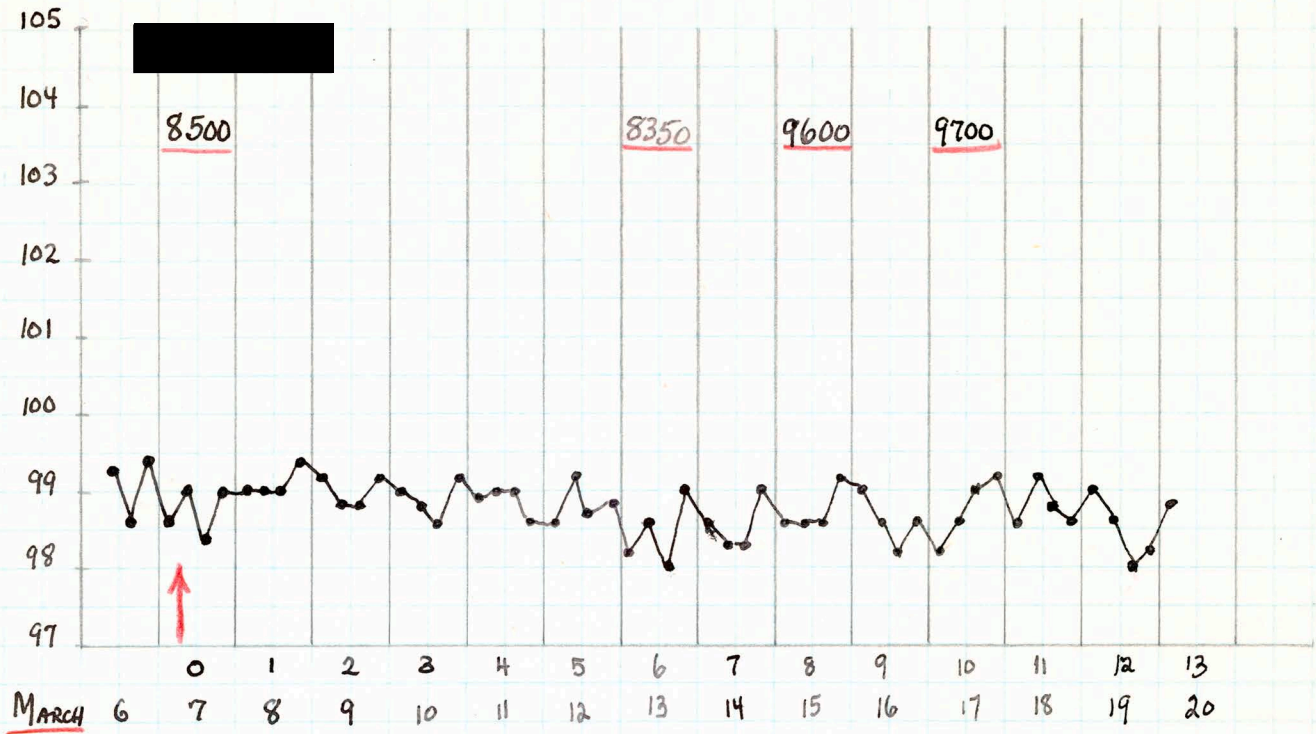
1cc 5TH PASSAGE VIRUS INTRACUTANEOUSLY

SICILIAN VIRUS - SICILIAN VIRUS ATTACK 4 MONTHS AGO



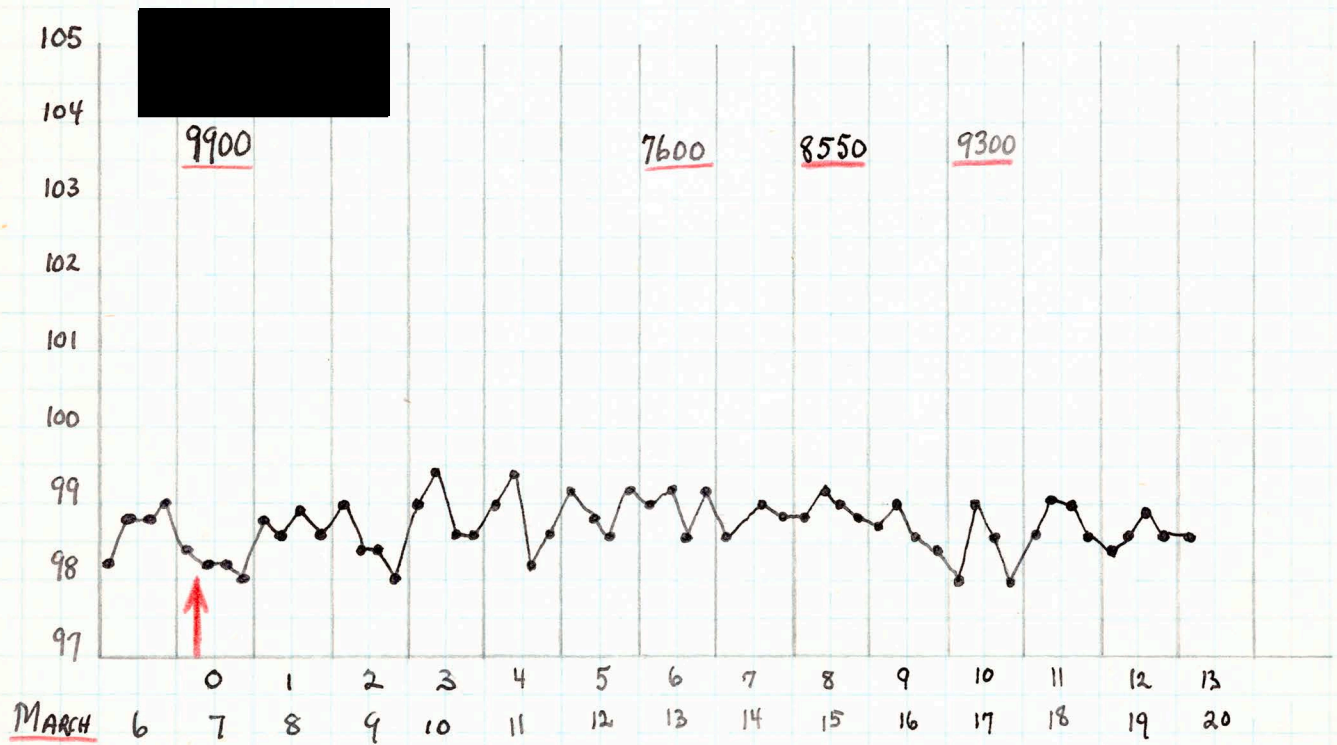
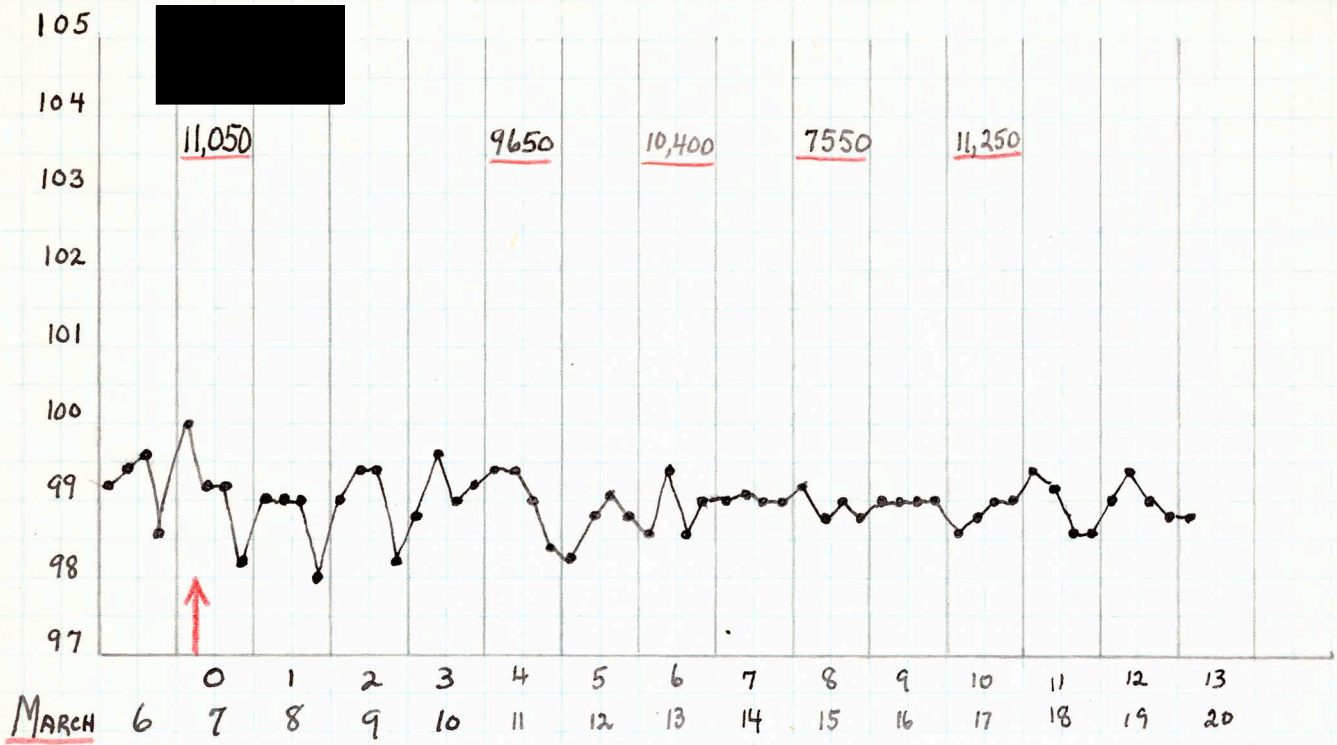
1 cc. 5TH PASSAGE VIRUS INTRACUTANEOUSLY

SICILIAN VIRUS - MIDDLE EAST VIRUS ATTACK 4 months ago



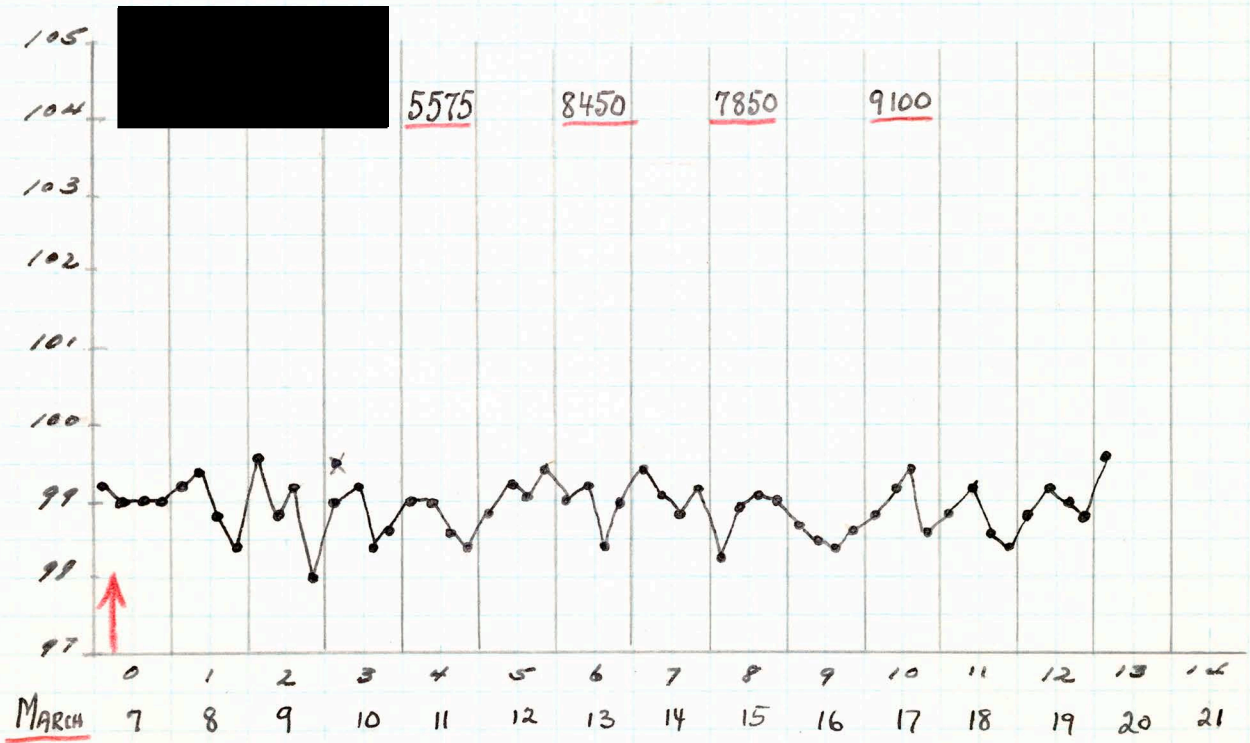
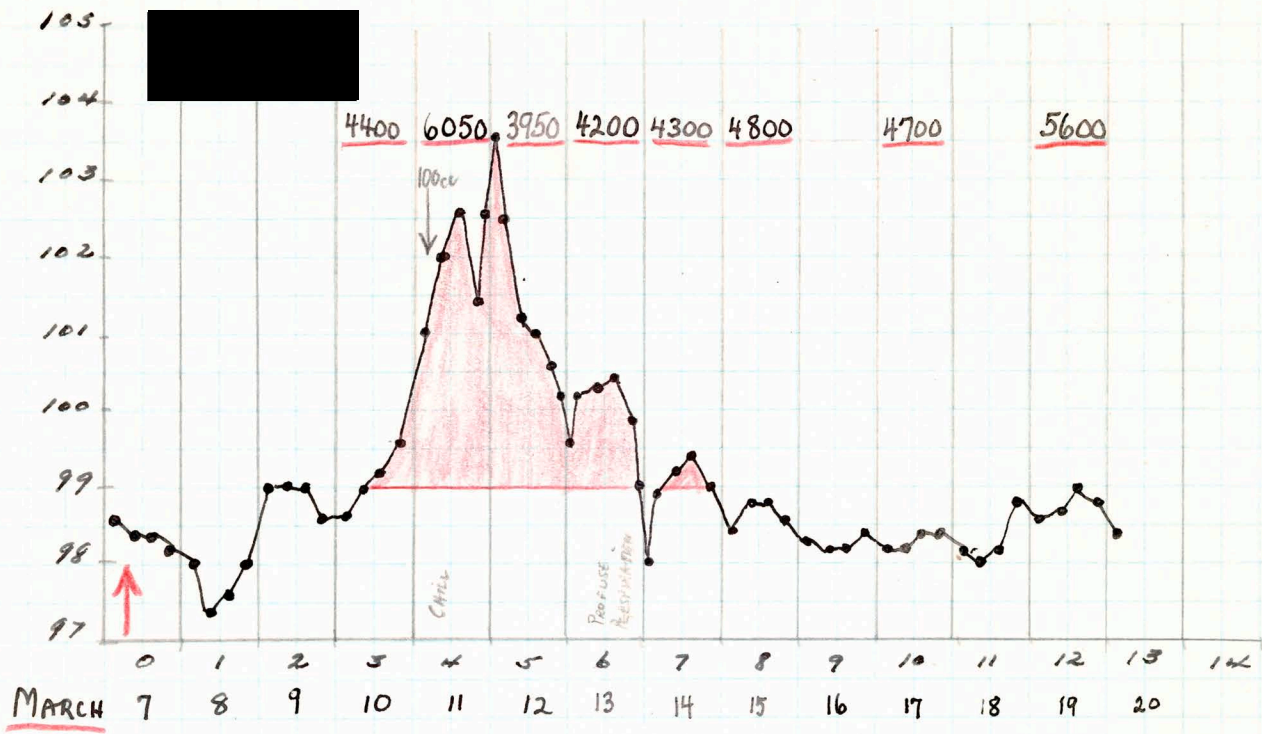
1cc 5TH PASSAGE VIRUS INTRACUTANEOUSLY

