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# PREVALENCE OF COMMUNICABLE DISEASES IN THE UNITED STATES

December 1-28, 1940

The accompanying table summarizes the prevalence of eight important communicable diseases, based on weekly telegraphic reports from State health departments. The reports from each State are published in the Public Health Reports under the section "Prevalence of Disease." The table gives the number of cases of these diseases for the 4-week period ended December 28, 1940, the number reported for the corresponding period in 1939, and the median number for the years 1935–39.

#### DISEASES ABOVE MEDIAN PREVALENCE

Influenza.—For the 4 weeks ended December 28 there were 126,111 cases of influenza reported, the weekly incidence rising from 9,663 to 38,056 during this period. These figures may be compared with reports of from 4,000 to 7,000 cases per week during the corresponding period in 1939, and from 1,000 to 2,000 cases per week during preceding nonepidemic years.

The current epidemic started in the Mountain and Pacific regions, with a peak in reported cases in California during the week ended December 14 (13,133 cases). Other States in those regions followed with peaks in Washington, Oregon, Wyoming, and Nevada during the week ended December 21. By the end of the period (week ended December 28) the disease had spread into the South Central regions and practically every State reported the maximum weekly incidence This was also the case in a few States in the South up to that time. Atlantic and North Central regions. The epidemic had apparently not reached the New England and Middle Atlantic regions. place of origin and its travel from west to east the present epidemic resembles those of 1932-33 and 1928-29. The major epidemic of 1918-19 began in the Northeast and traveled generally to the west and south, as have several of the succeeding minor epidemics.

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Later reports (week ended January 4, 1941) show a still further decline in the number of cases in the Mountain and Pacific regions, but each State in the South Central regions except Louisiana reported the highest weekly incidence up to that date; of approximately 56,500 cases reported from those regions, Texas reported 32,983, Kentucky, 9,601, and Arkansas, 6,516 cases. Virginia and South Carolina, in the South Atlantic region, continued to report an increase in the weekly incidence. The total number of cases reported for the week was 77,144.

Mortality records indicate that the cases have been of a mild type. A rise in the mortality rate from all causes in 92 large cities began in the latter part of November and while the weekly rates were slightly higher than those for the corresponding weeks in 1939 they compared very favorably with the preceding 3-year average rates. During the week ended December 21 when there were approximately 42,500 cases of influenza reported, the death rate for large cities was 12.2, as compared with 11.7, 12.0, and 12.0 for the corresponding week in 1939, 1938, and 1937, respectively; for the following week, when approximately 38,000 cases of influenza were reported, the death rate was 12.5 as compared with 12.4, 12.8, and 13.2 in the three preceding years.

Measles.—The number of cases of measles rose from approximately 13,000 during the preceding 4-week period to approximately 24,000 for the 4 weeks ended December 28. The number of cases was more than twice the number reported for the corresponding period in 1939, which figure (11,035) represented the 1935–39 median incidence for this period. The disease was unusually prevalent in the Middle Atlantic and North Central regions, only slightly above normal in the South Central regions, and comparatively low in all other regions. In the Pacific region the incidence of measles was unusually high during the years 1938 and 1939, but during the current year the number of cases has compared more favorably with the average of preceding normal years.

Poliomyelitis.—For the 4 weeks ended December 28 there were 260 cases of poliomyelitis reported, as compared with 265, 76, and 134 cases for the corresponding period in 1939, 1938, and 1937, respectively. While the incidence of this disease declined rapidly during the current period in all sections of the country, in the East North Central and South Atlantic regions, where the disease has been unusually prevalent, the incidence is still considerably above the normal seasonal expectancy. Each of the States in the East North Central region reported a relatively high incidence, while Virginia and West Virginia seemed mostly responsible for the excess in the South Atlantic region.

Number of reported cases of 8 communicable diseases in the United States during the 4-week period, Dec. 1-28, 1940, the number for the corresponding period in 1939, and the median number of cases reported for the corresponding period 1935-39

Division	Cur- rent period	1939	5-year me- dian	Cur- rent period	1939	5-year me- dian	Cur- rent period	1939	5-year me- dian	Cur- rent period	1939	5-year me- dian
	D	iphthe	ria.	In	fluenza	,1	N	1easles	; 2		ingoco eningi	
United States	1, 369	2, 355	2, 788	126, 111	23, 874	7, 736	23, 776	11, 035	11, 035	115	132	317
New England Middle Atlantle East North Central West North Central South Atlantic East South Central West South Central West South Central Mountain Pacific	17 173 205 82 321 146 304 46 75	28 271 378 135 658 298 384 88 115	52 349 477 -206 658 298 406 88 156	2, 318 33, 612 30, 401	113 337 542 10, 659 2, 950	494 316 2, 007 823 2, 554	9, 735 7, 626 1, 409 922 858	274 629	2, 849 1, 492 876 962 264 369 812		5 40 15 6 25 17 7 8	12 54 42 18 57 37 31 8
	Pol	iomyel	itis	Sca	rlet fev	er	s	mallpo	ox	Typhe typ	oid and hoid fe	l para- eve <b>r</b>
United States	260	265	201	11, 519	14, 672	17, 630	220	414	711	426	473	516
New England Middle Atlantic East North Central West North Central South Atlantic East South Central West South Central Mountain Pacific	1 12 110 39 39 12 14 9	10 35 23 50 24 18 21 40 44	2 9 23 19 22 18 21 5 34	858 2, 525 3, 722 1, 352 1, 148 730 362 332 490	4, 702 1, 852	3, 393 5, 623 2, 955 1, 246 588 725 735	0 79 107 1 0 13	0 48 165 4 3 57 117 20	0 79 233 4 2 32	87 48 84 26	18 69 68 22 89 25 115 33 34	89 78 55 92 39

 $<sup>^{\</sup>rm I}$  Mississippi, New York, and Pennsylvania excluded; New York City included.  $^{\rm I}$  Mississippi excluded.

#### DISEASES BELOW MEDIAN PREVALENCE

Diphtheria.—Diphtheria again registered a record low level. The 1.369 cases reported for the 4 weeks ended December 28 was the lowest recorded number for this period in the 12 years for which these data are available. The current incidence compares with a total of 2,355 cases for the corresponding period in 1939 and 2,788 cases for the same period in 1938. Each section of the country reported the lowest incidence in recent years.

Meningococcus meningitis.—The relatively low incidence of meningococcus meningitis which has prevailed throughout the year was maintained during the current 4-week period. The number of reported cases (115) was less than 90 percent of the number recorded for the corresponding period in 1939 and slightly more than one-third of the 1935-39 median figure for the period. The incidence was relatively low in all sections of the country; in the Middle Atlantic and South Atlantic regions the incidence was the lowest reported during this period in 10 years.

Scarlet fever.—The incidence of scarlet fever was the lowest recorded for this period in the 12 years for which these data are available.

number of cases (11,519) reported for the current 4-week period was less than 80 percent of last year's figure for the corresponding period and about 65 percent of the 1935-39 median incidence for the period. In the East South Central region the number of cases was slightly above the seasonal expectancy, but in all other regions the incidence was comparatively low.

Smallpox.—The incidence of smallpox was also relatively low. The total of 220 cases reported for the current 4-week period compared with 414, 711, and 1,338 cases reported during the corresponding period in 1939, 1938, and 1937, respectively. The situation was favorable in all sections of the country. While the number of cases reported in the East North Central section was considerably above that of last year, it stood at the 1935–39 median level.

Typhoid fever.—Typhoid fever also reached a new low level. The number of cases (426) reported for the 4 weeks ended December 28 was 10 percent less than the comparatively low incidence in 1939 and about 20 percent below the preceding 5-year median for this period. The only exception to the favorable situation was in the East South Central region, where an increase of approximately 25 percent over the seasonal expectancy was shown.

#### MORTALITY, ALL CAUSES

The average mortality rate from all causes in large cities for the 4 weeks ended December 28, based on data received from the Bureau of the Census, was 12.3 per 1,000 inhabitants (annual basis). The average rate for the corresponding period in 1937–39 was 12.2. Apparently the reported cases of influenza have been mild, for the death rate has been affected very little, if any, by the current epidemic.

# A STUDY OF CERTAIN FACTORS WHICH INFLUENCE THE DETERMINATION OF THE MOUSE PROTECTIVE ACTION OF MENINGOCOCCUS ANTISERUM <sup>1</sup>

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Following Miller's (1) introduction of mucin to increase the lethal action of Neisseria meningitidis for mice there have been a number of reports which demonstrate the mouse protective activity of antimeningococcus serum. Reference to these reports was made in a previous paper (2). Furthermore, it has been shown that the mouse protective activity of the serum is associated with the specific antibody. In 1937.

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Rake (3) reported that the Group I 2 protective power of antimeningococcus serum probably parallels the homologous specific antibody nitrogen content. Pittman, Branham, and Sockrider (4) showed by means of the "plate precipitation" reaction a close correlation between the amount of precipitation and the protective power of the antiserum. Subsequently, Rake and Scherp (5), using sera the precipitin content of which had been measured quantitatively by Scherp (6), demonstrated further the relation between the specific immune nitrogen content of the serum and the mouse protective capacity of the serum. absorption of the sera with specific polysaccharide removed 90 to 99 percent of the protective antibodies. In addition, Alexander (7) has presented findings which suggest that there is a broad correlation between the mouse protective activity and the therapeutic value of meningococcus antiserum. Recently, McLeod (8a) and Branham (8b) presented evidence that the recovery of several patients suffering from Group II infections was influenced by treatment with serum containing homologous specific antibodies.

At present it seems that the best method available for evaluation of meningococcus antiserum is the mouse protection test. It is evident that the agglutinin titer  $(56^{\circ}\text{C.})$ , which is now the criterion for approval, bears little relation to the protective action, excepting the fact that high protective action is not found when the agglutinin titer is low. The quantitative measurement of the specific immune nitrogen (6, 9) offers certain possibilities but there are two objections to this method. One is that carefully purified and standardized precipitinogens, which are necessary for the measurements, are not readily available; and the other is that an occasional horse serum contains a greater amount of precipitable immune nitrogen than mouse protective antibodies (4). The same objections are applicable to the turbidimetric method recommended by Little (10). The second objection is also applicable to the "plate precipitation" reaction.

In the performance of the mouse protection test, there has been lack of uniformity of techniques employed in the various laboratories. Consequently, the results obtained have not always been comparable. It therefore seemed desirable before adopting this method to make a detailed study of the different variables in order that the best procedures might be selected for the standard method. The results of the study are reported in this paper. It includes (a) the selection of a medium suitable for the maintenance of the virulence of the culture, (b) a comparison of the susceptibility of different cultural strains to antiserum, preparatory to the selection of a test culture, (c) a study of the susceptibility of mice as influenced by breed, sex, and weight,

<sup>&</sup>lt;sup>2</sup> In this paper the word "Group" is used to designate immunological subdivisions of meningococci on the basis of soluble specific substances (probably capsular substances). In Group I are included those strains known as Type I and Type III in the Gordon-Murray classification, and in Group II only those strains known as Type II in the same classification.

(d) a comparison of different solutions for suspension of the culture, (e) a study of mucin in relation to variability of different lots and methods of preparation, and (f) the selection of the time interval between inoculation of serum and culture.

Material and methods.—The general procedures for the plate precipitation test, for the mouse protection test, and for the Reed-Muench method (11) of calculating results of the protection test have been described in a previous paper (4). Certain changes which were made in these procedures are as follows: In the precipitation test, 0.8, 0.4, and 0.2 cc. amounts of serum were each added to 16 cc. of melted agar in place of 1.0, 0.5, and 0.2 cc. amounts. Neopeptone solution was used for the first suspension of the culture instead of Ringer's solution.

Mice were inoculated with 0.5 cc. of the dilutions of serum; hence the actual amount which a mouse received was one-half of the amount in 1 cc. of the designated dilution.

