

4hr Expt

<u>Generations</u>	<u>T4</u>	<u>T5</u>	<u>T6</u>
83	1092	673	213
89	2236	603	196
95	7686	594	266
101	9160	478	224
107	13,000	338	199

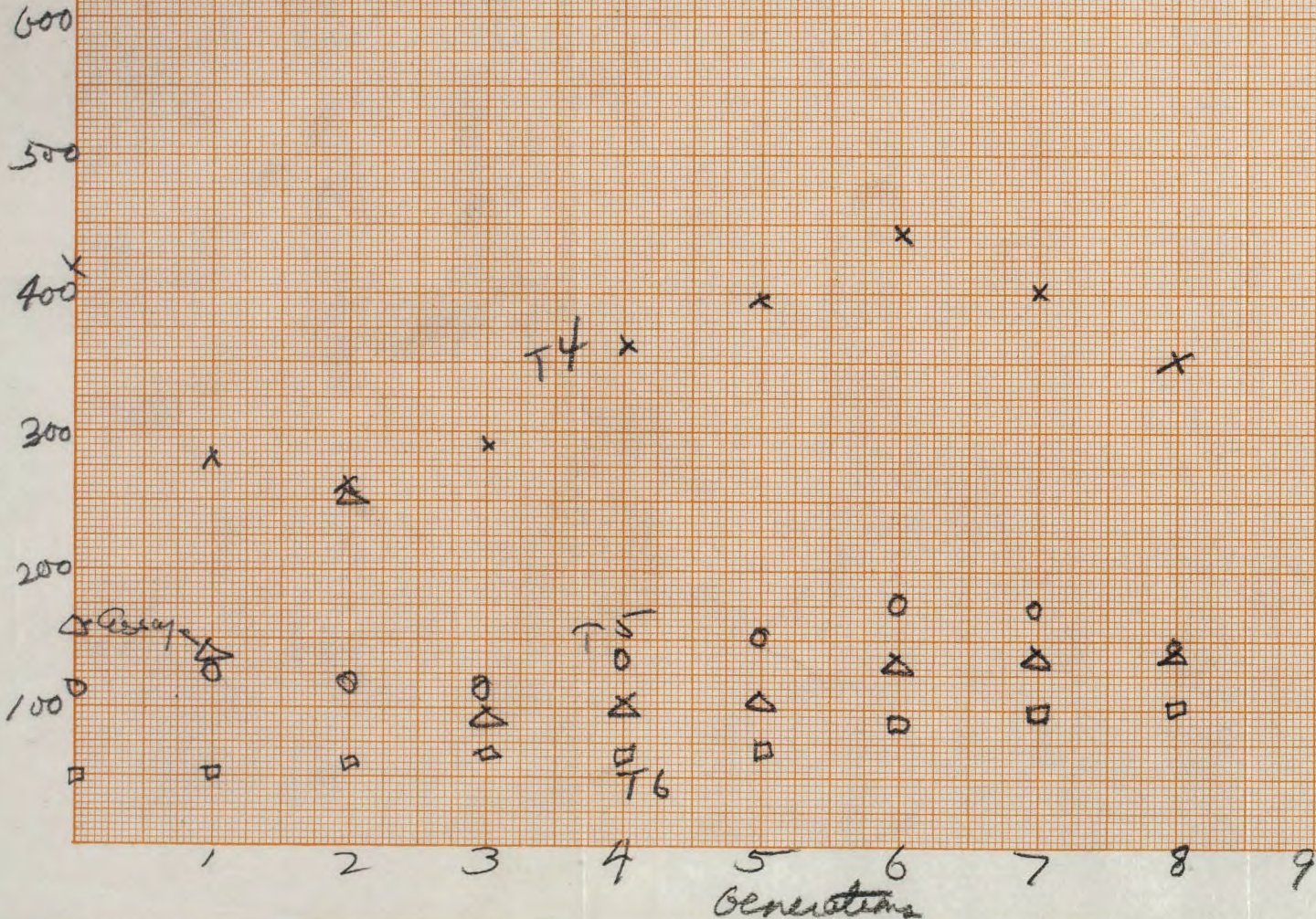
Below are a few examples of characteristics of earlier "falls"

Rate	Conc typ	Strain	generation fall began	generation fall ended	fraction left after fall
6hr	1000 γ	fast	31	45	0.5
2hr	200 γ	fast	38	45	0.4
6hr	1000 γ	hyper	50	80	0.1

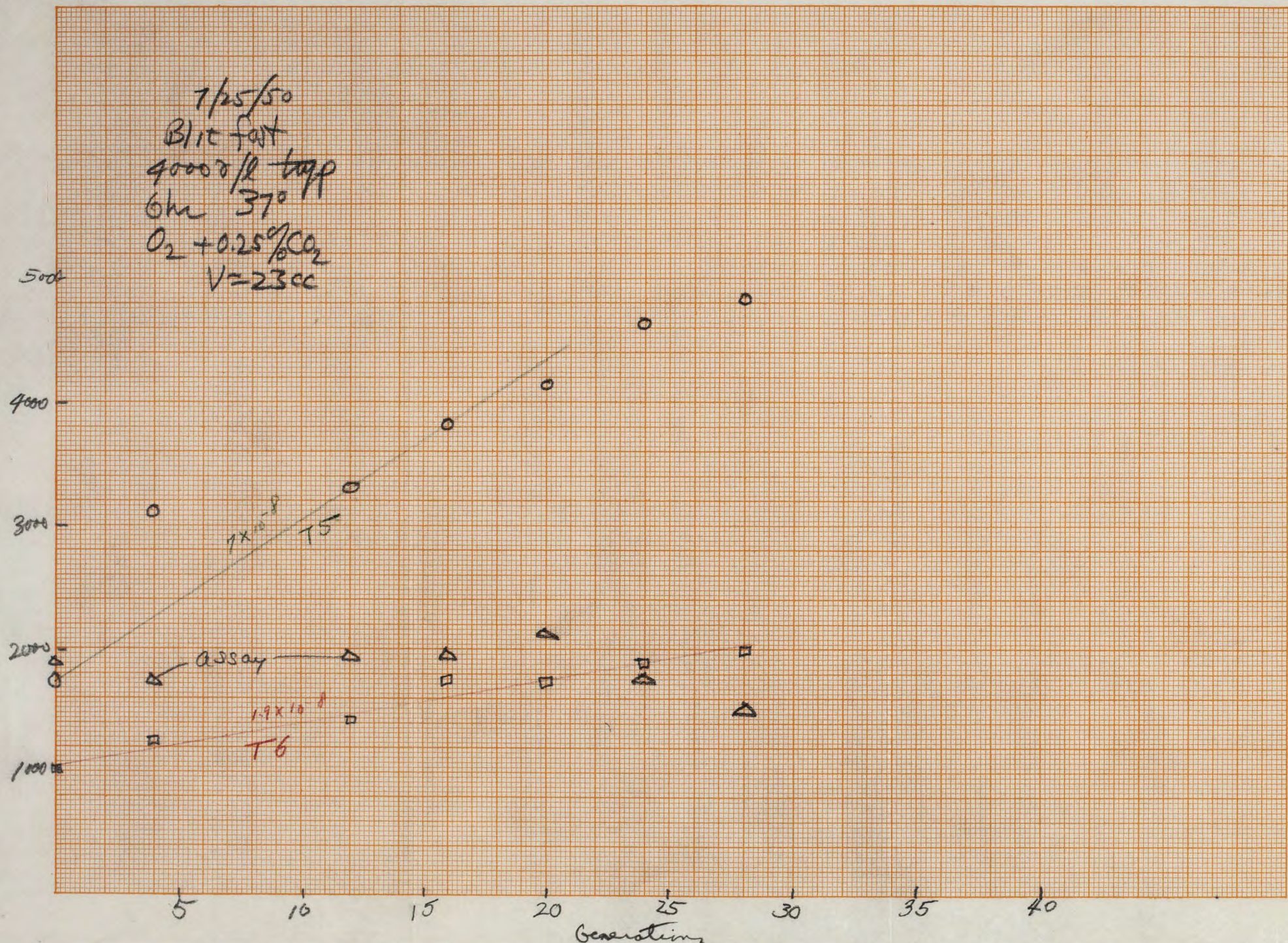
P.S. About you're coming east. I don't think the phase meetings will be worth the trip. Perhaps it would be better to spend the time together in Chicago — unless you have other business in NYC.

My sister is getting married the 27th of August. I'll probably be busy on the 26th and 27th entertaining relatives.

