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THE OFFICIAL UNITED STATES AND INTERNATIONAL UNIT FOR STANDARDIZING GAS GANGRENE ANTITOXIN (*VIBRION SEPTIQUE*)

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With a view to establishing an international standard for use in measuring the potency of gas gangrene antitoxin (*Vibrion septique*), cooperative tests were undertaken by the National Institute of Health, Washington, and six foreign laboratories. The project was organized and directed by the National Institute for Medical Research, London, in compliance with the request of the Permanent Standards Commission of the Health Organization of the League of Nations that the National Institute for Medical Research prepare a "standard for gas gangrene antitoxin (*Vibrion septique*) and arrange for preliminary laboratory investigations with a view to obtaining international agreement on a standard preparation and the definition of a unit in terms of it." The suggestion was made that the work proceed along lines similar to those adopted for gas gangrene antitoxin (*Perfringens*) (1).

In the United States a provisional unit for measuring the potency of gas gangrene antitoxin (*Vibrion septique*) had been established and was in use by the biological firms manufacturing the product (2). Great Britain (3), France (4), and the Argentine Republic (5) had also prepared standards for their own countries. The relationship of these units was found to be as follows (2):

American.....	1
British.....	1/2. 3
French.....	1/4. 4
Argentine Republic.....	1/3. 7

The National Institute for Medical Research, London, proposed that an amount exactly one-half the United States provisional unit be used as the basis of the international tests, leaving final decision

as to the size of the unit for a later meeting of the Permanent Standards Commission. This unit was designated as the provisional international unit, or the P unit. The proposal was acceded to by Dr. Weinberg, of the Pasteur Institute, Paris, and by this country.

The necessary reagents and protocols of tests were received from London in January 1934. The proposed international standard antitoxin was a dried antitoxin which had been received by the National Institute for Medical Research from Dr. Weinberg, and which had been placed in approximately 1-gram amounts into ampoules and subjected to dehydration over phosphorus pentoxide, *in vacuo*, until absolutely dry. The ampoules were filled with pure dry nitrogen, sealed, and stored in the dark at a temperature of -4°C . Tests by Dr. Hartley indicated that one provisional international unit was contained in 0.2377 mg of the dried serum. This was equivalent to 0.1566 mg of the provisional American standard.

The materials supplied for the international tests were described in the memorandum accompanying the protocols as follows:

1. A solution of gas gangrene antitoxin (*Vibrio septique*) prepared from the dry preparation made as a provisional standard at the National Institute of Health, Washington, U. S. A. (label marked 1).

In accordance with instructions received from Dr. McCoy, the contents of 1 ampoule were dissolved in 125 cc of a glycerol-saline solution (glycerol, 2 parts; physiological salt solution, 1 part). Of this solution, 1 cubic centimeter contains 100 P units.

2. A solution of gas gangrene antitoxin (*Vibrio septique*) prepared by dissolving an accurately weighed quantity of the proposed international standard preparation in glycerol (2 parts), saline (1 part) (label marked 2). According to our determinations, 1 P unit is contained in 0.2377 mg of the proposed international standard preparation. The volume of the solution was finally adjusted so that 1 cc contains 23.77 mg of the dry standard preparation, i. e., in this case also, 1 cubic centimeter contains 100 P units.

3. A dry preparation of gas gangrene toxin (*Vibrio septique*) (label marked 3). This has been prepared by precipitating the germ-free filtrate from a 3-day growth of the organism with ammonium sulphate, removing the precipitate, and drying it over phosphorus pentoxide * * *.

4. Two sealed ampoules each containing about 2.5 cc of a gas gangrene antitoxin (*Vibrio septique*) of unstated potency, for the purpose of trial assay (label marked 4).

The methods used at the National Institute for Medical Research, London, in carrying out the tests were described as follows:

The assay of the dry standard preparation of gas gangrene antitoxin (*Vibrio septique*) proposed for international adoption, in comparison with the American preparation, has been carried out by two methods:

- A. By the intravenous injection of mixtures of toxin and antitoxin into the tail veins of mice weighing from 17 to 20 gm; and

- B. By the intracutaneous injection of mixtures of toxin and antitoxin into the shaved, or depilated, flanks of white or light-colored guinea pigs weighing from 300 to 400 gm.

A. The mouse-intravenous method

By adopting the following procedure the "test dose" of the toxin supplied and the relative potencies of the glycerol solutions of the two preparations of gas gangrene antitoxin (*Vibrion septique*) may be determined in one and the same experiment:

1. 1 cc of the solution of the American provisional standard antitoxin is diluted to 20 cc.
2. 1 cc of the solution of the proposed international standard antitoxin is diluted to 20 cc.
3. A quantity of the dried toxin, V. S. 10, is quickly and accurately weighed out and dissolved in physiological salt solution, the final volume being adjusted so that 1 cc of the toxin solution contains 20 mg.
4. Mixtures of each of the antitoxin dilutions are made so that 0.5 cc of each mixture (the quantity injected intravenously into a mouse) contains 0.2 cc of the diluted antitoxin (1 P unit) + varying quantities of the toxin solution. Suitable quantities of the toxin for purposes of the test will probably be found to be between 3.5 and 4.5 mg * * *.

The mixtures are allowed to stand at room temperature for 1 hour and then 0.5 cc quantities are injected intravenously into groups of mice. The mice should be drawn from a uniform stock and should be as nearly equal in weight as practicable (17 to 20 gm). For the final comparison of the two solutions we have used groups of six or more mice for each mixture of toxin and antitoxin tested. The mice are kept under observation for 3 days; the majority of the animals which succumb to the injection die during the first 48 hours, but deaths occurring within 72 hours of the injection are regarded as significant.

In accordance with the protocols submitted, tests were performed with the reagents received. The comparative results of the British and American experiments to determine the "test dose" of the British toxin (V. S. 10) against 1 P unit of the 2 antitoxin solutions are shown in table 1. The "test dose" of the toxin used may be considered to be that amount of toxin which when mixed with one P unit of antitoxin and injected intravenously into a group of mice will kill some but not all the mice in the group. Experiment I shows this dose to be about 4.1 mg of the toxin used.

TABLE 1.—Comparison of American and P (provisional international) sera. Mouse intravenous method. Protocol of experiments made to determine the "test dose" of British toxin V. S. 10

I. DOSE OF TOXIN VARIED, ANTITOXIN CONSTANT

Toxin dose, mg	+0.2 cc=1 P unit (American solution)				+0.2 cc=1 P unit (proposed international solution)			
	Number of mice	Number dying	Number surviving	Proportion surviving	Number of mice	Number dying	Number surviving	Proportion surviving
4.7-----	6	6	0	1% (9%)	6	6	0	9% (9%)
4.4-----	6	5	1	3% (9%)	6	5	1	3% (9%)
4.1-----	6	3	3	3% (7%)	6	1	5	5% (9%)
3.8-----	6	0	6	9% (9%)	6	0	6	9% (9%)

IIA. DOSE OF TOXIN CONSTANT (4.1 MG), DOSE OF ANTITOXIN VARIED

Toxin dose, mg	Solution of American provisional standard antitoxin				Solution of proposed international standard antitoxin				
	P units	Number of mice			Proportion surviving	Number of mice			Proportion surviving
		Inoculated	Dying	Surviving		Inoculated	Dying	Surviving	
4.1-----	0.9	6	6	0	1% (3%)	6	5	1	3% (3%)
	1.0	6	3	3	3% (5%)	6	1	5	5% (5%)
	1.1	6	0	6	9% (9%)	6	0	6	9% (9%)

IIB. DOSE OF TOXIN CONSTANT (4.2 MG), DOSE OF ANTITOXIN VARIED

Toxin dose, mg	Solution of American provisional standard antitoxin				Solution of proposed international standard antitoxin				
	P units	Number of mice			Proportion surviving	Number of mice			Proportion surviving
		Inoculated	Dying	Surviving		Inoculated	Dying	Surviving	
4.2-----	0.9	6	6	0	1% (1/2)	6	6	0	9% (9/2)
	1.0	6	4	2	3% (1/2)	6	2	4	3% (1/2)
	1.1	6	0	6	9% (1 1/2)	6	1	5	5% (1 1/2)

¹ Figures in parentheses are the results of tests by the National Institute for Medical Research, London.

In experiment IIa, the "test dose" indicated in the preceding experiment was mixed with varying doses of antitoxin, and the results show that the dose of toxin may be slightly too low. In experiment IIb the dose of toxin was raised to 4.2 mg.

In the three experiments there is close agreement between the British and the American results (British results in parenthesis).

Similar tests were then carried out using the same reagents except that the American toxin was substituted for the British (table 2). Experiment I indicates that the "test dose" of the American toxin is about 5 mg. Experiment II, in which the dose of toxin was constant and the dose of antitoxin varied, shows that this amount may be a trifle high. Experiment III shows the results obtained when

one-half the P unit was tested against one-half the amount of toxin used in the preceding test, i. e., 2.5 mg, which is the "test dose" of toxin which had been in use in this country in determinations of the potency of antitoxins in terms of the American provisional unit. It may be noted that there are more survivals on proportionate doses with the more dilute reagents than with the more concentrated.

TABLE 2.—Comparison of American and P (provisional international) sera: Mouse intravenous method. Protocol of experiments made to determine the "test dose" of American toxin V. S. 1

I. DOSE OF TOXIN VARIED, ANTITOXIN CONSTANT

Toxin dose, mg	+0.2 cc=1 P unit (American solution)				+0.2 cc=1 P unit (proposed international solution)			
	Number of mice	Number dying	Number surviving	Proportion surviving	Number of mice	Number dying	Number surviving	Proportion surviving
5.6-----	6	6	0	0%	6	6	0	0%
5.3-----	6	6	0	0%	6	6	0	0%
5.0-----	6	3	3	50%	6	4	2	33%
4.7-----	6	0	6	100%	6	2	4	66%
4.4-----	6	0	6	100%	6	0	6	100%

II. DOSE OF TOXIN CONSTANT (5.0 MG), DOSE OF ANTITOXIN VARIED

Toxin dose, mg	Solution of American provisional standard antitoxin					Solution of proposed international standard antitoxin				
	P units	Number of mice			Proportion surviving	Number of mice			Proportion surviving	
		Inoculated	Dying	Surviving		Inoculated	Dying	Surviving		
5.0-----	0.9	6	6	0	0%	6	6	0	0%	
	1.0	6	5	1	16%	6	5	1	16%	
	1.1	6	2	4	66%	6	1	5	83%	

III. REPETITION OF ABOVE EXPERIMENT USING 1/2 UNIT OF ANTITOXIN AND 1/2 THE AMOUNT OF TOXIN (2.5 MG)

Toxin dose, mg	Solution of American provisional standard antitoxin					Solution of proposed international standard antitoxin				
	P units	Number of mice			Proportion surviving	Number of mice			Proportion surviving	
		Inoculated	Dying	Surviving		Inoculated	Dying	Surviving		
2.5-----	0.45	6	3	3	50%	6	5	1	16%	
	.5	6	2	4	66%	6	3	3	50%	
	.55	6	0	6	100%	6	0	6	100%	

A sample of an antitoxin of unknown potency was submitted for test to the various collaborating laboratories. The potency was indicated to lie between 100 and 200 provisional international units. A preliminary test was made against the British toxin and the final tests were made against both the British and American toxins. The results with both toxins are in close agreement, and the potency is shown to be between 160 and 170 units. The results returned by the other laboratories participating in the tests as reported by Dr. Hartley were 150-155, 154, 155, 160, 160, and 170.