Group I meningococci have been largely used throughout the study. In mouse protection tests, Strain No. 1027 was used unless otherwise indicated.

The media, cultures, mice, and mucin are described below under the respective headings.

#### CULTURE MEDIA

Many media have been used by different workers in the cultivation of meningococcus. In the majority of instances, however, the selection has been made largely on the basis of multiplication and longevity of the bacteria and not maintenance of virulence. In the performance of the mouse protection test it is essential that the cultural inocula contain uniformly highly virulent bacteria. We have studied the influence of eight media on the virulence of meningococci. Observations were made also for any change in agglutinability and in precipitation of immune serum in agar plates. Other media might well have been included in the study. The purpose was not, however, to determine how many media are suitable for maintenance of virulence but to select one which could be relied upon with some degree of certainty. The media which were employed are listed in table 1.

The cultures grown on these media had not previously been subjected to artificial cultivation. They were started on the different media either directly from spinal fluid or from a primary culture grown from spinal fluid. There were eight strains, all of Group I. No strains of Group II meningococcus in spinal fluid were available at the time for study. As quickly as possible after isolation, subcultures of each strain were stored in the ice box in the lyophile state. Cultures so preserved apparently retain their original characteristics. The dried cultures were used as controls in observing the changes in the bacteria after they had been maintained on the different media. The examinations were made after about 6 months and again after another 6 months.

The virulence of a culture was determined by inoculating mice with dilutions of the culture, in mucin, ranging from 10<sup>-3</sup> to 10<sup>-4</sup>. The highest dilution which killed more than half of the mice was considered the measurement of the virulence

of the culture. The original virulence of all eight cultures was  $10^{-8}$  or  $10^{-9}$ ; cultures which killed in dilutions between  $10^{-5}$  and  $10^{-7}$  will be reported as decreased in virulence; and cultures which failed to kill in the  $10^{-5}$  dilution will be reported as avirulent.

The agglutinability of a culture was tested with Types I and III and polyvalent (M17) antisera. The test tubes were incubated at 56° C. for 20 hours. The ability to precipitate immune serum was tested on agar plates containing 5 percent of serum M17.

The results of the study are summarized in table 1. On all eight media, the majority of the cultures remained highly virulent for 6 months. In this respect, however, the solid media were superior to the semisolid. Blood agar was the only one on which the same cultures showed no loss in virulence during this period. Yet on each of the other solid media, serum glucose, EDB (12), and egg, there was only one culture which showed any decrease in virulence. Furthermore, on each of these three media the meningococci remained alive longer between transfers than on blood agar. The egg slant was most favorable in this respect as the transfers were made only every 21 days. Although EDB is quite favorable for maintaining virulence, the difficulty of preparation makes it impractical for routine use.

Of the solid media, serum glucose was the only one on which maintenance was continued for 12 months. During the second 6-month period five cultures which had shown no change in virulence were carried. All decreased in virulence but none became avirulent. In the use of serum glucose agar we had been previously somewhat prejudiced owing to an immunological change in a Type IV culture when it was grown on this medium (15). With these eight Group I strains, however, there were no apparent immunological changes.

Of the whole, semisolid media were much better than the solid for maintaining the longevity of the cultures but they were inferior in maintaining virulence. The loss in virulence, however, was not rapid. More than half of the cultures after a year on the semisolid media killed mice in a dilution of  $10^{-5}$  or higher. Five were as virulent as when originally isolated. The latter had been grown on Hitchens' medium. One had been transferred every 21 days and the other four had been transferred every 2 or 3 days. Three of the four, however, had lost their ability to remain in suspension in saline.

From the work of others it appears that the rapidity with which a culture loses certain of its characteristics may be influenced by whether or not it has been previously subjected to artificial cultivation. Kirkbride and Cohen (16) compared the influence of serum glucose and Hitchens' media on the precipitative activity of cultures on immune serum agar plates. If stock cultures, routinely maintained on serum glucose, were grown on Hitchens' medium, there was a rapid loss in ability to precipitate immune serum. On the other hand, if recently isolated cultures were grown on Hitchens' medium,

TABLE 1.—Changes in 8 strains of meningococcus after being maintained on 8 media

					-		
Medical			6 months			12 months	
Michigan	ransier mervar	Virulence	Agglutination Precipitation	Precipitation	Virulence	Agglutination	Precipitation
5 percent rabbit blood agar slant.	24 or 48 hours	(6 unch)	}6 unch	6 unch	Discontinued		
5 percent horse serum 0.5 percent glucose agar slant	do	(6 unch	6 unch	} unch	(3 discontinued.	6 unch	5 un <b>ch.</b>
Dorset's egg slant	21 days	(6 unch	6 unch	5 unch1	Discontinued		
EDB (4 percent extract) agar slant (1?)	24 or 48 hours	(6 unch 1 decr	}7 unch	7 unch	Discontinued		
EDB (4 percent extract) 0.15 percent agar	21 days	6 unch 1 decr 2 avir	8 unch	8 unch	Discontinued		
	48 or 72 hours	6 unch	2 unch	8 unch	4 unch	2 unch	}8 unch.
percent NaCl) (18).	21 days	(7 unch	7 unch	}8 unch	1 unch 6 decr 1 avir	6 unch	}8 unch.
Special 0.15 percent agar (beef infusion, 0.15 percent agar, 0.5 percent NaCl, 0.02 percent KCl and 0.01 percent CaCl;) (14).	do.	5 unch 2 decr 1 avir	0 unch 2 sp	}8 unch	(3 decr.	6 unch. 2 sp.	}8 un <b>eh.</b>
Plain 0.15 percent agar (beef infusion, 0.15 percent agar, 0.5 percent NaCl).	do.	5 unch 1 decr 2 avir	7 unch	}8 unch	{	6 unch 2 sp.	}7 unch.
Task Hashenson Dilition of 10-1 to 10-1 billed more than held of more	If of mice						

Unch.=Unchanged—Dilution of 10-4 or 10-7 killed more than half of mice.

— Specis—Decensed—Highest dilution which killed more than half of mice was between 10-4 and 10-7.

Aptir = Aytulent—Dilution of 10-4 falled to kill mice.

Sp. = Spontaneous.

there was less change in this activity. Cohen (17) extended the study and reported that the virulence of a stock culture, routinely maintained on serum glucose, was definitely decreased when grown on Hitchens' medium.

It is of interest to note, under the conditions of our experiments, that there was no correlation between spontaneous agglutination and virulence of a culture. At the time of the first testing of the cultures, 12 were agglutinated spontaneously; 9 of these were highly virulent, 1 was moderately virulent, and only 2 were avirulent. At the time of the second testing, the same number were agglutinated spontaneously; 5 of these were highly virulent, 4 were moderately virulent, and 3 were avirulent. On the other hand, 12 cultures which were avirulent showed no spontaneous agglutination.

There was also no correlation between virulence and precipitation of immune serum in agar with the exception of one culture from the egg medium which was avirulent and caused no precipitation. Cultures which killed mice in dilutions of  $10^{-9}$  produced no better halos than cultures which failed to kill mice in dilutions of  $10^{-5}$ . No doubt, if the cultures had been completely avirulent, a difference would have been observed.

From the viewpoint of simplicity of preparation and preservation of virulence, our results indicate that, of the media studied, the rabbit blood agar and the serum glucose agar are the most satisfactory for the routine maintenance of cultures for the mouse protection test. We have chosen the blood agar. Transfers are made daily or every 2 days. In order to be absolutely certain that the cultures are kept at maximum virulence, they are passed through mice about every 2 weeks. The bacteria are recovered from the peritoneal exudate. In case a culture is to be used only occasionally, it is preserved in the lyophile state. Before using this culture, which has been resting, several daily or twice daily transfers are made on blood agar in order to restore its original vigor.

#### THE TEST CULTURE

In order to select a suitable strain of Group I meningococcus for measuring the mouse protective potency of antiserum, a number of strains were studied. It was found that, although of comparable virulence, the strains varied considerably in their susceptibility to antiserum. In some instances more than twice as much serum was required to protect mice against certain cultures as against others. This finding is in accord with data recently published by Branham (18).

Table 2.—The determination of the protective value of 3 sera with 2 strains of meningococci of unequal susceptibility to antiserum

No. of culture	Dilution of culture (1.0 cc.)		dilution of ich protected of mice	Potency— percentage of control, M19
1027 1041	2. 5×10 <sup>-4</sup> 2. 5×10 <sup>-4</sup>	No. 1 1:128 1:200	M19 1:75 1:107	170 187
1027 1041	2. 5×10 <sup>-4</sup> 4×10 <sup>-4</sup>	No. 2 1:335 1:460	1:135 1:240	174 191
1027 1041	2. 5×10 <sup>-4</sup> 3×10 <sup>-4</sup>	No. 3 1:124 1:170	1:154 1:215	80 79

Although our cultures were not equally susceptible, each usually reacted similarly to all antisera, so that when the potency of the unknown serum was expressed in the percentage value of the control, comparable values were obtained for the serum. These findings are illustrated in table 2. In the experiments two cultures, 1027 and 1041, were used in determining the potency of three antisera.

In each instance more of the unknown serum was required to protect against 1027 than against 1041. Likewise, corresponding amounts of the control serum M19 were required to protect against the respective strains. Hence, comparable values for the potency of each serum were obtained with both cultures. Similar results were obtained with several other cultures; dissimilar results were obtained with only one culture.

Our results indicate that a number of strains of Group I could be used with equal success in evaluating the mouse protective action of antisera. It is essential, however, that the culture be of highest virulence. Cultures only moderately virulent induce irregular results.

The culture selected for the mouse protection test.—Culture 1027 was selected for determining the potency of the antisera against Group I meningococcus. This culture grows well on artificial media, its virulence is such that less than 10 organisms in mucin are lethal for a mouse, and mice are protected against many lethal doses of it with Group I specific immune serum. Furthermore, the protective value of a serum obtained by the use of this culture, in almost every instance, is definitely correlated with the amount of specific antibody as measured by the plate precipitation reaction. In 1938, Pittman, Branham, and Sockrider (4), using this strain, reported a correlation between mouse protection and precipitins. This study has been continued and at present more than 100 different sera have been tested. Quly a few have shown a discrepancy.

#### MICE

The results of the study of influence of breed, sex, and weight of mice on the determination of the mouse protective action of antimeningococcus serum are given below.

Breed.—Mice of different strains and from different colonies of one strain were employed in the study. The most important factor seemed to be that the mice came from a pure, closely inbred line. different strains did vary in susceptibility to meningococci but mice from different colonies of the same strain likewise varied. difference in mice from the same strain is illustrated in table 3. mice were from two colonies bred at the National Institute of Health (NIH) and from a dealer; all were of Swiss origin, the stock of each line originally coming from the same source. It may be seen that about 35 percent less serum was required to protect the mice from the dealer than those from the NIH colony No. 1; and about 40 percent less to protect mice from the NIH colony No. 2 than those from the NIH colony No. 1.

TABLE 3.—The difference in susceptibility of Swiss mice from 3 colonies

Experiment No.	endpoint of	50 percent antiserum 19)	Percent varia- tion	Experiment No.	Calculated endpoint o (M	Percent varia- tion	
1	NIH (No. 1) mice 1:160 1:100	Dealer mice 1:215 1:135	34 35	3	NIH(No.1) mice 1:84 1 1:50	NIH(No.2) mice 1:117 1 1:72	39 44

Serum No. 5.
 Male mice, 16-18 grams, were used in experiments 1, 3, and 4; male and female mice, 18-19 grams, were used in experiment 2.
 3.5 and 4.25 percent mucin suspensions were used in experiments 1 and 2, and 3 and 4, respectively.

These results show some of the variations in susceptibility that may arise in mice of the same strain. It is obvious that in order to obtain consistent results in mouse protection tests only mice from a pure, closely inbred line should be used.

Sex.—In table 4 are given the results of six protection tests in which the susceptibility of male and female mice was compared. Five tests were made with a Group I culture and one with a Group The mice were of comparable weights, between 16 and The males were about 5 weeks old and the females were 18 grams. approximately a week older.

