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PLAGUE. SULFADIAZINE TREATMENT OF GUINEA PIGS INFECTED BY ARTIFICIAL METHODS OR BY FLEA TRANSMISSION

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Reports have been made of the therapeutic value of each of several sulfonamide compounds in human cases of plague (1-8), and the results of treatment of inoculated animals with sulfanilamide, sulfapyridine, or sulfathiazole (5,9-12). The reports of the clinical course of the disease in man when under therapy with these drugs have varied widely and a consensus regarding its efficacy does not seem to have been reached. Some of the divergence of opinion seems to arise from failure to consider its value in different stages or types of the disease. The results with experimental animals seem more consistently satisfactory, but the conditions of the experiments have been rather artificial in that the animals were inoculated subcutaneously and treatment was initiated before inoculation, contemporary with it, or at a predetermined number of hours thereafter.

The inconsistency of the results obtained in man with the different sulfa derivatives administered and the selectivity of the experimental methods used with the laboratory animals appeared to warrant further tests and a trial of sulfadiazine which is excreted more slowly than other forms of these compounds. A further stimulus to additional tests arose from the observation that the growth of *P. pestis* was inhibited when planted on 5 percent blood agar plates prepared from human blood containing 1.1 mg. percent of sulfonamide.

The experiments were designed to reproduce in guinea pigs, as closely as practicable, the course of the disease in man and to determine the therapeutic value of the drug when administered after the characteristic buboes had developed and before the occurrence of a septicemia. The bubo is a finding which can be recognized in both man and guinea pig and is the first criterion which suggests the diagnosis in many human cases.

Fifty guinea pigs were used in the tests. Twenty-six received sulfadiazine and 24 did not but were matched individually insofar as possible with a treated animal with regard to weight, general physical condition, method of inducing infection, characteristic clinical findings,

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and nonspecific administrations. Twenty-two of the 24 controls died of plague; another developed enormous buboes, became thin and was febrile, but lived for 34 days after inoculation when it was killed and the infection proved bacteriologically; the other animal died of hemorrhage on the third day after inoculation without showing definite evidence of plague. Seven of the 26 treated animals died. One of these 7 showed no evidence of infection when it died 11 days after inoculation but the pelves and ureters of the kidneys were stuffed with crystals of a sulfonamide. Another was progressing favorably toward recovery but developed an acute purulent cervical adenitis probably caused by injury from a capsule being pushed deep into its pharynx in order to assure its being swallowed.

All the animals used in the test were examined by necropsy and bacteriological methods before they were discharged, at death, or when killed after clinical recovery and survival for at least 21 days after inoculation. Among the animals which died while under treatment, it was noted that there were but few organisms obtained in preparations from their tissues, and that most of these were of involution forms in contrast to the large numbers of typical morphology obtained from the controls.

Preliminary series of animals were tested by the use of sulfathiazole; dosage was guided in frequency and amount by determination of the amount of sulfonamide which persisted in samples of heart blood withdrawn at varying periods after administration (Bratton and Marshall method). It was found difficult to maintain consistently in the blood an amount of drug which was believed necessary to accomplish good therapy. Sodium sulfadiazine was then tried and was found to effect rapidly a good blood level, but the level also declined rapidly. treatment adopted was an initial dose of 100 mg. of sodium sulfadiazine administered subcutaneously in aqueous solution and 100 mg. of sulfadiazine given in a capsule by mouth. This dose was followed by 100-mg. capsules, by mouth, as frequently and for as long a period as seemed indicated by the course of the infection in the individual animal. All but five of the animals receiving the sulfadiazine were given sodium bicarbonate in capsules by mouth in dosage approximately double that of the drug.

Drug treatment was instituted in 25 animals after a papule had developed at the site of inoculation or flea bite with a rise of temperature to 39° C. and an unmistakable bubo had developed in one or two contiguous lymph nodes. One animal developed a papule and fever but failed to show a definite bubo.

The tests were run in parallel on a treated animal and an untreated control, with but two exceptions in each of which there were two treated animals and one untreated as the control. The inequality in

the number of controls in this test resulted from the technical difficulties of obtaining comparable animals which had been infected by flea bites on the same day.

Two groups of ten each were inoculated intracutaneously on the abdomen with suspensions of a blood agar culture of *P. pestis*, strain No. 3035, grown 24 hours at 30° C. This strain had consistently killed guinea pigs, white rats, and white mice under various experimental conditions throughout the past 2 years. Approximately 1,500 organisms were given to each of one group; 12,000 to a second group;

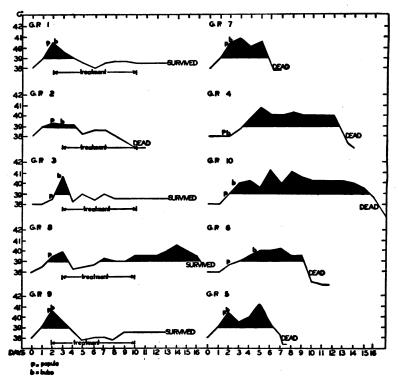


FIGURE 1.—Guinea pigs inoculated intracutaneously with 1500±P. pestis, strain B 3035, showing period and range of temperature, survivals, and deaths among those treated with sulfadiazine and those without treatment.

and 25,000 of a recently isolated strain, D 67, to the third group. The number of organisms was determined by turbidity standards and by plating out on blood agar.