TABLE 3.—*Determination of P units in a sample of Vibriion septique antitoxin of unknown potency*

I. PRELIMINARY TEST

Number of P units tested for—	Amount of $\frac{1}{50}$ dilution of antitoxin	4.1 mg British toxin			
		Number of mice injected	Number dying	Number surviving	Proportion surviving
	Cc				
200.....	0.10	2	2	0	$\frac{0}{2}$
166.....	.12	2	0	2	$\frac{2}{2}$
143.....	.14	2	0	2	$\frac{2}{2}$
125.....	.16	2	0	2	$\frac{2}{2}$
111.....	.18	2	0	2	$\frac{2}{2}$
100.....	.20	2	0	2	$\frac{2}{2}$

II. FINAL TEST

Number of P units tested for—	Amount of $\frac{1}{50}$ dilution of antitoxin	4.2 mg British toxin				5 mg American toxin			
		Number of mice injected	Number dying	Number surviving	Proportion surviving	Number of mice injected	Number dying	Number surviving	Proportion surviving
	Cc								
200.....	0.10	6	6	0	$\frac{0}{6}$	6	6	0	$\frac{0}{6}$
190.....	.105	6	6	0	$\frac{0}{6}$	6	6	0	$\frac{0}{6}$
180.....	.111	6	5	1	$\frac{1}{6}$	6	6	0	$\frac{0}{6}$
170.....	.117	6	2	4	$\frac{4}{6}$	6	5	1	$\frac{1}{6}$
160.....	.125	6	1	5	$\frac{5}{6}$	6	2	4	$\frac{4}{6}$
150.....	.133	6	0	6	$\frac{6}{6}$	6	0	6	$\frac{6}{6}$

The specimen contains 160-170 P units.

In addition to the tests on the reagents submitted, comparative tests were made to determine the potency of one commercial antitoxin using both the British and the American toxins. The results are shown in table 4.

TABLE 4.—Tests to determine the potency of a commercial *Vibrión septique antitoxin*

L. PROTOCOL OF JUNE 3, 1933, SHOWING RESULTS OF TEST ON SERUM 259 ACCORDING TO THE METHOD OF THE NATIONAL INSTITUTE OF HEALTH, WASHINGTON

Number of units tested for	Dilution of antitoxin	Amount of dilution	Amount of toxin (American)	Number of mice			Proportion surviving
				Injected	Dying	Surviving	
		Cc	Mg				
90.....	$\frac{1}{60}$	0.25	2.5	6	0	6	$\frac{9}{6}$
100.....	$\frac{1}{100}$.25	2.5	6	0	6	$\frac{9}{6}$
110.....	$\frac{1}{110}$.25	2.5	6	4	2	$\frac{2}{6}$
120.....	$\frac{1}{120}$.25	2.5	6	6	0	$\frac{9}{6}$
50 (U. S. standard).....	$\frac{1}{60}$.25	2.5	6	2	4	$\frac{4}{6}$

The serum contains 110 provisional American units.

I. PROTOCOL OF MAR. 15, 1934, SHOWING RESULTS OF TEST ON SERUM 259 ACCORDING TO THE PROPOSED INTERNATIONAL METHOD

Number of units tested for	Dilution of antitoxin	Amount of dilution	Amount of toxin (British)	Number of mice			Proportion surviving
				Injected	Dying	Surviving	
		Cc	Mg				
180.....	$\frac{1}{20}$	0.111	4.2	6	0	6	$\frac{9}{6}$
200.....	$\frac{1}{20}$.100	4.2	6	0	6	$\frac{9}{6}$
220.....	$\frac{1}{20}$.091	4.2	6	4	2	$\frac{2}{6}$
240.....	$\frac{1}{20}$.83	4.2	6	6	0	$\frac{9}{6}$

The serum contains 220 international units.

The first test had been carried out 9 months previously, using our own toxin and the method which was in use for determining the potency in terms of the American provisional unit. The second test was made with the British toxin according to the proposed international method. The results are in exact agreement, the serum containing 110 of the American provisional units and 220 of the proposed international units.

THE GUINEA PIG INTRACUTANEOUS METHOD

In addition to the mouse intravenous method it was recommended that the guinea pig intracutaneous method be used. This method was described as being "simple, convenient, and economical" and "yielding results comparable in accuracy with those obtained by the mouse intravenous method." White or light-colored guinea pigs weighing from 300 to 400 grams were recommended. As a result of the tests performed at the National Institute for Medical Research, London, it had been found that 0.5 mg of the toxin supplied (V. S. 10) produced a large necrotic lesion, while 0.05 mg produced a small reaction and 0.025 mg failed to produce any reaction. It was recommended that each observer decide for himself as to the most desirable end point. The antitoxin dilutions were to be made up as for the mouse test (1 to 20, so that 1 cc contained 5 units) and the toxin

dilution to such a strength that 1 cc contained 30 mg of the British toxin.

The tests were carried out as in the case of the mice, first determining the "test dose" of toxin against one provisional unit of antitoxin using varying doses of toxin. With the "test dose" of toxin established, this amount of toxin was tested against 1.1, 1.0, and 0.9 units of antitoxin. The mixtures of toxin and American standard antitoxin were injected on one flank of the guinea pig and the mixtures of toxin and the proposed international standard antitoxin on the other flank. Observations were made in 24 and 48 hours. The results are shown in table 5.

TABLE 5.—Comparison of American and P (provisional international) sera Guinea pig intracutaneous method. Protocol of experiments made to determine the "test dose" of British toxin V. S. 11 and American toxin V. S. 1

(a) TOXIN DOSE VARIED, ANTITOXIN DOSE CONSTANT

I. BRITISH TOXIN V. S. 11

Antitoxin solution	Amount	Mixture prepared, volume 2 cc		Dose of mixture injected	Result after 48 hours
		Toxin	Antitoxin		
American.....	Cc	Mg	1 c c = 5 units.....	0.2 cc.....	{ Large reaction; necrosis. Rather large reaction; necrosis. Moderate reaction; necrosis. Slight reaction. No reaction. Do.
	0.9	27			
	.85	25.5			
	.8	24			
	.75	22.5			
Proposed international.....	.7	21	1 c c = 5 units.....	.2 cc.....	{ Large reaction; necrosis. Rather large reaction; necrosis. Moderate reaction; necrosis. Slight reaction. No reaction. Do.
	.65	19.5			
	.9	27			
	.85	25.5			
	.8	24			

II. AMERICAN TOXIN V. S. I

Antitoxin solution	Amount	Mixture prepared, volume 2 cc		Dose of mixture injected	Result after 48 hours
		Toxin	Antitoxin		
American.....	Cc	Mg	1 c c = 5 units.....	0.2 cc.....	{ Marked reaction; necrosis. Do. Moderate reaction; necrosis. Slight reaction. No reaction. Do.
	0.9	27			
	.85	25.5			
	.8	24			
	.75	22.5			
Proposed international.....	.7	21	1 c c = 5 units.....	.2 cc.....	{ Marked reaction; necrosis. Do. Moderate reaction; necrosis. Slight reaction. No reaction. Do.
	.65	19.5			
	.9	27			
	.85	25.5			
	.8	24			

TABLE 5.—Comparison of American and P (provisional international) sera. Guinea pig intracutaneous method. Protocol of experiments made to determine the "test dose" of British toxin V. S. 11 and American toxin V. S. 1—Continued

(b) TOXIN DOSE CONSTANT, ANTITOXIN DOSE VARIED

AMERICAN TOXIN V. S. 1

Antitoxin solution	Mixture prepared, volume 2 cc			Volume of mix- ture in- jected	Result after 48 hours
	Toxin	Antitoxin			
		Cc	P units		
American.....	Mg 24	0.90	4.50	0.2 cc.....	Large reaction; necrosis. Do. Moderate reaction. Slight reaction. No reaction. Large reaction; necrosis. Do. Moderate reaction. Slight reaction. No reaction.
		.95	4.75		
		1.00	5.00		
		1.05	5.25		
		1.10	5.50		
Proposed international.....	24	.90	4.50		
		.95	4.75		
		1.00	5.00		
		1.05	5.25		
		1.10	5.50		

As in the case of the mouse tests, the tests were carried out first with the British toxin (a second lot V. S. 11) and then with the American toxin. The results with the two antitoxins against each toxin checked closely with those obtained by the National Institute for Medical Research, London.

The intracutaneous test was also used in determining the strength of the antitoxin of unknown potency submitted for test. The American toxin was used in this test. The results (table 6) confirmed those obtained in mice.

TABLE 6.—The intracutaneous guinea pig method. The sample of *Vibrion septique* antitoxin of unknown potency (American toxin)

No.	Toxin 24 mg in 1 cc	Antitoxin 1/20 dilution	Salt solution	Volume of mixture injected intracutane- ously	Actual quanti- ties injected		Result after 48 hours
					Toxin	Anti- toxin	
		Cc	Cc	Cc	Mg	Cc	
1.....	0.8	0.555.....	0.645.....	0.2	2.4	0.00278.....	Large reaction, necrosis. Do. Moderate reaction. Slight reaction. No reaction. Moderate reaction.
2.....		.585.....	.615.....			.00293.....	
3.....		.625.....	.575.....			.00313.....	
4.....		.675.....	.525.....			.00338.....	
5.....		.714.....	.496.....			.00367.....	
Control.....	.8	5 P units in 1 cc.	0	.2	2.4	0.5 unit	

The sample of antitoxin contains 160 to 170 P (proposed international) units per cc.

The results obtained in the above tests were submitted to the meeting of the Permanent Commission on Biological Standardization of the Health Organization of the League of Nations at Copenhagen, in August 1934, as were those of the six other participating laboratories. At this meeting the following resolution was adopted:

"The Commission recommends:

"1. That the dry preparation of gas gangrene antitoxin (*Vibrio septique*), prepared at the National Institute for Medical Research, London, from material supplied by Dr. Weinberg of the Pasteur Institute, Paris, be accepted as the international standard for this antitoxin and that the specific antitoxic activity contained in 0.2377 mg of the dried standard preparation be defined as the international unit."

(The antitoxin in 0.2377 mg of the international unit is equivalent to one-half the amount of antitoxin contained in the provisional American unit, viz 0.1566 mg.)

In accordance with the international agreement regarding the size of the unit, the following statement was issued by the National Institute of Health, Washington:

NATIONAL INSTITUTE OF HEALTH,
Twenty-fifth and E Streets NW., Washington, D. C.,
September 30, 1934.

It is proposed to adopt as the official unit for the measurement of the potency of gas gangrene antitoxin (*Vibrio septique*) the equivalent of the International unit adopted by the Permanent Commission on Biological Standardization of the Health Organization of the League of Nations. This unit is one-half that previously proposed as a provisional unit for the United States (Pub. Health Rep., 1934, 49, 251) and is that amount of antitoxin contained in 0.1566 mg of the dried standard serum prepared at the National Institute of Health. The dried serum as dissolved and diluted for distribution contains 100 units in 1 cc.

The standard unit will be distributed on special request addressed to the director of the National Institute of Health.

It is expected that this unit will be employed by all producers not later than January 1, 1935.

G. W. McCoy,
Director, National Institute of Health.

Following the adoption of a unit one-half that previously in use as a unit of measurement for this country, the question arose whether it would be advisable to carry out the test as it had been done previously, merely changing the factor used in calculating the number of units, or whether it would be better to use the international method. Our method in the mouse test was based on the use of such a dilution of the antitoxin that 1/2 the unit (1/4 the former provisional unit) was contained in 0.25 cc of a 1/50 dilution of the glycerinated antitoxin. By chance the "test dose" of the standard toxin against this dose of antitoxin was 0.25 cc of a 1/100 dilution (10 mg to 1 cc) of the toxin. The

sum of the doses of toxin and antitoxin was therefore exactly 0.5 cc, a convenient dose for the intravenous inoculation of mice. The international method was based on the use of such a dilution of the standard antitoxin that 1 unit was contained in 0.2 cc of a 1/20 dilution of the glycerinated antitoxin. The test dose of the British toxin against this amount of toxin was 4.2 mg, or 2.1 cc of a 1/50 dilution of the toxin (20 mg per cc). The doses of toxin and antitoxin were therefore twice as concentrated in the international tests as they were in the American tests.

In order to determine the effect of varying the dilutions of toxin and antitoxin in multiple proportions, tests were carried out in mice and also in rabbits. The antitoxin was diluted 1/25, 1/50, and 1/100 and the toxin was diluted so that 1 c c contained 20 mg, 10 mg, and 5 mg. The results of the test in mice are shown in table 7.