It is shown that on an average about twice as much serum was required to protect the females as the males—in three instances twice as much was required, in one almost three times, and in another a little less than twice. It can be readily seen how irregular results may arise from the use of mice of both sexes if they are not equally distributed.

Table 4.—The influence of sex on the protective action of meningococcus antiserum

Group of culture	Dose of culture	Concentra-	Serum No.	Calculated 50 point of	percent end- serum with
	(1.0 cc.)	(percent)		Male mice	Female mice
I (1027)	1x10 <b>→</b>	3. 5	M19 No. 6	1:160 1:103	1:80 <1:40
	1x10→	3.5	M19 No. 7	1:144 1:120	1:91 1:43
	2x10-4	4. 25	M19	1:105	1:48
II (963)	2x10-4	4. 25	M19	1:13. 5	1:6. 5

Weight of mice 16-18 gm.

Mucin No. 35,689, plus 0.5 percent NaCl and 0.5 percent dextrose.

Weight.—The influence of the weight of mice on the determination of the protective action of antiserum is illustrated by two experiments, the results of which are recorded in table 5. In the first experiment, mice weighing 15 to 17 grams were compared with mice weighing 21 to 23 grams. The calculated dilutions of serum which protected 50 percent of the mice were 1:138 and 1:168, respectively. In the next experiment 15 to 16 gram mice were compared with In this instance with the wider variation in 22 to 24 gram mice. weight there was a corresponding variation in protection. percent protection endpoints of the serum were 1:156 and 1:308, respectively.

TABLE 5.—The influence of weight on the protective action of meningococcus antiserum

Waisht of miss (seems)		Diluti	on of se	rum M	19-0.5	cc. ino	culum		Calculated	
Weight of mice (grams)	1:50		1:100		1:,200		1:400		5C percent endpoint	
15 to 17	6S 6S		3S 6S	3D	28 28	4D 4D	18	5D 6D	1: 138 1: 168	
15 to 16	4S 68	2D	58 48	1D 2D	28 48	4D 2D	2S 4S	4D 2D	1: 156 1: 308	

Equal number of males and females inoculated with each dilution of serum.

Test dose of culture = 1 cc. of 2.5×10<sup>-4</sup> dilution:

Mucin No. 35,689, 3.5 percent concentration, plus 0.5 percent NaCl and 0.5 percent dextrose.

The results of the work with mice show that inherent susceptibility. sex, and weight are influential factors in determining the protective action of antiserum. Variations in any one of the factors may give rise to irregular results. By controlling these variables, however, we have found that it has been possible to duplicate the results of a mouse protection test with as few as 24 or 30 mice for a serum.

In the use of mice, one other factor should be mentioned. It is environmental temperature. We have noted that during the hot summer months in Washington mice are much less resistant to meningococci than during the winter. Dilutions of serum which will protect

half of the mice against a given dose of culture during the winter fail to protect any against the same dose during hot weather. Furthermore, the survival time after infection is also decreased. In order to obtain results in the summer comparable to those obtained in the winter, it is necessary to use a smaller test dose of culture or to decrease the concentration of mucin. Colvin and Mills (19), in work with controlled environmental temperatures, have demonstrated that mice kept at 90° F. are less resistant to hemolytic streptococci than mice kept at 65° F.

#### SOLUTION FOR SUSPENSION OF CULTURE

In preparing a culture for the mouse protection test it is the general practice to suspend it first in a clear solution in order that an estimate of the number of organisms can be made by comparing it with a turbidity standard. A number of solutions have been employed by different workers. The list includes distilled water, saline, Ringer's, Locke's (plus 0.1 percent gelatin), and meat infusion broth. We have compared the influence of each of these, with the exception of the broth, on the viability of meningococci. Solutions of proteose peptone and of neopeptone were also included. The latter solutions contained 1 percent peptone and 0.5 percent sodium chloride.

The procedure for the test was as follows: A suspension of a 5-hour-old blood agar culture was prepared in neopeptone solution. Its density corresponded to that of 500 parts per million of silica (20). One cubic centimeter of the suspension was added to 4 cc. of each of the solutions to be studied. These dilutions were designated as  $1\times10^{-1}$ . Serial dilutions  $\times$  10 were made as quickly as possible to  $1\times10^{-3}$  in the respective diluents. Beginning with the  $1\times10^{-5}$  dilution, 0.2 cc. of each was cultured in special 0.15 percent agar simultaneously with the making of the dilution. After an interval of 30 minutes and again after 2 hours subcultures were made from the dilutions. The growth which took place in the subcultures was read after 24 hours of incubation; the tubes were examined again after 72 hours in order to make sure that no growth had occurred in the tubes that had been negative on first reading.

After the last subcultures had been made from the different dilutions, all of the latter tubes were placed in the incubator for 24 hours, after which time 1 loopful was cultured on a blood agar plate.

The results of one experiment in which distilled water, saline, Ringer's, and neopeptone solutions were used are given in table 6.

From the dilutions in neopeptone solution, comparable amounts of growth were obtained in the subcultures made simultaneously with the dilution of the culture and 30 minutes and 2 hours after diluting. This solution was apparently in no way injurious to the bacteria. Moreover, multiplication took place in each of the original dilutions.

On the other hand, in the presence of each of the other three solutions, there was rapid dying of the bacteria. Distilled water was the most injurious, saline was less, and Ringer's was least. But even

in the presence of the latter there was marked reduction in the number of bacteria within 30 minutes. Furthermore, in certain other experiments the number of bacteria recovered in the subcultures from Ringer's solution made immediately after diluting were less than from corresponding dilutions in neopeptone solution.

Table 6.—The survival of meningococci in distilled water, saline, Ringer's, and neopeptone solution

<b>-</b>	Time interval		Gr	owth of s	ubcultur	es from d	lilutions		
Diluent	between diluting and subculturing	10 -1	10 -2	10 -3	10 -4	10 -5	10 →	10 -7	10 -
Distilled water	Simultaneous	++++	++++ +++ -	++	= (1)	+;+ - -	_(7) 	.(1) _ _	17117
0.85 percent NaCl	Simultaneous  30 min  2 hrs  24 hrs.*		++++ ++++ -	++++ ++ -	+++ (1)	+++ (11) - -	++ (10) - -	(4) (1) -	=======================================
Ringer's	Simultaneous 30 min 2 hrs 24 hrs.*	 ++++ ++++	++++ ++++ +	++++ ++ -	++ + -	++++ (10) - -	++ (3)	+ (1) -	(3) 
Neopeptone	Simultaneous	++++	++++	 ++++ ++++	++++ ++++ ++++	++++ ++++ ++++ ++++	++ ++ ++ +++	+ + + +++	(7) (3) (8) +++

= Number in parentheses indicates actual number of colonie
 = One loopful of a dilution cultured on blood agar plate.

In other experiments which included Locke's solution containing 0.1 percent gelatin and proteose peptone solution, it was observed that the survival of the meningococci in Locke's solution was similar to that in Ringer's solution, while in proteose peptone solution it was similar to that in neopeptone solution. In the original dilutions in proteose peptone solution the growth was more luxuriant than in the presence of neopeptone.

Our observations on the viability of meningococci in distilled water, saline, and Ringer's solution, in general, are in agreement with the findings of Flexner (21) which were reported in 1907. He found that meningococci suspended in distilled water fragmented more rapidly than in saline, that saline was directly injurious to the organism, and that viability was more prolonged in Ringer's solution than in saline.

On account of the rapidly injurious action of distilled water, saline, Ringer's, and Locke's solution on meningococci, it would seem advisable not to use any one of them for the suspension of meningococci for the mouse protection test.

On the other hand, neopeptone and proteose peptone apparently are not only not harmful to the bacteria but are favorable for multi-

plication. We have selected neopeptone for use in our routine mouse protection tests. It should be mentioned, however, that in freshly prepared solutions of neopeptone, growth is not always initiated in the higher dilutions of the culture. In this case, if the solution is permitted to stand at room temperature for a week or two, this inhibitory action disappears and inocula of less than 10 meningococci will promote multiplication.

#### MUCIN

Mucin has been one of the most troublesome variables in the performance of the mouse protection test. During the early work, investigators found that different lots of mucin prepared by the same manufacturer varied tremendously in their influence on the lethal action of meningococci in mice. At times the use of a mouse protection test as a means of evaluating antimeningococcus serum seemed almost hopeless. Recently, however, better preparations of granular mucin have been available, but even these may vary in influence on lethal action of meningococci and on primary toxicity for mice. The difference in the effect of two lots of mucin on the viability of meningococci will be demonstrated below.

Different methods of preparing suspensions of mucin for the mouse protection test have been employed by various workers. Some have used distilled water suspensions with or without the addition of dextrose, while others have added 0.5 percent sodium chloride. Both the autoclave and the Arnold sterilizer have been used in the sterilization of the suspensions. In the present work we have determined the viability of meningococci in suspensions of mucin prepared by the different methods. We have also observed the influence on the mouse protection test of distilled water and saline suspensions of mucin, and of suspensions sterilized in the autoclave and in the Arnold sterilizer. Data on the influence of viscosity of mucin and the proper concentration of mucin for the mouse protection test are given.

The viability of meningococci in suspensions of mucin.—Two lots of mucin, Nos. 31,111 and 35,689, were used. Suspensions of No. 31,111 were dark, slightly viscous, and contained many hard particles. Suspensions of No. 35,689 were creamy, very viscous, and contained practically no hard particles.

A 5 percent suspension of each lot was prepared in distilled water, sterilized in the autoclave, and adjusted to pH 7.4. Each solution was divided into four portions. To one nothing was added, to the second 1 percent dextrose, to the third 0.5 percent sodium chloride, and to the fourth both 1 percent dextrose and 0.5 percent sodium chloride.

In each of the different preparations serial dilutions  $\times$  10 of a 5-hour-old blood agar culture suspended in neopeptone solution (density 500 p. p. m.) were made. Similar dilutions were made in neopeptone solution. After the dilutions had stood for 2 hours at room temperature, 0.1 cc. of each was cultured in special 0.15 percent

Then the dilutions were placed in the incubator and after 48 hours 1 loopful of each was cultured on a blood agar plate.

The results of an experiment with the two lots of mucin prepared by the different methods are given in table 7.

In the presence of all preparations containing mucin No. 31,111 there was dving of the bacteria. It was most rapid in the distilled water suspension. In the presence of dextrose or sodium chloride the death was slower while in the presence of both dextrose and sodium chloride conditions were still more favorable for the bacteria. though the time interval for the subculturing in this experiment was 2 hours, in preliminary tests it was found that even within 30 minutes in a suspension of this mucin containing dextrose there was a decided decrease in the number of bacteria.

In the presence of mucin No. 35,689 (table 7) there was less dying of the bacteria than in presence of the other lot of mucin but only when both dextrose and sodium chloride were added were the conditions most favorable. However, there was multiplication of the bacteria in all dilutions of each preparation of this lot of mucin except in the distilled water preparation.

TABLE 7.—The influence of dextrose and sodium chloride on the viability of meninaococcus in mucin

	Growth in subcultures from culture dilutions in mucin													
Diluent	Ai	ter 2 hou	ırs—0.1 c	ec.1	After	48 hours	incubat	ion—1 lo	pful ³					
	5×10−6	5×10− <sup>7</sup>	5×10-8	5×10-	5×10-5	5×10 <b>−</b> 6	5×10 <sup>-7</sup>	5×10-8	5×10-0					
Neopeptone	++++	++++	+	±			++++	++++	++++					
Mucin No. 31,111	++++ ++++ ++++	_ ++ ++++	=	= 1 = 1	- ±	- -	=	=	=					
0.5 percent NaCl	++++	++++	++	-	+++	-	-	-	-					
Mucin No. 35,689	++++ ++++ ++++	++++ ++++ ++++	- - ++++ ++++	- - - ++++	± ++++ ++++	_ ++++ ++++	 ++++ ++++	_ ++ ++++ ++++	_ + ++++					

Subcultures were made in special 0.15 percent agar and incubated for 4 days.
 Subcultures were made on blood agar plates.