SICILIAN VIRUS - MIDDLE EAST VIRUS ATTACK 4 MONTHS AGO



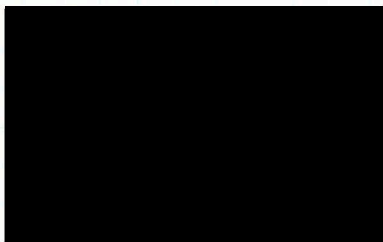
1cc 5TH PASSAGE VIRUS INTRACUTANEOUSLY

SICILIAN VIRUS - 101mm FILTRATE 2 weeks ago



1cc 5TH PASSAGE VIRUS INTRACUTANEOUSLY

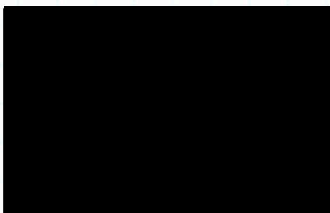
1. IMMUNITY OF SUBJECTS WHO DID NOT DEVELOP DISEASE FOLLOWING INOCULATION OF ULTRAVIOLET IRRADIATED VIRUS



1cc irradiated virus on Feb. 15, 1944

1cc " " on JAN. 24, 1944

2. IMMUNITY OF SUBJECTS WHO DID NOT DEVELOP DISEASE FOLLOWING INOCULATION OF CONVALESCENT SERUM - VIRUS MIXTURE

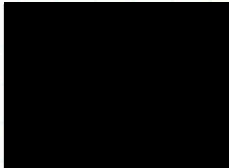
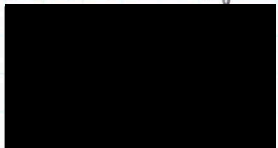


CONVALESCENT SERUM - VIRUS ON FEB. 28, 1944

" " " " DEC. 20, 1943

3. TEST FOR PRESENCE OF VIRUS IN SERUM OF PATIENT WHO SUCCEMPTED FOLLOWING INOCULATION WITH 207 μ l FILTRATE

THIS SERUM - from [redacted] - USED AS VIRUS IN THESE TESTS. CONTROL SUBJECTS HAD PREVIOUSLY BEEN INOCULATED WITH 1:10,000 (SLOAN) and 1:100,000 (ROSS) dilutions of virus ON FEB. 28, 1944

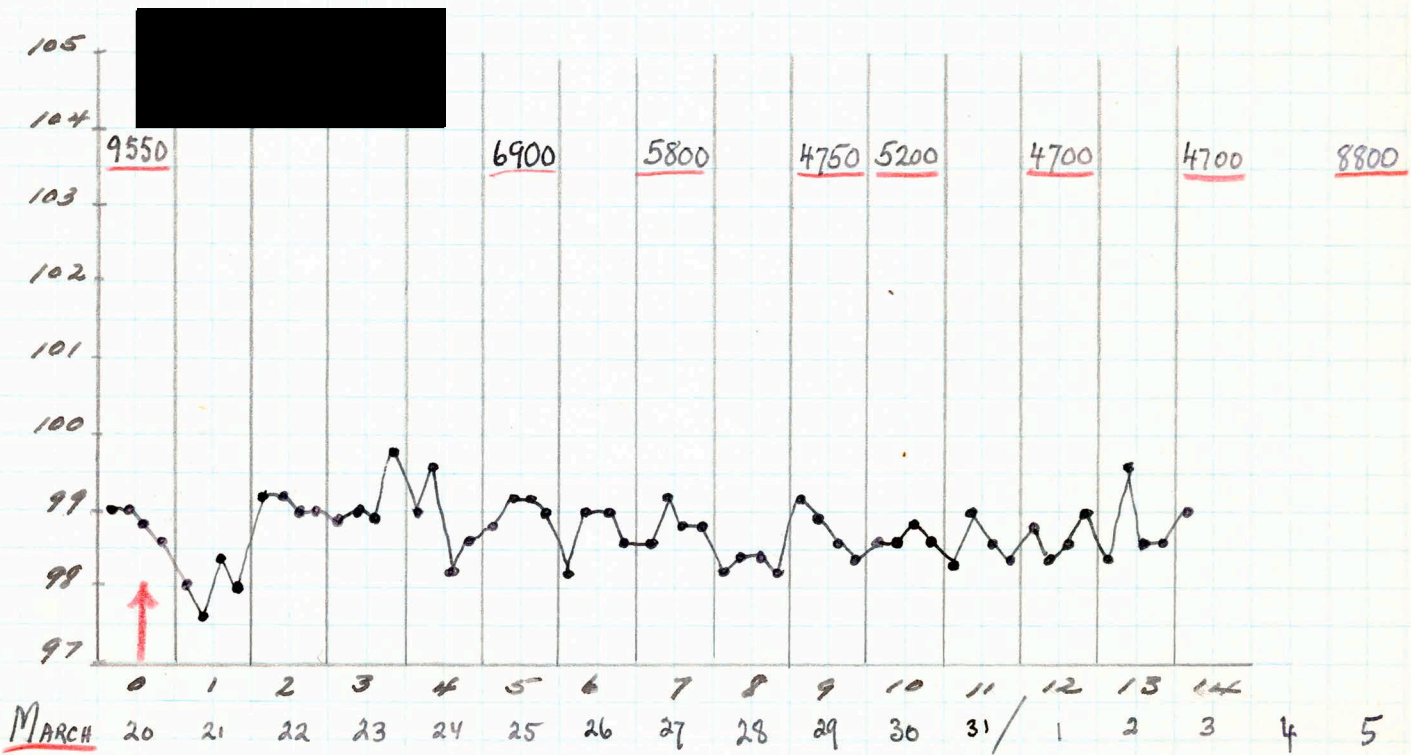
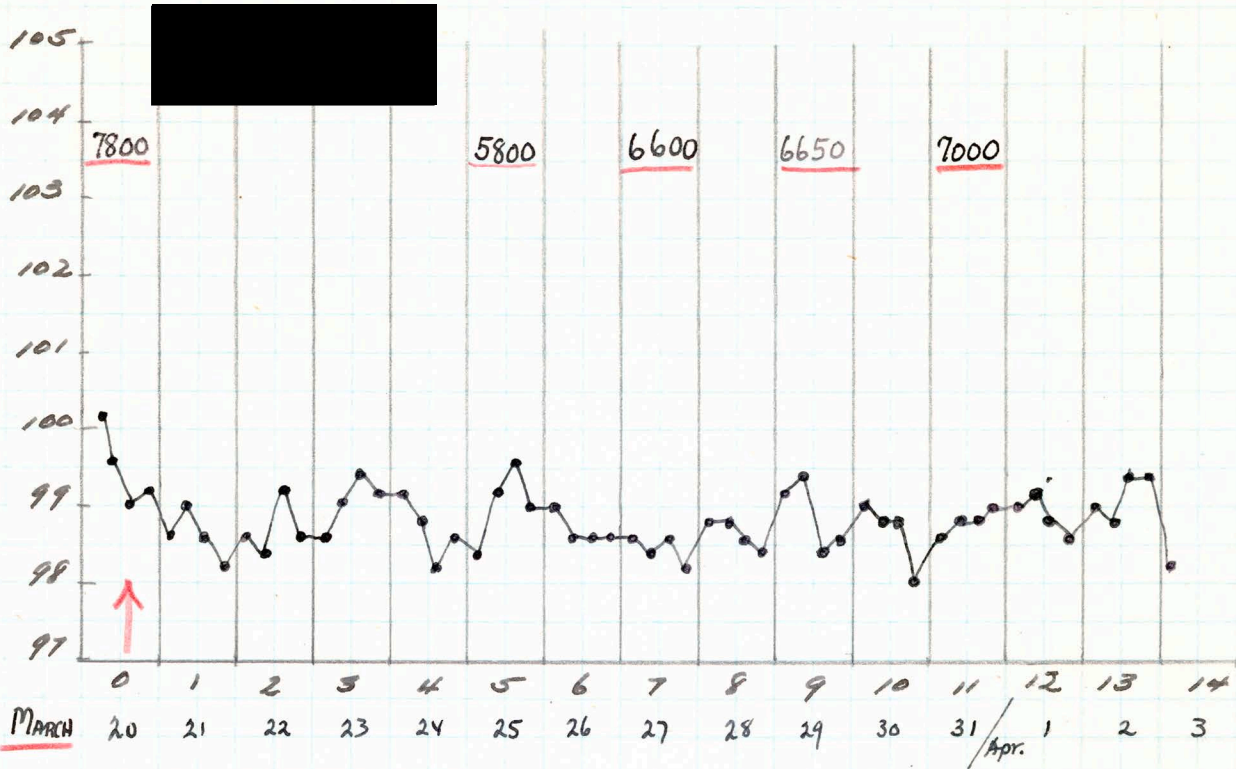


VIRUS - ALL SUBJECTS INOCULATED WITH 1cc OF [redacted] SERUM - 0.2cc i.cut. in each of 5 sites 2 P.M.

3-21-44 - No reaction at any of inoculated sites

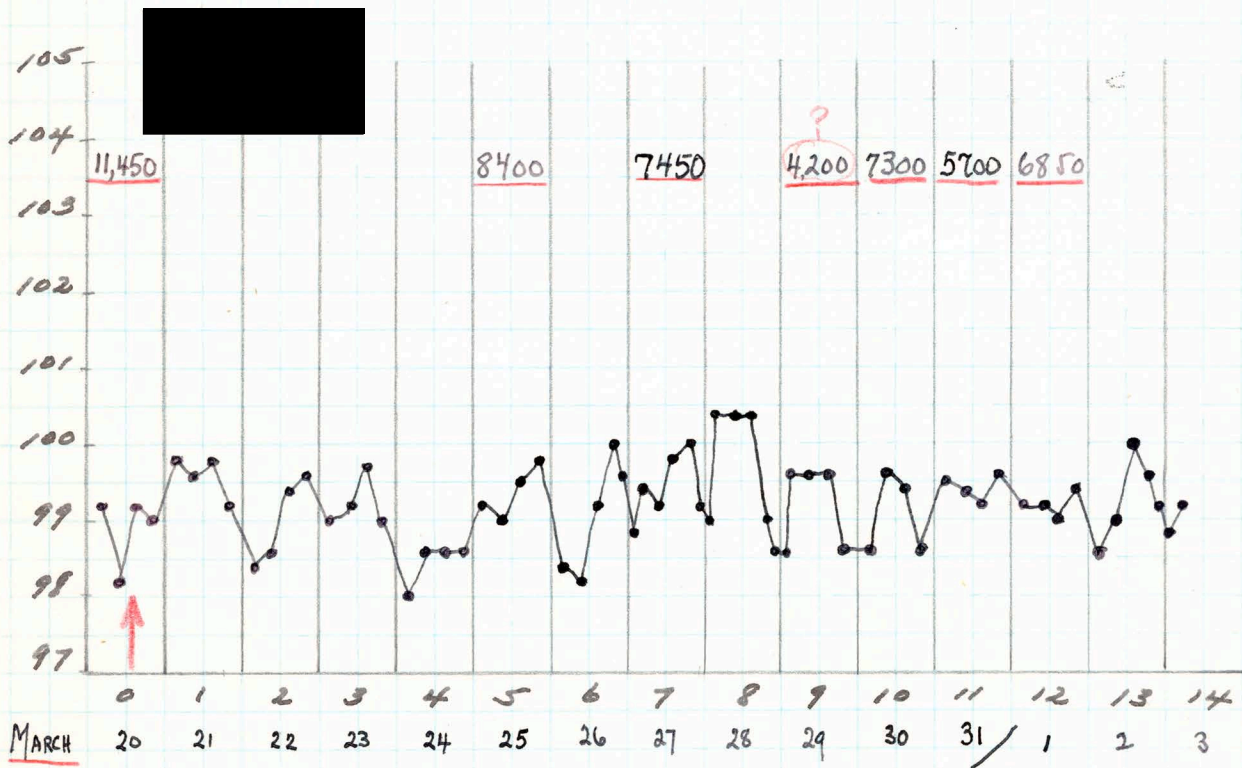
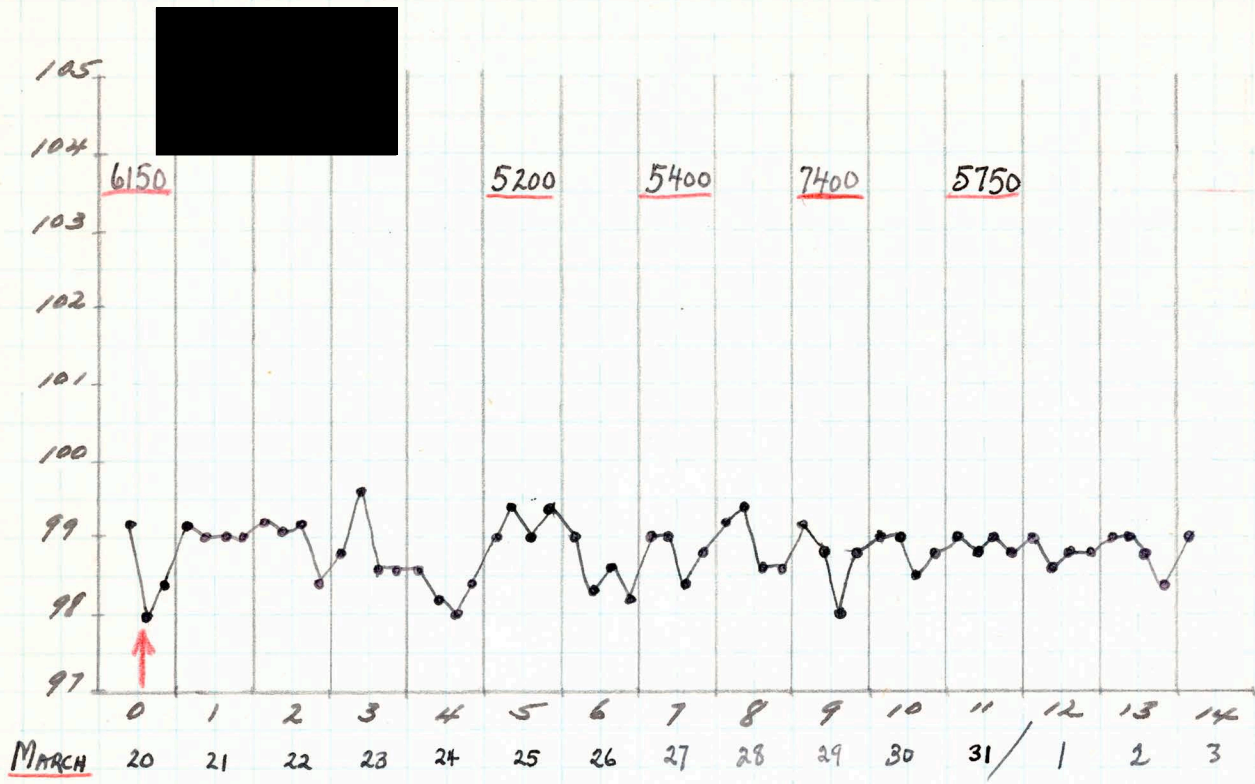
SICILIAN VIRUS -