TABLE 7.—*Effect of diluting the toxin and antitoxin in multiple proportions*

Antitoxin			Toxin		Number of mice	Proportion surviving
Dilution	Amount of dilution	Number of units	Dilution	Amount of dilution		
	<i>C c</i>		<i>Mg per c c</i>	<i>C c</i>		
1/100	0.25	1/4	5	0.25	6	3/6
1/50	.25	1/2	10	.25	6	3/6
1/25	.25	1	20	.25	6	0/6

A larger proportion of mice inoculated with the more concentrated reagents died—in this case all the mice in the group of 6 inoculated with the mixture of the 1/25 dilution of antitoxin and the dilution of toxin containing 20 mg per cubic centimeter. A similar tendency was shown in tests II and III, table 2. The discrepancy is probably not sufficiently great, however, to warrant a definite fixing of one particular method of making the dilutions of the reagents to the exclusion of the others in the case of mice. A test carried out with a commercial antitoxin, however, in which a range of doses of antitoxin was tested against the "test dose" of toxin indicated that it was difficult to establish a definite end point with the more dilute solutions. The dilution 1/50 of the antitoxin, using 0.5 unit in the mouse test, appears to be entirely satisfactory.

There were marked discrepancies in the results in rabbits, on the other hand, when dilutions of toxin and antitoxin were varied in multiple proportions. Dilutions of the reagents were made as for the mice. The results are shown in table 8.

TABLE 8.—Effect of diluting the toxin and antitoxin in multiple proportions in rabbits

Antitoxin			Toxin		Number of rabbits	Proportion surviving
Dilution	Amount of dilution per kg	Number of units per cc	Dilution	Dose per kg		
	<i>Cc</i>		<i>Mg per cc</i>	<i>Mg</i>		
1/100	1.0	1	5	6.5	4	4/4
1/100	1.0	1	5	6.0	4	4/4
1/100	1.0	1	5	5.5	4	4/4
1/100	1.0	1	5	5.0	4	4/4
1/50	1.0	2	10	13	7	1/7
1/50	1.0	2	10	12	7	3/7
1/50	1.0	2	10	11	7	5/7
1/50	1.0	2	10	10	3	3/3
1/25	1.0	4	20	26	4	0/4
1/25	1.0	4	20	24	4	0/4
1/25	1.0	4	20	22	4	0/4
1/25	1.0	4	20	20	3	2/3

Using 1 cc per kg of a 1/50 dilution of the standard antitoxin against doses of toxin ranging from 11 to 13 mg per kilogram, some animals died and some survived. When the concentration of the reagents was doubled, all of the animals died; when it was halved, all of the animals survived. The more dilute solution of the antitoxin (1/100) neutralized completely the corresponding doses of toxin so that there remained unneutralized less than one minimal lethal dose, the more concentrated solution of the antitoxin (1/25) failed to neutralize completely the corresponding doses of toxin so that one or more minimal lethal doses remained unneutralized.

In view of the above results and for the reason that the rabbit and mouse tests were correlated in our original work in establishing the provisional unit, it seems desirable to continue the use of the 1/50 dilution of the glycerinated antitoxin (2 international units per cc).

It may therefore be concluded that the most satisfactory results with our standard reagents may be obtained if the test is carried out as formerly, multiplying the number of units obtained thus by the factor 2. The glycerinated solution of the antitoxin is diluted 1/50, and in the mouse test 0.25 cc of this dilution (0.5 unit) is tested against the "test dose" 2.5 mg of the standard toxin. In the rabbit test 1 cc per kilogram of the same dilution (2 units) is tested against a "test dose" of 12 to 13 mg per kilogram of the standard toxin.

SUMMARY

As one of the participants in a project to establish an international unit for use in determining the potency of gas gangrene antitoxin (*Vibrio septique*), a comparison was made of the proposed international antitoxin and the American standard antitoxin, using reagents supplied by the National Institute for Medical Research,

London. The proposed international unit was contained in 0.2377 mg of a dried serum preparation and this was equivalent to 0.1566 mg of the American standard antitoxin—one-half the American provisional unit.

Protocols of tests made at the National Institute for Medical Research, London, were submitted to the various countries participating in the tests. The correctness of the British assay of the international unit was confirmed by tests on mice (intravenous inoculation) and guinea pigs (intracutaneous inoculation), testing with both the British and American toxins.

Results obtained in evaluating a specimen of antitoxin of unknown potency agreed well with those obtained by the six other laboratories participating in the tests. A commercial antitoxin was also evaluated, using both the American and British toxins with identical results.

The proposed international unit, equivalent to one-half of the American provisional unit, was adopted as the international unit of gas gangrene antitoxin (*Vibrio septique*) at a meeting of the Permanent Commission on Biological Standardization of the Health Organization of the League of Nations in Copenhagen, Denmark, in August 1934. The international unit was adopted as the United States official unit in September 1934.

As the result of tests on mice and rabbits for the purpose of determining the best application of the international unit in the measurement of the potency of gas gangrene antitoxin (*Vibrio septique*) it was shown that the most satisfactory results with the American reagents were obtained by performing the tests as formerly in determining provisional units and multiplying by the factor 2.

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THE INFLUENCE OF VITAMIN B₂ ON HEMATOPOIESIS IN EXPERIMENTAL ANEMIA OF THE ALBINO RAT

By MAURICE I. SMITH, *Principal Pharmacologist*, and E. F. STOELMAN, *Junior Pharmacologist, United States Public Health Service, National Institute of Health*

Anemia is often a marked feature in clinical pellagra. Experimentally, Rhoads and Miller (1) observed anemia in dogs on diets producing black-tongue. The work of Castle and Strauss (2) suggests a close relationship between the extrinsic factor in pernicious anemia

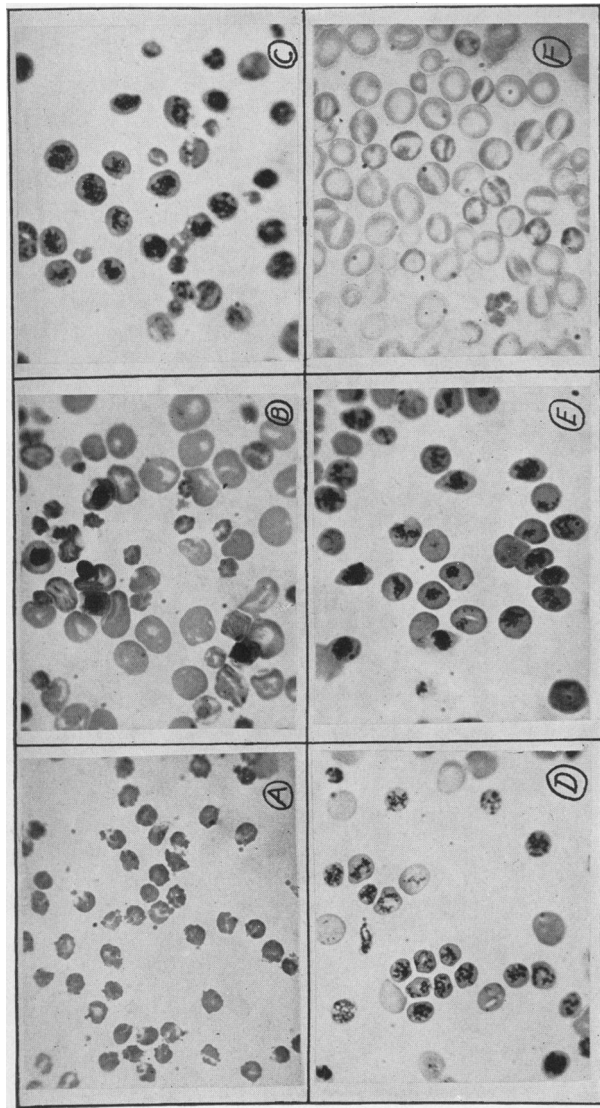
and vitamin B₂. Guha and Mapson (3) observed a diminution of erythrocytes in rats on a diet deficient in B₂, which they state was restored to normal by feeding autoclaved marmite (a yeast extract). Wills and Bilimoria (4) reported an anemic condition in monkeys on a diet deficient in vitamin B which was corrected by the addition of marmite to the diet. Certain reports to the effect that potent antianemic liver extracts are also active with respect to B₂ further suggest a possible relationship. There is, however, lack of agreement on this point (5).

We have attempted to approach this problem by comparing the rate of hemoglobin and red-blood cell regeneration in animals with a standard degree of anemia under adequate dietary conditions with that under conditions of B₂ deficiency. Albino rats were used because this species is the most suitable for rigidly controlled nutritional investigations. The type of experimental anemia employed in this work was that produced by means of phenylhydrazine. We have found this substance particularly suitable for our purpose, because of its specificity and the quantitative relationship between dosage and effect.

PHENYLHYDRAZINE ANEMIA IN THE WHITE RAT ON AN ADEQUATE DIET

The destructive action of phenylhydrazine on the blood has been well known. Bodansky (6) studied the effect of a series of hydrazine compounds in dogs and rabbits and concluded that phenylhydrazine was the most effective anemia-producing agent. We are not aware of any detailed study of the effects of phenylhydrazine in the rat which, in the course of this work, has been found to react in certain essentials quite differently from other animal species reported heretofore.

The maximum tolerated dose of phenylhydrazine hydrochloride upon intravenous injection in the normal rat in 1 percent solution is about 80 mg per kilo. While a single injection of such a dose will produce a moderate degree of anemia, it has been found that three successive daily injections of smaller doses, such as 40 to 60 mg per kilo, will produce a rather severe degree of anemia, with the hemoglobin and red cells reduced to about one-third of the normal. Such doses have no apparent toxic effects other than the destructive action on the blood and a temporary loss of weight. The blood-cell characteristics are somewhat as follows: Twenty-four hours after the last injection, Wright's preparations show numerous microcytes and poikilocytes. The cells present marked crenation, and various stages of disintegration (plate I, A). The following day polychromatic macrocytes make their appearance. Anisocytosis, poikilocytosis, polychromatophilia, and considerable numbers of nucleated red cells may now be seen (plate I, B). In a cresyl blue preparation many reticulo-



4, Blood film 24 hours after last of three intravenous injections of phenylhydrazine, showing various stages of disintegration of red blood cells. Wright's stain. $\times 800$. B, Blood film one day later. Microcytes, poikilocytes, polychromatophilic macrocytes, and erythroblasts. Wright's stain. $\times 800$. C, Preparation showing many reticulocytes along with microcytes, macrocytes, and erythroblasts. Third day. Cresyl blue counterstained with Wright's stain. $\times 800$. D and E, Eighty to ninety percent reticulocytes 3 to 4 days after the last injection of phenylhydrazine. Cresyl blue and Wright's stain. $\times 800$. F, Wright's preparation 4 days after the last injection of phenylhydrazine. Cells nearly normal except for some anisocytosis and polychromatophilia. $\times 800$. (All reduced approximately one-third.)

cytes may be seen among the microcytes (plate I, C).¹ The reticulocytes now usually rapidly increase in number so that on the third or fourth day as much as 90 percent of the circulating red-blood cells may be reticulocytes (plate I, D and E). Wright's preparations at