In other experiments 1 percent proteose peptone was added to water suspensions of each of the two lots of mucin. In the presence of each there was no injurious action on the bacteria. Even in the presence of mucin No. 31,111 there was multiplication of the bacteria in all dilutions.

Although the test-tube experiments with suspensions of mucin containing dextrose but not sodium chloride indicated that such suspensions were injurious to the bacteria, this action was further demonstrated in a mouse protection test. Two sets of mice were inoculated,

45 minutes apart, with the same dilutions of an antiserum. The mice of the first set were inoculated with a culture immediately after it had been suspended in the mucin solution (No. 31,111) containing 1 percent dextrose, and the mice of the other set were inoculated with the same cultural suspension after it had been held about 45 minutes. The dilution of serum which protected 50 percent of the first mice was 1:47, and in the second case it was 1:90; that is, only one-half as much serum was required to protect the mice against the bacteria that had been exposed to the injurious action of the suspension of mucin for 45 minutes. In a similar experiment in which the mucin suspension contained 0.5 percent sodium chloride, the dilutions of serum which protected the mice of each set were comparable.

Sterilization of mucin.—Whether the mucin is sterilized in the autoclave or in the Arnold sterilizer does not seem to make any particular difference in the results of the mouse protection test. The results of two experiments performed at different times are as follows: In the first, the calculated dilutions which protected 50 percent of the mice were 1:163 and 1:150, respectively. In the second, the dilutions were 1:147 and 1:168, respectively.

With autoclaved mucin, it has been noted that certain preparations are slightly toxic for mice. This toxicity, however, disappears if the preparation is allowed to stand for about 2 weeks before use. When sterilizing in the autoclave, great care is taken to prevent the pressure going over 15 pounds (121° C.). Thirty minutes is used for amounts of 1,000 cc., and 20 minutes for 500 cc. or less. For sterilization of the same amounts in the "Arnold," we have used, respectively, 1 hour and 30 minutes on 3 successive days. With "Arnold" mucin we have had some difficulty with lack of sterilization.

Viscosity of mucin.—It is interesting to note that although the suspensions of mucin sterilized differently induced similar results in the mouse protection test, they do differ in viscosity. Preparations of 3.5 percent mucin sterilized in the autoclave have been found to have an average viscosity of 9.0, while like preparations sterilized in the "Arnold" have a viscosity of about 12.0.

The fact that viscosity alone is not the determining factor of mucin which increases the lethal action of meningococci for mice was very strikingly demonstrated with a preparation that had been improperly sterilized in the "Arnold" due to escape of steam. (The stopper holding the thermometer in the top of the sterilizer had inadvertently been left out of place.) This mucin was used on the final day of sterilization in a mouse protection test. Although no contaminating organisms were seen in the peritoneal smears of the mice which died, 2 weeks later the mucin contained a toxigenic gram-positive sporeforming rod. The viscosity of this preparation was 18.6. In the

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same experiment a preparation of autoclaved mucin of the same concentration with a viscosity of 8.7 was also used. With the "Arnold" mucin the 50 percent endpoint dilution of serum M19 was 1:159, and with the autoclaved mucin it was 1:177.

The comparable results obtained in the mouse protection tests with preparations of mucin which contain the same concentration of mucin but which differ in viscosity substantiate the conclusion of Anderson and Oag (22) that it is not viscosity per se which enables a substance to increase the lethal action of meningococcus for mice.

Concentration of mucin.—It has not been possible to use a fixed concentration of mucin for the mouse protection test. The amount has had to be adjusted for the particular lot of mucin, for the susceptibility of the mouse as influenced by breed and weather, and the immunological group of the test culture.

For Group I cultures, with mice from the open market and mucin lot No. 31,111, a 5 percent suspension of mucin was used with a  $1\times10^{-3}$  to  $5\times10^{-4}$  dilution of the culture. With mucin lot No. 35,689, it was necessary to decrease the dilution of the culture to about  $1\times10^{-4}$ , the other factors remaining constant. With the shift to inbred Swiss mice the concentration of mucin was reduced to 4 percent and later to 3.5 percent during the summer. With 3.5 percent mucin the dilution of culture has varied from 1 to  $2.5\times10^{-4}$ .

For Group II cultures it has been necessary to use a slightly higher concentration of mucin than with Group I cultures.

It is believed that the minimum concentration of mucin that will render at least 10 highly virulent meningococci lethal for more than half of the mice inoculated, should be used.

#### TIME INTERVAL BETWEEN INOCULATION OF SERUM AND OF CULTURE

In a study of the mouse protection test, Branham (23) reported in 1935 that, on the whole, the best protection was obtained when serum was given 4 hours preceding the culture. The other time intervals which she used were 1, 8, 12, and 24 hours preceding the culture. In her work relatively large doses of serum, at times as much as 0.5 cc. of undiluted serum, were used to protect against large numbers of meningococci which were not suspended in mucin. Other workers have given the serum 30 to 60 minutes before the culture, while still others have given it either simultaneously or mixed with the culture. In our more recent work a time interval of 1 hour has been used. This interval was selected after studying the influence of inoculating the serum 4 hours and 1 hour before and simultaneously with the culture.

It was found that the selection of the best time interval between inoculation of serum and culture was dependent on the size of the test dose of culture. With large amounts of culture and corresponding amounts of serum the best protection was obtained when the serum was given 4 hours in advance. With a small number of bacteria and high dilutions of serum the best protection was obtained when the inoculations were made simultaneously. With intermediate-size inocula the greatest protection was obtained with the hour interval. These findings are illustrated in tables 8, 9, and 10.

31 TABLE 8 .- The influence of time interval between inoculations of a large amount of serum and varying amounts of culture

					Dilu	tion of	lum	Calcu-			
Experi- ment No.	Serum 1	in	Cime terv iours	al	1:1	.00	1:1,	000	1:10	.000	lated 50 percent endpoint
				s	D	8	D	8	D	of culture	
1	M19	{	(2)	4	3 0	5 8	4 2	4	5 7	3 1	1:1,000 1:2,500
2	No. 9 Concentrated	{	(2)	1	2 0 2	6 8 6	8 3 3	0 5 5	8 7 4	0 1 4	1:210 1:1, 900 1:2, 600

<sup>1</sup> Each serum diluted 1:5, 0.5 cc. inoculated.

Table 9.—The influence of time interval between inoculations of small amounts of serum and a constant amount of culture

		I	)ilu	ion of	serum	1-0.5	e. inoc	ulum	Calcu-
Time interval	Amount of culture		1:2	200	1:4	100	1:8	300	lated 50 percent endpoint
			8	D	s	D	s	D	of serum
4 hours 1 hour None 2	}1.0 cc. of 2 x 10 <sup>-3</sup>	{	5 6 6	3 2 2	4 4 5	4 4 3	0 1 4	8 7 4	1:310 1:365 1:540

Serum No. 9 concentrated.

Equal distribution of male and female mice; weight: males 18-20 gm.; females 16-18 gm.

Table 10.—The influence of time interval of inoculation of serum on the amount required to protect against approximately 100,000 m. f. d. of culture

	:	Dilutio	n of se	rum M	f 1 <del>9—</del> 0.	5 ec. in	oculun	a	50 per-	
Time interval		50	1:100		1::	200	1:-	400	cent end- point of	
		D	s	D	s	D	S	D	serum	
4 hours	3 4 3	2 2 3	3 5 2	3 1 4	3 4 2	3 2 4	1 0 3	5 6 3	1:114 1:175 1:100	

<sup>1</sup> Serum and culture inoculated simultaneously. Test dose of culture=1 cc. of 2.5 x 10-4 dilution. Equal number of male and female mice, 16-17 gm.

<sup>&</sup>lt;sup>1</sup> The serum and culture were inoculated simultaneously.

Male mice 18-20 gm.

Mucin No. 35,689, 3.5 percent concentration, plus 0.5 percent NaCl and 0.5 percent dextrose.

<sup>2</sup> Serum and culture injected simultaneously.

In table 8 are given the results of two experiments. The mice were inoculated with 0.1 cc. of serum and varying amounts of culture. In each instance the mice were protected against the largest amount of culture when the serum was given 4 hours in advance and against the least amount when serum and culture were given simultaneously.

In the next experiment recorded in table 9, a moderately high dilution of culture,  $2.5 \times 10^{-5}$  (approximately 10,000 m. f. d.) was inoculated with varying dilutions of serum. Under these conditions the best protection was obtained when the serum and culture were given simultaneously and least when serum was given 4 hours in advance.

In table 10 are given the results of an experiment with intermediate inocula. In this case the optimum protection was obtained when the serum was given 1 hour in advance, and poorest when given with culture. The test dose of culture contained approximately 100,000 m. f. d. This size inoculum has been advocated by several workers. It has not seemed advisable to use a larger dose because of the primary toxicity of meningococci for mice.

It may also be noted in table 10 that in case of the simultaneous inoculations, the protective action of varying dilutions of serum was not clearly differentiated. This finding has been observed in other experiments. Furthermore, with certain concentrated sera the occurrence of a marked prozone has been observed following simultaneous inoculation of serum and culture. The least occurrence of prozones has been noted when the serum was given 4 hours in advance.

The above results suggest that to obtain the maximum mouse protection with an antiserum, the choice of the time interval to be used between inoculation of serum and culture is dependent on the size of the test dose of culture. If a test dose of 100,000 m. f. d. is used the best results are apparently obtained when the serum is given 1 hour preceding the culture.

#### SUMMARY

The results of the comparative study of different variables or factors which influence the determination of the mouse protective action of antimening ococcus serum indicate that:

- 1. Five percent rabbit blood agar is a satisfactory medium for the maintenance of the virulence of meningococci. Serum glucose agar, EDB agar, and Dorset's egg medium are almost as favorable as the blood agar. On semisolid media the bacteria remain alive longer but there is a gradual loss in virulence.
- 2. Although different Group I strains of meningococcus differ in susceptibility to antiserum, it appears that comparable values in relation to control scrum may be obtained with a number of strains.

- 3. Mice of different strains and from different colonies of the same strain may vary in susceptibility to meningococci. The important factor is that they come from a pure, closely inbred line. Furthermore, with closely inbred mice it has been demonstrated that about twice as much serum is required to protect females as males and that wide variations in weight cause irregular results.
- 4. In distilled water, saline, Ringer's or Locke's solution, meningococci die very rapidly and the use of either for the first suspension of the culture is contraindicated. Neopeptone or proteose peptone solutions are not injurious to the bacteria and may be used for the suspension fluid.
- 5. Different lots of mucin vary in influence on the viability of meningococci.

Preparations of mucin which contain both sodium chloride and dextrose appear to be the most favorable for the bacteria.

Comparable results are obtained with suspensions of mucin sterilized in the autoclave and in the Arnold sterilizer though they do differ in viscosity.

6. To obtain the maximum protection with antiserum, the time interval between inoculation of serum and culture is dependent on the size of the test dose of culture. With a test dose of 100.000 m. f. d., it appears that the best protection is obtained when the serum is inoculated 1 hour in advance of the culture.

#### REFERENCES

 Miller, C. P.: Experimental meningococcal infection in mice. Science 78: 340 (1933).
 Branham, S. E., Pittman, M., Rake, G., and Scherp, H. W., A proposed mouse protection unit for antimeningococcus serum. Proc. Soc. Exp. Biol. and Med., 39: 348 (1938).

(3) Rake, G.: The mouse protection test in the standardization of antimeningococcus serum. Canad. Pub. Health J., 28: 265 (1937).
(4) Pittman, M., Branham, S. E., and Sockrider, E. M., A comparison of the precipitation reaction in immune serum agar plates with the protection of mice by antimeningococcus serum. Pub. Health Rep., 53: 1400 (1938).
(5) Rake, G., and Scherp, H. W.: Protective antibodies effective against Type I maningococcus infection in mice. Abstract. J. Ract. 38: 118 (1030).

meningococcal infection in mice. Abstract. J. Bact., 38: 118 (1939).