TEST FOR IMMUNITY FOLLOWING
ULTRAVIOLET IRRADIATED VIRUS ON FEB. 15, 1944

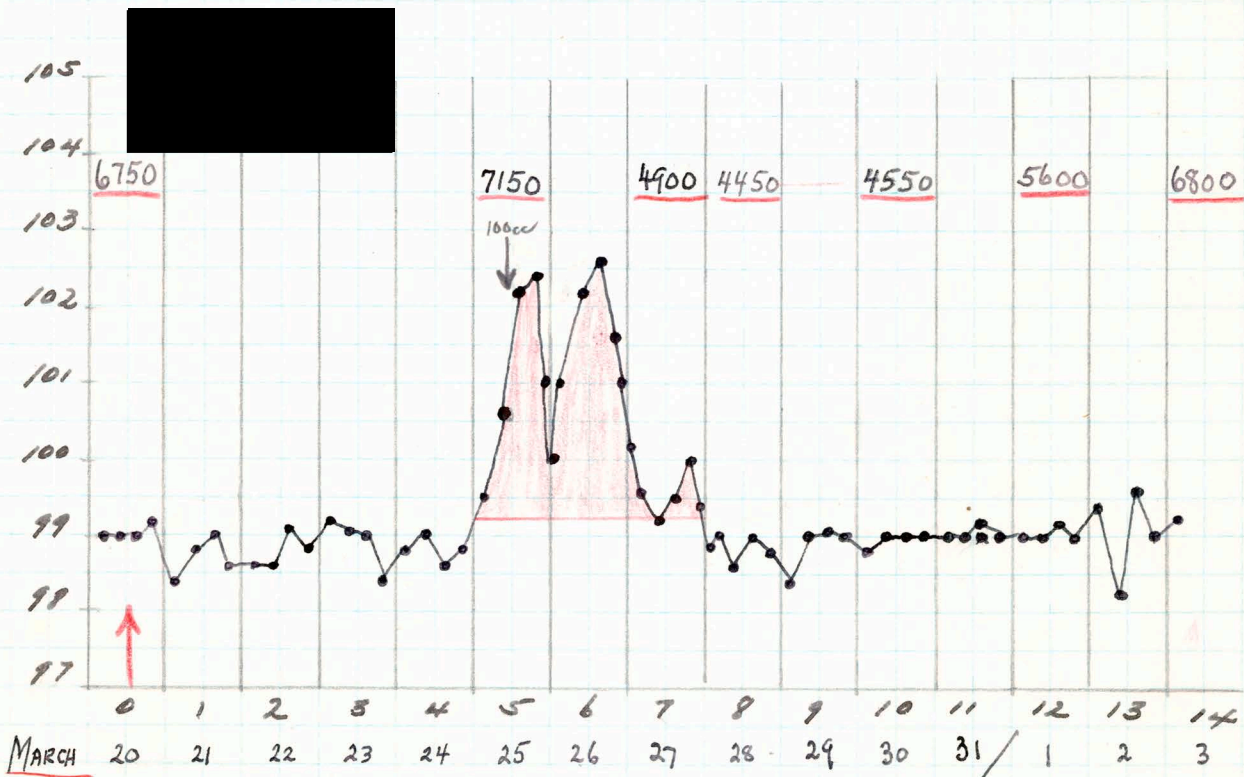
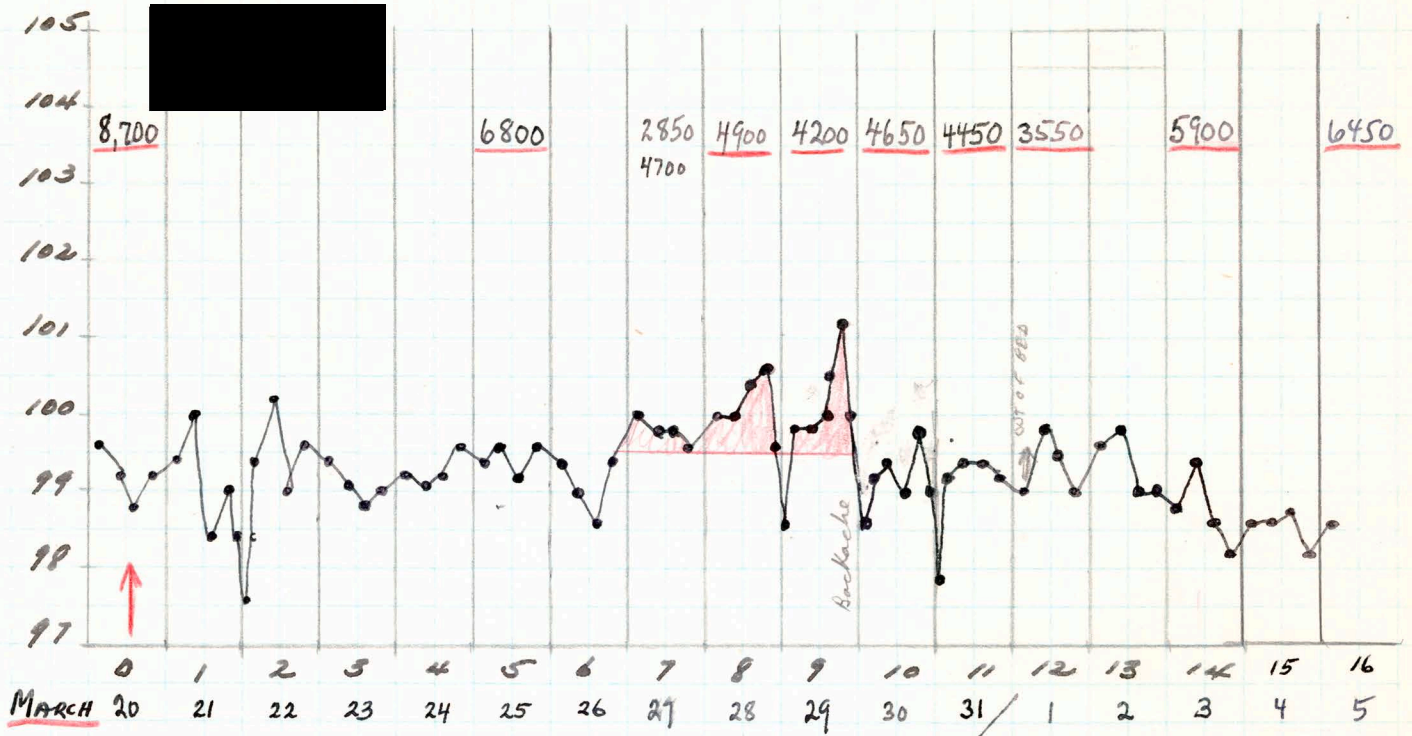


SICILIAN VIRUS -

TEST FOR IMMUNITY FOLLOWING
ULTRAVIOLET IRRADIATED VIRUS ON JAN. 24, 1944

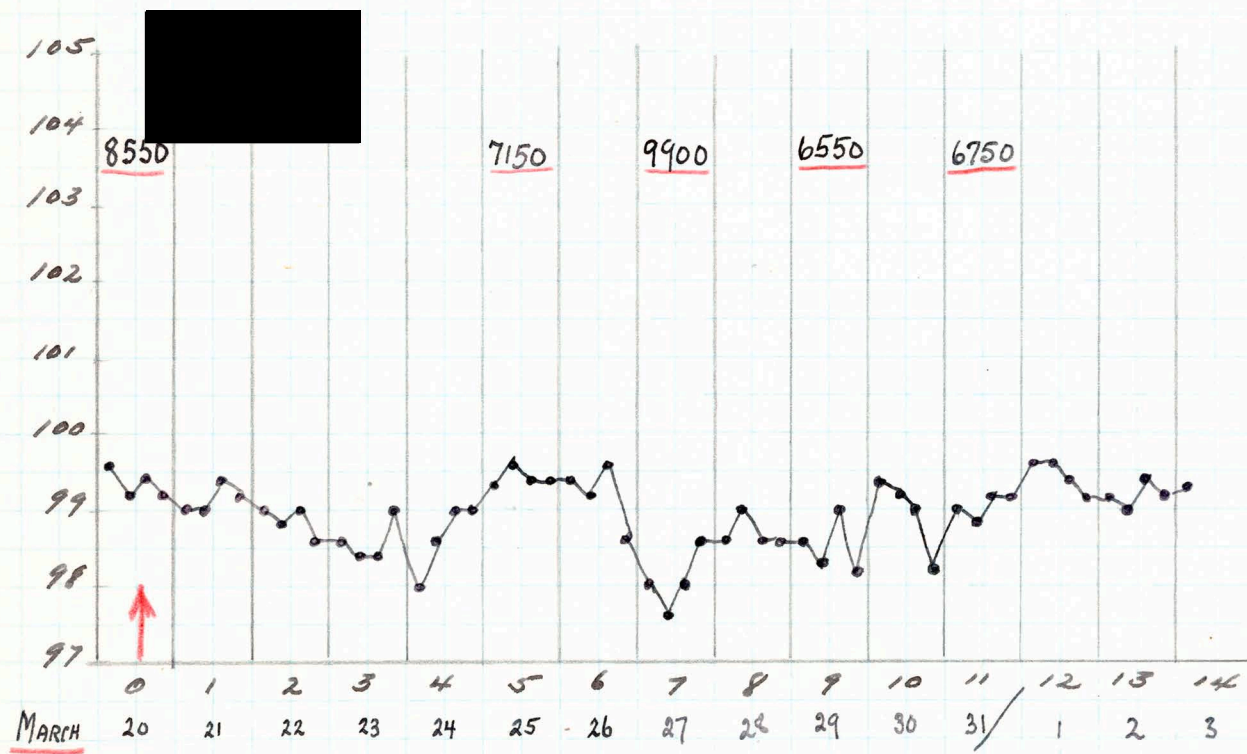
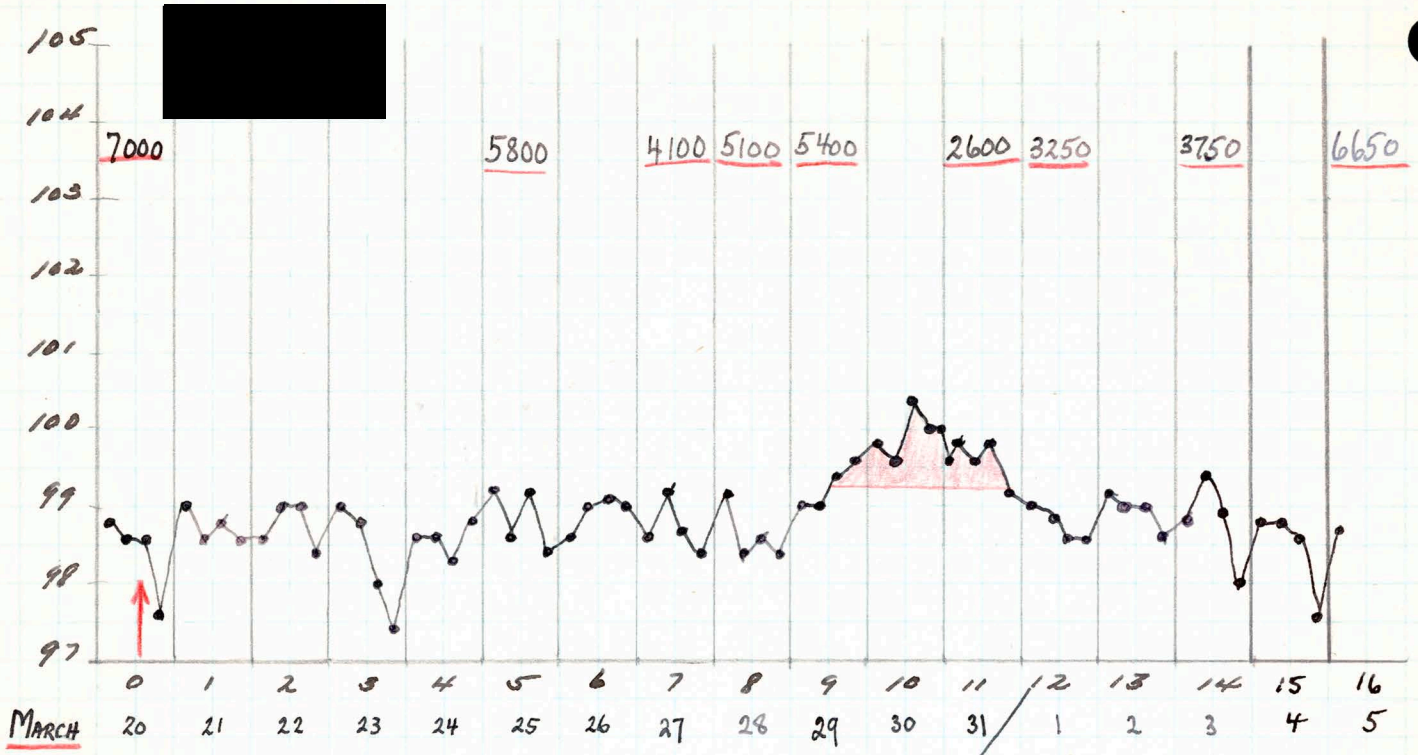