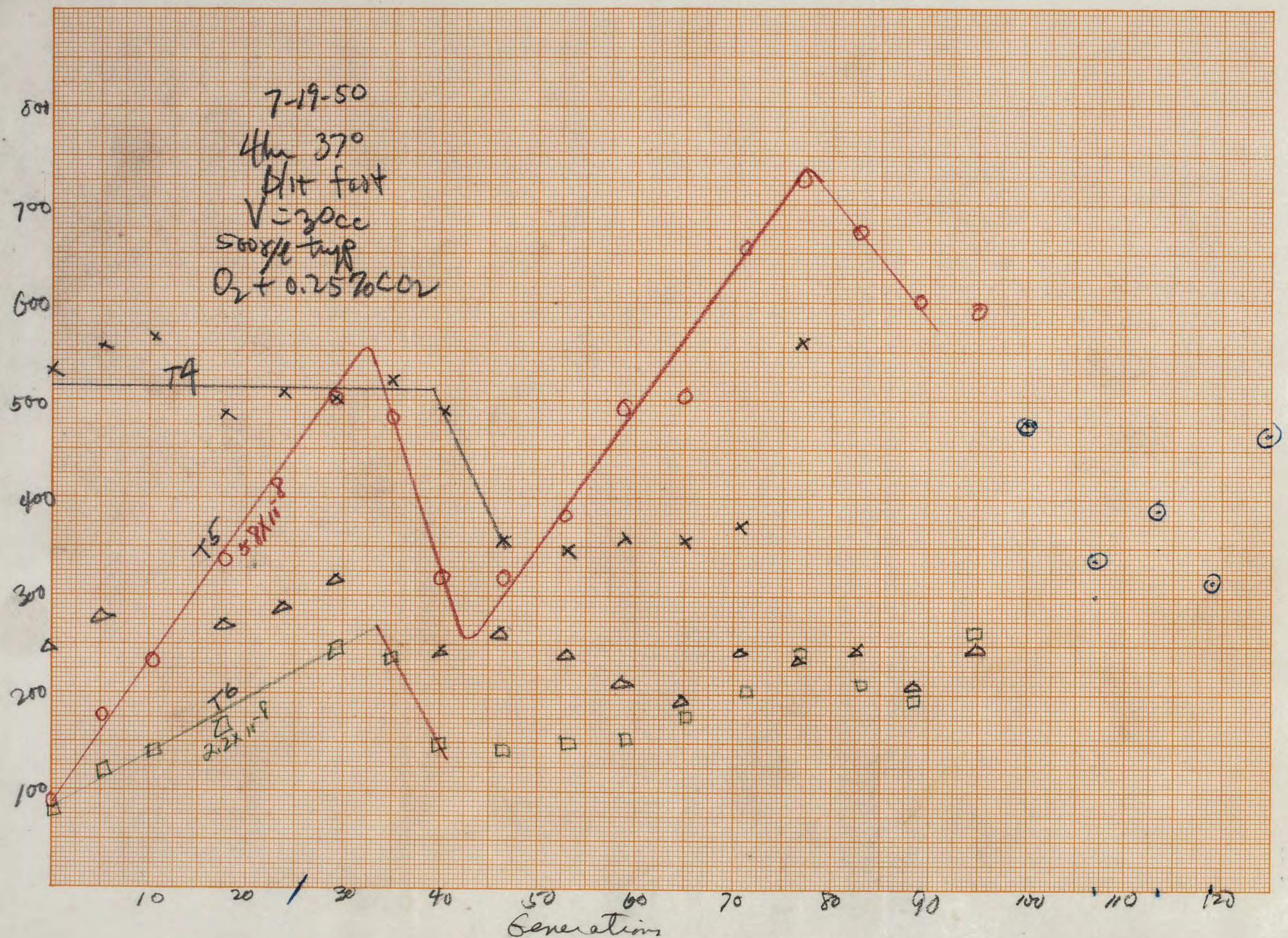
7-27-50
Blr fast
24hr 37°
500 ml top
V=80cc
O₂ + 0.25% CO₂



7/25/50
Blit fast
40000/r tapp
6hr 37°
O₂ + 0.25% CO₂
V = 23cc



X
↑
T4



2 hr samples
on sun
used sample 8/5/50
6 hr after
above

Expt of 8/2/50
collected in ice
samples
divided in 2
"cold" centrifuged
immediately
"Warm" incubated
at 31° for 14 hr

24 hr samples
are diluted
in F6 1/2

λ	2hr	6hr	4hr cold	24hr cold	4hr warm	24hr warm
350	.020	.036	017	025	034	026
300	.039	.137	032	064	060	068
290	.103	.575	076	162	125	208
280	.167	.945	122	260	212	350
270	.180	.970	140	279	273	390
260	.169	.850	142	267	305	376
250	.148	.695	147	274	342	365
240	.153	.680	200	409	482	505
230	.290	1.21	402	900	960	1.07

167
103
64

945
5.75
3.70

970
5.75
395

140
76
64

945 = 472
2
208

not
typical
at 4 hr warm
rate should
be [140]
+60
190
observed
200

Absorption of inorganic phosphorus estimation

2900 Å
2800
2700
:
2400

0.220
.2765
.295
↓
.137

(about 1/2
of value for
2800 Å)

4 hr, 5000/l on O₂

Generations	T4	T5	T6	Assay
101	9160	478	224	2.2 x 10 ⁸
107	13700	338	202	2.2 x 10 ⁸
113	23,300	350	294	1.5 x 10 ⁸
119	36,000	314	265	0.87 x 10 ⁸
125	46,300	414		1.88 x 10 ⁸

2.4 hr O₂ 5000/l tryp

8	355	146	104	1.45 x 10 ⁸
9	259	156	113	1.62 x 10 ⁸
switched to 12 hrs after this point - calling the pt 0 gen				
2	537	238	174	2.5 x 10 ⁸
4	555	310	202	1.66 x 10 ⁸
6	577	327	231	1.38 x 10 ⁸

these are 12 hr generations

6 hr high tryp (40000)

32	—	5080	1750
36	—	3670	1510
40	—	3440	1530
44	—	3180	1460
48	—	3530	1570

6 hr	
2.0 x 10 ⁹	
1.66 x 10 ⁹	
1.9 x 10 ⁹	
1.7 x 10 ⁹	
1.9 x 10 ⁹	

5a

If we keep a strain of bacteria growing in the growth tube of the chemostat and another bacterial strain is generated from it ~~at~~ through spontaneous mutations ~~the~~ which has the same growth rate as the original strain ($\lambda^* = \lambda$) ~~will~~ against [i.e. in the absence of selection for or against the mutant] the density of the mutant bacterial density n^* of the mutant strain should increase linearly with time i.e.

$$\frac{dn^*}{dt} = \frac{\lambda^*}{\tau} n^* ; \text{ as long as } n^* \ll n$$

where n^* is the density of the mutant population, ~~and~~ n the density of the ~~original~~ parent population ^{of parent strain} and λ the ~~frequency~~ ^{number} of the occurrence of a ~~particular~~ ^{mutant produced} ~~product~~ ^{per generation} per bacterium. This should hold ^{time} (as long as $n^* \ll n$) ~~if~~ ^{unless the back mutation rate is very}

If the ~~mutant~~ ^{growth rate of the} mutant strain is less than the ~~large~~ growth rate of the parent strain ($\lambda^* < \lambda$) i.e. if the mutant is selected against ~~the~~ ~~the~~ ~~density~~ then after an initial rise

the density ^(~~is~~) of the mutant population in the growth phase of the chemostat should ~~be~~ - after an initial rise - remain constant at a level given by

$$n^* = \frac{d}{\mu - d} K_m$$

~~Of these two cases of steady state the former one is more suitable for determining~~

~~If we are interested in determining the mutation rate to the more resistant selection is the more favourable one since ~~the~~~~

Of the various mutations occurring in the growing bacterial population mutants resistant to a bacterial virus are particularly easy to score with great accuracy and we ~~used~~ ^{used} ~~it~~ in our experiment mutants ~~of the~~ ~~of~~ the cold strain of our cold strain which were resistant to the Bacterial viruses T4 or T5 or T6.

When we grew ~~our~~ the strain B/11 in the chemostat ~~with~~ with a high ^{conc} ~~conc~~ of the pyrophosphate conc in the nutrient in the storage tank but a ^{appropriate} low conc of lactate

so that lactate rather than tryptophane may be the controlling growth factor we found that after a short initial period the number of ~~the density~~ ~~densities~~ of the bacterial density was for all three mutants ~~fastest~~ of all three and the mutant which grows to T₄ or T₅ or T₆ regardless of mutant T₂

remained constant from day to day, the level corresponding to a ~~the~~ selection factor $(\frac{\lambda - \lambda^*}{\lambda})$ of a few per cent. —

~~It is not trapped by tryptophane~~
~~It is perhaps not too surprising~~

~~One can perhaps understand that under conditions of starvation for ^{the main} carbon source~~
~~in that most mutants grow most~~

~~It might be that~~

~~It might be these organisms~~

~~Perhaps it might be so that most mutants are selected ~~apparently~~ from slower~~

~~than the wild. It is conceivable that practically all mutational steps or ~~at least~~ which lead away from the "wild type" ~~from~~~~

~~lead to mutants which grow slower under conditions of "starvation" for ^{the main} carbon source~~

~~We should not however expect~~
~~the mutant~~ There is however no
reason to expect that mutants
in general ~~cannot~~ grow slower than
the "wild type"

If ~~that~~ we grow our strain ~~in the~~ in the chemostat with a
high conc of lactate in the medium in
the storage tank, ~~and~~ use pyrophosphate
as a growth factor and ~~we~~ have the
generation time T well above the minimum
generation time of 20 min (subject of holds
for high pyrophosphate concentrations) then
there is no reason to expect ~~no~~
in general mutants to grow slower ⁱⁿ
than the parent strain. ^{growth is slow down by being}
~~mutants should have some direct effects~~ ^{of the growth by being}
~~likely to affect the uptake or~~ ^{the pyrophosphate can}
~~utilization of pyrophosphate then~~
~~are not of course expected~~
In this case ^{in the chemostat} we would expect ~~a~~
~~direct~~ the mutation to affect the
growth rate only if the mutation
affect the uptake or utilization
of pyrophosphate

Fig 8 shows the ^{Bacterial density per} ~~number of~~ mutations resistant to T5 ~~as~~ ~~function of~~ present in the population of the bacterium as a function of the number of generations through which the parent strain has passed (~~number of~~ number of generations = $\frac{t}{\tau_w}$) for two experiments

in which the generation times were either 2 hrs, 6 hrs and 12 hrs respectively. ~~The~~ The slopes of the straight lines give the mutation rates per 10^8 generation as 2.5, 7.5 and 12.5 ~~per~~ $\times 10^8$ per ~~generation~~ generation per bacterium. —

We see that the mutation rate per gen is 3 times as high for $\tau=6$ hrs as it is for $\tau=2$ hrs and 6 times " " for $\tau=12$ hrs as it is for $\tau=2$ hrs.

What appears to remain constant is the number of mutations produced per unit time and bacterium. ~~Worked~~

$$\frac{\mu}{\tau_{\text{gen}}} = \frac{2.5}{2} 10^{-8} = 1.25 10^{-8} \text{ per hour and bacterium}$$

This is contrary to what had been generally expected ~~of~~

epitheloid

1) ~~Fibroblast~~ [Correll] ^{cell} with embryonic

b) no embryonic juice

Is another epithelium.

Doljansky

see Laser (Lippmann)

Edmund Mayer
Stanford Univ.