(6) Scherp, H. W.: A quantitative study of the precipitative reaction between Type I polysaccharide and antimeningococcal horse sera. J. Immunol., **37**: 469 (1939).

(7) Alexander, H. E.: Response to antiserums in meningococcic infections of human beings and mice. Am. J. Dis. Child., 58: 746 (1939).

(8) (a) McLeod, J. H.: A report of clinical experience with meningococcus meningitis. Read before the American Academy of Pediatrics, Washington, D. C., April 5, 1940.

(b) Branham, S. E.: The role of the laboratory in the therapy of meningococcus meningitis. Read before the American Academy of Pediatrics, Washing-

ton, D. C., April 5, 1940.

(9) Scherp, H. W., and Rake, G.: Standardization and concentration of antimeningococcus horse serum (Type I). J. Exp. Med., 63: 547 (1936). (10) Little, P. A.: On a rapid method for the standardization of antimeningococcal

horse serum Type I. J. Immunol., 35: 117 (1938).

(11) Reed. L. J., and Muench, H.: A simple method of estimating fifty percent endpoints. Am. J. Hyg., 27: 493 (1938).

(12) Murray, E. G. D., and Ayrton, R.: Observations on the growth of meningococci in vitro in relation to virulence. J. Hyg., 23: 23 (1924).
(13) Hitchens, A. P.: Advantages of culture mediums containing small percentages of agar. J. Infect. Dis., 29: 390 (1921).

(14) Recommended procedures of the American Public Health Association. In press.

(15) Unpublished.

(16) Kirkbride, M. B., and Cohen, S. M.: Precipitation reactions of meningococcus strains with immune serum in agar plates in relation to antigenic activity. Am. J. Hyg., 20: 444 (1934).

(17) Cohen, S. M.: A study of the virulence of meningococcus strains and of the

protective activity of antimeningococcus sera. J. Immunol., 30: 203

(18) Branham, S. E.: The effect of sulfapyridine and sulfanilamide with and without serum in experimental meningococcus infection. Pub. Health Rep., 55: 12 (1940).

(19) Colvin, J. W., and Mills, C. A.: Ease of body heat loss and resistance to infection. Science, 90: 275 (1939).

(20) Standard methods of water analysis. Am. Pub. Health Assoc., New York,

1925, p. 4.
(21) Flexner, S.: Contributions to the biology of Diplococcus intracellularis.

(21) Flexner, S.: Contributions to the biology of Diplococcus intracentularis.
J. Exp. Med., 9: 105 (1907).
(22) Anderson, C. G., and Oag, R. K.: The effect of gastric mucin on the pathogenicity of the meningococcus and other organisms: with particular reference to its fractionation. Brit. J. Exp. Path., 20: 25 (1939).
(23) Branham, S. E.: Protection of mice against meningococcus infection by polyvalent antimeningococcic serum. Pub. Health Rep., 50: 768 (1935).

## WHAT INDUSTRY HAS LEARNED ABOUT LEADERSHIP 1

### ITS APPLICATION TO PUBLIC HEALTH

By S. J. FOSDICK, Pittsburgh, Pa.

During the last quarter century industry has become increasingly concerned about the worker's job satisfaction. In order to promote a greater degree of satisfaction, it has fostered a number of programs to improve working conditions and help workers adjust themselves to their jobs. Among the developments toward this end are welfare departments, personnel departments, wage plans, annual income guarantees, and many other similar devices.

As recently as 1934, however, Dr. Robert Hoppock, in his illuminating study, "Job Satisfaction," pointed out that probably one out of every three workers is more or less dissatisfied with his job. Even though the dissatisfaction is frequently the result of the worker's own incapacity, industry continues its effort to remove it. But Schopen-

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EDITORIAL NOTE.—A health officer's success as administrator of the public health program depends in large measure upon his ability as a leader. As the executive, responsible for the maintenance of staff morale and a high level of performance by his subordinates, the health officer requires the abilities, attitudes, and other traits that characterize good leaders.

Industry has found it profitable to study intensively the characteristics of its best leaders and to institute programs to improve the qualities of leadership possessed by its executives and subexecutives. Many of the principles found useful in industry can be practiced to advantage by health officers and other supervisory public health workers. Hence this paper, prepared by a man who has concentrated on the development of leaders in his own organization, is presented.

hauer taught, as you will recall, that every satisfaction leads to a new struggle, implying that continued satisfaction is impossible without annihilating desire. Experience has proved that the dissatisfactions of yesterday adjust to become routine operations today, and new dissatisfactions will appear tomorrow.

As a result, industry has been seeking some common factor upon which it could concentrate to secure a greater continuing satisfaction, and perhaps serve to "annihilate desire" in this particular. The most powerful common factor seems to be that of leadership. There is nothing new in this, except that in industry, as well as in other fields, it has heretofore been considered that leadership was a characteristic impossible to develop in those not born with it. But, just as it is no longer thought that salesmen are born and not made, it is now agreed that men may at least be trained to be better leaders. Given certain fundamental capacities, their ability as leaders may be improved through training.

You may all be thinking at this point, "What has this to do with me and public health?"—and well you might. I had much the same feeling when I was asked to prepare this paper. But reflection will show you, I am sure, that training for leadership has a very great deal to do with the men and women in public health work.

Let us consider for a moment the training that most public health officers have had. Throughout their training, their internship, and their clinical work, the whole emphasis has been on the relationship of the doctor to his patient—individual to individual. If most doctors are like the ones that I have known they have learned to impose their opinions upon their patients and have not found it necessary to "sell" their patients. It is, "Stop smoking or expect to have trouble,"—not a sales talk on the desirability of discontinuing smoking. This may be necessary, and I have no criticism of it. But when doctors become public health officers with staffs of people to handle, possibly this method of handling the human relationship problem may not be so successful. Frequently the energies of both the physician and his staff are vitiated by conflicting personalities.

Leadership has been defined by a very competent woman in the retail industry as the functioning of personality so as to secure immediate and spontaneous cooperation. I am sure that you will all agree that immediate and spontaneous cooperation in any field of human endeavor is a matter eagerly to be sought and greatly to be desired. Before discussing the qualities of good leadership, let us examine the idea of leadership as it has been recast under present social conditions. Too many of us think of leadership in terms of our school-day reading—the hero fighting on a battlement while his troops fall around him, Bismarck with his iron will and his iron fist welding a nation out of unrelated principalities, the Little Corporal from Corsica plunging

Europe into blood, captains of industry and finance building great empires of economic wealth, zealots leading important movements in the sweeping progress of civilization.

But the leadership we speak of here is warmer, more human. It is a quality that can be developed in any intelligent person willing to work patiently toward understanding and consideration. It is a persuasive and impelling force rather than an arbitrary and coercive one, and is motivated by deep-seated sincerity, integrity, kindliness, and tolerance. It cannot fail to make one who practices it a more valuable social being in the strife-torn world.

The purpose of this paper is to describe some of the characteristics of good leadership which industry has discovered in the process of training its own executives. Roughly, we can divide leadership into two parts, one, methods, and the other, relationships with subordinates. Two of the principal divisions under methods might be termed methods of praise and criticism and methods of direction.

Methods of praise and criticism are important because it is in this respect that supervisors most frequently fail in their handling of people. In your thinking about these problems, it is vitally important that you compare the way you like to be handled with the way in which you are handled. This will give you a desirable point of view in the handling of those you supervise. People, as a rule, respond to the same general sort of treatment, because the response engendered is usually emotional and not reasoned. The treatment that I dislike from my superiors is usually exactly the same as the treatment for which my subordinates criticize me. To put it more simply, it is nothing but the Golden Rule: "Do unto others as you would have them do unto you."

Upon reflection you will probably agree that there is more criticism given than praise. This situation tends to encourage a belief on the part of the worker that his supervisor is not fair. Constant criticism without the leaven of properly administered praise leads to discouragement. The worker says, "Is there anything I do right?" and answers his own question, "There are many things that I do right and do well, but my supervisor is not big enough to recognize them."

Well-merited praise should be given as freely as well-founded criticism. But criticism should never be given in the presence of others, nor should either praise or criticism be general. Praise should be given only for specific accomplishment. Criticism should be directed at a specific act. We have found by many surveys that workers frequently say, "A pat on the back is frequently worth more than money in the pay envelope."

It has been determined that criticizing attitude is a mistake. If one criticizes a person's attitude, it tends to disintegrate his person-

ality. Probably no one would argue the statement that most personalities are to some degree disintegrated and that individuals are always struggling, consciously or unconsciously, to achieve better integration. Everyone likes things that flatter him, that build up his good opinion of himself; and everyone is afraid of what hurts him. These feelings are both attempts at personality integration. Criticism of an act does not necessarily hinder integration or promote disintegration of a personality, but in making the criticism it is always advisable to mention something praiseworthy at the same time. In this way the criticism loses its sting but not its force. I know of no kind of situation in which this is not both profitable and desirable.

For example, you might say to a visiting nurse, who is not doing an entirely satisfactory piece of work, "Miss Jones, you have one of the best records in our department for the number of calls which you make each day and for the way in which you handle your patients. But much of this good work is offset by the fact that your reports are prepared in such a manner that it is almost impossible for us to make the proper use of them." From this point, you can develop your criticism more fully. The criticism should close with a word of encouragement and praise. This method will establish in the mind of the worker that you are being fair and have considered her good qualities as well as those about which you are concerned. If the interview is properly developed, she should leave your office more enthusiastic about her job than before, more determined to do a better job, and convinced that she must improve.

In this connection, let me quote you a statement which, I am told, hung in the office of a well-known department store for a long period of time. It is, perhaps, a little too sweeping in its statement; nevertheless the idea is an important one.

Unless a reproof or criticism is given in such a way as to leave the employee more interested and enthusiastic than before, the exercise of authority has been detrimental to this Company and the person administering it has proven himself unfit to manage men.

In the criticism of conscientious people you must be extremely careful. Always try to put yourself in the other person's position. In the interview, if possible, develop an atmosphere of helpfulness and of the mutuality of the problem involved. For instance, to refer again to the case of the conscientious nurse who was careless about her reports, it would be in accord with this idea for you to say, "I know how much bother these reports are. This is one thing I have to keep constantly schooling myself on, because I do not like to make them out either. But you will agree that they are of great importance; so all of us must give them our best attention and make them out completely and fully."

Another question that has provoked much discussion in conferences considering leadership is, "What is the best time of day to give a subordinate a serious reprimand?" In speaking of "serious reprimand" we mean one that may lead eventually to dismissal of the employee, a final warning. The head of a Baltimore store which is doing a remarkable job on this type of leadership training became so much interested in the question that he asked Dr. Adolph Meyer, the eminent psychiatrist at Johns Hopkins University, this question. Dr. Meyer replied, after some consideration, that he felt the best time to give a criticism of this type was the first thing in the morning. At that time the person administering the reprimand as well as the person receiving it is fresher and more relaxed. This serious blow can then be received with the least permanent damaging effect. However, this is not always possible. This employer said, "I cannot run my store that way. There are too many pressing duties at that time." Dr. Meyer replied, "That is something else. You asked me the best time of day for the individual, and, in my opinion, it is the first thing in the morning."

Other things being equal, we are in agreement that the morning is the best time, but these are other points to be considered: How do you feel? How does the person to be criticized feel? Can the matter wait? What effect will it have on other people? When do you have the time?

In giving criticism it is not necessary to make the person admit his fault. It is not desirable to remind employees of their past errors except in the case of a final warning. Then it might be well to total up the score. It should never be given angrily or with any show of irritation. It should be calm and objective. I have dwelt at considerable length on this matter of criticism, but I think it is of the greatest importance in handling people.