SICILIAN VIRUS - CONTROLS FOR IMMUNITY TESTS



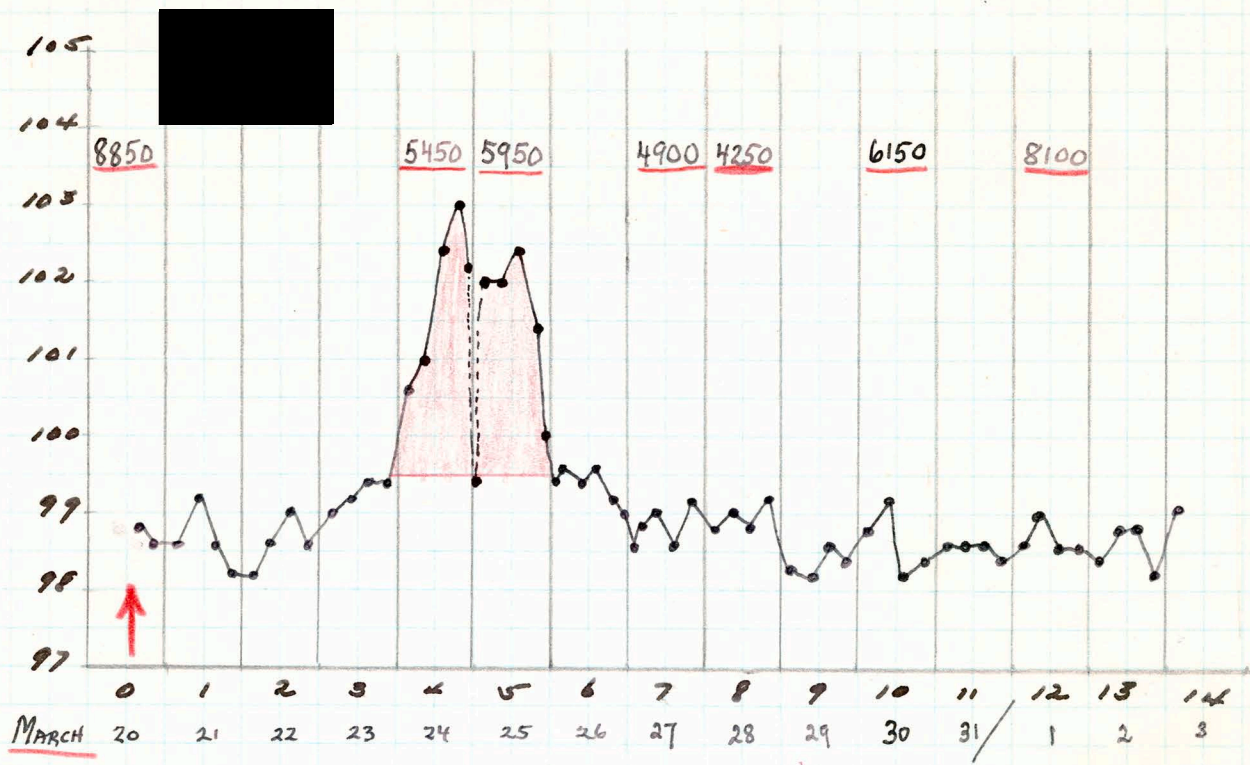
SICILIAN VIRUS -

TEST FOR IMMUNITY FOLLOWING
CONVALESCENT SERUM-VIRUS MIXTURE ON FEB 28 '44



SICILIAN VIRUS -

TEST FOR IMMUNITY FOLLOWING
CONVALESCENT SERUM-VIRUS MIXTURE ON DEC. 20, 1943



PATIENT	DATE	PROCEDURE	CEPH Floc.
[REDACTED]	2-2	INOCULATED P. FALCIPARUM	
	2-4		0
	2-6	THICK SMEAR POSITIVE	0
	2-8	FIRST FEVER. CINCHONINE STARTED	#
	2-10		##
	2-11	THICK SMEAR NEGATIVE AFEBRILE	0
	2-12		0
	2-14		#
	2-17		#
	2-20		#
	2-29		0
	3-7		0
	[REDACTED]	2-8	INOCULATED P. FALCIPARUM
2-10			0
2-11			+
2-12		THICK SMEAR POSITIVE	
2-14			#
2-16		FIRST FEVER. THERAPY STARTED	#
2-20		AFEBRILE. THICK SMEAR NEG	
2-22			#
2-23			##
3-1		##	
[REDACTED]	2-8	INOCULATED P. FALCIPARUM	
	2-11		#
	2-13	THICK SMEAR POSITIVE	
	2-14	FIRST FEVER	
	2-15	THERAPY STARTED	##
	2-17	AFEBRILE	+
	2-19	THICK SMEAR NEG	
	2-20		##
	2-22		#
	2-23		##
	3-1		##

PATIENT	DATE	PROCEDURE	CEPH FLOC
[REDACTED]	2-14	INOCULATED P. FALCIPARUM	
	2-19		+
	2-22		0
	2-23	THICK SMEAR POSITIVE	+
	2-24	FIRST FEVER. THERAPY STARTED	
	2-25		+
	2-28	AFEBRILE	+
	2-29	THICK SMEAR NEGATIVE	
3-6		0	
[REDACTED]	2-17	INOCULATED P. FALCIPARUM	
	2-19		0
	2-21	THICK SMEAR POSITIVE	
	2-22	FIRST FEVER	0
	2-23	THERAPY STARTED	0
	2-24		+
	2-25		0
	2-27	AFEBRILE AND THICK SMEAR NEG	
	2-28		0
3-6		++	
[REDACTED]	2-17	INOCULATED P. FALCIPARUM	
	2-19		0
	2-22	THICK SMEAR POSITIVE	0
	2-23		0
	2-24	FIRST FEVER	
	2-27	LARGE DOSES OF ATABRINE FOR FULMINATING INFECTION	
	2-28		+
	3-6		+++

PATIENT DATE PROCEDURE GEPA FLOC.

[REDACTED]	1-27	INOCULATED P.VIVAX	
	2-3	THICK SMEAR POSITIVE. FIRST FEVER	○
	2-6	THERAPY STARTED	
	2-8	A FEBRILE	○
	2-11	THICK SMEAR NEG	
	2-13		○
	2-15		○
	2-18	THICK SMEAR POSITIVE AND RECURRENCE OF FEVER	○
	2-22		+++
	2-25	ATABRINE STARTED	
	2-26		+++
	2-29		+++
	3-7		+++

[REDACTED]	1-31	INOCULATED P.VIVAX	
	2-4	THICK SMEAR POSITIVE	
	2-6	FIRST FEVER	
	2-7		○
	2-9	THERAPY STARTED FEVER NOT INTERRUPTED BY R	+++
	2-10		+++
	2-13		+++
	2-18		+++
	2-22		+++

[REDACTED]	2-18	INOCULATED P.VIVAX	
	2-19		○
	2-22		○
	2-23	THICK SMEAR POSITIVE	● ○
	2-24	FIRST FEVER	
	2-25		○
	2-26		+
	← 2-27	THERAPY STARTED	+++
	2-28		+++
	3-2		+++

PATIENT	DATE	PROCEDURE	CEPH FLOC.
[REDACTED]	2-31	INOCULATED P. VIVAX	
[REDACTED]	2-4	THICK SMEAR POSITIVE AND FIRST FEVER	
[REDACTED]	2-7	THERAPY STARTED	0
[REDACTED]	2-9	THICK	+++
[REDACTED]	2-10	THICK SMEAR NEGATIVE	
[REDACTED]	2-11	A FEBRILE	+++
[REDACTED]	2-13		+++
[REDACTED]	2-17		+++
[REDACTED]	2-19		+
[REDACTED]	2-25	RE-INOCULATED	
[REDACTED]	2-29	THICK SMEAR POSITIVE RECURRENCE OF FEVER	⊕+
[REDACTED]	3-3		+++
[REDACTED]	2-26	+	+
[REDACTED]	2-27	INOCULATED P. VIVAX	
[REDACTED]	3-1		0
[REDACTED]	3-2	THICK SMEAR POSITIVE	
[REDACTED]	3-3		++
[REDACTED]	3-4		+++
[REDACTED]	3-5	FIRST FEVER	
[REDACTED]	3-6		+++

HAD SORE THROAT ON 2-8 AND WAS THOUGHT AT THE TIME TO HAVE INFECTIOUS MONONUCLEOSIS. HOWEVER SMEARS WERE NEVER TYPICAL AND SHEER CELL AGGLUTINATION WAS REPEATEDLY NEGATIVE.



Unpublished data
Third (New York University) Research Service
Goldwater Memorial Hospital

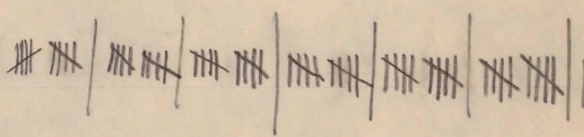


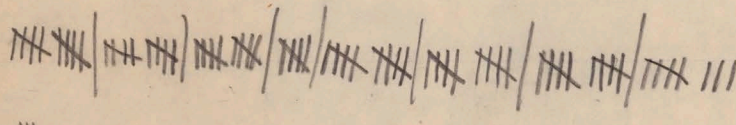
Sicilian

JAN. 7, 1944

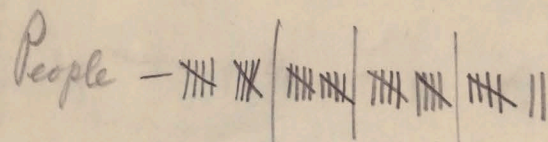
M. E.

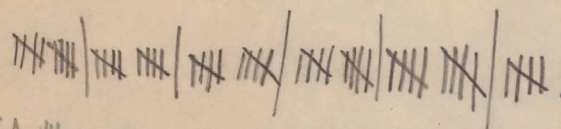
presumptuous
impertinent
insolent

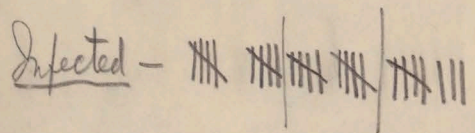
Inoculations -  = 61

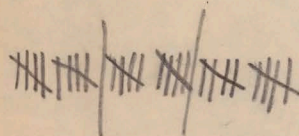
 = 73

S.A. III

People -  = 37
+ 4 (ME imm)

 = 56
S.A. III 4 Sic

Infected -  = 28

 = 30

III

Leuk. only

Total inoculated = 96

Mosquito subj. = 6

137 inoculations

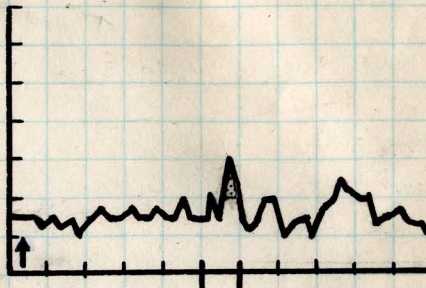
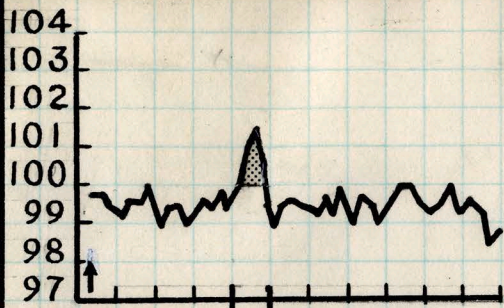
58 exp. S.F. exd. M.E. up to Feb. 18 35

137

VARIOUS TYPES OF FEVER PRODUCED BY EXPERIMENTAL INOCULATION
OF PHLEBOTOMUS FEVER VIRUS IN HUMAN BEINGS

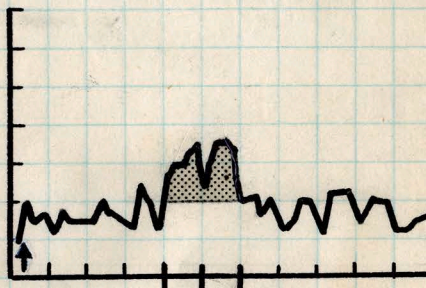
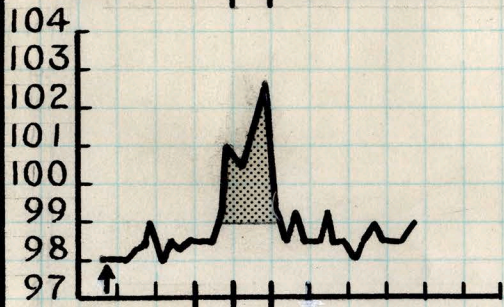
1 DAY