Brues; Paul Weiss;

Cyranovitch

Wesley T. Pratt Jr, Minnesota

Ann. of exp. Biology

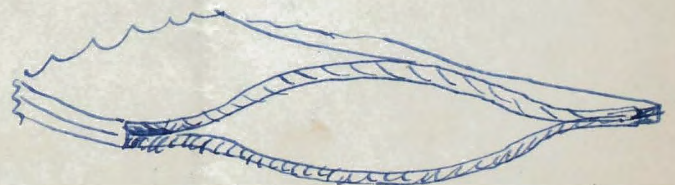
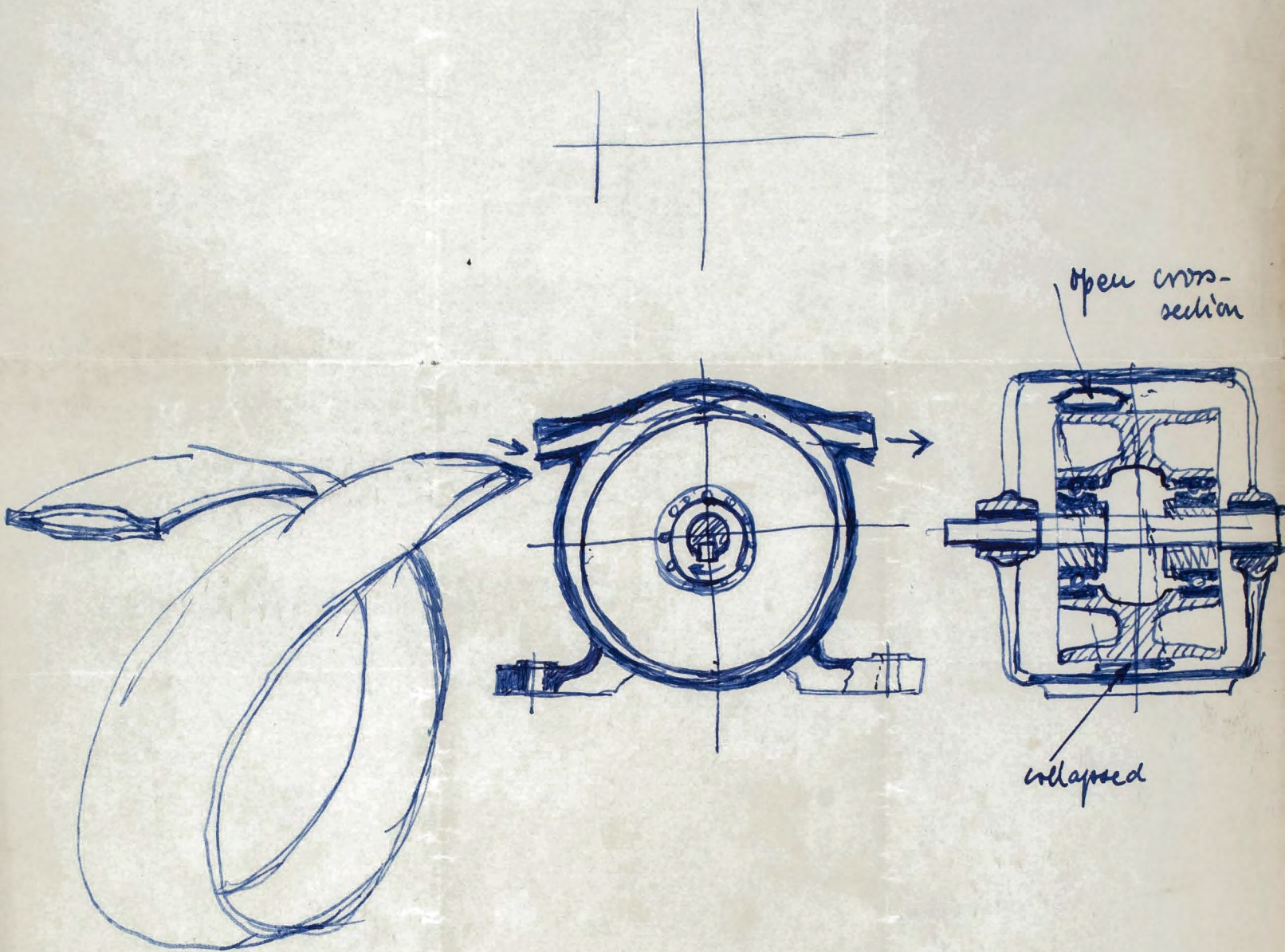
Vol. 107 No. 1 Febr 48 p. 38.

(Check embryo on eggs)

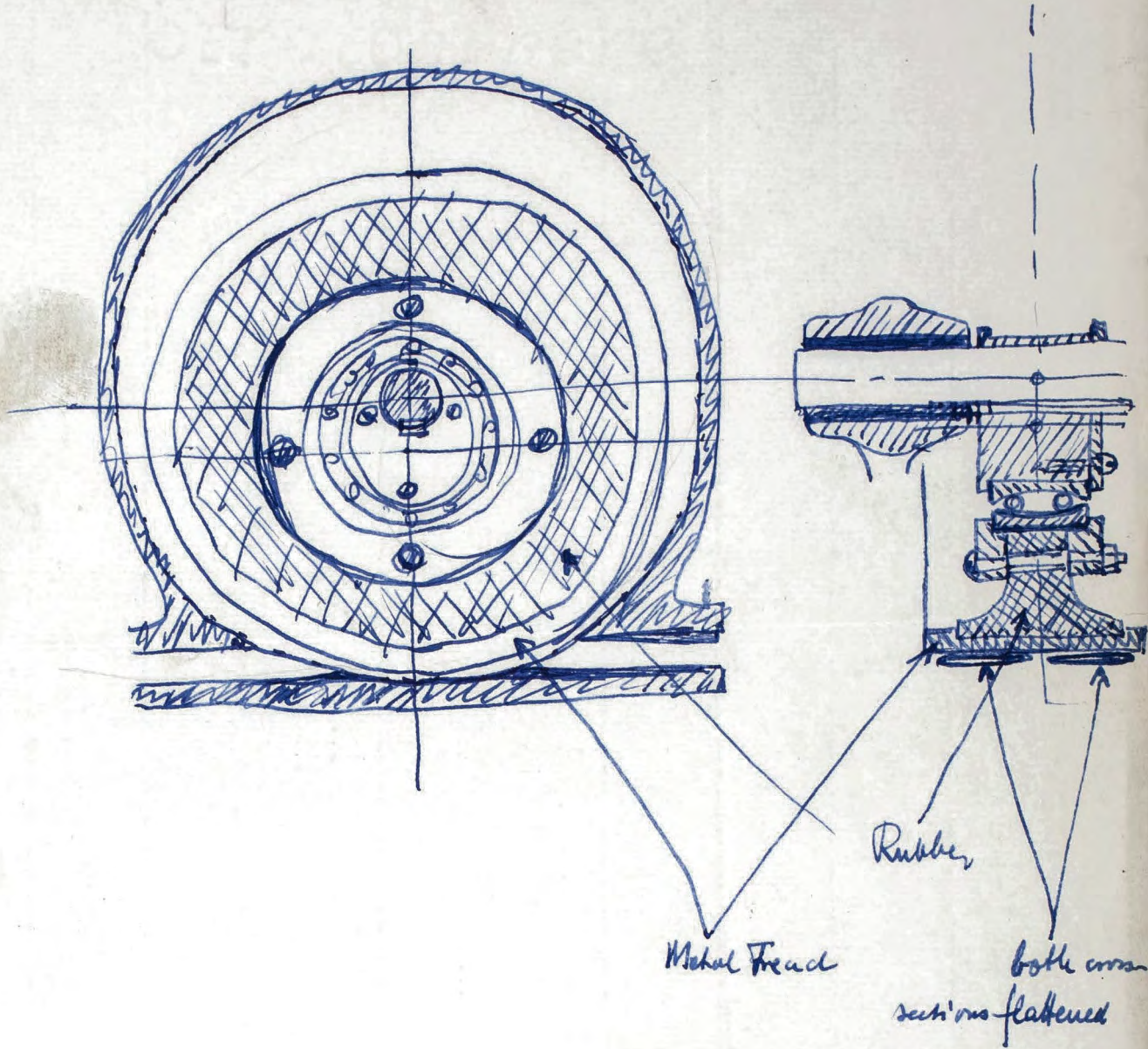
Tabulae Zoologicae
Vol XIX Pars I (Cellula) Giffenrijck Sr. W. Junk
1839 Den Haag.

V. P. Potter [Advances in Embryology Vol 4
Interference. 1944]

Plant tissue cultures
Howard. Dept. of Biology



Cross section



Metal Flange

Rubber

both cross sections flattened

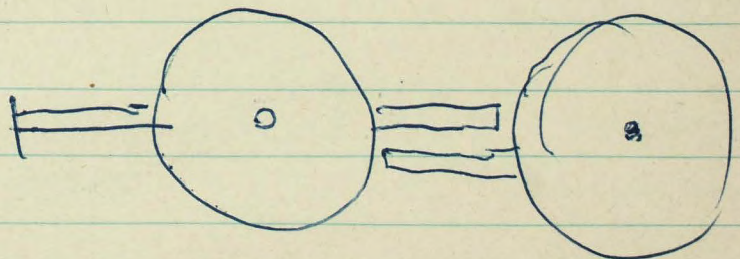
Trypaphysia description

2900 Å 0.220

2800 Å 0.276

2700 Å 0.295

2400 Å 0.137 (about $\frac{1}{2}$ of value
at 2800 Å)