How do you go about encouraging your people? Do you ever let them make errors? Most of the human race learn to some degree through trial and error. Although our best leaders think that this is a desirable situation, certainly no industry could afford to let an employee make a serious error that would cost a large sum of money. Neither could you afford to allow an employee to make an error that might cost lives. But in any job there are many details such that an occasional error in carrying them out is unimportant. Let the worker try his wings. After the error is made, point out how the thinking was faulty and encourage him to try again. Do not stifle the initiative of your people by too close supervision in every small detail. Insofar as possible, let them work in the manner that is the easiest for them so long as the results you require are obtained.

When one of your subordinates comes to you with a suggestion or a plan, follow it as closely as possible. Give him full credit for the

suggestion. Do not be afraid that you will not get credit for your work. Give every credit possible to the subordinate and give it freely and openly. I have known executives who, presented with a plan by a subordinate, would change the plan in some nonessential way and then adopt it as theirs. This is not good leadership. If the plan is good in the main, let it go through as suggested. Later it may be necessary to make some small adjustment. Never be fearful that your subordinates will get credit that is rightfully yours. Be generous to a fault with them on this score. In the long run, you are judged by the results of your department, and you do get the larger and longer range credit.

Do your people ask questions, and do you answer them carefully, completely, exactly, and patiently? Industry has found that the worker asks too few questions, primarily because he does not realize his need for more complete instruction.

Let us discuss briefly the problems raised by the second part of our technique of good leadership, that of the relationship between you and the subordinate. Which type of error is usually more serious for an executive to make in his manner toward his subordinates, being too cold and brusque, or being too free and easy? We believe the more serious error is that of being too cold and brusque. Naturally, it is desirable that an executive maintain a manner of gracious friend-liness toward his subordinates. It is better for him to err on the side of being free and easy, as coldness repels rather than attracts.

Above all, an executive must be honest with his subordinates. If you have unjustly criticized a subordinate, good leaders say it is desirable to admit that error. This does not weaken authority. Say frankly, "I did not understand all the problems involved, and find that you were quite right in what you did."

When a subordinate comes to you with a problem that seems serious to him, treat it just as seriously yourself, regardless of how petty or how small it may seem to you. It is advisable in such a case to admit the seriousness of the problem, and then you can point out with consideration those elements in the problem which indicate that it is not of the importance which the employee felt it to be.

Be sure, in laying down rules, that you give reasons for them. When you explain these reasons to your subordinates, show that their point of view has been taken into consideration and that the rule is desirable for other reasons which overbalance this consideration. When an employee understands the reasoning behind rules, he is much more likely to follow them carefully and with greater good will and cheerfulness. In taking over a new piece of work, do not let yourself fall into the error of making radical and rapid changes. Study a new job slowly and as you become familiar with all of its

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elements and with people whom you are to supervise, make the necessary changes.

You should also keep in mind that only rarely will you become conscious of dissatisfaction on the part of your staff. The reasons for this are obvious. The only way in which you will know is by developing and practicing the art of good leadership so that your people may be frank and candid with you.

When a new member is added to your staff, take the time and trouble to see that he is well established and that he exactly and fully understands the duties of his job and knows what is expected of him. If duties, responsibilities, the amount of interest required, the exactitude necessary, are carefully described to a new employee, it makes a tremendous impression on him. He cannot help but feel that he is now associated with an organization headed by a sympathetic, understanding, and capable executive.

In industry, it is rarely advisable to mingle socially with one's subordinates outside of business hours. Joking and teasing of subordinates is never permissible. Such actions almost always tend to lower the respect of the individual for his superior. Uniform courtesy at all times is an emphatic rule of those who have executive responsibility, not the cold courtesy of lip service alone, but the courtesy that springs from a genuine interest and a genuine consideration for others.

I hope that the few suggestions I have made on what industry has learned about leadership will start you thinking on the subject. Industry has found leadership to be of great importance, and proficiency in it to be of great value. You will go far toward putting into practice the fundamental principles of good leadership if you will remember the admonitions expressed in these fragments of ancient sayings: "Wisdom is the principal thing; therefore get wisdom, and with all thy getting, get understanding" and "You may destroy a man's gold, his land, and his house, but yet be forgiven. But if you destroy his self-respect, you have destroyed what makes it possible for him to walk among men."

### COURT DECISION ON PUBLIC HEALTH

State law for the control of contagious abortion in cattle held valid.—
(Virginia Supreme Court of Appeals; Stickley v. Givens, State Veterinarian, 11 S.E.2d 631; decided November 25, 1940.) A suit was brought by a dairyman against the State Veterinarian of Virginia to enjoin the enforcement of a 1938 Virginia statute (ch. 439) providing for, among other things, the prevention, control, and eradication of Bang's disease. Some of the statute's provisions pertaining to Bang's disease related to the making of the agglutination test and to the condemnation, quarantine, appraisal, and slaughter of cattle found

infected. The plaintiff, as the result of the testing of his cattle for Bang's disease and the subsequent designation of a number of them as infected, challenged the validity of the statute. His contention was that the statute was in violation of the Federal and State constitutions in that it deprived him of his property without due process of law and denied to him the equal protection of the laws. The supreme court of appeals, however, took the view that the challenged act was a reasonable exercise of the police power of the State and that it was not violative of either the Federal or State constitution.

## DEATHS DURING WEEK ENDED JANUARY 4, 1941

[From the Weekly Health Index, issued by the Bureau of the Census, Department of Commercel

	Week ended Jan. 4, 1941	Corresponding week, 1940
Data from 88 large cities of the United States: Total deaths. Average for 3 prior years. Deaths under 1 year of age Average for 3 prior years. Data from industrial insurance companies: Policies in force. Number of death claims Death claims per 1,000 policies in force, annual rate.	9, 251 9, 280 585 560 64, 796, 540 10, 108 8, 1	9, 250 10, 027 566 607 66, 416, 327 10, 204 8. 0

# PREVALENCE OF DISEASE

No health department, State or local, can effectively prevent or control disease without knowledge of when, where, and under what conditions cases are occurring

# **UNITED STATES**

# REPORTS FROM STATES FOR WEEK ENDED JANUARY 11, 1941 Summary

A total of 89,828 cases of influenza was reported for the current week, as compared with 77,144 for the preceding week. This represents an increase of 12,684 cases, or 16 percent, as against 31,669, or 70 percent, for the prior week. The number of cases for the current week is the largest number reported for any week since December 1932 and the largest number for the comparable week during the past 10 years, or since 1929, when approximately 140,000 cases were reported.

The peak week in the 1928-29 epidemic, with nearly 196,000 cases, was the week ended January 5, while in the 1932-33 epidemic the largest number of cases (90,102) was reported for the week of December 31, 1932.

For the current week, decreases were recorded only for the Mountain and Pacific States. Although all other areas showed increased incidence, the only significant changes were in the South Atlantic region, where the number of cases rose from 4,308 to 13,629, and in the New England States, with 2,563 cases as compared with 149 last week. Three States accounted for most of the increase in the South Atlantic group (Virginia, from 1,752 to 4,200; South Carolina, from 1,581 to 3,686; and Georgia, from 788 to 5,002), while increases in Maine (from 40 to 1,345) and New Hampshire (from 0 to 1,000) accounted principally for the rise in the New England area.

For the week ended January 13, 246 cases of influenza were reported in Alaska.

Of the other 8 communicable diseases reported weekly, measles, poliomyelitis, and whooping cough were above the 5-year (1936-40) median expectancy.

One case of Rocky Mountain spotted fever was reported in Oklahoma, 1 case of undulant fever each in Mississippi and Utah, 1 case of encephalitis each in Maryland and Oklahoma, 2 cases of tularemia in North Carolina and 25 cases of endemic typhus fever, all in the Southern States except 1 case in Ohio and 1 in California.

For the current week the Bureau of the Census reports 9,803 deaths in 88 major cities of the United States, as compared with 9,251 for the preceding week and with a 3-year (1938-40) average of 9,319.

Telegraphic morbidity reports from State health officers for the week ended January 11, 1941, and comparison with corresponding week of 1940 and 5-year median. In these tables a zero indicates a definite report, while leaders imply that, although none were reported, cases may have occurred.

	D	iphthe	ria	I,	nfluenza	•	1	deasles	,	Men ir	ingitis igococo	men-
Division and State	Week	ended	Me-	Week	ended	Me-	Week	ended	Me-	Week	ended	Me-
	Jan. 11, 1941	Jan. 13, 1940	dian 1936– 40	Jan. 11, 1941	Jan. 13, 1940	dian 1936- 40	Jan. 11, 1941	Jan. 13, 1940	dian 1936– 40	Jan. 11, 1941	Jan. 13, 1940	dian 1936- 40
NEW ENG.												
Maine New Hampshire Vermont Massachusetts Rhode Island Connecticut	0 0 1 0 0	0 0 9 0 0	4 0 7 0 4	1, 345 1, 000 76 31 111	32	6	70 3 76 509 0 11	73 12 3 185 226 161	73 22 11 269 74 161	0 0 4 0 0	0 0 1 1 1	0 0 1 0 1
MID. ATL.  New York  New Jersey  Pennsylvania	17 3 14	25 7 28	37 15 39	1 94 30	1 13 18	1 17 18	2, 135 876 2, 129	369 23 50	389 41 135	2 2 1	1 0 9	10 3 6
E. NO. CEN.  Ohio <sup>2</sup>	8 16 12 7 1	28 14 41 11 2	34 25 41 11 2	795 176 37 82 61	88 25 36 15 48	14 25 36 2 48	645 63 1, 109 1, 775 404	22 7 33 384 221	70 11 48 384 221	1 1 4 1 0	0 1 0 0	6 3 7 2 1
W. NO. CEN. Minnesota	1	5	4	3	la	2	2	255	122	0		1
Missouri North Dakota South Dakota Nebraska Kansas	14 7 2 2 0 2	11 14 0 5 4 11	6 21 0 1 4 11	470 95 83 3, 163	11 18 42 13	118 11 1 1 32	138 40 11 2 10 148	49 5 1 6 8 141	34 7 5 6 8 18	0 0 0	0 0 0 1 0	1 1 0 0 0
SO. ATL.  Delaware.  Maryland 22  Dist. of Col.  Virginia  North Carolina 2  South Carolina 2  Florida 2	0 6 1 10 8 22 5 16 3	0 5 3 25 19 41 7 13 5	2 8 10 25 15 33 7 13	19 92 90 4, 200 430 40 3, 686 5, 002 70	37 10 869 37 211 3, 948 2, 192 28	24 2 52 26 652 136 11	26 12 4 198 61 94 70 26 6	1 1 0 25 8 67 3 26 11	2 98 7 71 17 98 7 26 11	0 1 0 5 0 2 2 2 0	0 0 1 2 0 2 1 0	0 0 1 3 4 3 1 0 2
KentuckyTennesseeAlabama <sup>2</sup> Mississippi <sup>3</sup>	10 7 7 9	19 6 15 9	18 18 19 8	5, 950 4, 719 2, 201	21 184 1, 360	65 184 352	241 61 68	5 74 50	83 67 50	1 1 3 1	2 4 2 0	6 4 5 1
W. SO. CEN. Arkansas Louisiana <sup>2</sup> Oklahoma <sup>4</sup> Texas <sup>2</sup>	7 4 6 32	18 13 17 57	16 13 15 67	4, 664 4, 983 2, 550 33, 283	638 32 263 895	203 36 183 716	30 1 11 50	4 2 9 307	23 13 216	1 0 1 2	0 0 0 0	0 1 0 2
MOUNTAIN  Montana Idaho Wyoming Colorado New Mexico Arizona Utah 1 Nevada	2 0 5 2 4 0	1 1 1 14 1 2 0	1 2 1 12 3 6 0	1, 065 29 1, 207 1, 376 122 1, 118 1, 564 153	17 3 24 80 6 242 458	17 3  21 6 117	7 1 5 108 196 101 12 0	11 48 9 43 5 10 149	9 48 4 28 29 3 72	0 0 1 0 0	1 0 0 1 0 0	1 0 0 1 1 0
PACIFIC Weakington	0	7	1	260		1	50	999	141	2	1	0
Washington Oregon California *	1 10	7 7 18	7 32	578 2, 725	274 223	39 86	60 62	141 326	27 326	0	0 1	0 3
Total	284	543	676	89, 828			11,717	4, 568	5, 203	40	33	106
2 weeks	576	1, 031	1, 353	166, 972	22, 146	6, 273	19, 484	7, 451	8, 412	68	58	201

Telegraphic merbidity reports from State health officers for the week ended January 11, 1941, and comparison with corresponding week of 1940 and 5-year median—Con.