6%



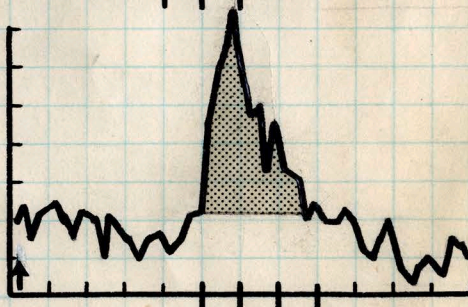
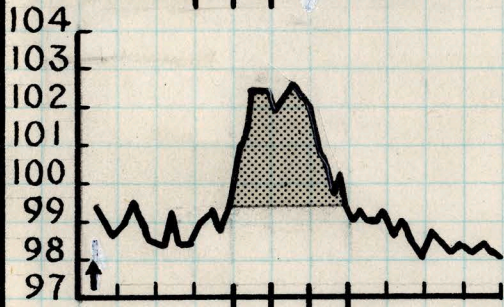
2 DAY

22%



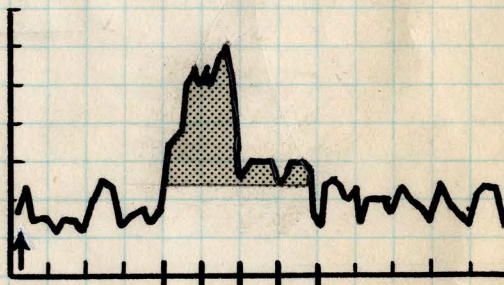
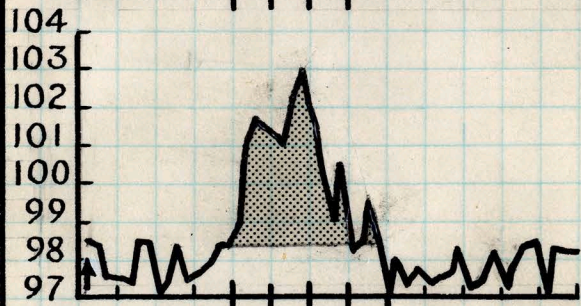
3 DAY

43%



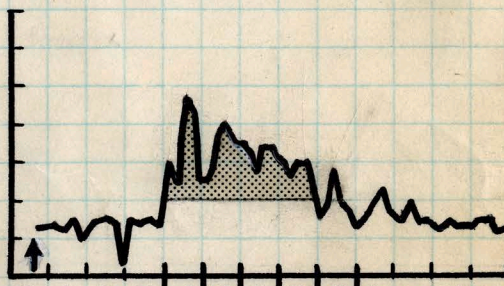
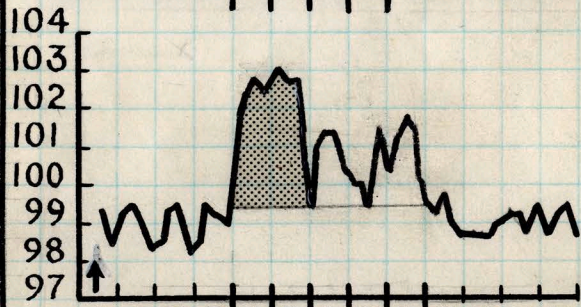
4 DAY

20%



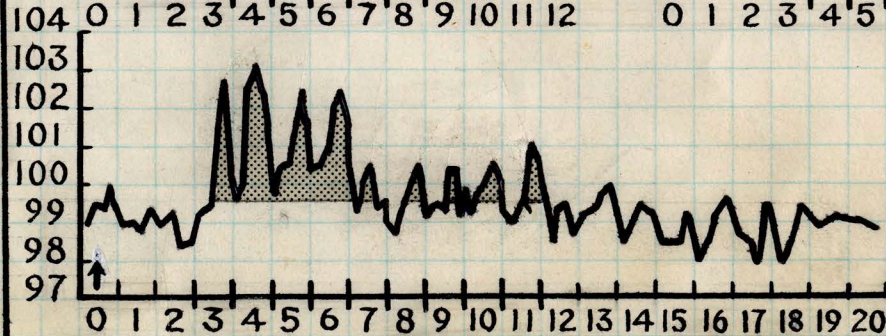
5 DAY

4%

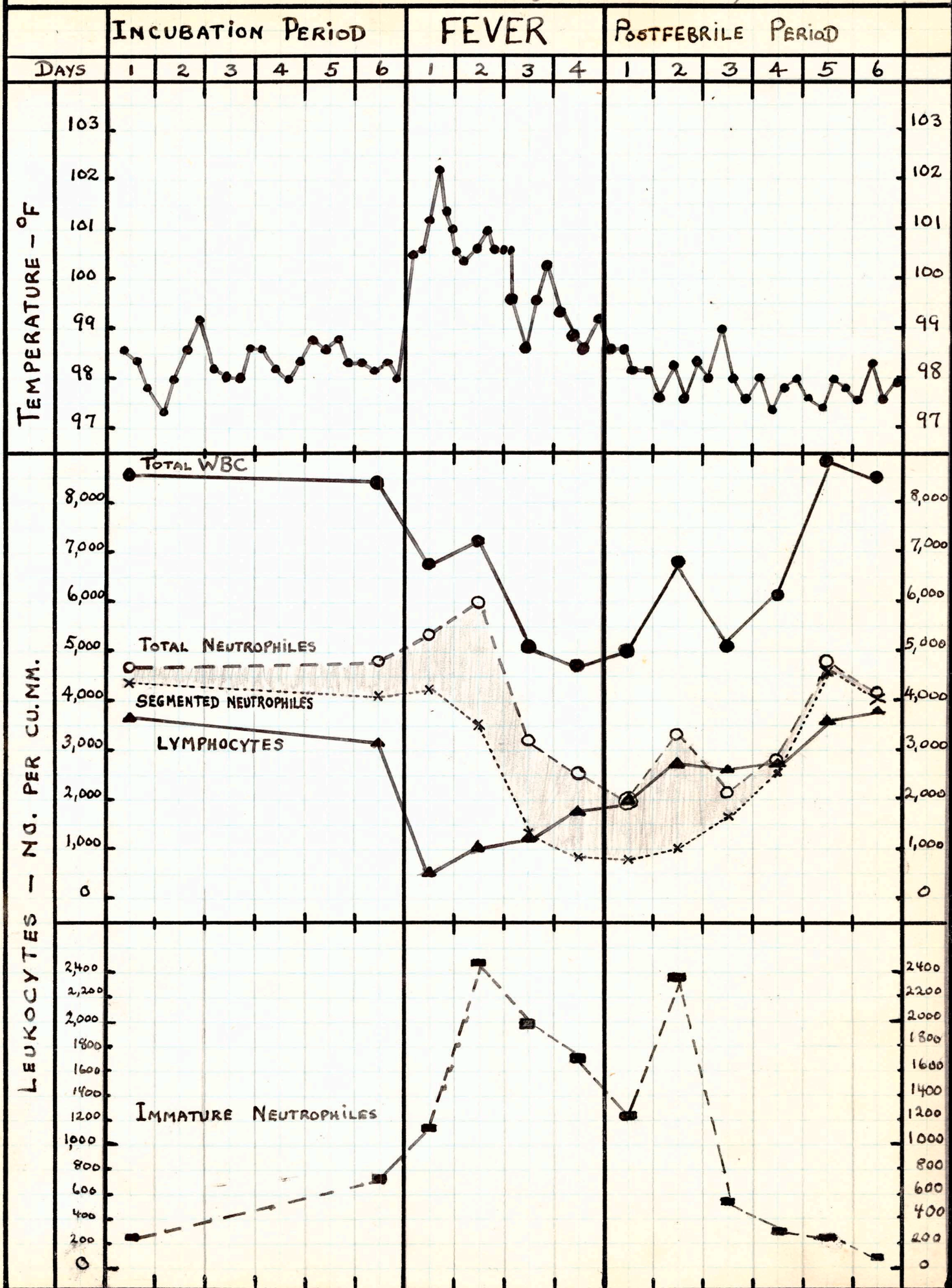


9 DAY

2%



EXPERIMENTAL SANDFLY FEVER - F. GR. ^{SPLEBOTOMUS} COLORED MALE, - AGE 50

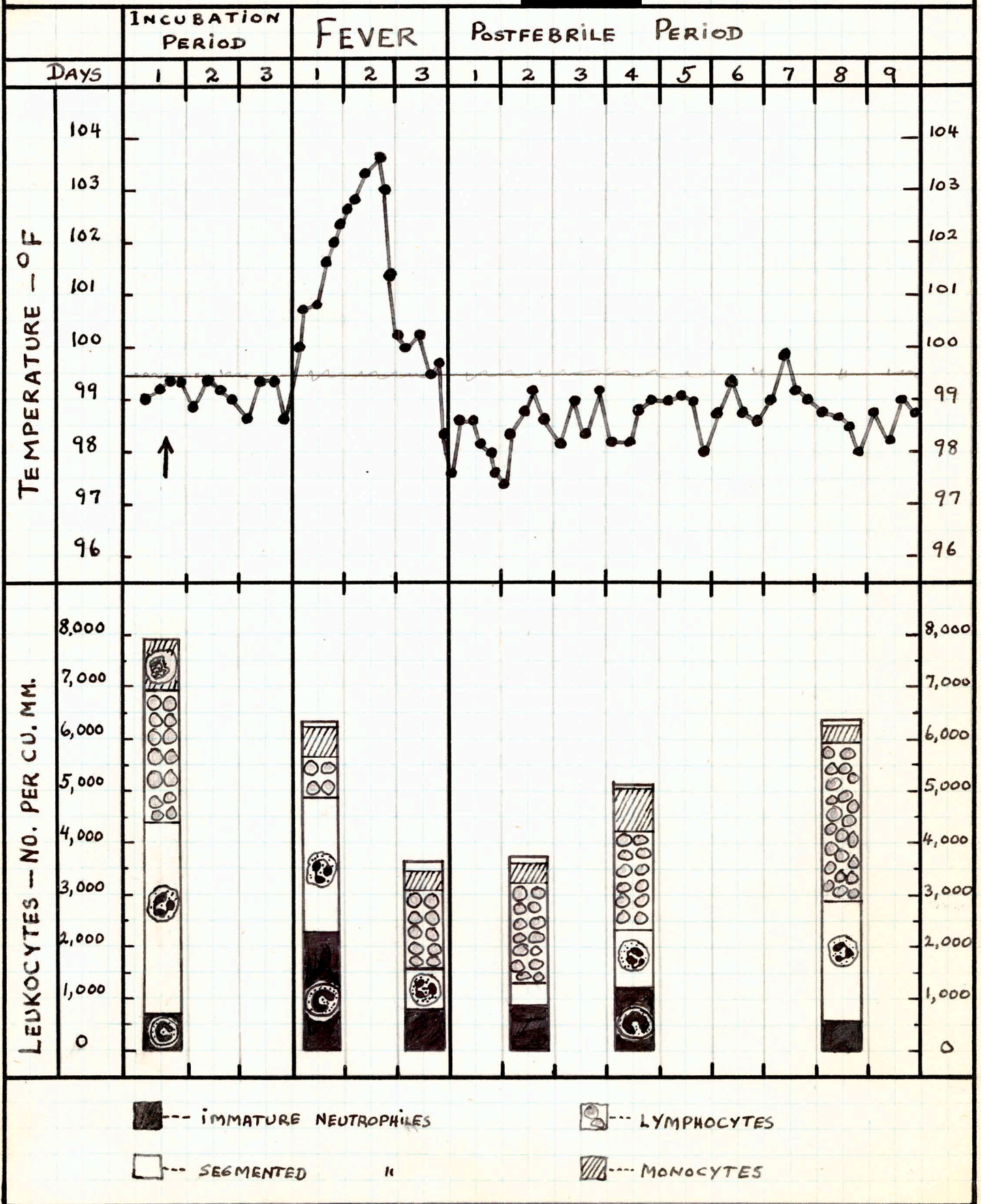


	12/8/13	12/13	12/14	12/15	12/16	12/17	12/18	12/19	12/20
Total WBC	8,550	8,400	6,850	7,250	5,100	4,800	5,000	6,800	5,100
" neutrophils	4,702 (55%)	4,872 (58%)	5411 (79%)	6,017 (83%)	3315 (65%)	2592 (54%)	2,050 (41%)	3400 (50%)	2193 (43%)
" lymphocytes	3,676 (43%)	3,192 (38%)	548 (8%)	1,015 (14%)	1275 (25%)	1776 (37%)	2,050 (41%)	2720 (40%)	2601 (51%)
" immature neutrophils	256 (3%)	756 (9%)	1164 (17%)	2465 (34%)	1989 (39%)	1728 (36%)	1,250 (25%)	2380 (35%)	561 (11%)
" monocytes	171 (2%)	168 (2%)	753 (11%)	217 (3%)	510 (10%)	384 (8%)	550 (11%)	612 (9%)	153 (3%)
Segmented neutrophils	4446 (52%)	4116 (49%)	4247 (62%)	3552 (49%)	1326 (26%)	864 (18%)	800 (16%)	1020 (15%)	1632 (32%)

	12/21	12/22
Total WBC	6,100	8,950
" neutrophils	2828 (48%)	4928 (55%)
" lymphocytes	2745 (45%)	3,580 (40%)
" immature neutrophils	305 (5%)	268 (3%)
" monocytes	244 (4%)	90 (1%)
Segmented neutrophils	2623 (43%)	4654 (52%)