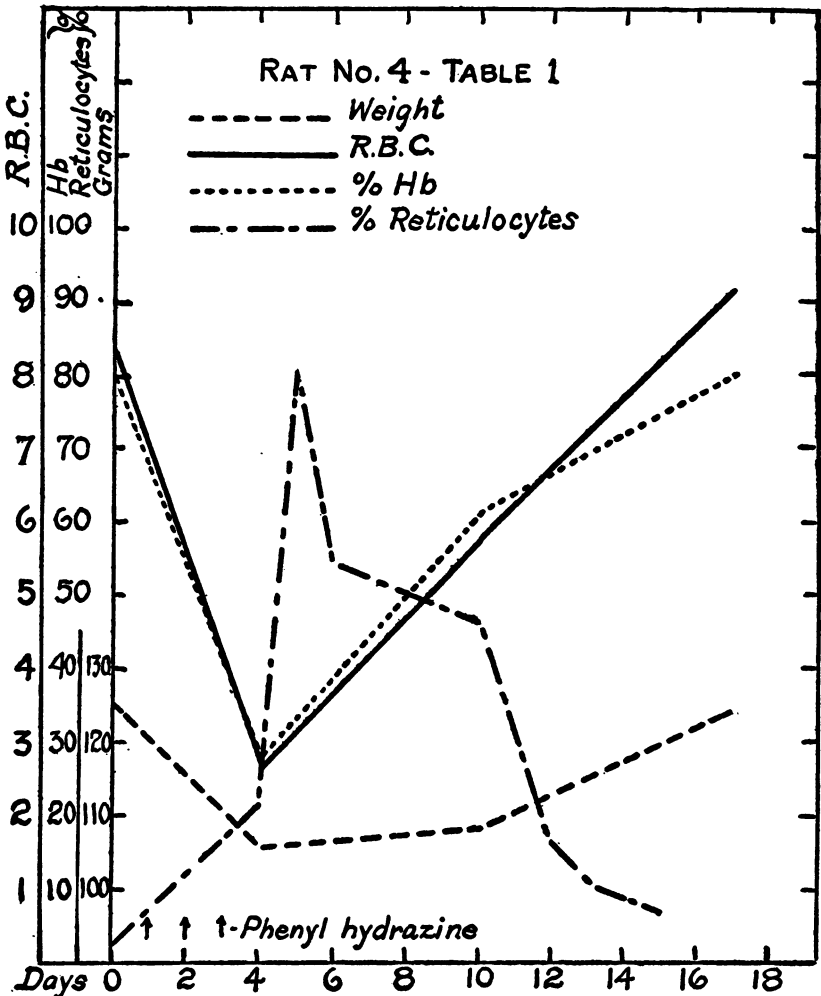


FIGURE 1.—Blood regeneration in phenylhydrazine anemia of rat on standard adequate diet.

this time usually present a nearly normal appearance, except for some polychromatophilia (plate I, F). Recovery now proceeds rapidly so that by the eighth or tenth day the blood picture presents a normal appearance, the reticulocytes being usually less than 1 percent.

¹ Reticulocytes were stained essentially by the method of Osgood and Wilhelm (9).

TABLE 1.—*Hematopoiesis in the white rat on a standard adequate diet following a standard dose of phenylhydrazine hydrochloride*

No.	Normal			Extent of anemia			Recovery				Maximum percentage of reticulocytes observed during recovery
	Weight	RBC	Hb	Weight	RBC	Hb	Days	Weight	RBC	Hb	
	<i>Grams</i>		<i>Per-cent</i>	<i>Grams</i>		<i>Per-cent</i>		<i>Grams</i>		<i>Per-cent</i>	
1	164	10.46	98	129	3.18	30	4	140	4.91	58	14 percent on fourth day.
2	185	10.30	95	150	3.97	35	10	184	8.41	88	
3	210	10.59	89	180	3.72	55	12	218	9.03	90	30 percent on fifth day.
4	126	8.29	79	106	2.58	27	6	106	5.75	61	
							13	123	9.22	79	80 percent on third day. 90 percent on fourth day.
5	130	8.73	74	114	2.49	33	7	130	8.12	87	
6	202	10.50	90	186	3.52	36	8	218	8.26	96	

¹ Killed on fifth day.

The progress of the anemia as judged by hemoglobin ² and red cell regeneration is shown in table 1, from which it will be seen that recovery is in evidence as early as the fourth day, and is nearly complete in 7 to 10 days. The color index tends to mount in the course of the anemia, and more especially during the recovery phase, indicating a relatively more rapid regeneration of hemoglobin than erythrocytes.

PHENYLHYDRAZINE ANEMIA IN RATS ON B₂ DEFICIENT DIET

In this series of experiments the extent of phenylhydrazine anemia and blood regeneration was studied in a group of rats that had been kept on a B₂ deficient diet for varying periods up to 80 days. The diet differed from that in the preceding experiments in that it contained no known source of vitamin B, while that of the preceding group contained 5 percent of dried brewer's yeast. The B₂ deficient diet of this group consisted of the following:

Casein, purified.....	18
McCullum's salt mixture (185(7)).....	4
Cod-liver oil.....	2
Olive oil.....	8
Corn starch.....	68

100

To meet the B₁ requirements, 0.04 mg of the crystalline antineuritic vitamin hydrochloride was injected intravenously at 4- to 5-day intervals. This is more than 5 times the minimal B₁ requirement of the rat.³ On this diet rats maintain body weight with but slight losses, and almost uniformly develop typical skin lesions, in from

¹ Newcomer hemoglobinometer.

² We are indebted to Dr. R. R. Williams, of Columbia University, for a generous supply of the B₁ crystals. An assay of this material by the method used in this laboratory (8) showed that it had an activity of 0.007 mg as the minimum curative dose, which is sufficient to bring about a remission in beri-beri rats for about 5 days.

6 to 12 weeks. The chief characteristics of the skin lesions are bleeding fissures in the corners of the mouth, and erythema, edema, exudation, desquamation, and even ulceration of the toes and feet. The addition of active B₂ preparations for a few days causes resumption of growth and prompt healing with disappearance of the lesions.

Rats of the above description appear to be somewhat more susceptible to the toxic effects of phenylhydrazine. Consequently, several rats receiving doses well tolerated by the animals of the preceding group died during the injection period. In the successful experiments, the dose of phenylhydrazine was reduced somewhat, so that in the experiments summarized in table 2 usually three doses of 30 to 50 mg per kilo each were given on successive days.

TABLE 2.—*Hematopoiesis in the white rat on B₂-deficient diet following a standard dose of phenylhydrazine hydrochloride*¹

No.	Days on deficient diet	Preanemic state			Extent of anemia			Recovery				Maximum percentage of reticulocytes observed during recovery
		Weight	RBC	Hb	Weight	RBC	Hb	Days	Weight	RBC	Hb	
		<i>Grams</i>		<i>Per-cent</i>	<i>Grams</i>		<i>Per-cent</i>		<i>Grams</i>		<i>Per-cent</i>	
1	47	130	10.28	90	114	3.12	32	{ 7 13	----- 112	6.22 9.72	71 86	82 percent on fourth day.
2	60	94	8.22	74	90	4.79	38	{ 7 15	----- 100	5.54 9.32	58 76	63 percent on fifth day.
3	67	122	10.12	78	114	4.78	35	{ 6 14	----- 122	4.55 7.95	62 81	60 percent on fourth day.
4	77	126	9.92	82	118	2.80	32	{ 9 14	----- 122	5.87 8.22	62 80	90 percent on third, fourth, and fifth days.
5	80	90	8.72	74	86	6.02	47	{ 11 19	----- 84	6.21 8.32	68 70	36 percent on sixth day.
6	32	164	11.46	94	148	4.35	35	11	158	8.54	92	3 percent on third and fifth days.
7	36	104	13.38	100	92	3.77	27	7	94	7.29	72	21 percent on seventh day.
8	41	142	12.04	108	118	2.78	27	13	94	7.73	72	17 percent on second, fourth, and seventh days.
9	50	130	9.74	86	106	2.79	35	15	105	8.02	56	26 percent on seventh day.

¹ Experiments 6-9 also restricted iron intake. See text.

Experiments 1-5 (table 2) show the condition of the blood in rats on B₂-deficient diet up to 80 days, the extent of anemia produced by phenylhydrazine, the extent of recovery from this anemia, and the highest degree of reticulocytosis observed during recovery. Experiments 6-9 give similar data for a group of rats subsisting on a restricted iron intake in addition to the B₂ deficiency. The restriction was effected by omitting iron citrate from salt mixture 185 of the synthetic diet. The restriction was only partial, however, for analysis of the constituents of the synthetic diet indicated 20 mg of iron per 100

gm of casein and 2.4 mg of iron per 100 gm of starch, with negligible traces for the other ingredients.⁴ Calculation showed that the rats of this group received about 0.5 mg of iron per day as against some 5 mg in the unrestricted group.

Analysis of the data in table 2 indicates that absence of B₂ from the diet neither affects the normal production of red blood cells or hemo-

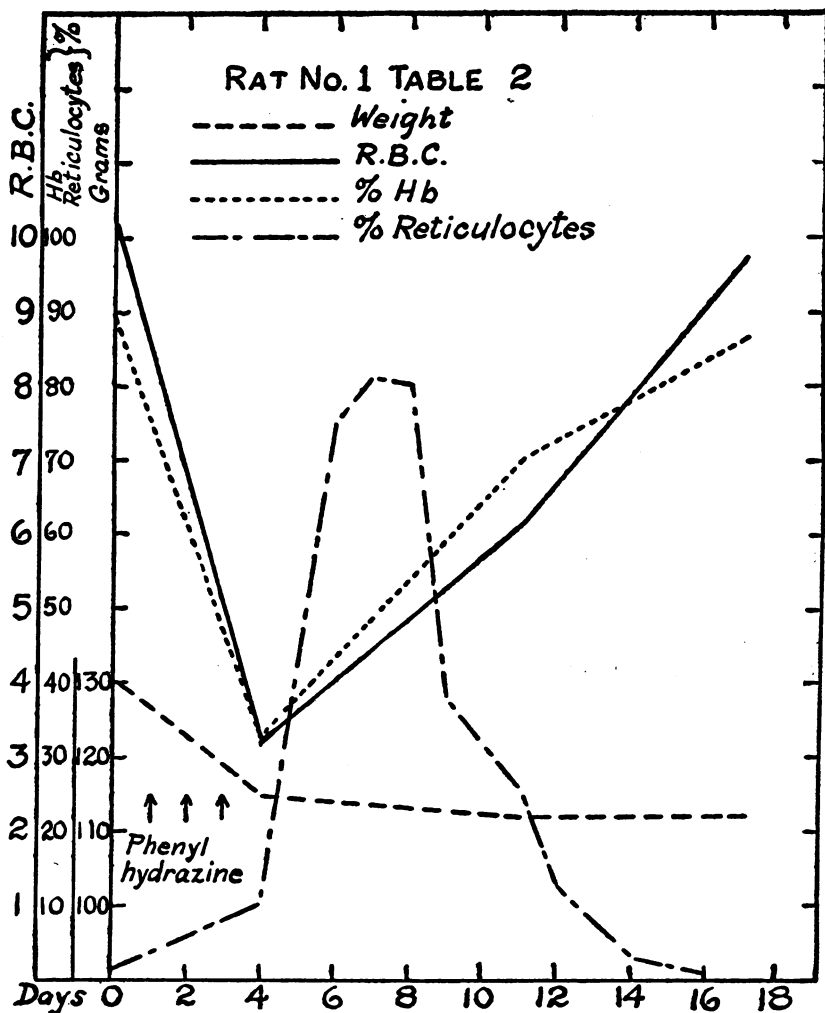


FIGURE 2.—Blood regeneration in phenylhydrazine anemia of rat on B₂ deficient diet.

globin in the rat, nor does it seriously impair the regeneration of hemoglobin and red cells destroyed by means of phenylhydrazine. There is also the same general tendency for the color index to mount during the recovery phase in this group as in the preceding group on the adequate diet. A comparison of the recovery phase in tables 1 and 2

⁴ We are indebted to Dr. E. Elvove, of this laboratory, for the iron analyses.

would, however, seem to suggest a somewhat retarded rate of regeneration especially in the group deficient in B₂ with restricted iron intake. Reticulocytosis also appears to have been less pronounced in this group of animals. In any event, the effect of B₂ deficiency is only slight, and the conclusion seems justified that vitamin B₂ is not concerned with hematopoiesis in the white rat.

SUMMARY AND CONCLUSIONS

A standard procedure for the production of a moderately severe anemia in the albino rat by means of phenylhydrazine is described.

In the normal rat this anemia is transient in character, complete recovery taking place in about 10 days. The early stages of recovery are accompanied by pronounced reticulocytosis.

Elimination of the B₂ vitamin from the dietary of the rat does not materially affect the progress of recovery from the standard phenylhydrazine anemia.

Vitamin B₂ does not appear to be concerned with hematopoiesis in the albino rat.