1941, and compar	1			T	y we		7			1		<del>оп.</del>
	Pe	oliomye	litis	s	carlet	fever	1	Smallp	οx	Typh ty	oid an phoid f	d para- ever
Division and State	Week	ended	Medi-	Week	ende	_ Medi-	Week	ended	Medi-	Week	ended	Medi-
	Jan. 11, 1941	Jan. 13, 1940	an 1936- 40	Jan. 11, 1941	Jan. 13, 1940	936- 40	Jan. 11, 1941	Jan. 13, 1940	an 1936- 40	Jan. 11, 1941	Jan. 13, 1940	an 1936- 40
NEW ENG.												
Maine New Hampshire Vermont Massachusetts Rhode Island Connecticut	0 0	0 0	0 0 0	5 3 15 136 4 27	26 0 5 128 4 72	16 8 6 260 26 77	0 0 0	0 0 0	0 0 0 0	1 0 0 1 0 2	0 0 0	0 0 2 0 0
New York New Jersey Pennsylvania	0 1	1 1 0	0 0	316 173 268	419 224 308	564 164 536	0.0	0 0 0	0 0 0	9 1 4	9 2 1	8 3 13
E. NO. CEN.  Ohio <sup>2</sup> Indiana Illinois Michigan <sup>3</sup> Wisconsin	3	1 0 0 1 1	1 0 0 0 1	251 101 340 250 129	354 150 433 321 133	433 174 548 500 288	0 5 2 4 12	1 4 0 1 6	9 5 14 1 6	6 2 3 0 0	4 1 4 2 0	4 1 4 2 0
W. NO. CEN.	2	١,	0	21	124	136	12	13	18		١,	١.
Minnesota. Lowa	0 1 0 0 0 1	0 0 0 0 0	0	61 54 75 8 11 38 92	91 82 20 22 39 135	156 193 35 26 44 160	12 4 1 0 1 1 2	31 1 1 1 0	18 18 19 12 5 3 20	0 1 2 0 0 0	1 0 0 0 1 1	1 0 1 0 0 1 2
Delaware	0 0 1 0 2 0 0 2	0 1 0 2 1 3 0 0	0 0 0 0 0 1 0	13 50 22 59 48 76 9 24	17 66 13 54 66 84 17 27	15 66 222 54 66 63 11 23 8	0 0 0 0 0 2 0	0 0 0 1 0 0 0	0 0 0 0 0 0 0 0 0	0 1 2 4 3 6 0 4 2	0 3 0 2 3 3 3 4	0 2 1 2 2 3 1 2
E. SO. CEN. Kentucky	2	2	0	86	70	70	0	0	2	1	0	7
Tennessee Alabama <sup>2</sup> Mississippi <sup>3</sup>	1 0 0	0 1 0	0 1 0	57 26 10	67 41 13	48 24 10	0 0 0	0 0 0	0 0 0	1 1 1	0 1 1	7 2 2 2
W. SO. CEN. Arkansas Louisiana 2 Oklahoma 4 Texas 2	0 0 2 1	2 0 0 4	1 0 0 1	13 8 39 46	9 19 35 61	18 18 36 111	1 0 0 0	3 0 5 2	3 1 3 12	. 2 2 9	4 7 2 12	2 7 2 12
MOUNTAIN  Montana Idaho Wyoming Colorado New Mexico Arizona Utah 2 Nevada	1 0 1 0 0 0	0 1 0 0 0 0	0 1 0 0 0 0	16 25 2 31 5 2 7	52 12 5 27 14 7 24	56 19 10 50 24 11 33	0 0 0 3 0 0	0 0 0 33 1 1 1	9 14 1 15 0 0	0 0 0 1 2 2 0 1	0 0 1 1 0 2 0	0 0 0 0 5 1
PACIFIC Washington Oregon California 2	0	0 0 16	0 0 3	38 7 129	49 23 161	56 63 218	2 1 0	0 3 0	8 12 12	1 4 7	1 : 1 1	2 3 2
Total	37	42	22 3	, 211	4, 134	6, 186	53	110	315	89	78	122
2 weeks	101	85	43	, 874	7, 731	11, 210	93	184	591	159	159	220

Telegraphic morbidity reports from State health officers for the week ended January 11, 1941, and comparison with corresponding week of 1940 and 5-year median—Con.

		oping ugh			oping igh	
Division and State	Week	ended	Division and State	Week ended		
	Jan. 11, 1941	Jan. 13, 1940		Jan. 11, 1941	Jan. 18, 1940	
NEW ENG.			so. ATL.—continued			
Maine	12 269	65 21 40 152 16	South Carolina 3 Georgia 3 Florida 3	75 10 11	10 14 7	
Connecticut	108	87	E. SO. CEN.		_	
MID. ATL.  New York  New Jersey  Pennsylvania	540 164 651	487 115 414	Kentucky Tennessee Alabama <sup>2</sup> Mississippi <sup>3</sup>	74 33 42	0 17 13	
E. NO. CEN.	l		W. SO. CEN.			
Ohio <sup>2</sup>	121	149 43 119 101 101	Arkansas Louisiana <sup>3</sup> Okiahoma <sup>4</sup> Texas <sup>3</sup>	24 8 12 232	3 2 0 94	
W. NO. CEN.  Minnesota Iowa Missouri Morth Dakota	14	72 9 3 13	Montana. Idaho. Wyoming. Colorado. New Mexico. Arizona.	8 9 28 22 26	3 6 6 15 14 37	
South Dakota Nebraska Kansas	2	4 1 36	Utah <sup>8</sup>	34 0	79	
So. ATL.  Delaware  Maryland <sup>2 2</sup> Dist. of Col	20 92 10	3 80 5	Washington	45 5 410	49 27 183	
Virginia	151	29	Total	4, 776	2, 794	
West Virginia <sup>3</sup>	42 327	11 39	2 weeks	8, 102	4, 871	

New York City only.
 Typhus fever, week ended Jan. 11, 1941, 25 cases as follows: Ohio, 1; Maryland, 1; North Carolina, 3;
 South Carolina, 5; Georgia, 6; Florida, 2; Alabama, 1; Louisiana, 1; Texas, 4; California, 1.
 Period ended earlier than Saturday.
 Rocky Mountain spotted fever, week ended January 11, 1941, Oklahoma, 1 case.

# WEEKLY REPORTS PROM CITIES

### City reports for week ended December \$8, 1940

This table summarizes the reports received weekly from a selected list of 140 cities for the purpose of showing a cross section of the current urban incidence of the communicable diseases listed in the table.

State and city				Mea- sles	Pneu- monia	Scar- let	Small- pox	Tuber- culosis	Ty- phoid	Whoop- ing	Deaths,
	theria cases	Cases	Deaths	C8.808	deaths	fever cases	pox	deaths	fever cases	cases	CSUSES
Data for 90 cities: 5-year average Current week 1.	166 57	529 4, 869	99 90	1, 295 2, 689	826 518	1, 364 8 <b>3</b> 0	23 11	346 278	21 18	994 987	
Maine: Portland	0	1	. 0	0	5	2	0	0	0	8	82
New Hampshire: Concord	٥		0	0	1	2	0	اها	0	0	10
Manchester	Ŏ		0	Ō	3	14	Ò	Ŏ	Ô	Ŏ	23
Nashua	0		0	0	0	0	0	0	0	0	6
Vermont: Barre	0		0	0	0	0	0	0	0	0	2
Burlington	0		0	0	0	1	0	9	Õ	Ó	2 10
Rutland Massachusetts:	0		0	0	0	0	0	1	0	0	2
Boston	0		3	71	19	88	0	11	0	98	210
Fall River	0		0	0	0	.9	0	1	0	7	38
Springfield Worcester	ŏ		ŏ	64	6	14 29	ő	4	ŏ	ŏ	38 30 60
Rhode Island:	_				1			ا ما			
Pawtucket Providence	0		0	0	3 4	1	0	0	0	0 5	15 66
Connecticut:			1	-		- 1	-		-	_	
Bridgeport	0	1	1 0	1	2 0	1	0	0	0	1 5	84 45
Hartford New Haven	ŏ	i	ŏ	1	ĭ	5	ŏ	ō	ŏ	7	40
Ī		1						1			İ
New York: Buffalo	o		1	40	6	20	0	3	0	25	138
New York	19	82	5	720	55	120	Ō	51	2	122	1, 482
Rochester	8		0	2	2 2	2 8	0	1	Õ	12 8	50 43
New Jersey:			-			1			-		
Camden	8	1	0	67 54	5 5	4 21	8	0	0	0 8	81 119
Newark Trenton	ŏ		ŏ	0	ĭ	19	ŏ	ī	ŏ	2	22
Pennsylvania:		7					ا ا	17		82	465
Philadelphia Pittsburgh	2	"	0	354 2	13 18	58 15	0	16	1	47	175
Reading	ō		ŏ	74	ĭ	0	0	ŏ	0	5.	21
Scranton	0			1		0	0		0	0	
Ohio:	,	ł					_ [		_		
Cincinnati	1 0	22	1 0	10 58	22 8	8 17	0	9 4	0	9 51	146 207
Cleveland Columbus	2	3	3	9	7	6	ŏl	2	ŏ	3	79
Toledo	ō	1	0	1	2	8	0	7	0	10	86
Indiana: Anderson	اه		ol	o	3	1	اه	ol	اه	0	14
Fort Wayne	Ŏ		0	Ō	1 7	6	0	O	Ŏ.	Ō	28
Indianapolis	3 0		3	1 0	7 5	17 2	0	1 0	0	8	122 13
Muncie South Bend	ŏl		ŏ	ŏ	6	ől	ŏ	ŏ	ŏ	ŏ	26
Terre Haute	0		1	i	2	Ó	0	0	0	0	20
Illinois:	o	- 1	o	اه	2	8	o	ol	ol	اه	6
Chicago	11	12	4	516	38	131	Ó	34	0	74	691
Elgin Moline	0		0	1 0	0	1 1	0	0	8	0	8 12
Springfield	ŏ		ŏ	ĭ	4	5	ŏ	ŏ	ŏ	ŏ	28
Michigan:	اه	4	0	473	25	71	7	18	اه	82	259
Detroit Flint	ŏ	*	ől	4	5	1	6	16	öl	0	209 24
Grand Rapids	Ŏ		ŏ	2	2	8	ŏ	ŏ	ŏ	17	41
Wisconsin: Kenosha	0		اه	o	اه	اه	o	اه	اه		3
Madison	Ó.		Ó	Ó	1	5	0	Ŏ.	0 1	2	4
Milwaukee	0		8	22	0	23	8	1 0	Ö	16 0	122 12
Racine											

Figures for Wilmington, N. C., and Boise estimated; reports not received.