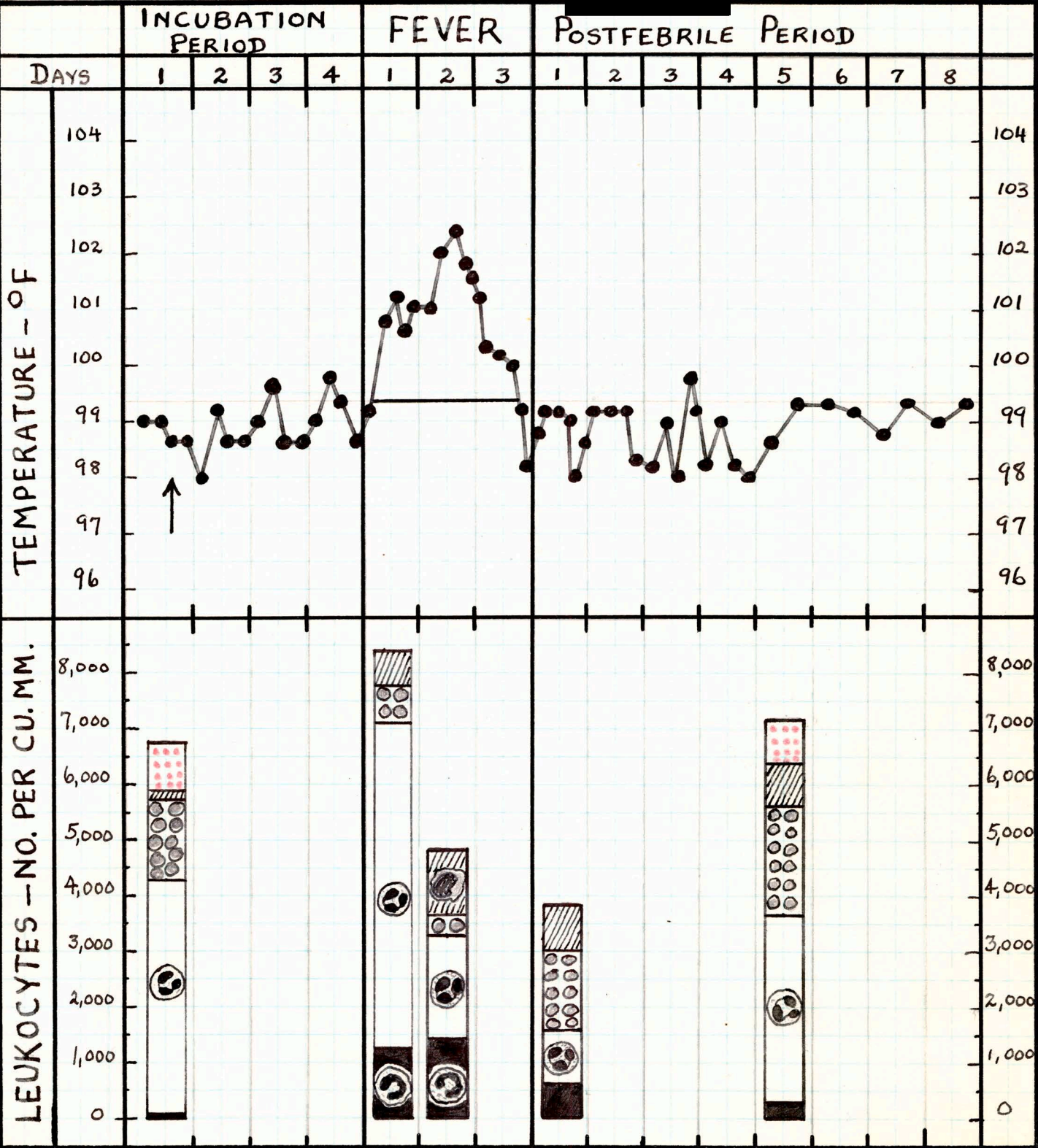
PHLEBOTOMUS

EXPERIMENTAL SANDFLY FEVER - [REDACTED] WHITE MALE, AGE 18



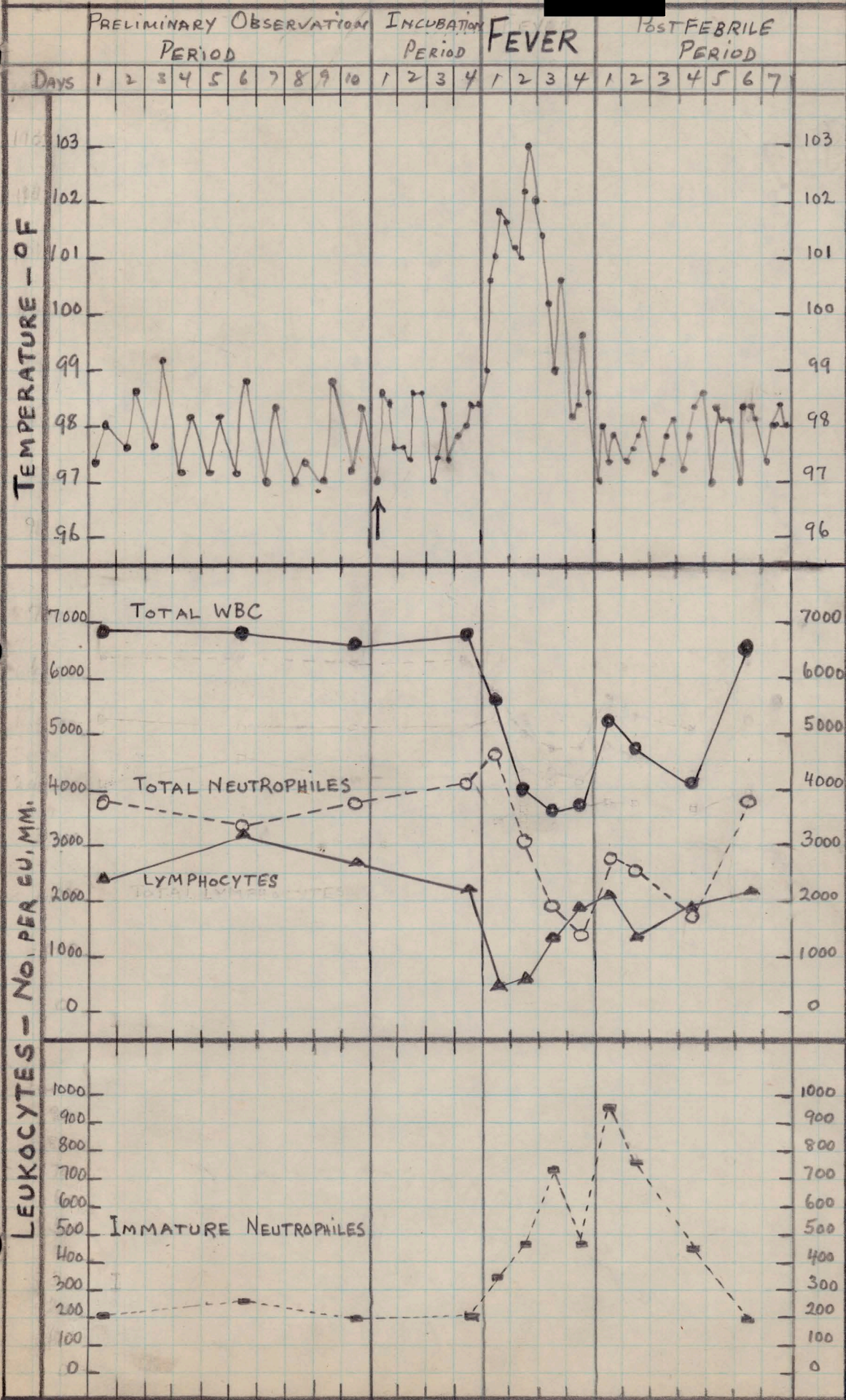
A.O.

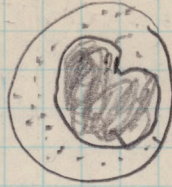
EXPERIMENTAL SANDFLY FEVER [REDACTED] WHITE FEMALE, AGE 45



■ ... IMMATURE NEUTROPHILES ◯ ... LYMPHOCYTES
 □ ... SEGMENTED " ▨ ... MONOCYTES ◻ ... EOSINOPHILES

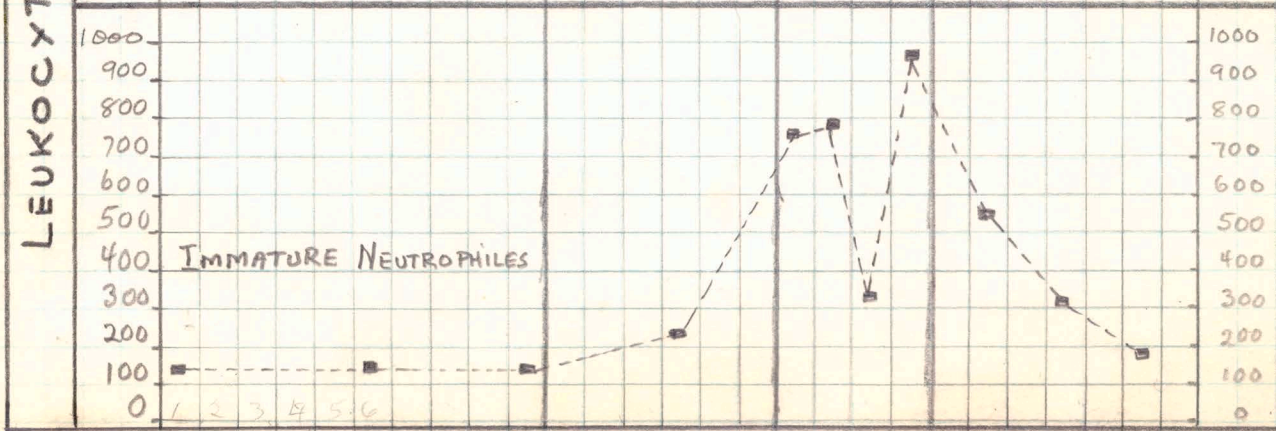
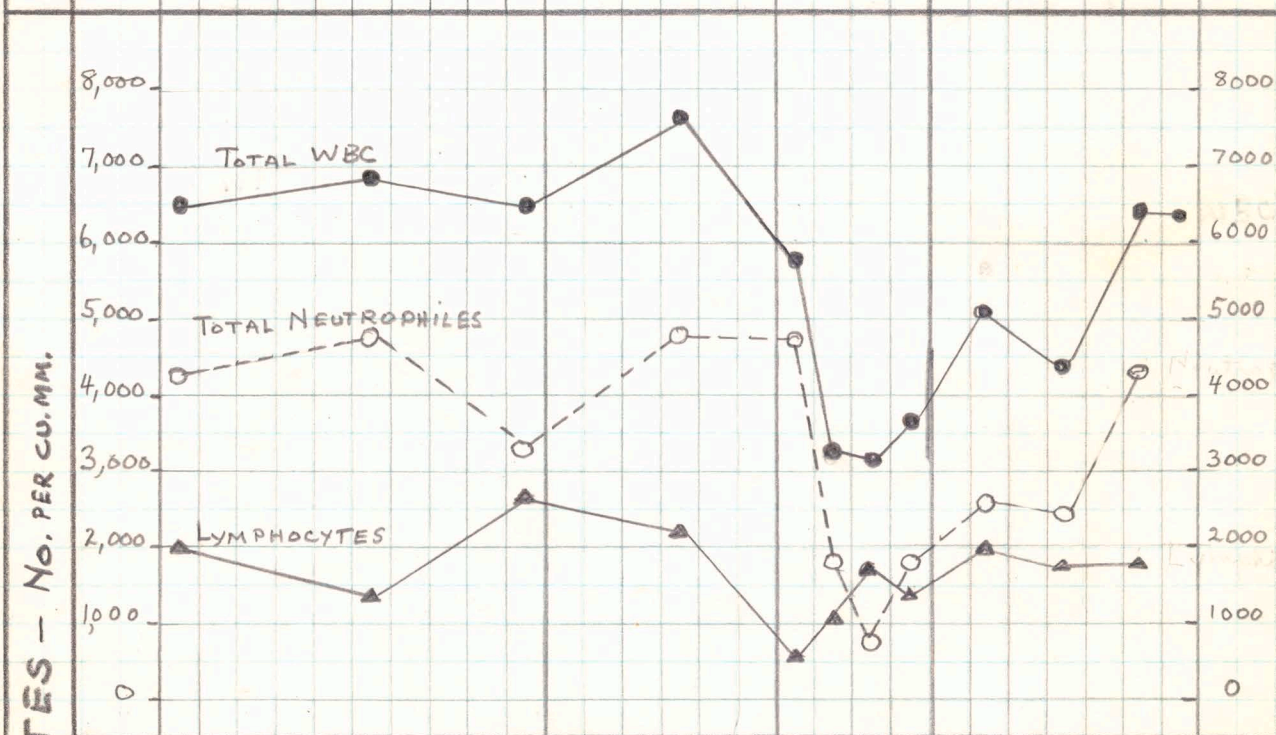
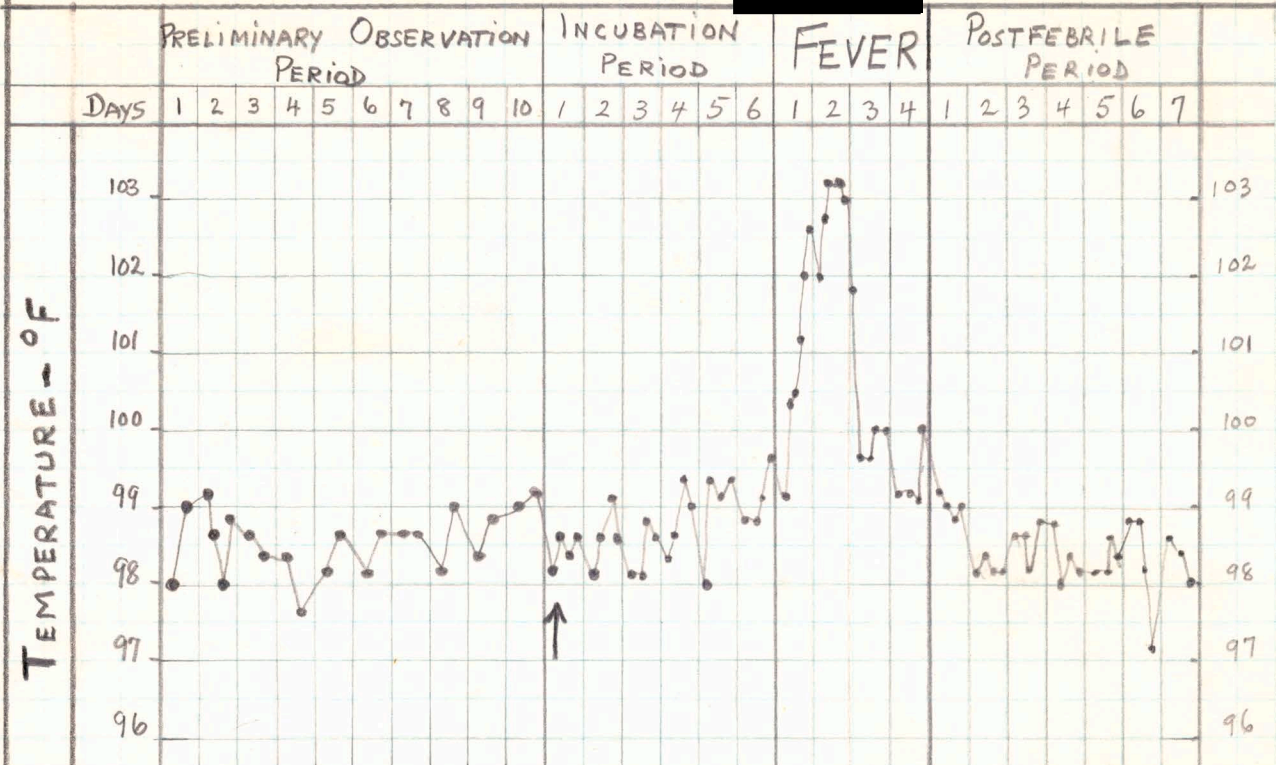
EXPERIMENTAL SANDFLY FEVER - [REDACTED] WHITE MALE, AGE 31