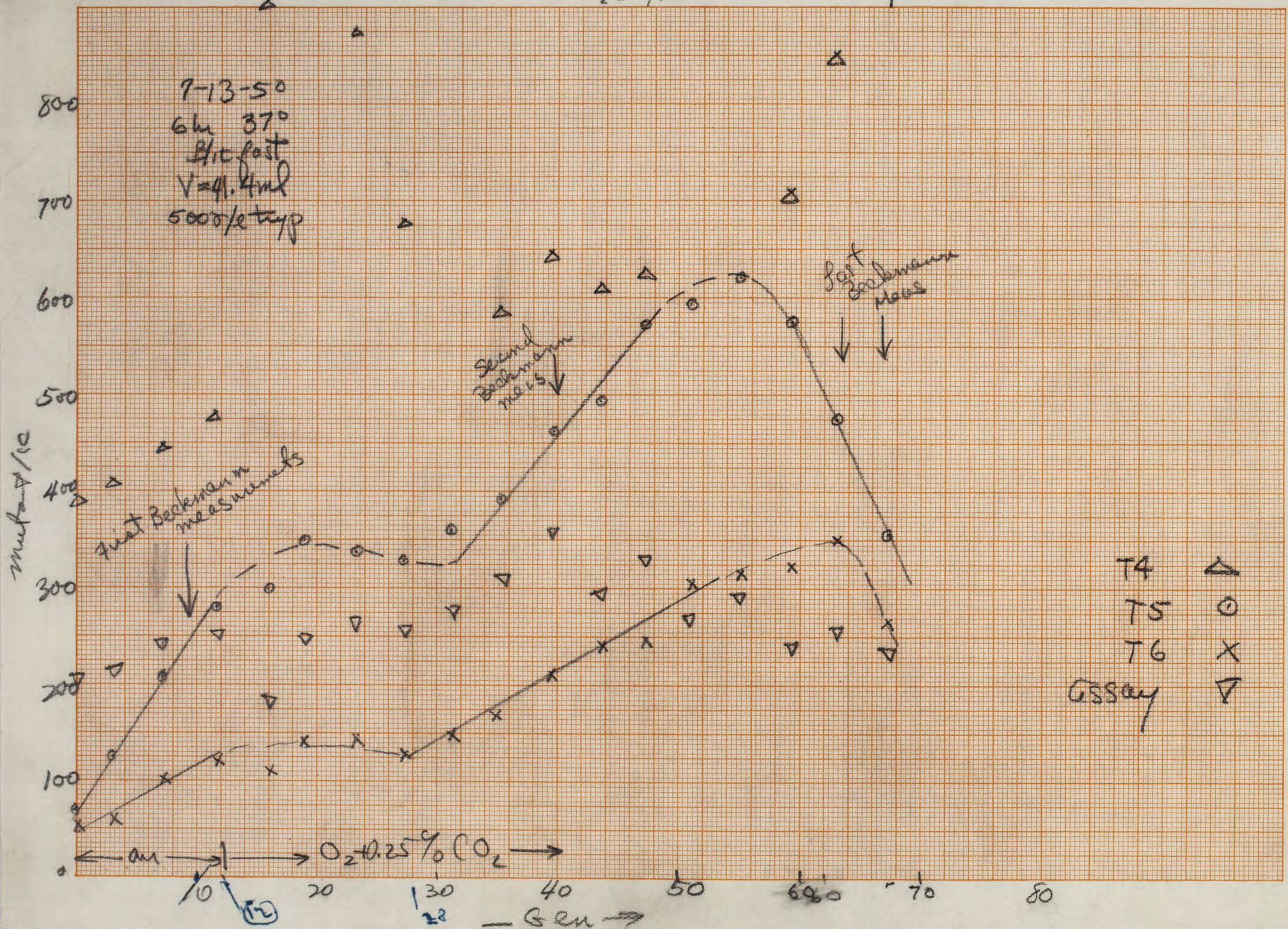


$14/2.5 = 5.7$

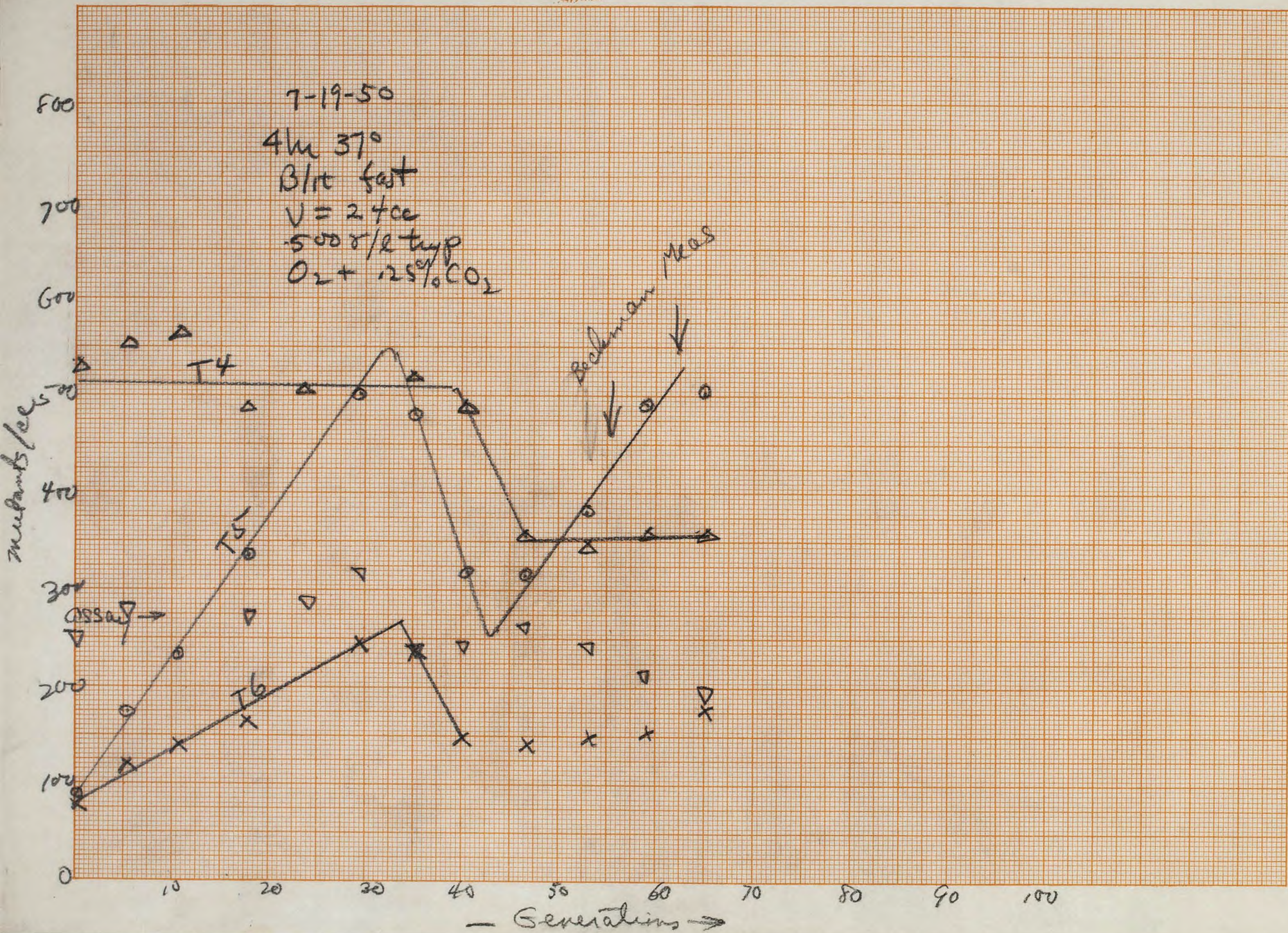
$\frac{325}{550} = 14$

1211
 ↑

7-13-50
 6h 37°
 3/10 foot
 V=4.4 ml
 5000/e typ



- T4 \triangle
- T5 \circ
- T6 \times
- GSSay ∇



July 13, 1950

First Beckmann Meas
Undiluted samples vs dist water

	undiluted		diluted 1:1 with H ₂ O		
	2hr	6hr	2hr	6hr	6hr
350	.103	0.116	.059	.055	.055
340	.108		.062	.058	.058
330	.113		.065	.060	.060
320	.120		.068	.064	.064
300	.155	0.210	.082	.091	.088
290	.288	0.626	.156	.296	.291
280	.420	0.980	.231	.479	.296
270	.460	1.00	.253	.485	.479
260	.451	0.85	.242	.403	.485
250	.410	0.645	.208	.291	.47
240	.385	0.600	.185	.236	
230	.710	1.20	.158	.355	
226	1.10	1.58	.135	.730	

Absorption of bacteria alone at this density and absorption of F given below

λ	Bacteria	F
290	0.25	0.05
280	0.31	0.06
270	0.36	0.06
260	0.39	0.06

Recent results on reverse side

110 620

231
150
71

July 30, 1950

Bacteria in water vs water

λ	<u>2hr</u>	<u>24hr</u>	<u>4hr</u>	6hr (4000 γ /l dil $\frac{1}{8}$ in 1/20)
350	.148	.181	.148	.179
300	.200	.240	.199	.243
290	.254	.303	.255	.300
280	.320	.371	.320	.360
270	.370	.424	.367	.400
260	.392	.454	.396	.424
250	.390	.450	.390	.430
240	.398	.470	.400	.470
230	.560	.655	.552	.682
225	.664	.783	.663	.840

Filtrates vs F med

λ	<u>2hr</u>	<u>24hr</u>	<u>4hr</u>	6hr dil $\frac{1}{8}$ in F	6hr (old) of 7/22	6hr old of 7/28
350	.0175	.022	.010	.015	.018	.010
300	.052	.092	.045	.060	.062	.043
290	.215	.440	.215	.400	.132	.117
280	.370	.760	.376	.673	.238	.217
270	.405	.775	.417	.685	.1288	.259
260	.350	.660	.353	.560	.290	.256
250	.263	.475	.275	.385	.1290	.242
240	.226	.475	.277	.312	.332	.251
230	.433	1.08	.585	.700	.591	.420
225	1.03	1.43	1.17	1.3	—	—