# City reports for week ended December 28, 1940—Continued

	Diph-	Infl	uenza	Mea-	Pneu-	Scar- let		Tuber-	Ty- phoid	Whoop-	Deaths,
State and city	theria cases	Cases	Deaths	sles cases	monia deaths	fever cases	pox cases	culosis deaths	fever cases	cough	all causes
Minnesota:			0								
Duluth Minneapolis	0		1	0 2 0	1 1	1 15	3 0	0 2	0	5	24 113
St. Paul	0		0	0	3	3	0	1	0	20	72
Cedar Rapids Davenport	Ò			1		0	0		0	0	
Des Moines Sioux City	0		0	0	0	5 11	0	0	0	0	31
Waterloo Missouri:	0			0		3	0		0	1	
Kansas City St. Joseph	0	3	0	3	16 11	6	0	0	0	6	108 23
St. Louis North Dakota:	4	5	0	0	20	26	0	7	1	5	274
Fargo Grand Forks	1 0	11	0	0	0	0	0	0	0	6	5
Minot South Dakota:	0		0	0	0	0	0	0	0	0	7
Aberdeen Sioux Falls	0			0		4 2	0		0	1 0	7
Nebraska: Lincoln	1			1		5	0		0	2	
Omaha Kansas:	Ō		0	Ō	2	2	Ŏ	0	Ŏ	2	47
Lawrence	0	2	0	6 2	1 3	0 2	0	0	0	0 2	8 25
Topeka Wichita	ŏ	144	ŏ	ō	4	õ	ŏ	ŏ	ĭ	12	32
Delaware:	0		o	4	6	0	0	0	0	2	30
Wilmington Maryland:	l	3	0	3	12	20	0	5	0	49	217
Baltimore Cumberland	1 0		Ö	0	1	0	Ó	0	0	0	9
Frederick Dist. of Col.:	0		0	0	0	0	0	0	0	0	9
Washington Virginia:	0	6	0	5	16	16	0	9	0	16	207
Lynchburg Norfolk	8	25	0	0 2	2 5	0	0	0	0	0	10 30
Richmond Roanoke	1 0		1 0	1 19	2 2	6 4	0	3 2	1 0	0 5	61 22
West Virginia: Charleston	0	3	0	0	0	0	0	1	0	0	12
Huntington Wheeling	0			0	2	1 0	0		0	1 7	16
North Carolina: Gastonia	0			0		0	0		0	1	
Raleigh	ŏ		0	Ŏ	0	2	Ŏ	0	0	4	1
Wilmington Winston-Salem	ı		Ō	0	2	0	0	0	0	10	31
South Carolina: Charleston	0	18 12	0	15 8	3 0	1 0	0	0	0	3 2	23 2
Florence Greenville	ŏ		ŏ	î	8	ĭ	ŏ	ŏ	ŏ	ō	22
Georgia: Atlanta	ļ	49	o	0	4	10	o o	8	0	.10	86 6
Brunswick Savannah	1 0	18	0 2	0 4	0 2	0 2	0	ŏ	ŏ	ĭ	29
Florida: Miami	2	5	2	Q	1	0	0	1	1	0	35
Tampa	0		0	0	1	0	0	0	0	0	23
Kentucky: Ashland		7	o	0	0	0	0	0	Q	1	4
Covington Lexington	0		0	3 66	2 4	1 0	0	0 2	0	0 13	12 16
Tennessee: Knoxville	2	3	0	0	4	4	0	0	0	4	32
Memphis Nashville	Õ	29	Ŏ 2	13 0	3 2	6	0	2	0	7	63 42
Alabama:		3	0	15	3	2	0	2	0	0	52
Birmingham Mobile				10							
374	0	71	4	1	2	1 3	0	2	1	0	36
Montgomery Arkansas:				1 0	2	3	0	2	Ô	ő	30

City reports for week ended December 28, 1940-Continued

State and city	Diph-	Infl	uenza	Mea-	Pneu- monia	Bear- let	Small-	Tuber- culosis	Ty- phoid	Whoop-	Deaths,
State and city	ca.ses	Cases	Deaths	CASES	deaths	fever cases	CBSCS	deaths	fever cases	cases	CRUSOS
Louisiana: Lake Charles New Orleans Shreveport Oklahoma:	0 1 0	45 645	1 8 0	0	1 20 2	0 1 0	0 0	1 10 3	0 0 2	0 1 0	155 20
Oklahoma City. Tulsa Texas:	2 0	43	0 1	0 1	1	6	0	0	0	0	37 8
Dallas	4 0 0 1 1	15  691 181	3 1 0 0 14	0 2 1 0 1	5 4 5 14	0 3 0 8 0	0 0 0 0	0 3 4 6	0 0 0 1 0	0000	86 50 21 83 110
Montana: Billings Great Falls Helena Missoula Idaho:	0 0 0	11 80 86	0	0 0 0	3 2 0 2	1 1 0 0	0 0	0 0 0 1	0 0 0 0	0000	10 13 7 10
Boise	0 1 0	405	0 2 0	2 26 16	1 8 1	0 9 1	0 1 0	1 4 0	0 0 1	0 8 1	12 84 8
Albuquerque Utah: Salt Lake City.	0		4	1	4	0	0	1	0	3	59
Washington: Seattle Spokane Tacoma Oregon:	0 0 0	219	7 7 7	1 0 2	5 4 2	2 3 0	0	2 0 1	0	7 0 4	113 71 58
Portland California: Los Angeles Sacramento San Francisco	Ť	407 1, 519 96 371	13 3 4	2 0 1	16 9 7	1 14 2 1	0 0 0	0 14 2 8	1 0	26 0 13	105 372 46 187

State and city		ngitis, ococcus	Polio- mye- litis	State and city		Meningitis, meningococcus		
	Cases	Deaths	C8S88		Cases	Cases Deaths		
Connecticut: Bridgeport. New York: New York. Pennsylvania: Philadelphia. Ohio: Cincinnati. Cleveland Indiana: Indianapolis. Illinois: Springfield Michigan: Detroit.	1 1 0 0 0	0 0 0 0 0	0 0 0 1 1 1 1	Wisconsin: Milwaukee North Dakota: Fargo. Maryland: Baltimore. District of Columbia: Washington Virginis: Norfolk Louislana: Shreveport Washington: Seattle.	0 1 1 0 1 0	0 0 1 1 0 1	1 0 0 0 0	

Encephalitis, epidemic or lethargic.—Cases: New York, 2.
Pellagra.—Cases: Charleston, S. C., 1; Atlanta, 2; Savannah, 1; Birmingham, 1.
Typhus fever.—Cases: Atlanta, 2; Savannah, 1; Miami, 1; Mobile, 2; Montgomery, 2; New Orleans, 1;
Los Angeles, 1.

# FOREIGN REPORTS

#### CANADA

Provinces—Communicable diseases—Week ended December 7, 1940.— During the week ended December 7, 1940, cases of certain communicable diseases were reported by the Department of Pensions and National Health of Canada as follows:

Disease	Prince Edward Island	Nova Scotia	New Bruns- wick	Que- bec	Onta- rio	Mani- toba	Sas- katch- ewan	Alber- ta	British Colum- bia	Total
Cerebrospinal meningitis. ChickenpoxDiphtheriaDysentery		2 19 34	1 2	1 227 69 4	534 8	63 9	57 3	128 2	2 135	16 1, 163 122 4
Influenza Measles Mumps		223 197	3	122 87	87 254 100	43 135 13	103	61 7	899 164 5	1, 252 1, 039 218
Pneumonia Poliomyelitis		8		2	14 3	1	3 2	1	9	31 7
Scarlet fever	1	21 9	5 9	114 34	102 43	10 2	10 8	15 5	82	309 111
phoid fever		3	1	16 166	1 156	19	24	18	6 15	24 396

Vital statistics—Second quarter 1940.—The Bureau of Statistics of Canada has published the following preliminary statistics for the second quarter of 1940. The rates are computed on an annual basis. There were 21.5 live births per 1,000 population during the second quarter of 1940 as compared with 21.3 during the second quarter of 1939. The death rate was 9.4 per 1,000 population for the second quarter of 1940 and 9.9 for the second quarter of 1939. The infant mortality rate was 56 per 1,000 live births in this quarter as compared with 60 for the same quarter of 1939. The maternal death rate was 3.7 per 1,000 live births for the second quarter of 1940 and 4.7 for the same quarter of 1939.

The accompanying tables give the numbers of births, deaths, and marriages, by Provinces, for the second quarter of 1940 and deaths by causes in Canada for the second quarter of 1940 and the corresponding quarter of 1939.

(125)

## Number of births, deaths, and marriages, second quarter 1940

Province	Live births	Deaths (exclusive of still- births)	Deaths under 1 year of age	Maternal deaths	Marriages
Canada   Prince Edward Island   Nova Scotia   New Brunswick   Quebec   Ontario   Manitoba   Saskatchewan   Alberta   British Columbia	61, 157 544 3, 164 3, 028 21, 467 17, 325 3, 648 4, 543 4, 024 3, 414	26, 778 253 1, 367 1, 217 8, 132 9, 227 1, 545 1, 536 1, 544 1, 957	3, 448 34 176 225 1, 530 730 183 242 206 122	229 3 6 15 88 58 16 10 21	30, 988 128 1, 530 1, 062 9, 510 10, 936 2, 120 1, 709 1, 431 2, 312

<sup>&</sup>lt;sup>1</sup> Exclusive of Yukon and the Northwest Territories.

#### Deaths, by cause, second quarter 1940

G	(88	ada i ond rter)	Province								
Cause of death	1939	1940	Prince Edward Island	Nova Scotia	New Bruns- wick	Que- bec	Onta- rio	Mani- toba	Sas- katch- ewan	Alber- ta	British Colum- hia
Automobile accidents Cancer Cerebral hemorrhage, cerebral embolism.	314 3, 095	331 3, 290	1 26	17 196	23 128	83 900	148 1, 200	15 182	14 180	10 196	20 282
and thrombosis	525 454 63	549 387 35 2, 770	12 2 23	37 7 136	44 15 2 95	111 228 21 534	214 68 3 1, 355	33 19 5 152	35 25	31 16 3 130	32 7 1 196
Diseases of the heart Homicides Influenza	4, 747 28 1, 209	4, 858 36 534 42	36 9	199 33		1, 169 4 205	2, 035 14 115	303 2 35	255 7 54	259 7 55	422 2 16
Measles Nephritis Pneumonia Poliomyelitis	1, 726 1, 596 12	1, 700 1, 452 4	16 15 1	82 90	59 96	18 797 <b>423</b>	11 476 453 1	8 52 89 2	1 72 104	60 99	86 83
Puerperal causes Scarlet fever Suicides	280 41 297	229 31 298	3 2	6	15 1 6	88 19 46	58 8 117	16 1 16	10 29	· 21 27	12 51
Tuberculosis Typhoid fever	1,682 47	1, 527 58	13	100 1	85 7	676 30	271 5	102	61	81	138
Other violent deaths Other specified causes Unspecified or ill-	1, 039	1, 010 7, 350	5 86	57 377	38 366	220 2, 451	399 2, 246	54 433	55 461	76 438	106 <b>492</b>
defined causes	118	163 124	3	14 9	31 14	56 53	20 10	7	6 11	19 12	7

<sup>1</sup> Exclusive of Yukon and the Northwest Territories.

#### **JAMAICA**

Communicable diseases—4 weeks ended December 21, 1940.—During the 4 weeks ended December 21, 1940, cases of certain communicable diseases were reported in Kingston, Jamaica, and in the island outside of Kingston, as follows:

Disease	Kings- ton	Other localities	Disease	Kings- ton	Other localities
Chickenpox Diphtheria Dysentery Erysipelas	3 3 14 1	19 1 10 9	Leprosy	22 6	6 5 77 34

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# REPORTS OF CHOLERA, PLAGUE, SMALLPOX, TYPHUS FEVER, AND YELLOW FEVER RECEIVED DURING THE CURRENT WEEK

NOTE.—A cumulative table giving current information regarding the world prevalence of quarantinable diseases appeared in the Public Health Reports of December 27, 1940, pages 2408-2412. A similar table will appear in future issues of the Public Health Reports for the last Friday of each month.

#### Plague

Union of South Africa—Orange Free State—Bothaville District.—During the week ended November 16, 1940, 6 deaths from pneumonic plague were reported in Bothaville District, Orange Free State, Union of South Africa.

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