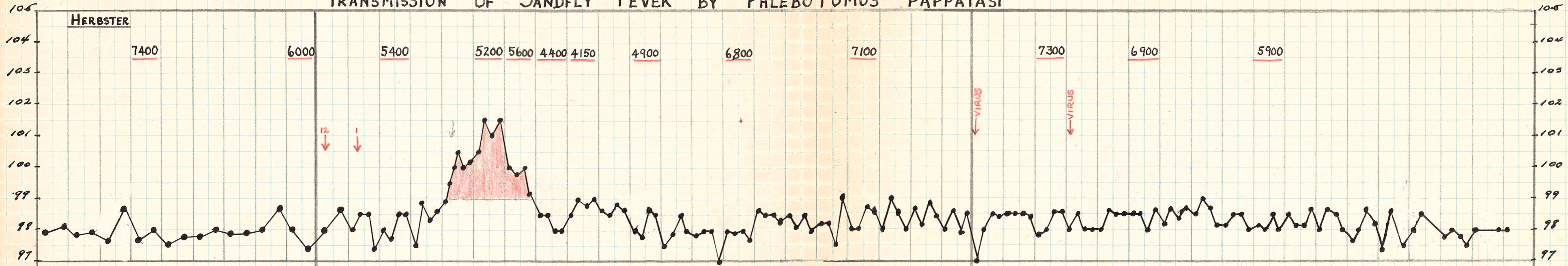


EXPERIMENTAL SANDFLY FEVER - [REDACTED]

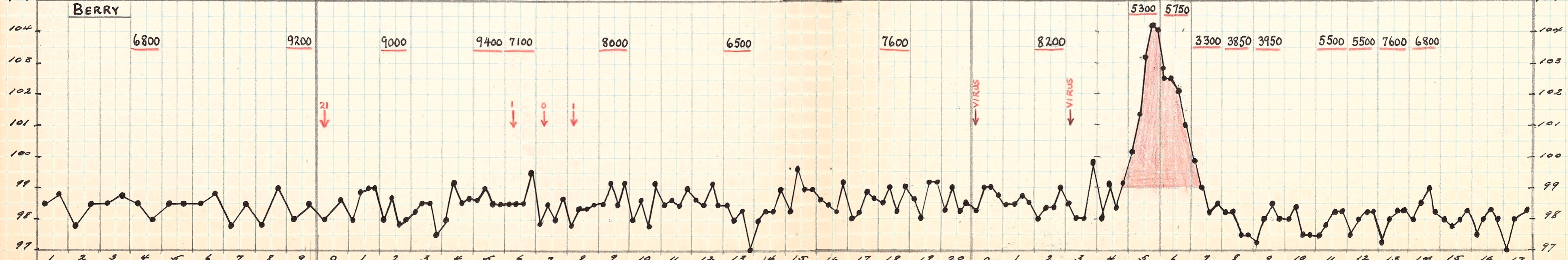
WHITE MALE, AGE 33



TRANSMISSION OF SANDFLY FEVER BY PHLEBOTOMUS PAPPATASI

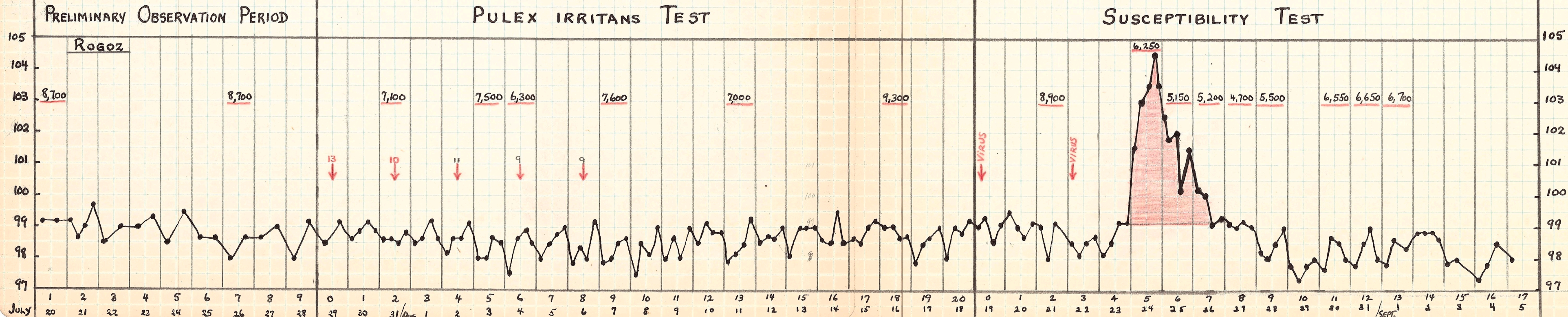
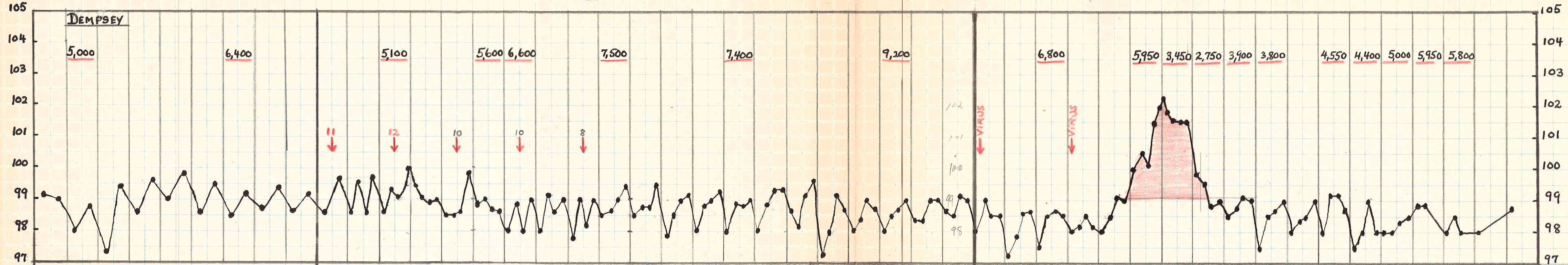


PRELIMINARY OBSERVATION PERIOD PHLEBOTOMUS PAPPATASI TEST IMMUNITY TEST

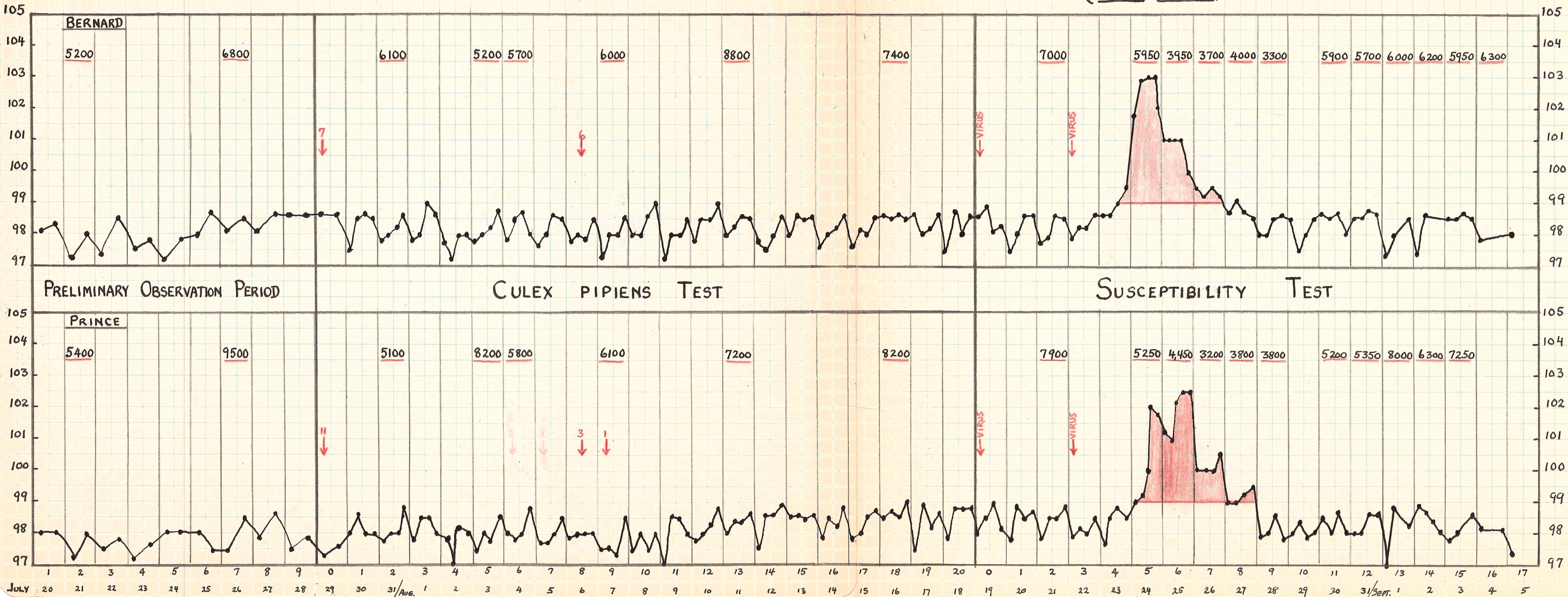


July 20 21 22 23 24 25 26 27 28 29 30 31/Aug 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31/Sept. 1 2 3 4 5

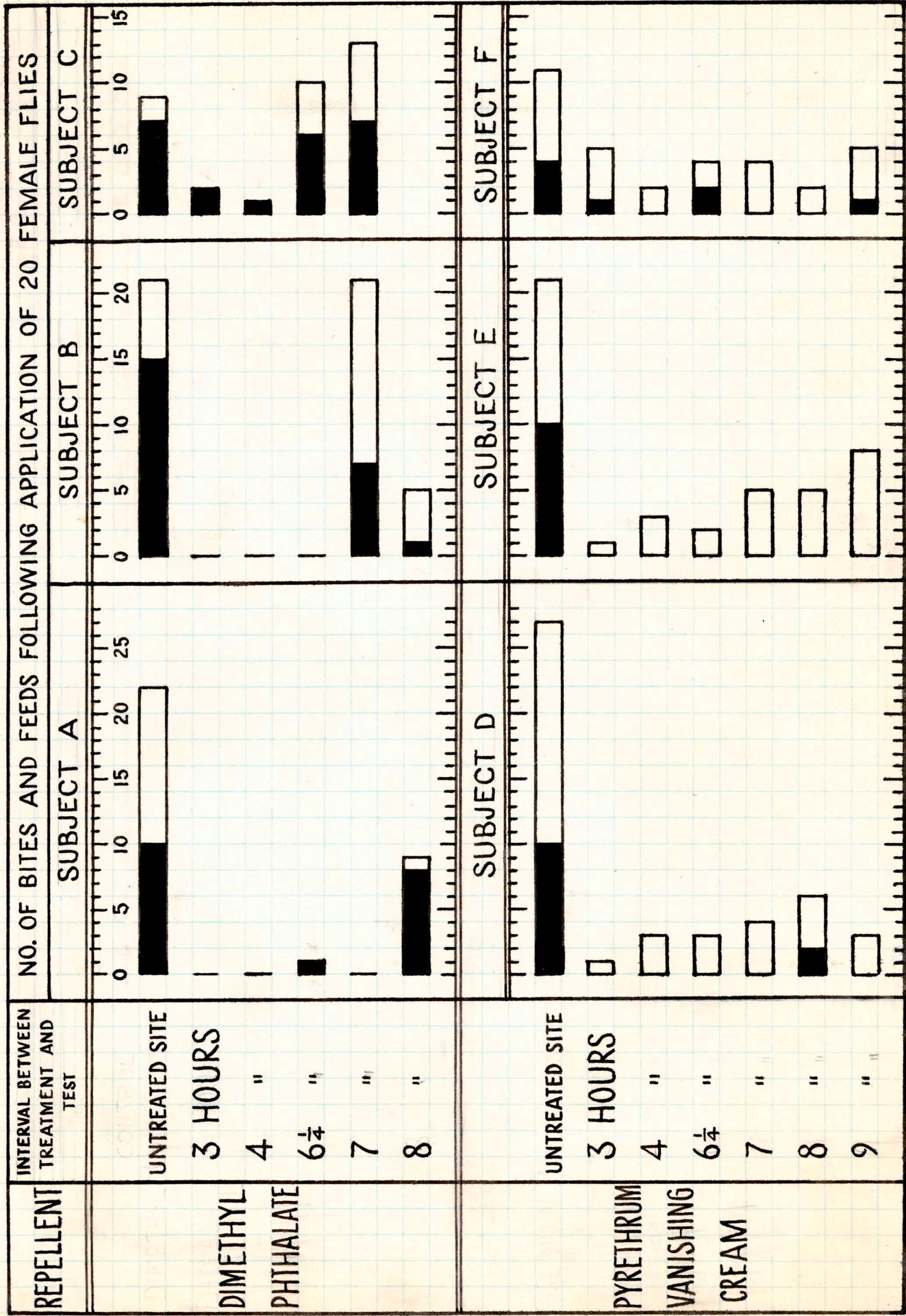
CAN HUMAN FLEAS (PULEX IRRITANS) TRANSMIT SANDFLY FEVER?



ATTEMPTED TRANSMISSION OF SANDFLY FEVER BY MOSQUITOES (*CULEX PIPIENS*)



PROTECTIVE EFFECT OF TWO REPELLENTS AGAINST PHLEBOTOMUS PAPATASI



TOTAL BAR = NO. OF BITES; SHADED AREA = NO. OF FEEDS, (NO. OF FLIES ENGORGED)

ERYTHROCYTE SEDIMENTATION RATES IN PATIENTS WITH
EXPERIMENTAL SANDFLY FEVER

ROURKE-ERNSTENE METHOD WITH CORRECTION FOR
CELL VOLUME (Normal 0.08 - 0.35 mm./min.)