Chromostat II
System

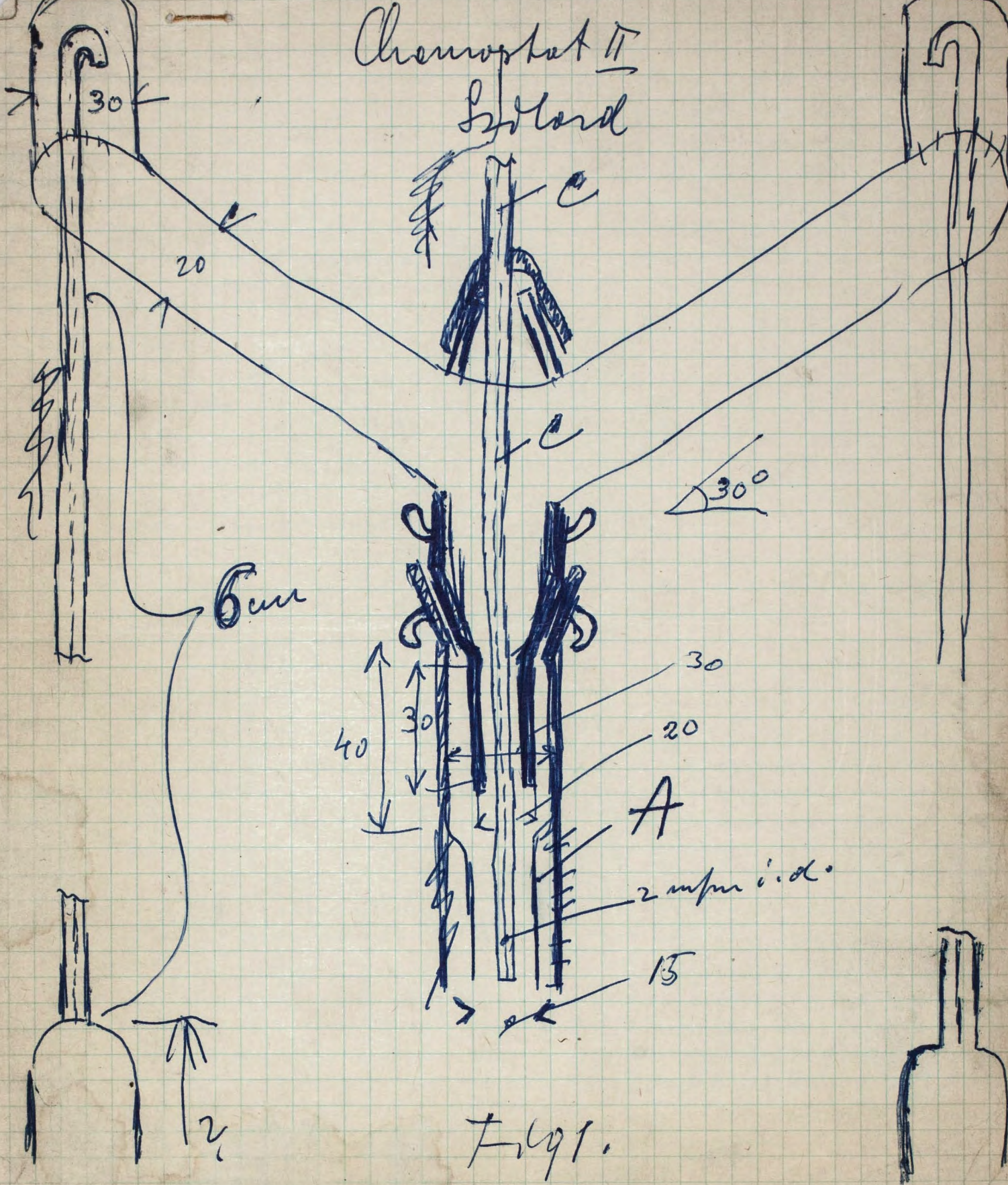
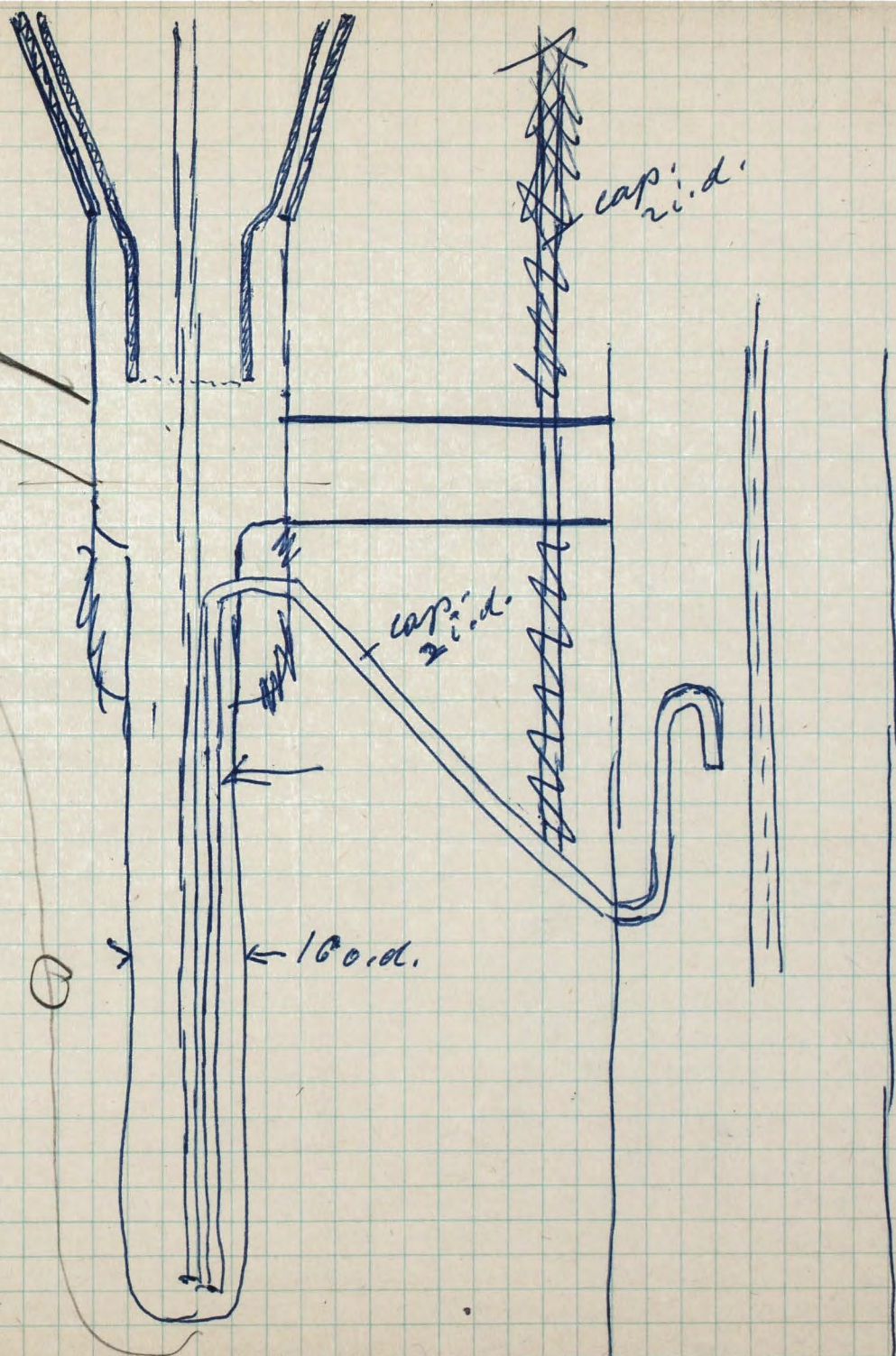
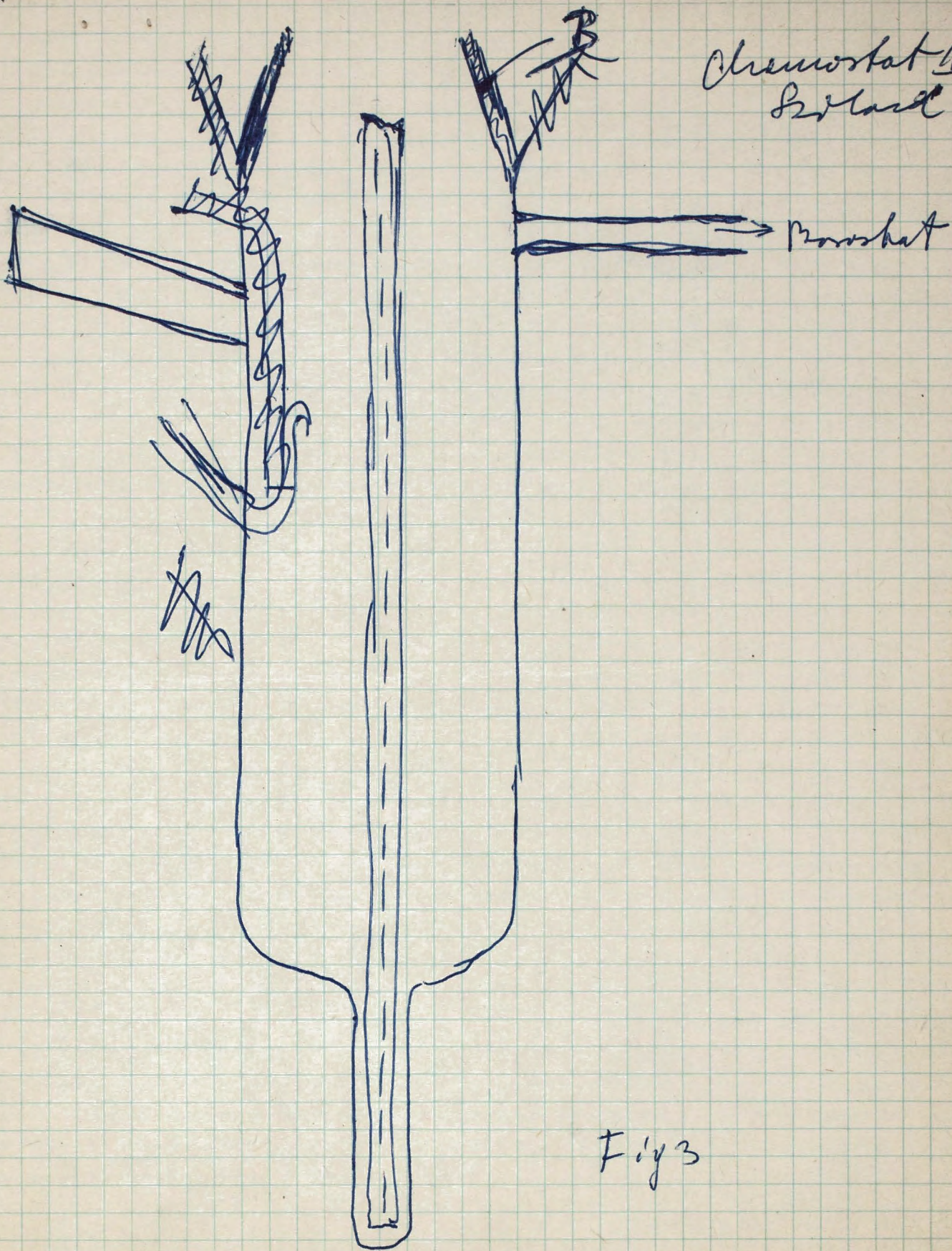


Fig 1.