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COURT DECISION ON PUBLIC HEALTH

Chronic benzol poisoning held compensable under workmen's compensation act.—(Minnesota Supreme Court; *Funk v. Minnesota Mining & Mfg. Co. et al.*, 256 N. W. 889; decided October 26, 1934.) Subdivision 9 of section 4327, Mason's Minnesota Statutes 1927, provided that, for the purposes of the workmen's compensation act, only the diseases enumerated in such subdivision should be deemed to be occupational diseases and hence compensable. The said subdivision contained a schedule in two columns. In column one there was a description of the diseases and in column two a description of the processes which raised a presumption that the disease was due to the nature of the employment in which the employee was engaged. Paragraph 7 of column 1 of subdivision 9 read:

Poisoning by nitro and amido-derivatives of benzine (dinitro-benzol, anilin and others), or its sequelae.

Opposite this paragraph in column 2 was the following:

Any process involving the use of a nitro or amido-derivative of benzene or its preparations or compounds.

In the instant case the State industrial commission awarded compensation for chronic benzol poisoning. The employee had been engaged in the production of rubber cement tape. Benzol was placed in a mixing churn and rubber and other solid ingredients were added to the contents of the churn which continued the mixing process for about 24 hours. The compound or preparation was then drawn into five-gallon cans and taken to a box or hopper where it was spread upon the material to be made into tape. The employee was engaged, among other duties, in pouring the preparation into the box or hopper. In so doing he was exposed to the fumes from the benzol preparation and was disabled.

On appeal to the supreme court that court stated that the sole question presented was whether chronic benzol poisoning came within the definition contained in paragraph 7 above set out. In affirming the decision of the industrial commission the appellate court said:

It is conceded that the word "benzine" was used by the legislature as meaning benzene or benzol, C_6H_6 , and not C_nH_{2n+2} , the ordinary benzine. The effects of poisoning from the fumes of the benzol derivatives are usually acute, while those from benzol are characterized as chronic. Did the legislature intend to distinguish between benzol and its derivatives and to include the latter and exclude the former as a cause of occupational disease? It is conceded that benzol poisoning is a typical occupational disease in the general sense, and might well have been included in the coverage of subdivision 9. If the legislature intended to exclude benzol, there was no rational basis for the exclusion. We think the language of column 1 should be interpreted liberally in connection with that in column 2, where "its" evidently refers to benzol and "preparations" apparently would cover mixtures such as that which caused respondent's disability. Therefore benzol poisoning is properly compensable when occurring under the circumstances set out in column 2. The decision of the commission is affirmed and the writ discharged.

DEATHS DURING WEEK ENDED DEC. 8, 1934

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

	Week ended Dec. 8, 1934	Correspond- ing week, 1933
Data from 86 large cities of the United States:		
Total deaths.....	8,383	8,601
Deaths per 1,000 population, annual basis.....	11.7	12.0
Deaths under 1 year of age.....	592	623
Deaths under 1 year of age per 1,000 estimated live births.....	55	53
Deaths per 1,000 population, annual basis, first 49 weeks of year.....	11.3	10.9
Data from industrial insurance companies:		
Policies in force.....	67,105,185	67,326,257
Number of death claims.....	12,321	13,845
Death claims per 1,000 policies in force, annual rate.....	9.6	10.7
Death claims per 1,000 policies, first 49 weeks of year, annual rate.....	9.8	9.8

¹ Data for 81 cities.

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

CURRENT WEEKLY STATE REPORTS

These reports are preliminary, and the figures are subject to change when later returns are received by the State health officers

Reports for Weeks Ended December 15, 1934, and December 16, 1933

Cases of certain communicable diseases reported by telegraph by State health officers for weeks ended Dec. 15, 1934, and Dec. 16, 1933

Division and State	Diphtheria		Influenza		Measles		Meningococcus meningitis	
	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933
New England States:								
Maine.....	2	3	1	15	48	1	1	0
New Hampshire.....		1			7	12	0	0
Vermont.....	6	3			7	59	0	0
Massachusetts.....	21	26			195	482	3	1
Rhode Island.....	5	2			3	9	1	1
Connecticut.....	1	10	6	4	314	17	1	0
Middle Atlantic States:								
New York.....	37	54	161	128	787	584	3	3
New Jersey.....	32	25	64	20	54	99	0	0
Pennsylvania.....	75	51			989	327	2	3
East North Central States:								
Ohio.....	97	65	60	101	271	120	1	3
Indiana.....	31	65	46	61	232	39	0	2
Illinois.....	48	52	21	11	778	34	1	4
Michigan.....	16	26	19	4	191	37	0	0
Wisconsin.....	4	13	15	17	353	161	1	3
West North Central States:								
Minnesota.....	31	8			812	8	2	1
Iowa.....	15	18	31	1	784	30	0	1
Missouri.....	62	80	78	6	120	112	0	2
North Dakota.....	11	10	11	2	124	33	0	1
South Dakota.....		22			38	217	1	0
Nebraska.....	9	6			42	13	1	1
Kansas.....	8	29		1	207	43	1	0
South Atlantic States:								
Delaware.....	2	1			1	1	0	1
Maryland.....	19	25	12	20	81	21	0	0
District of Columbia.....	9	10	1	1	5	25	0	1
Virginia.....	52	67			165	87	4	0
West Virginia.....	47	47	94	14	236	6	2	2
North Carolina.....	53	60	22	14	505	503	4	2
South Carolina.....	5	14	419	459	4	125	0	0
Georgia.....	13	35				299	2	1
Florida.....	15	22		3	8		1	1

See footnotes at end of table.

Cases of certain communicable diseases reported by telegraph by State health officers for weeks ended Dec. 15, 1934, and Dec. 16, 1933—Continued

Division and State	Diphtheria		Influenza		Measles		Meningococcus meningitis	
	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933
East South Central States:								
Kentucky.....	36	60	38	25	152	4	2	1
Tennessee ¹	30	44	59	113	76	269	0	1
Alabama ²	20	33	56	47	44	114	2	1
Mississippi ³	15	19					0	0
West South Central States:								
Arkansas.....	15	18	44	32	10	294	0	0
Louisiana.....	30	30	14	11	19	1	0	0
Oklahoma ⁴	10	70	98	53	1	39	0	0
Texas ⁵	88	207	288	143	19	193	1	0
Mountain States:								
Montana.....	8	7	14	5	81	2	1	2
Idaho.....					5	10	0	0
Wyoming.....	1				15	34	0	0
Colorado.....	7	12			287	4	2	0
New Mexico.....		10	2		49	74	0	0
Arizona.....	2	5	18	20	5	8	0	0
Utah ⁶	1		2	2	12	129	0	1
Pacific States:								
Washington.....	3	6		3	37	219	1	0
Oregon.....		1	36	17	27	18	0	0
California.....	63	32	41	48	171	137	2	3
Total.....	1,055	1,404	1,671	1,301	8,371	5,048	43	43

Division and State	Poliomyelitis		Scarlet fever		Smallpox		Typhoid fever	
	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933
New England States:								
Maine.....	2	0	35	10	0	0	3	1
New Hampshire.....	0	1	8	12	0	0	0	1
Vermont.....	0	0	27	15	0	0	1	3
Massachusetts.....	0	0	170	222	0	0	2	3
Rhode Island.....	0	0	13	17	0	0	0	1
Connecticut.....	0	0	39	55	0	0	1	0
Middle Atlantic States:								
New York.....	1	7	429	466	0	0	16	8
New Jersey.....	0	1	129	146	0	0	5	5
Pennsylvania.....	2	1	542	418	0	0	20	12
East North Central States:								
Ohio.....	4	4	549	553	5	2	19	7
Indiana.....	6	1	203	183	3	4	4	4
Illinois.....	0	1	556	379	3	3	16	7
Michigan.....	1	1	283	293	0	0	7	11
Wisconsin.....	2	1	487	101	17	64	2	0
West North Central States:								
Minnesota.....	0	0	187	100	6	3	0	2
Iowa ⁷	1	0	60	87	1	1	3	4
Missouri.....	0	0	84	131	2	3	9	1
North Dakota.....	0	0	59	34	0	0	1	1
South Dakota.....	0	0	19	11	6	0	1	0
Nebraska.....	0	0	29	28	20	2	1	5
Kansas.....	1	0	77	115	2	7	2	5
South Atlantic States:								
Delaware.....	0	1	20	6	0	0	0	1
Maryland ⁸	1	0	117	80	0	0	6	9
District of Columbia.....	0	0	17	14	0	0	0	1
Virginia.....	0	0	119	128	14	0	12	16
West Virginia.....	1	0	153	144	0	0	21	7
North Carolina ⁹	0	0	84	131	0	1	4	6
South Carolina ⁹	0	0	3	21	0	0	1	6
Georgia ¹	0	3		27	0	0	11	11
Florida.....	0	0	4	5	0	0	2	2

See footnotes at end of table.

Cases of certain communicable diseases reported by telegraph by State health officers for weeks ended Dec. 15, 1934, and Dec. 16, 1933—Continued

Division and State	Poliomyelitis		Scarlet fever		Smallpox		Typhoid fever	
	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933	Week ended Dec. 15, 1934	Week ended Dec. 16, 1933
East South Central States:								
Kentucky.....	0	0	66	114	0	0	11	5
Tennessee ¹	1	0	61	129	1	2	12	6
Alabama ²	0	1	22	37	1	0	15	4
Mississippi ³	1	1	24	25	1	3	2	0
West South Central States:								
Arkansas.....	0	0	19	15	9	2	13	3
Louisiana.....	0	1	21	14	0	30	8	15
Oklahoma ⁴	0	1	27	47	1	0	15	5
Texas ⁵	0	0	78	122	1	12	42	35
Mountain States:								
Montana.....	3	0	37	15	1	18	2	2
Idaho.....	0	0	2	8	0	1	6	0
Wyoming.....	0	0	18	12	1	0	0	0
Colorado.....	0	0	245	19	1	3	1	6
New Mexico.....	0	0	20	33	0	0	10	6
Arizona.....	0	0	10	13	0	0	6	1
Utah ¹	0	0	37	10	1	11	1	0
Pacific States:								
Washington.....	8	4	44	37	52	2	2	3
Oregon.....	1	1	52	44	0	9	2	2
California.....	14	6	260	205	16	8	5	20
Total.....	50	37	5,527	4,831	165	191	323	255

¹ New York City only.

² Week ended earlier than Saturday.

³ Typhus fever, week ended Dec. 15, 1934, 31 cases, as follows: Maryland, 1; North Carolina, 1; South Carolina, 2; Georgia, 13; Alabama, 6; Texas, 9.

⁴ Rocky Mountain spotted fever, week ended Dec. 15, 1934, Tennessee, 1 case.

⁵ Exclusive of Oklahoma City and Tulsa.

SUMMARY OF MONTHLY REPORTS FROM STATES

The following summary of cases reported monthly by States is published weekly and covers only those States from which reports are received during the current week.