<u>DATE</u>	<u>NAME</u>	<u>DAYS</u> ^{after onset} _{of fever} <u>of FEVER</u>	<u>CORRECTED E.S.R.</u> - mm/min.
JAN. 24	[REDACTED]	3 rd	0.72 → 0.88 10 th
		"	0.63 → 0.46 10 th
JAN. 25		3 rd	0.62 → 0.56 9 th
		4 th	1.35 - 1.14 9 th
		5 th	0.62
JAN. 26		1 st (not S.F. - 1 day later)	0.58 → 0.97 7 th
JAN. 31		2 ND	0.64 → 0.56 9 th
		10 th	1.14
		3 rd	0.22
		10 th	0.46
		9 th	0.56
		10 th	0.88
FEB. 3		1 st	1.04
FEB. 7		5 th	0.90
		9 th	0.56
	5 th	0.30	
	5 th	1.22	

FEB. 17

6TH

0.10

7TH

0.75

7TH

0.97

Before fever

0.32

" "

0.50

FEB. 18

1ST

0.20

(Oralate)

FEB. 19

2ND

0.26

2ND

0.54

FEB. 23

6TH

0.64

6TH

0.45

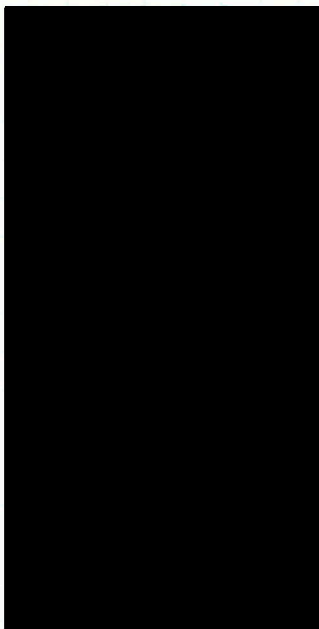
6TH

0.64

PHOSPHATASE DETERMINATIONS ON PATIENTS

WITH EXPERIMENTAL SANDFLY FEVER

JAN. 25, 1944



		<u>Bodansky Units per 100cc</u>	<u>Inorganic P</u>
1 st day	- frozen -	4.4	3.2
3 rd	" - " -	4.5	3.0
"	" - " -	5.4	2.9
"	" - fresh -	2.1	3.7
4 th	" - " -	3.1	3.6
5 th	" - " -	2.7	3.3
Normal control	- frozen -	3.7	3.7

CEPHALIN - CHOLESTEROL (HANGER) FLOCCULATION TEST

Jan 24, '44

		16hrs	24	40	48	64
72	[REDACTED]					
±		3 RD DAY S.F. - fresh	- 0	0	0	0 ±
+		" " " - "	- 0	0	0	0 ±
#		" " " - " - hemolyzed	- #	#	#	# #
#		2 ND " " - frozen	- ±?	0-#	+	+ #
#		<u>INFECT. HEPAT.</u> - first day jaundice - frozen	- ###	##	##	##
0		- normal control - fresh	- 0	0	0	0 0
0		- control	- 0	0	0	0 0

DESIGNATION OF READINGS

- 0 = uniform suspension
- ± = fine granularity without sedimentation - uniformly dispersed
- + = coarse granularity with sedimentation
- # = large floccules - sedimented ^{and in suspension} - but considerable opalescence remaining
- ## = well packed - sedimented ppt. - only faint opalescence remaining
- ### = complete flocculation - su. liquid clear.

Note - On Jan. 22 Dr. Wasserman tested two sera for me

		24hrs	48hrs
	1 st day S.F. - fresh	- 0	0 (±?)
	" " - frozen	- 0	0 (±?)

CEPHALIN-CHOLESTEROL FLOCCULATION TEST (HANGER)

JAN. 25, 1944

66 hrs			15 hrs.	24	40	48
#	[REDACTED]	1 st day S.F. - fresh	- 0	0	#	#
0	[REDACTED]	5 th day S.F. - "	- 0	0	0	0
0	[REDACTED]	3 rd " S.F. - " Some hemolysis	- 0	0	0	0
0	[REDACTED]	1 st " S.F. - frozen	- 0	0	0	0
+	[REDACTED]	1 st " S.F. - "	- 0	0	+	+

MUSTART - HEPATITIS - FROZEN

#	Pool I	[REDACTED]	bled 6/14 - frozen 6/16	- 0	0	0	0
0	[REDACTED]	[REDACTED]	bled 7/7 - frozen 7/8	- 0	0	0	0
0	[REDACTED]	[REDACTED]	bled 7/7 - frozen 7/8	- 0	0	0	0
#	[REDACTED]	[REDACTED]	bled 7/7 - frozen 7/8	- +	+	#	#
#	[REDACTED]	[REDACTED]	bled 6/26 - frozen 6/27	- #	#	#	#
#	[REDACTED]	[REDACTED]	hemolysed; bled 7/2 - frozen 7/2	- #	#	#	#
#	[REDACTED]	[REDACTED]	hemolysed, fatty; bled 7/1 - frozen 7/2	- +	+	#	#
0	[REDACTED]	[REDACTED]	fatty; Bled 6/18 - frozen 6/19	- 0	0	0	0
0	[REDACTED]	[REDACTED]	bled 7/7 - frozen 7/8	- 0	0	0	0
#	[REDACTED]	[REDACTED]	Bled 6/18 - frozen 6/19	- #	#	#	#
#	[REDACTED]	[REDACTED]	Bled 6/14 - frozen 6/16	- #	#	#	#

+	[REDACTED]	- 39 days - 8/27 - frozen (HEP 9/28)	- 0	0	0	0
0	[REDACTED]	- 34 " 9/22 - " (HEP 10/25)	- 0	0	0	0
#	[REDACTED]	- 19 " 9/7/43 - "	- +	+	#	#

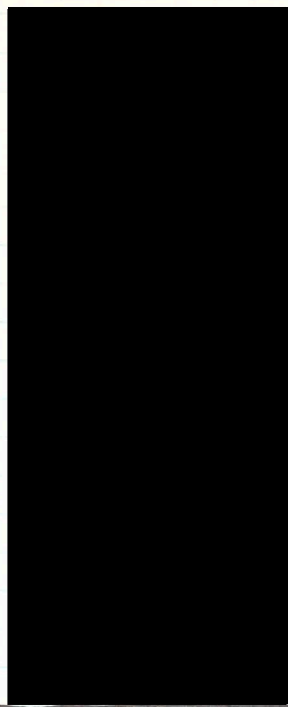
0	[REDACTED]	- frozen - normal serum control	- 0	0	0	0
0	SALINE control		- 0	0	0	0

Mixtures set up at 5:40-5:50 P.M.

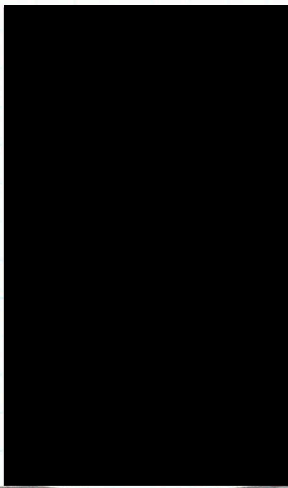
CEPHALIN - CHOLESTEROL FLOCCULATION TEST (HANGER)

JAN. 27

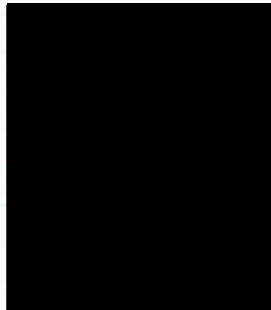
All sera frozen in CO₂



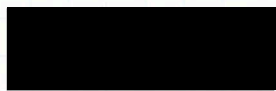
	18 hrs.	24	45	70
(7-19-43)	- #	#	#	
(7-20-43) - fatty	- #	#	#	
(7-21-43) sl. hem.	- +	#	#	
(8-2-43) sl. fatty	- #	#	#	
(8-24-43) mod. hem.	- 0	0	0	±
" sl. hem.	- 0	0	0	+
" " "	- 0	0	+	#
" mod "	- ±?	0?	+	#
" " "	- +	+	#	



(10-10-43) sl. fatty	- 0	0	+	#
"	- ?	0?	0	+
(10-12-43)	- #	#	#	
(10-16-43) sl. hemol.	- +	#	#	
(10-20-43)	- 0	0	0	+
(11-18-43)	- 0	0	0	0 0



(11-22-43)	- 0	0	0	0 0
(8-31-43)	- 0	0	#	###
(8-20-43)	- +	+	#	
(9-3-43) very fatty	- ?	0	#	



- Normal control - 0 0 0 0

SALINE CONTROL - 0 0 0 0

Mixtures at 4:30 P.M.

CEPHALIN - CHOLESTEROL FLOCCULATION TEST (HANGER)

JAN. 28, 1943
1944

ALL SERA FROZEN IN CO₂

			<u>21</u>	<u>46</u>	
NORMAL PREINOCULATION SERA	[REDACTED]	(7-5-43)	- #	#	
	[REDACTED]	" fatty	- #	#	
	[REDACTED]	"	- #	#	
	[REDACTED]	(7-20-43) sl. fatty	- + #	#	
	[REDACTED]	" fatty	- #	#	
	[REDACTED]	" sl. "	- #	#	
	[REDACTED]	" " "	- #	#	
		"	- 0	+	# (70 hrs)
<hr/>					
	[REDACTED]	(8-11-43) fatty	- #	#	
<hr/>					
PREINOCULATION SERA	[REDACTED]	(10-11-43)	- +	#	
	[REDACTED]	(10-6-43)	- +	#	
	[REDACTED]	"	- 0	±	# (70 hrs)
	[REDACTED]	"	- + #	#	
	[REDACTED]	" 12-7-43	- #	#	
<hr/>					
	[REDACTED]	- NORMAL CONTROL	- 0	0	± (70 hrs)
	SALINE	CONTROL	- 0	0	0

Mixtures made at 3:45 P.M.

+ marked flocculation in 2 hours

CEPHALIN-CHOLESTEROL FLOCCULATION TEST (HANGER)

JAN. 28

CONTROL TESTS ON FRESHLY ^{DRAWN} "NORMAL" SERA
AND ONE ACUTE S.F. SERUM

	<u>21</u>	<u>46</u>
[REDACTED]	- 0	0
[REDACTED]	- 0	0
[REDACTED]	- 0	0
[REDACTED]	- 0	0
(S.F. - 1 st day)	- 0	+
SALINE CONTROL	- 0	0

Mixtures set up at 4 P.M.

FLOCCULATION TEST

FEB. 3, 1944

	<u>24</u>	<u>48</u>	72
[REDACTED] AN. 11 - U.R. Infection) - frozen	- 0	0	0
[REDACTED] B. 3 - 1 st day S.F. - fresh	- 0	0	0
[REDACTED] " - " " - "	0	±	+
[REDACTED] " - " " - "	0	+	+
SALINE CONTROL	- 0	0	0

Mixtures set up at 5 P.M.

CEPHALIN - CHOLESTEROL FLOCCULATION TEST (HANGER)

FEB. 7

All freshly drawn sera (except positive control)

	24	48	72
- S.F. - 5 th day	0	0	0
- " - " "	0	±	+
- " - 10 th "	0	+	+
- " - 5 th "	0	0	0
- " - 9 th "	0	0	0
- F.U.O.	0	±	+
- SUSP. INF. MONONUCLEOSIS	+	±	±
- Positive JAUNDICE SERUM	###*	###	###
SALINE CONTROL	0	0	0

Mixtures set up at 2:45 PM

* ### in 2 hours

FLOCCULATION TEST

FEB. 11

	24hrs	48
- [REDACTED] - 1 ST DAY (fresh)	0	+
SALINE CONTROL	0	0

CEPHALIN - CHOLESTEROL FLOCCULATION TEST

FEB. 17

TESTS ON FRESH SERA

	<u>17 hrs</u>	<u>24</u>	<u>48</u>	<u>72</u>
[REDACTED] - 2 days after inoc - before fever -	0	0	0	0
[REDACTED] - " " " " " -	0	0	0	0
[REDACTED] - S.F. - 6 th day -	0	0	0	0
[REDACTED] - " - 7 th " -	0	0	±	+
[REDACTED] - " - 7 th " -	0	±	+	±
POSITIVE CONTROL -	+++			
SALINE CONTROL -	0	0	0	0

Mixtures set up 4:15 P.M.

FEB. 18

TESTS ON FRESH SERA

	<u>20</u>	<u>44</u>
[REDACTED] - S.F. - 1 st day -	0	0
[REDACTED] - S.F. - " " -	0	0
[REDACTED] - S.F. - 1 st day 1st -	0	0
SALINE CONTROL -	0	0

Mixtures set up 6 P.M.

TESTS ON FRESH SERA

	<u>24</u>	<u>48</u>	<u>FEB. 23</u>
[REDACTED] S.F. - 6 th day -	0	0	
[REDACTED] F. - " " -	0	0	
[REDACTED] - " " -	0	0	
SALINE CONTROL -	0	0	

SET UP
AT 2:30 P.M.

SUMMARY OF CEPHALIN-CHOLESTEROL FLOCCULATION TESTS ON PATIENTS WITH EXPERIMENTAL PHLEBOTOMUS FEVER

~~#~~ ALL TESTS ON FRESHLY DRAWN SERA

NAME	DAYS FROM ONSET OF FEVER	Result	
		24	48
[REDACTED]	1 ST	0	0 to ±
	5 TH	0	0
	4 TH	0	±
	1 ST	0	+
	10 TH	0	+
	1 ST	0	±
	15 TH	0	0
	1 ST	0	+
	5 TH	0	±
	1 ST	0	0
	5 TH	0	0
	3 RD	0	0
	3 RD	0	0
	3 RD	0	0
	9 TH	0	0
1 ST	0	+	
7 TH	0	±	
6 TH	0	0	
7 TH	±	+	
Before fever		0	0
1 ST		0	0
6 TH		0	0
Before fever		0	0
1 ST		0	0
6 TH		0	0
1 ST		0	0
6 TH		0	0