Armed at II
for work

Fig 2





Chernostat II
Sokolov

Moshkat

Fig 3

Chernostat II
Sizland

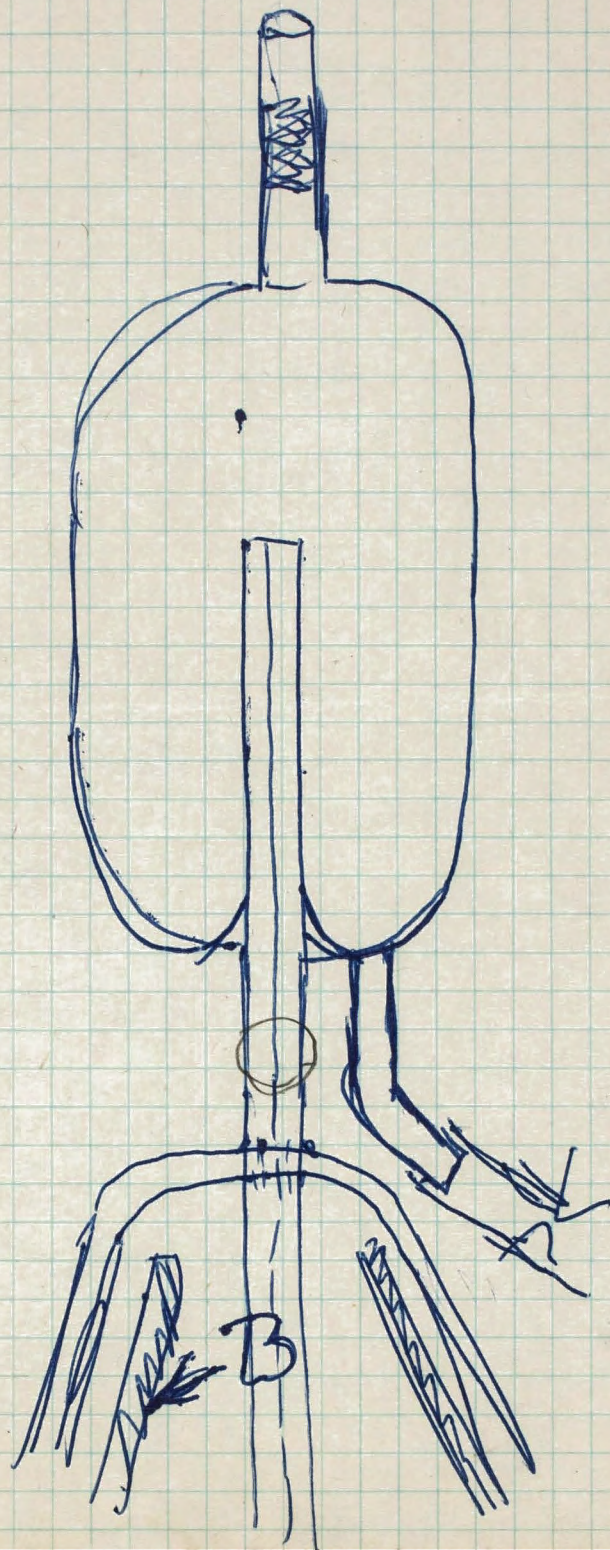


Fig 4

Chromostat &
Barometer

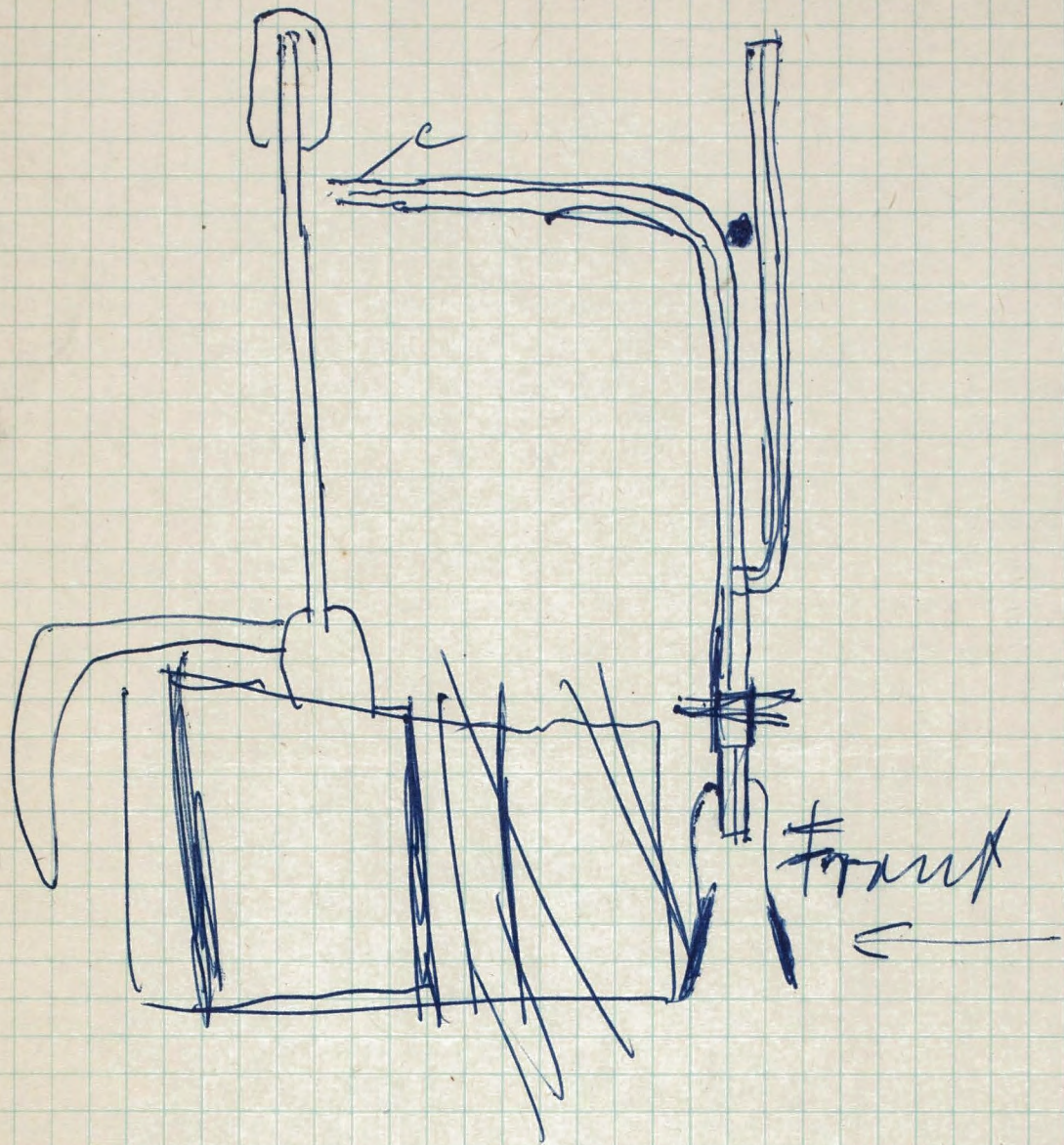


Fig 5.

Chemostat II
Lillard

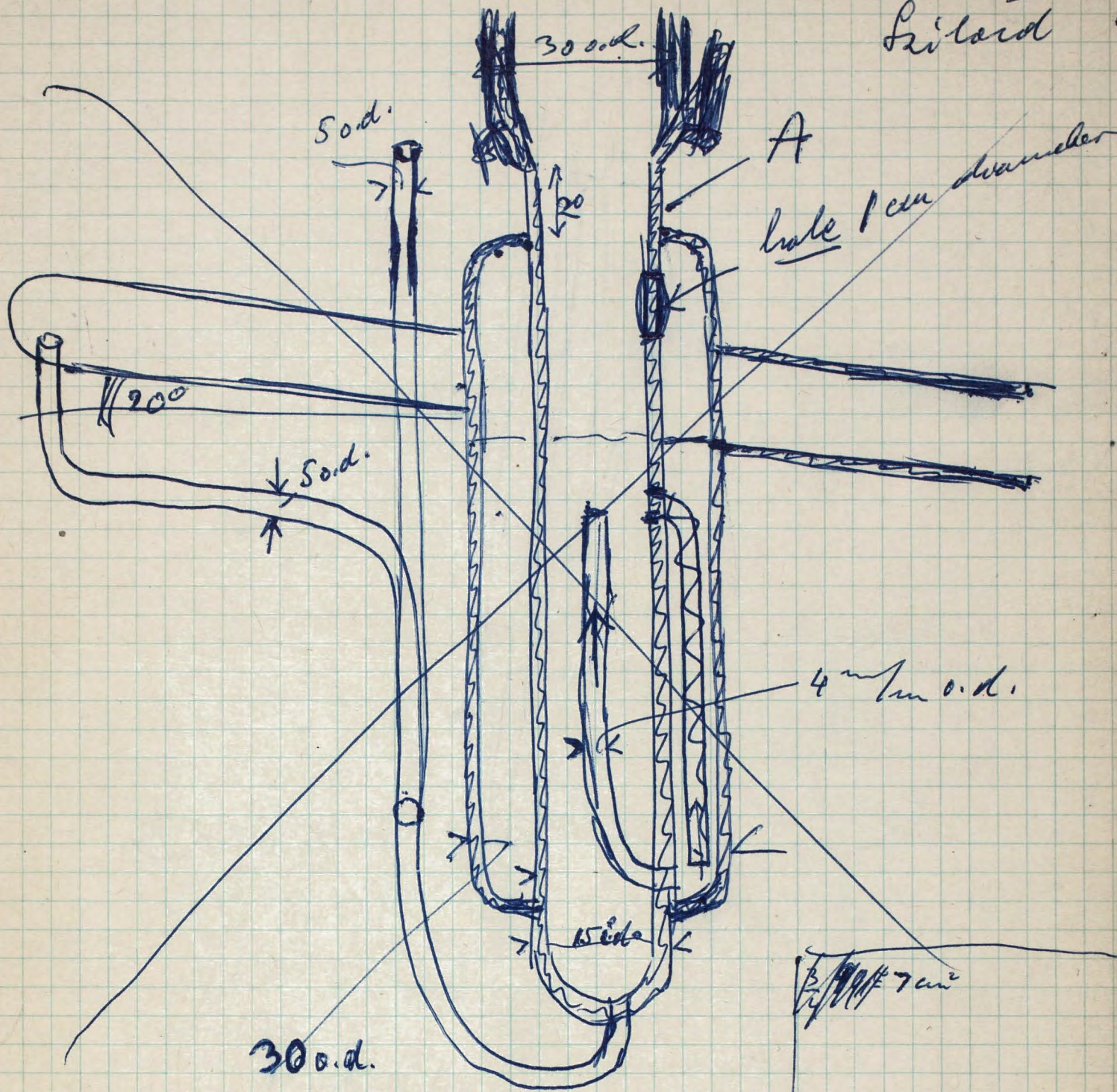
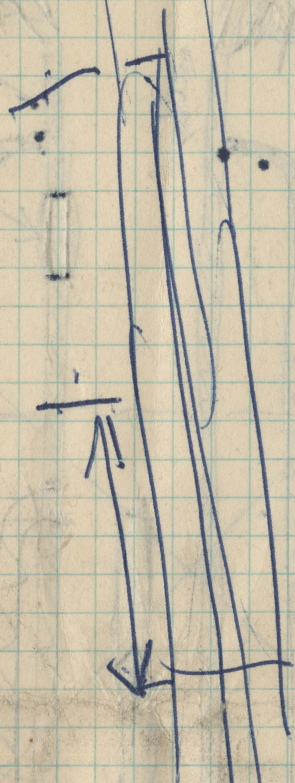
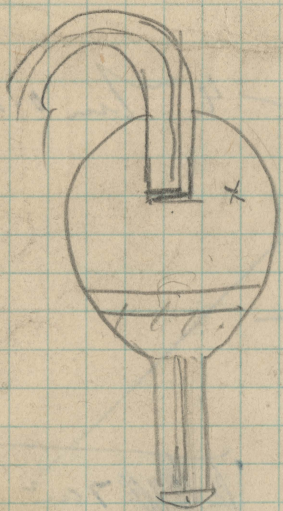