State	Menin- gococ- menin- gitis	Diph- theria	Infl- uenza	Malaria	Measles	Pe- lagra	Polio- mye- litis	Scarlet fever	Small- pox	Ty- phoid fever
<i>August 1934</i>										
Colorado.....	2	15		2			4	58	3	57
<i>September 1934</i>										
Colorado.....	1	25			29		3	104	8	25
Georgia.....	3	67	107	517	11	21	3	56	0	248
<i>October 1934</i>										
Arkansas.....					4				2	
Puerto Rico.....		54	74,764	1,242	113		0		0	11
<i>November 1934</i>										
California.....	3	207	117	12	696	6	100	767	10	56
Florida.....		69	1	102	19	2	1	27	0	8
Iowa.....	2	51	4		905		5	285	5	11
Missouri.....	6	322	187	65	411		5	408	17	121
New Jersey.....	5	92	88	1	172		1	451	0	25
New Mexico.....	4	27	16	5	190		2	101	1	83
New York.....	16	179		12	3,119		8	1,473	0	50
Ohio.....	3	440	115	3	652		17	2,390	7	31
Tennessee.....	1	254	172	119	57	9	6	388	0	61

August 1934		November 1934—Continued		November 1934—Continued	
Colorado:	Cases	Chicken pox—Con.	Cases	Puerperal septicaemia:	Cases
Anthrax.....	1	Ohio.....	2,697	New Mexico.....	1
Chicken pox.....	5	Tennessee.....	189	Ohio.....	4
Dysentery.....	1	Dengue:		Rabies in animals:	
Impetigo contagiosa.....	1	Florida.....	51	California.....	51
Lethargic encephalitis.....	5	Diarrhea and enteritis:		Missouri.....	15
Mumps.....	23	Ohio (under 2 years).....	13	New Jersey.....	13
Undulant fever.....	1	Dysentery:		New York ¹	1
Vincent's infection.....	1	California (amoebic).....	7	Rabies in man:	
Whooping cough.....	123	California (bacillary).....	39	Missouri.....	2
<i>September 1934</i>		Florida.....	1	Relapsing fever:	
Colorado:		Missouri.....	40	California.....	1
Chicken pox.....	30	New Jersey.....	7	Scabies:	
Dysentery.....	1	New Mexico.....	12	Tennessee.....	3
Impetigo contagiosa.....	1	New York (amoebic).....	4	Septic sore throat:	
Lethargic encephalitis.....	1	New York (bacillary).....	91	California.....	16
Mumps.....	17	Ohio.....	01	Missouri.....	37
Vincent's infection.....	13	Tennessee.....	12	New York.....	23
Whooping cough.....	68	Food poisoning:		Ohio.....	263
Georgia:		California.....	21	Tennessee.....	18
Chicken pox.....	9	Ohio.....	17	Tetanus:	
Dengue.....	9	German measles:		California.....	8
Dysentery (amoebic).....	8	California.....	51	Missouri.....	2
Dysentery (bacillary).....	16	New Jersey.....	33	New Jersey.....	1
Hookworm disease.....	164	New Mexico.....	16	New York.....	7
Lethargic encephalitis.....	2	New York.....	127	Tennessee.....	1
Mumps.....	26	Ohio.....	168	Trachoma:	
Paratyphoid fever.....	1	Tennessee.....	2	California.....	17
Rabies in man.....	1	Granuloma, coccidioidal:		Missouri.....	5
Septic sore throat.....	35	California.....	4	New Jersey.....	1
Tularaemia.....	2	Impetigo contagiosa:		New Mexico.....	1
Typhus fever.....	48	Tennessee.....	11	Ohio.....	1
Undulant fever.....	9	Jaundice, epidemic:		Tennessee.....	55
Whooping cough.....	109	California.....	1	Trichinosis:	
<i>October 1934</i>		Lead poisoning:		California.....	9
Arkansas:		New Jersey.....	2	Iowa.....	2
Chicken pox.....	3	Ohio.....	10	New Jersey.....	1
Mumps.....	5	Lethargic encephalitis:		New York.....	23
Trachoma.....	11	California.....	4	Tularaemia:	
Tularaemia.....	1	Iowa.....	1	Iowa.....	4
Undulant fever.....	1	Missouri.....	3	Missouri.....	4
Whooping cough.....	111	New Jersey.....	2	New York.....	13
Puerto Rico:		New York.....	8	Tennessee.....	1
Chicken pox.....	26	Ohio.....	1	Typhus fever:	
Dysentery.....	67	Tennessee.....	2	Florida.....	1
Filarisis.....	3	Milk sickness:		New York.....	2
Mumps.....	27	New Mexico.....	1	Undulant fever:	
Ophthalmia neonatorum.....	3	Mumps:		California.....	11
Puerperal septicaemia.....	8	California.....	411	Iowa.....	18
Tetanus.....	9	Florida.....	7	Missouri.....	7
Tetanus, infantile.....	9	Iowa.....	203	New Jersey.....	1
Trachoma.....	31	Missouri.....	61	New Mexico.....	1
Whooping cough.....	155	New Jersey.....	364	New York.....	26
<i>November 1934</i>		New Mexico.....	9	Ohio.....	3
Actinomycosis:		Ohio.....	330	Vincent's infection:	
California.....	1	Tennessee.....	30	Iowa.....	1
Chicken pox:		Ophthalmia neonatorum:		New York ¹	53
California.....	1,035	California.....	2	Tennessee.....	6
Florida.....	7	Missouri.....	1	Whooping cough:	
Iowa.....	485	New Jersey.....	2	California.....	276
Missouri.....	269	New York.....	10	Florida.....	32
New Jersey.....	1,255	Ohio.....	80	Iowa.....	51
New Mexico.....	30	Tennessee.....	1	Missouri.....	413
New York.....	2,597	Paratyphoid fever:		New Jersey.....	919
		California.....	5	New Mexico.....	45
		New York.....	3	New York.....	2,640
		Ohio.....	1	Ohio.....	505
		Tennessee.....	1	Tennessee.....	185

¹ Exclusive of New York City.

DENGUE IN SOUTHEASTERN STATES

During the week ended December 8, 1934, 81 cases of dengue were reported in Georgia, and 47 cases were reported during the week ended December 15.

The following table shows the number of cases of dengue reported in Florida for the weeks ended December 1 and December 8, 1934:

Locality	County	Number of cases, week ended—	
		Dec. 1, 1934	Dec. 8, 1934
Miami.....	Dade.....	3	3
Orlando.....	Orange.....	3
Pensacola.....	Escambia.....	1
.....	Hillsboro.....	1
.....	Lee.....	1
Total.....	7	5

WEEKLY REPORTS FROM CITIES

City reports for week ended Dec. 8, 1934

[This table summarizes the reports received regularly from a selected list of 121 cities for the purpose of showing a cross section of the current urban incidence of the communicable diseases listed in the table. Weekly reports are received from about 700 cities, from which the data are tabulated and filed for reference.]

State and city	Diphtheria cases	Influenza		Measles cases	Pneumonia deaths	Scarlet fever cases	Small-pox cases	Tuberculosis deaths	Typhoid fever cases	Whooping cough cases	Deaths, all causes
		Cases	Deaths								
Maine:											
Portland.....	0	0	0	0	2	1	0	1	1	7	33
New Hampshire:											
Concord.....	0	0	0	0	0	1	0	0	0	0	10
Manchester.....	0	0	0	0	1	0	0	0	0	0	15
Nashua.....	2	0	1	0	0	0
Vermont:											
Barre.....	0	1	0	0	0	0	0	1	0	3	5
Burlington.....	2	0	0	0	0	4	0	0	0	0	12
Massachusetts:											
Boston.....	8	2	1	14	35	0	0	13	2	34	193
Fall River.....	2	0	27	2	0	0	1	0	0	2	27
Springfield.....	0	0	12	0	3	0	1	0	0	6	31
Worcester.....	0	0	1	3	14	0	1	0	0	18	35
Rhode Island:											
Pawtucket.....	0	0	0	2	0	0	0	0	0	0	8
Providence.....	1	1	0	2	8	0	3	0	0	17	64
Connecticut:											
Bridgeport.....	0	0	2	3	4	0	1	0	0	2	24
Hartford.....	0	0	161	4	0	0	4	0	0	0	45
New Haven.....	0	0	5	3	1	0	1	0	0	0	35
New York:											
Buffalo.....	2	1	31	9	20	0	5	0	0	22	137
New York.....	26	58	11	48	172	0	78	4	4	231	1,528
Rochester.....	0	0	57	3	12	0	1	0	0	23	59
Syracuse.....	0	0	1	3	7	0	0	0	0	6	35
New Jersey:											
Camden.....	0	3	0	1	5	4	0	0	0	10	33
Newark.....	1	16	1	3	6	16	0	5	1	45	88
Trenton.....	0	1	0	4	3	12	0	3	0	1	48
Pennsylvania:											
Philadelphia.....	12	16	3	9	42	0	22	0	0	171	528
Pittsburgh.....	11	0	19	15	29	0	6	3	3	24	139
Reading.....	0	0	1	2	1	0	0	0	0	8	23
Scranton.....	1	17	3	0	0	1
Ohio:											
Cincinnati.....	13	1	3	1	14	29	0	1	0	5	144
Cleveland.....	9	30	0	9	18	28	0	12	0	47	180
Columbus.....	8	1	1	9	1	40	0	3	0	1	67
Toledo.....	0	3	3	24	5	20	0	5	0	6	66
Indiana:											
Fort Wayne.....	4	0	2	1	0	0	3	1	0	0	21
Indianapolis.....	8	0	1	19	24	0	1	0	0	16
South Bend.....	0	0	27	0	2	0	0	0	0	1	13
Terre Haute.....	2	0	0	0	2	0	0	0	0	0	14
Illinois:											
Chicago.....	15	6	1	84	53	286	0	38	2	40	760
Springfield.....	1	0	2	4	0	1	0	6	18
Michigan:											
Detroit.....	7	19	0	24	12	84	0	18	1	47	234
Flint.....	0	0	1	1	1	12	0	4	0	6	21
Grand Rapids.....	0	0	0	1	0	12	0	1	0	7	29

City reports for week ended Dec. 8, 1934—Continued

State and city	Influenza		Measles cases	Pneumonia deaths	Scarlet fever cases	Smallpox cases	Tuberculosis deaths	Typhoid fever cases	Whooping cough cases	Deaths, all causes
	Cases	Deaths								
Wisconsin:										
Kenosha	0	0	1	1	10	0	0	0	8	7
Milwaukee	1	0	45	7	210	1	4	0	71	88
Racine	0	0	1	0	9	0	0	0	12	17
Superior	0	0	0	0	1	1	0	0	0	8
Minnesota:										
Duluth	0	0	107	3	0	0	1	0	0	18
Minneapolis	1	1	184	5	27	0	1	0	11	88
St. Paul	0	1	3	4	11	1	1	0	13	49
Iowa:										
Davenport	0	0	22	0	1	0	0	0	0	0
Des Moines	3	0	0	0	11	0	0	0	0	29
Sioux City	3	0	0	0	2	0	0	0	2	2
Waterloo	0	0	247	2	2	0	0	0	0	0
Missouri:										
Kansas City	1	3	1	20	14	0	8	0	1	108
St. Joseph	3	0	0	1	2	0	0	0	0	6
St. Louis	19	1	0	14	19	0	5	6	9	214
North Dakota:										
Fargo	0	1	0	0	7	0	0	0	5	6
Grand Forks	0	0	1	0	6	0	0	0	0	0
South Dakota:										
Aberdeen	0	0	3	0	1	1	0	0	4	0
Nebraska:										
Omaha	12	0	4	6	17	0	1	0	3	59
Kansas:										
Topeka	0	0	0	0	0	0	0	0	0	4
Wichita	1	0	0	3	2	0	0	0	0	23
Delaware:										
Wilmington	0	0	0	2	2	0	3	0	4	48
Maryland:										
Baltimore	1	2	0	4	22	47	0	12	36	225
Cumberland	0	0	0	1	2	0	1	0	0	10
Frederick	0	0	0	1	0	0	1	0	0	4
District of Columbia:										
Washington	11	0	5	13	24	0	9	0	4	151
Virginia:										
Lynchburg	1	0	4	2	2	0	1	0	6	13
Norfolk	1	0	1	4	2	0	1	0	4	35
Richmond	3	0	0	5	3	0	2	0	0	44
Roanoke	1	0	0	4	8	0	1	0	0	19
West Virginia:										
Charleston	2	2	1	12	2	13	0	1	2	25
Huntington	6	0	0	0	2	0	0	0	0	0
Wheeling	0	0	0	1	11	0	0	0	9	12
North Carolina:										
Raleigh	1	0	0	1	0	0	2	0	2	12
Wilmington	0	0	2	2	1	0	0	0	0	10
Winston-Salem	3	1	0	2	5	0	0	0	24	11
South Carolina:										
Charleston	0	26	1	0	4	0	3	0	0	34
Columbia	0	0	0	4	0	0	1	0	0	28
Greenville	0	0	0	0	0	0	0	0	3	3
Georgia:										
Atlanta	2	43	1	0	9	4	5	0	2	78
Brunswick	0	0	0	0	1	0	1	0	0	8
Savannah	0	3	1	0	3	0	2	0	2	23
Florida:										
Miami	2	0	0	0	1	0	0	0	0	28
Tampa	1	0	0	0	1	0	2	0	0	25
Kentucky:										
Ashland	2	2	0	0	1	0	0	0	0	1
Lexington	2	0	0	1	0	0	1	2	0	17
Louisville	12	2	2	3	6	22	3	1	12	73
Tennessee:										
Memphis	7	1	0	0	10	6	3	2	7	89
Nashville	4	0	0	8	6	0	2	3	1	50
Alabama:										
Birmingham	6	1	1	0	9	3	0	2	1	80
Mobile	2	1	0	0	0	0	1	0	0	27
Montgomery	1	0	0	0	1	0	0	0	0	0
Arkansas:										
Fort Smith	0	0	0	0	0	0	0	0	0	0
Little Rock	0	0	0	1	5	0	1	0	0	5
Louisiana:										
New Orleans	26	8	2	0	15	2	0	12	0	167
Shreveport	0	0	7	3	1	0	1	0	0	49

City reports for week ended Dec. 8, 1934—Continued

State and city	Diphtheria cases	Influenza		Measles cases	Pneumonia deaths	Scarlet fever cases	Small-pox cases	Tuberculosis deaths	Typhoid fever cases	Whooping cough cases	Deaths, all causes
		Cases	Deaths								
Texas:											
Dallas.....	10	1	1	0	7	9	0	1	0	0	60
Fort Worth.....	1		0	0	6	6	0	4	0	3	46
Galveston.....	0		0	0	1	1	0	1	1	0	19
Houston.....	10		1	0	11	1	0	7	0	0	81
San Antonio.....	0		1	1	9	7	13	4	0	0	63
Montana:											
Billings.....	9			9		3	0		0	0	
Great Falls.....	0		0	0	1	1	0	0	0	0	5
Helena.....	0		0	6	0	0	0	0	0	0	5
Missoula.....	0		0	0	1	0	0	0	0	0	7
Idaho:											
Boise.....	0		0	0	1	0	0	0	0	0	8
Colorado:											
Denver.....	2	49	1	183	11	108	0	2	0	4	86
Pueblo.....	3		0	0	2	6	0	0	0	0	11
New Mexico:											
Albuquerque.....	0		0	0	0	2	0	5	0	6	9
Utah:											
Salt Lake City.....	0		0	12	5	28	0	0	0	25	33
Nevada:											
Reno.....	1		0	0	1	1	0	0	0	0	6
Washington:											
Seattle.....	0			2		4	6		0	0	
Spokane.....	0	1	1	5	1	1	0	1	0	0	45
Tacoma.....	0		0	0	2	0	4	0	0	1	31
Oregon:											
Portland.....	0	2	1	1	5	15	0	1	0	0	82
Salem.....	0			0		1	0		0	3	
California:											
Los Angeles.....	9	26	1	5	14	63	0	17	1	3	309
Sacramento.....	3	1	1	0	3	2	0	2	1	0	31
San Francisco.....	1	4	3	2	7	18	0	11	0	9	165

State and city	Meningococcus meningitis		Polio-myelitis cases	State and city	Meningococcus meningitis		Polio-myelitis cases
	Cases	Deaths			Cases	Deaths	
Massachusetts:				North Dakota:			
Boston.....	1	0	0	Fargo.....	0	1	0
New York:				Maryland:			
Buffalo.....	1	0	0	Baltimore.....	0	0	1
New York.....	2	2	1	Georgia:			
Ohio:				Atlanta.....	2	0	0
Cincinnati.....	2	0	0	Utah:			
Cleveland.....	0	0	2	Salt Lake City.....	0	1	0
Indiana:				Washington:			
Indianapolis.....	0	1	0	Spokane.....	1	1	0
Illinois:				California:			
Chicago.....	2	1	0	Los Angeles.....	0	0	2
Michigan:				Sacramento.....	0	0	4
Grand Rapids.....	0	0	1				
Wisconsin:							
Milwaukee.....	0	0	5				

Dengue.—Cases: Charleston, S. C., 2; Atlanta, 2; Savannah, 44; Miami, 3.

Lethargic encephalitis.—Cases: Springfield, Mass., 1; Newark, N. J., 1; Trenton, 1; Pittsburgh, 1; Detroit, 1.

Pellagra.—Cases: Charleston, S. C., 2; Atlanta, 1; Savannah, 2; Louisville, 1; Montgomery, 1; Sacramento, 1; San Francisco, 1.

Typhus fever.—Cases: Savannah, 2; Dallas, 1.

FOREIGN AND INSULAR

INDIA

Vital statistics—First quarter ended March 31, 1934.—The following are vital statistics for India for the first quarter, ended March 31, 1934:

Population.....	263, 626, 328	Deaths from—	
Live births.....	2, 286, 733	Cholera.....	14, 946
Live births per 1,000 population.....	34. 7	Dysentery and diarrhea.....	56, 160
Stillbirths.....	44, 671	Plague.....	44, 714
Stillbirths per 100 live births.....	1. 9	Respiratory diseases.....	129, 178
Deaths.....	1, 552, 005	Smallpox.....	25, 186

PUERTO RICO

Notifiable diseases—4 weeks ended December 1, 1934.—During the 4 weeks ended December 1, 1934, cases of certain notifiable diseases were reported in the municipalities of Puerto Rico, as follows:

Disease	Cases	Disease	Cases
Chicken pox.....	16	Pellagra.....	1
Diphtheria.....	39	Pink eye.....	7
Dysentery.....	57	Ringworm.....	2
Erysipelas.....	3	Syphilis.....	38
Influenza.....	171	Tetanus.....	4
Malaria.....	1, 409	Trachoma.....	5
Measles.....	62	Tuberculosis.....	959
Mumps.....	25	Typhoid fever.....	8
Ophthalmia neonatorum.....	4	Whooping cough.....	153

CHOLERA, PLAGUE, SMALLPOX, TYPHUS FEVER, AND YELLOW FEVER

From medical officers of the Public Health Service, American consuls, International Office of Public Hygiene, Pan American Sanitary Bureau, health section of the League of Nations, and other sources. The reports contained in the following tables must not be considered as complete or final as regards either the list of countries included or the figures for the particular countries for which reports are given:

CHOLERA

[C indicates cases; D, deaths; P, present]

Place	September 1934				October 1934				November 1934											
	July 1-26, 1934		Aug. 26, 1934		6		13		20		27		3		10		17		24	
	Apr. 20-27, 1934	May 27-30, 1934	July 1-26, 1934	July 27-30, 1934	Aug. 26, 1934	1	8	15	22	29	6	13	20	27	3	10	17	24		
Ceylon: Colombo.....																				
China:																				
Amoy.....																				
Canton.....																				
Fort Bayard.....																				
Hankow.....																				
Shanghai.....																				
Tientsin.....																				
India.....																				
Assam.....																				
Bombay Presidency.....																				
Bombay.....																				
Calcutta.....																				
Chittagong.....																				
Madras Presidency.....																				
Madras.....																				
Negapatam.....																				
Punjab.....																				
Rangoon.....																				
Vizagapatam.....																				
India (French):																				
Chandernagor.....																				
Karikal.....																				
Mahe.....																				
Pondichery.....																				

! Suspected.

! Includes 4 imported cases.

! Imported.

Week ended—

CHOLERA, PLAGUE, SMALLPOX, TYPHUS FEVER, AND YELLOW FEVER—Continued

CHOLERA—Continued

[C indicates cases; D, deaths; P, present]

Place	Apr. 26- May 26, 1934	May 27- June 30, 1934	July 1-25, 1934	July 26- Aug. 25, 1934	Week ended—														
					September 1934						October 1934						November 1934		
					1	8	15	22	29	6	13	20	27	3	10	17	24		
Indo-China (see also table below):																			
Bacieu.....		2																	
Pnom-Penh.....				2															
Poulo Condor Island.....	1																		
Philippine Islands: Rizal Province—Maula.....	2	3	1																
Siam.....		1																	
On vessels:																			
S. S. <i>Viking II</i> at Calcutta from Aden.....	1																		
S. S. <i>Cape Ortega</i> at Calcutta from Bombay.....																			
S. S. <i>Jeladara</i> at Calcutta from Rangoon.....		1																	
S. S. <i>Khasara</i> at Calcutta from Karachi.....		2																	
S. S. <i>Empura</i> at Port Swettenham.....			1																
S. S. <i>Arends</i> at Rangoon from Calcutta.....					1														
Place	June 1934			July 1934			August 1934			September 1934			October 1934						
	1-10	11-20	21-30	1-10	11-20	21-31	1-10	11-20	21-31	1-10	11-20	21-30	1-10	11-20	21-31				
Indo-China (French) (see also table above):																			
Cambodia.....	1		2		1			2											
Cochin-China.....	1		2		1			1											
		10	9	6	2			1											
		8	4	4	2			1											

* Reports incomplete.

PLAGUE¹

[C indicates cases; D, deaths; P, present]

Place	Apr. 20-26, 1934	May 27-June 30, 1934	July 1-26, 1934	July 27-Aug. 25, 1934	Week ended—													
					September 1934				October 1934				November 1934					
					1	8	15	22	29	6	13	20	27	3	10	17	24	
Angola. (See table below.)																		
Argentina. (see also table below):																		
Santa Fe:																		
Santiago de Estero Province: ²																		
Azores. (See table below.)																		
Belgian Congo..... C		6																8
Bolivia. (See table below.)																		
Brazil:																		
British East Africa (see also table below):																		
Kenya..... C	2	9	1	5														
Uganda..... C	67	215	131	101						1	1	1						
British East Africa (see also table below):																		
Kenya..... C	64	208	122	101						12	15	12	12	14	18	22	23	23
Uganda..... C	9	9	1	1						14	15	12	14	18	22	22	23	16
Ceylon: Colombo..... D	9	1	1	1						1								1
Ceylon: Colombo..... D	1									1								1
China (see also table below):																		
Fort Bayard..... ³																		
Manchuria..... ⁴																		
Tungshai Island..... D																		
Dutch East Indies:																		
Java—Batavia..... C																		
Java—Batavia..... D																		
West Java..... D	1,609	1,273	1,148	1,721						485	467	374	434	441				
West Java..... D	1,609	1,273	1,144	1,720						485	467	374	434	441				
Ecuador. (See table below.)																		
Ecuador..... D																		

¹ Including plague in the United States and its possessions.² According to a newspaper report, 1 case of bubonic plague occurred on Nov. 22, 1934, in Santa Fe, Argentina. No official report of this case has been received.³ A report dated May 17, 1934, states that 15 deaths from plague occurred up to that date in Santiago de Estero Province, Argentina.⁴ Plague has been reported in Brazil as follows: Alagoas State, week ended Dec. 16, 1934, 5 cases; Ceara State, week ended Dec. 5, 1934, 2 cases of plague with other suspected cases.⁵ During the week ended June 2, 1934, suspected cases of plague were reported in Fort Bayard, Kwangchowwan Territory, China.⁶ A report dated Oct. 30, 1934, states that from June to Oct. 25, 1934, deaths from plague had been reported in Manchuria, China, as follows: Fengtien Province, Liaooyuan, 30, Shuangshan, 21, Tungliao, 41; Kirin Province, Changling, 12, Chienan, 26, Fuyi, 32, Hsinling City, 1, Nungau, 108.⁷ Imported.

Rangoon.....	O	3	8	1	1	2	1	1	1
Plague-infected rats Indo-China (see also table below):		3	3						
Bentre.....	O			1	1				
Longxuyen.....	O			1	1				
Pnom-Penh.....	D	2	4	1	1				
Sadee.....	O		1						
Saigon and Cholon.....	O	3	4	1	1		1		1
Vinhlong.....	O		1						
Iraq: Baghdad.....	O	1	3	1					
Libya.....	D		6						
Madagascar. (See table below.)	D		1						
Morocco: Tangier.....	C								4
Peru (see also table below): Salaverry—Plague-infected rats			1						3
Senegal. (See table below.)									
Sierra Leone.....	O	1	2						
South West Africa †									
Tunisia: Tunis—Plague-infected rats			4				2		1
Union of South Africa: Orange Free State.....	C								
United States:									
California:									
Human plague—Tulare County	O		1						
Plague-infected ground squirrels:									
Kern County.....		26							
Modoc County.....			36	2					
Tulare County.....		33	6						
Oregon: Lake County.....	O	1							
On vessel: S. S. <i>Barjora</i> at Rangoon from Moulmein.....	D						1		1
	D								

† From January to June 30, 1934, 26 cases of plague were reported in Ovamboland, South-West Africa.