Fig 2



Brachy

R.F. Kimball

Oak Ridge

Nov 11 (1)

[Sanction Nov 11 4]

Bundled

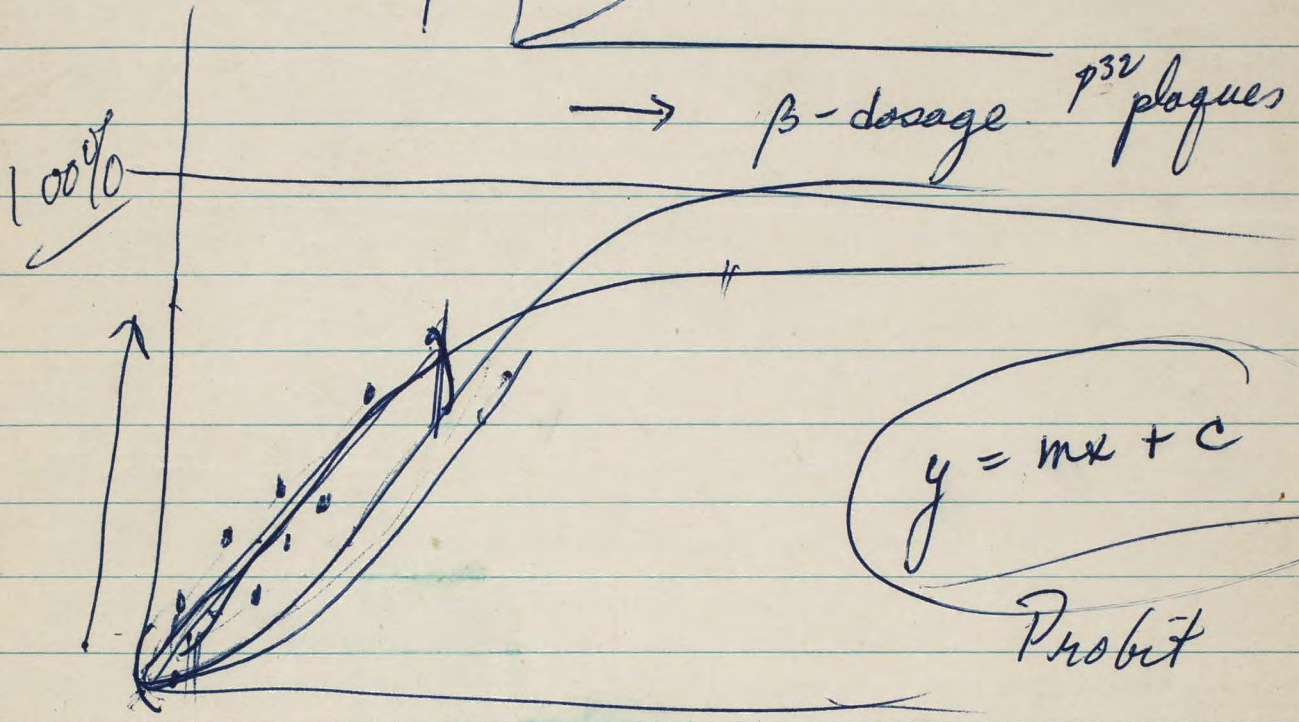
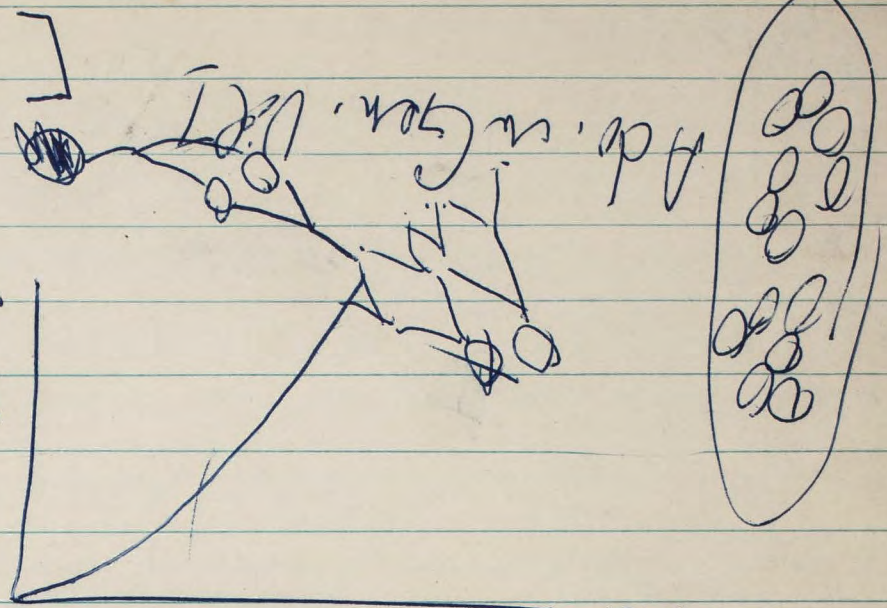
Person []

Spelman (bushu)

Parameter conditions

File
Standard

D. after out.



219
202

210x5
1050

(4)

24
192

up in new porce

125 x 40 x 40

Parameter (function)
10⁻⁷cc

Oct 27/48

: after ~~growth~~ mixed infection
 in B₁ (T₂ x T₄ cross) growth in B₂/₁, and
 then growth in B₄/₁, and last step is
 growth in B₄, then goes to T₁
 and plaque so obtained is "T₂" used
 in new cross with T₄

4.) Plating of ~~the~~ of plaque
 with lysate No. 1 from B₂ grown in B
 (plaque a) on various B₂ + B

~~4x6.~~ 4x6.) Plating of T₂ x T₄ cross on one
 B₂ + B (which registers plaque a!)

5.) Conditioning T₂ x T₄ cross by
 growing alternately in B₂ and B₄

death rate [and mut. rate] for ~~the~~
~~with X-rays~~ B and B₂ natural
 rate and X-rays.

Outis serum search for mutants
 which "escape" immunity

Phosphor

Inductively Standard

1000 ^{rpm} parts per million rpm water



Constant voltage devices

15, 30, 60, 120, 250, [110 and 6 Volts output]

Direct readings [Barrier layer cell]

~~balanced circuit~~ balanced circuit ~~is not~~ is ~~advantage~~ advantage
use of galvanometer below 500 ~~ohms~~ Ω

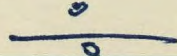
Lamp [6 Volt] Microscope illuminator

P.E. Platinum filament lamp

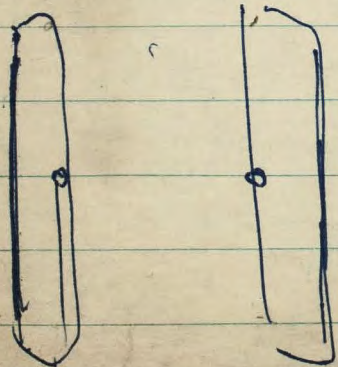
Solder lamp

Blue folder useful, to improve
scattered light

Galvanometer 3×10^{-7} Amp full scale
P.E. multiple reflection [243 p. 9] $\times 100$



Balanced circuit if constant
current constancy of lamp

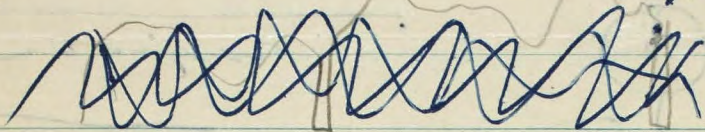


Resonating and outbeling.