‡ Includes 1 plague-infected wood rat.

CHOLERA, PLAGUE, SMALLPOX, TYPHUS FEVER, AND YELLOW FEVER—Continued

PLAGUE—Continued

[C indicates cases; D, deaths; P, present]

Place	May 1934	June 1934	July 1934	August 1934	Septem-ber 1934	Octo-ber 1934	Place	May 1934	June 1934	July 1934	August 1934	Septem-ber 1934	Octo-ber 1934
Angola.....	2	2	2	6	11		Peru (see also table above).....		1		1		3
Argentina (see also table above).....		3	5				Senegal:		33	55	52	17	13
Azores.....							Dakar ¹⁰	9	27	47	47	19	11
Bolivia.....	P						Diourbel ¹⁰	8		5	5	1	
British East Africa (see also table above):							D		2		3	1	
Kenya.....	5	6		13	5		Louga ¹⁰						
Uganda.....	90	21	10	103	54		D	2					
China: Kwangchowan.....		12	4	3			D	18	33	42	42	18	8
Ecuador.....		7					Rufisque ¹⁰						
Indo-China (see also table above):							Sobkhotane ¹⁰	9					
Cambodia.....	5	4	2	3	3	4	Thisis ¹⁰	26	64	50	39	6	6
Cochin-China.....	3	9	8	2	291	444	Tivaouane ¹⁰	35	32	27	82	26	21
Madagascar (central region).....	87	71	96	160	253	422							
	83	71		168									

¹⁰ Reports incomplete.

SMALLPOX

Place	Week ended—													
	September 1934						October 1934						November 1934	
	1	8	15	22	29	6	13	20	27	3	10	17	24	
Algeria:														
Algiers Department.....														
Constantine Department.....														
Oran Department.....														
Angola. (See table below.)														
Belgian Congo ¹ (see also table below)														
Bolivia. (See table below.)														
Brazil:														
Pernu Alegre (alestrim).....														
Sergipe State.....													P	

July 29-Aug. 25, 1934
 July 1-28, 1934
 May 27-June 30, 1934
 Apr. 26-May 26, 1934

India (Portuguese)	C																				
Indo-China (see also table below):					27																
Haiphong	C	17	7			1															
Pnom-Penh	C	5	24	17	9	3															
Tourane	D	27	88	13	5				6	1	4	2	1								3
Iraq	C	2																			
Amara Ijwa	C																				
Arbil	C				15																
Baghdad	C		1																		
Basra	C	5				2															
Italy:																					
Genoa	C					8	2														
Milan	C							1													
Ivory Coast. (See table below.)	C																				
Japan	C	12	7	6																	
Aomori Prefecture	C			3																	
Kobe	C			1																	
Nagasaki	C	1																			
Osaka	C	2																			
Liberia:																					
Mexico:																					
Chihuahua	C					1															
Guadalajara	D	1																			2
Mazatlan	C																				
Mexico, D. F.	C	14	12	5	6	2	3	2	1												5
Monterrey	C				1																
Piedras Negras	C	1																			
Rerita	C	11																			
San Luis Potosi	C	2				1															
Tehuacan	D																				
Torreon	D	1				1															
Morocco. (See table below.)	C																				
Mozambique. (See table below.)	C	1, 277	136																		
Nigeria	C	15	9	2	3	1			163	3											
Lagos	C																				24
Nyasaland. (See table below.)	C	2	7		3																
Palestine	C	1																			
Pensia	C																				
Teheran	D							1													
Peru. (See table below.)	C	1																			1
Poland	C																				
Portugal. (See also table below):	C	8	6																		
Lisbon	C																				
Oporto	C								1												
Portuguese East Africa. (See table below.)	C	60	85	121	30	10															
Salvador	C	3	96	6					2												
Shanghai	C								32												13
Sierra Leone	C																				

¹ For 2 weeks.

² A report states that from February to Sept. 10, 1934, 233 cases of smallpox with 79 deaths had been reported in Sonora, Liberia. All sanitary measures have been taken.

³ A report dated Aug. 27, 1934, states that smallpox has appeared in the suburbs of Mazatlan, Sinaloa, Mexico; the report also states that 104 deaths from smallpox have occurred in Tepic, Oaxaca, Mexico.

⁴ Imported.

⁵ For 3 weeks.

CHOLERA, PLAGUE, SMALLPOX, TYPHUS FEVER, AND YELLOW FEVER—Continued

SMALLPOX—Continued

[C indicates cases; D, deaths; P, present]

Place	Week ended—																
	September 1934							October 1934							November 1934		
	1	8	15	22	29	6	13	20	27	3	10	17	24				
Sierra Leone.....					242		89	125									
Spain.....					9		5	8									
Straits Settlements: Singapore.....					4		3	8									
Sudan (Anglo-Egyptian).....					1			1									
Syria:																	
Belrut.....																	
Damascus.....																	
Provinces.....																	
Trans-Jordan.....																	
Turkey.....																	
Union of Soviet Socialist Republics. (See table below.).....																	
Union of South Africa.....																	

‡ For 2 weeks.

§ Imported.

On vessels:

S. S. <i>Ku</i> Sang at Hong Kong from Amoy.....	Present.	May 9, 1934
S. S. <i>Tsingara</i> at Hong Kong.....	Present.	May 16, 1934
S. S. <i>Britannia</i> at Port Said from Liverpool.....	1 case.	May 31, 1934
S. S. <i>Rohna</i> at Penang from Madras.....	1 case.	June 14, 1934
S. S. <i>Rohna</i> at Penang.....	1 case.	June 28, 1934
S. S. <i>Rohna</i> at Penang from Madras.....	1 case.	July 12, 1934

On vessels—Continued.

S. S. <i>Tacoma</i> at Molli from Dairen.....	1 case.	July 28, 1934
S. S. <i>Ehiopta</i> at Rangoon from Madras.....	1 case.	Sept. 3, 1934
S. S. <i>Ussuri Maru</i> at Kobe from Dairen.....	1 case.	Sept. 24, 1934
S. S. <i>Rohna</i> at Penang from Madras.....	1 case.	Oct. 4, 1934
S. S. <i>Erinpura</i> at Rangoon from Madras.....	1 case.	Oct. 8, 1934

Place	1934												
	May	June	July	August	Septem- ber	October	Place	May	June	July	August	Septem- ber	October
Angola.....	35	110						Ivory Coast.....		4	2	2	
Belgian Congo (see also table above).....	67	360		204	183		Morocco.....		2	6			
Bolivia.....		82		136	70		Mozambique.....		2	3	2	8	
Cameroun (French).....		1					Nyasaland.....		3	3	4	4	6
Chosen.....	65	25	18	5	15		Peru.....	86	63	31	16	29	17
Dahomey.....	5	83		31			Portugal (see also table above).....	16	20	24	100	12	
Ecuador.....		1					Portuguese East Africa.....	110	40	70	60	36	
Finland.....		1					Turkey.....	1	6	4	6	2	
France.....	10	3		3	2		Turkey Socialist Republics.....	34	5	3	4	6	1
Guatemala.....				1			Union of Soviet Socialist Republics.....	6	6	17	4	4	2
Indo-China (see also table above).....	692	215	192	150	87	202		574	403				
	76	39	28	39	18	39							

TYPHUS FEVER

Week ended—

Place		Apr. 29—May 27, May 26, June 30, 1934		July 1, June 28, 1934		1934																
		1934		1934		August		September			October		November									
Algeria:						4	11	18	25	1	8	15	22	29	6	13	20	27	3	10	17	
Algiers Department.....	C		18	1						1									8	2	1	1
Constantine Department.....	C	41	91	32	6	3	6	3	1	3	1	1	1						3	3		
Bone.....	C			1					1													
Constantine.....	C		5	2				10											2			
Oran Department.....	C																					
Basutoland.....	C		208	135	52	30	26		13	19	13	12	6	4	7	6	3	4	4	4		1
Belgian Congo.....	C																					
Belgian Congo (see table below).....	C	2	6	2																		
Belgian East Africa.....	C	45	19	2					1										2	5	10	
Bulgaria.....	C	1,192	1,044	1,044	260	278	300															
Chili.....	C																					
Consepcion.....	C																					
Luquén.....	C																					
Santiago.....	C																					
Atacapica Province, ¹ Tacopilla.....	C	164	321	365	73		112															
Valparaiso.....	C																					
		11	22	30	2	8	7	4	3	8	1	6	2	2	2	6	3	8	10	9	6	

¹ From Apr. 18 to May 27, 1934, 256 cases of typhus fever with 7 deaths were reported in Belgian Congo.² For 9 weeks. ³ Imported. ⁴ A report dated July 13, 1934, states that 41 cases of typhus fever with 7 deaths have been reported in the villages of Urmagana and Fachtes, Tarapaca Province, Chile.

Irish Free State:																					
Cork County—Castletown.....						5					1										
Waterford County—Lismore.....					7												1				
Wicklow County—Aldmore.....																					
Italy:																					
Lephorn.....									1												
Palermo.....						1					2										
Japan:																	1				
Aomori Prefecture.....								3		1											
Kobe.....																					
Nagasaki.....																1					
Latvia (See table below.).....																					
Lithuania.....				12	4	17	3	2				2							2	1	
Mexico:																					
Guzdalaers.....	D																				
Mexico, D. F.....	D	64	108	38	8	12	5	9	4	11	6	5	8	6							
Saltillo.....	D										2										
San Luis Potosi.....	D			1									1								
Torreón.....	D			1												1				1	
Morocco:																					
Morocco.....	C	34	45	18	36	6			1	2	1				1						
Palrest.....	C		4						1	2					1						
Palestine.....	C																				
Persia.....	C	230	217	112	9	21	13	6	17	5	6	10	9	4	8	1				2	
Peru.....	C	36	22	25	1	4	1	2	2	3	3	1									
Peru (See table below.).....																					
Poland.....	C	428	284	148	20	20	25	12	10	13	9	8	13	13	9	13	13	14	12	16	
Portugal (see also table below): Oporto.....	D	28	23	12	1	1	1	4			1									3	
Rumania (See table below.).....	C									1											
Scotland.....	C	3																			
Spain: Catalonia.....	C		27																		
Straits Settlements: Singapore.....	C																				
Syria: Beirut.....	C				1																
Trans-Jordan.....	C	6	16	4	3	1															
Tunisia.....	C																				
Tunis.....	C	2	22	1						1											
Provinces.....	C	90	99	46												15	2		4	16	1
Turkey. (See table below.).....																					
Union of Soviet Socialist Republics. (See table below.).....																					
Union of South Africa. (See table below.).....																					
Yugoslavia. (See table below.).....																					

* Imported.

* Includes 1 imported case.

																							1	2
Gambia:																								
Fathurst	C																							
St. Mary's Island	C																							
Gold Coast:																								
Kokobee ¹	C			1																				
N Kaw Kaw	C																							
Ivory Coast:																								
Abidjan	C		3																					
Arboville	D		2																					
Bingerville	C																							
Bobo-Dioulasso	C					1																		
Rubino	D																							
Rubino	D		2																					
Tirailleur	C																							
Zuenoula ⁴	C																							1
Niger Territory:																								
Maradi	C																							
Zinder	D																							
Senegal: Matam	D																							
Sudan (Anglo-Egyptian): Wau	C																							

1 A report dated Nov. 28, 1934, states that 3 deaths from yellow fever had been reported in Restrepo, Intendencia of Meta, Colombia.
 2 Suspected.
 3 During the week ended Nov. 24, 1934, 1 case of yellow fever was reported in Kokobee, Saltpond district, Gold Coast.
 4 During the week ended Dec. 8, 1934, 1 case of yellow fever was reported in Zuenoula, Ivory Coast.