~~But~~ border coils, balanced circuit, Brown recorder if sensitive enough

otherwise:

A.C. amplifier (chopped light) ~~with~~



Vacuum phototube or multiplier

If ~~A.C.~~ amplifier zero would shift this is no trouble for recording but detaches relays. - If DC amplifier is used but multiplier tubes, zero does not shift. -

UV measurement

Gen. El. 4 Watt germicidal Lamp + Ballast for it 50 μ P 25 with starter.

Gen El. -

use with multiplier

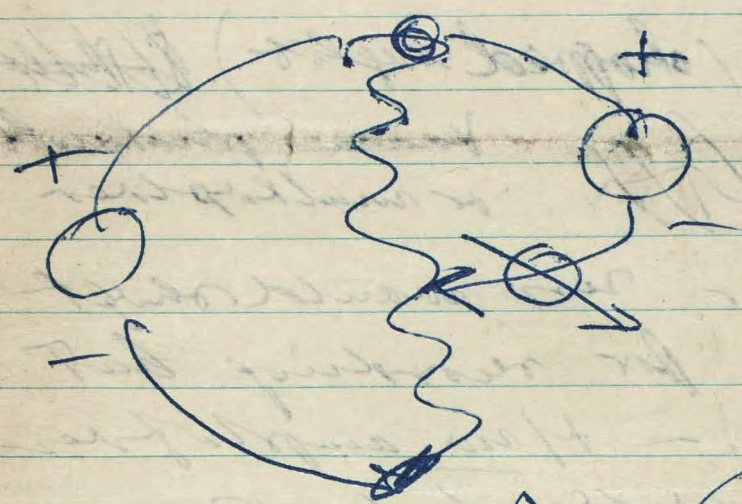
a.) ~~with~~ photomultiplier + galvanometer or

b.) also amplifier (A.C) from Lamp ~~with~~

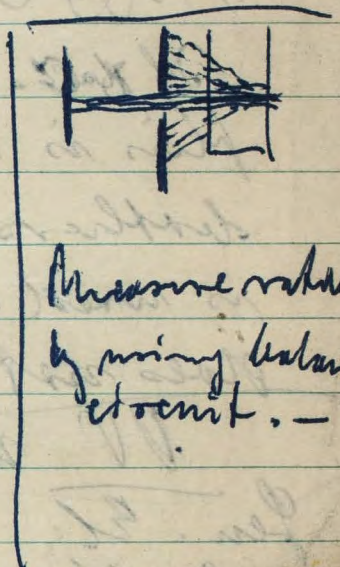
check lamp and amplifier with chopped ~~dividing~~ disk

by passing, H-Neg use Wood filter

if no amplifier is used for
multipliers use balanced circuit
to make sure that lamp fluctuations
are out,



General Radio }
Slide wire }
range 0-5

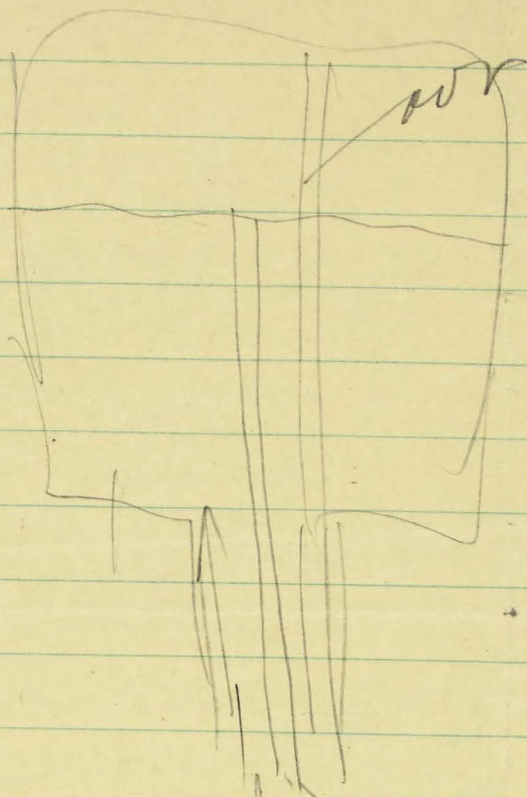


146 m
7 to 11 63 m
135 m

If U.V. lamp current through a.c.
amplifier (Baker) (ready at photo -
voltage) could be used current
enough to make such tube
luminous. Best unnecessary (X)
very well suited through
but gain does not change

Pyran
1 lbs 5 Gallons

Call 10-11
on Monday



15 mins

15 min
each

Add ^{Na} carbonate to get red Ca water

~~Add above acid Hydroxy acid to wash water~~

filter, wet cake Imps also alk
for 2 hours

filter again in alcohol
and heat to boil in alk.
filter ~~pyr~~ on wax paper

Klett & Funessee
Photo et Chemie.

Klett Wetz Co
N.Y.

Standard Li. Glass and
Optics Co N.Y.
automatic pipette

[
Frank [Essex] Conn. •
* Humphreys •
Wesley ~~Walt~~ •
Lansville Ky
Friday pm [Moreland]

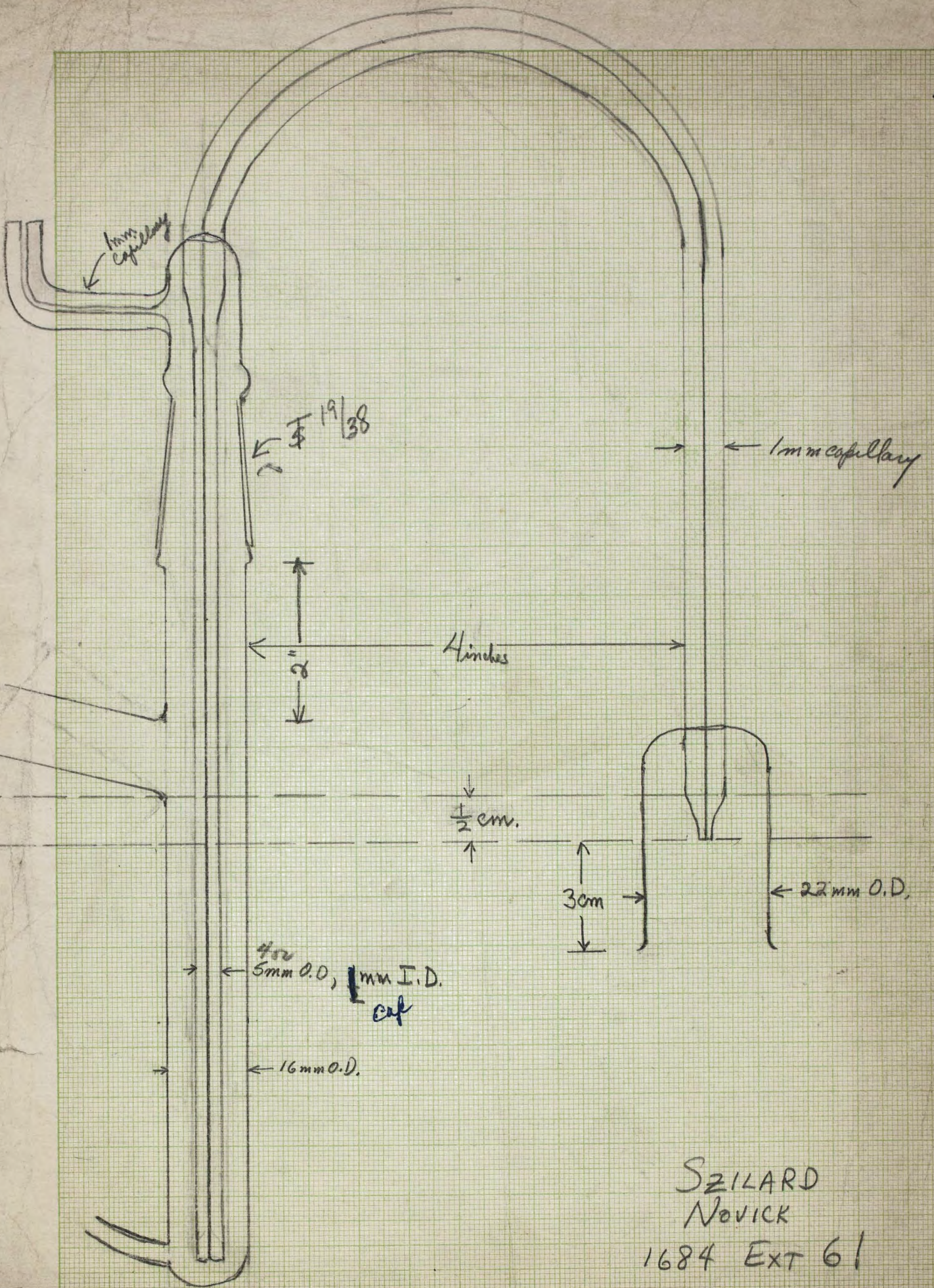
Woolley ^{Nov. Vol 67}
Proc. Am. Soc. Trop. Med. & Hyg. March 1946

~~Burnett~~ Austr. J. Exp. Biol. Med. Sci.
1937, 15, 227

Hark et al. J. Biol. Chem. 1946, 165, 241

Woolley & Green J. Prot. 1947 54 63

Green, Woolley
Journ of Exp. Med. [Dr. Winslow Fox]
1947, 86, 63
(430)



SZILARD
NOVICK
1684 EXT 61

Experiments

Oct 2, 1948

MS 42320
RID 3-3056

See in N.Y.

- 1.) Mark Udvaros *adversus*, *foldo*
- 2.) Davis - mutants that resist to wild type -
- 3.) Huddelberger about selection methods
- 4.) Withkin ~ Doremann [set B/r]
- 5.) Seymour Cohen in Philad.
- 6.) Stewart in Manchester ~ Bethe in "Hunca"

-
- 1.) cross T_2 and T_2h
grow in $B/2$ plate on $B + B/2$ see
if there is an increase in opaque plaques
"before" and after
 - 2.) also with T_1 cross T_1h

~~XXXXXXXXXX~~

3. (improvement $T_2 - s$) cross $T_2 \times T_4$
grow in $B/2$ plate in $B/4$. Prob
no plaques and mix virus of these.
Use this as T_2 and cross with T_4 ; determine
ratio of " $T_2 \times T_4$ and "growth" in $B/2$
- 3b.) refinement of the above (several
multiple infections in a $B/2$ plate.)