Twenty animals were infected by the bites of fleas (*Xenopsylla cheopis*) which had previously fed on guinea pigs artificially inoculated with the same strain of *P. pestis*.

Blood cultures and determinations of the drug content were made on samples of heart blood from 20 of the treated animals. A check on the drug content in the blood of 6 others was made by micro methods on blood from a vein in the foot. It was found that frequent heart puncture and withdrawal of a sample of 2 cc. of blood weakened the animal even though dextrose-saline solution was returned intraperitoneally; hence the frequency of sampling was lessened somewhat with succeeding groups and discontinued toward the last of the series. However, among the 15 animals inoculated artificially and treated, there were cultures of 43 blood samples of which 1 was positive; among the corresponding 15 untreated controls, cultures were made of 38 samples and 14 were positive. Among 5 of the animals infected by fleas and treated, there were cultures of 4 samples, 3 of which were positive. One of these was positive before treatment began. Among the 5 untreated controls, there were 4 blood cultures and all

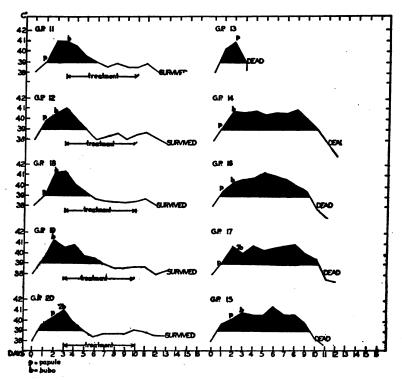


FIGURE 2.—Guinea pigs inoculated intracutaneously with 12,000±*P. pestis*, strain B 3035, showing period and range of temperature, survivals, and deaths among those treated with sulfadiazine and sodium bicarbonate and those without treatment.

were positive. The blood level of the drug in 44 specimens of heart blood collected at different periods from animals while under treatment varied from 1 mg. percent to 10 mg. percent with a median level of approximately 5.5 mg. percent, and approximately this same level was obtained in the peripheral blood from the foot.

Figures 1, 2, and 3 record the results of the individual animals of the groups which were inoculated intracutaneously with varying numbers of *P. pestis*, and show the temperature range, time of appearance

of the papule and bubo, period of treatment, and final results among those treated with the drug and the comparable untreated controls.

Figures 4, 5, and 6 record similar findings among the animals which were infected by the bites of fleas.

The protocols of the details of treatment, observations, and final results for each animal are appended.

It will be seen that among animals inoculated in a manner to simulate a flea bite with numbers of *P. pestis* varying from approximately 1,500 to 25,000 there were 13 of 15 untreated controls which died of

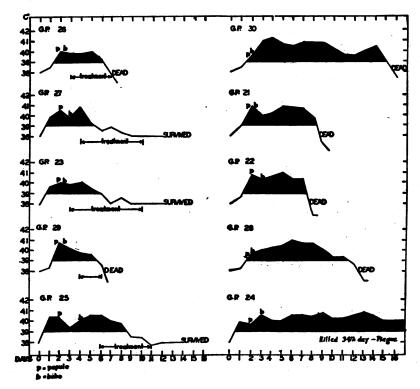


FIGURE 8.—Guinea pigs inoculated intracutaneously with 25,000±P. pestis, strain D 67, showing period and range of temperature, survivals, and deaths among those treated with sulfadiazine and sodium bicarbonate and those without treatment.

plague. (One was ill with plague and killed on the thirty-fourth day after inoculation; another died of hemorrhage without evidence of plague.) All of these developed a rise in temperature of from 39° to 41° C., and a papule or bubo or both. These findings continued through several days until about 20 to 30 hours before death when the temperature fell critically. Among fifteen animals similarly inoculated and treated with the drug, the temperature and findings developed but persisted for a shorter period, in most instances with a gradual return to normal with the exception of three which died. One of

these, No. 2, showed no evidence of plague at its necropsy, and probably died of a toxic condition caused by the drug. One death, No. 26, occurred in one of the few treated animals which developed a septicemia.

Nine untreated controls which were infected by flea bites developed the findings described above and died of plague, while among 11 animals similarly infected and treated with the drug, 4 died. One of these, No. 32, died of an acute cervical adenitis probably of traumatic origin. Treatment of another, No. 36, was delayed for 5 days after

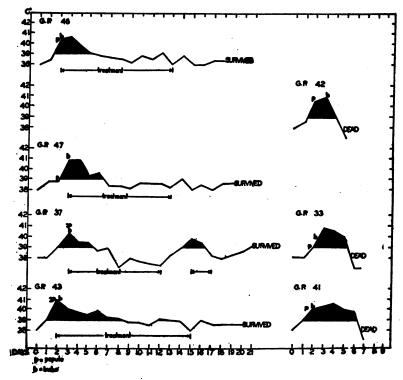


FIGURE 4.— Guinea pigs infected by bites of fleas infected with *P. pestis*, strain B 3035, showing period and range of temperature, survivals, and deaths among those treated with sulfadiazine and sodium bicarbonate and those without treatment.

inoculation while waiting for the development of a bubo, and septicemia had developed before treatment was commenced. Three of these animals infected by flea bites developed septicemia either before or shortly after treatment was begun. It was noted that the development of septicemia occurred more frequently and death occurred after a shorter course of the disease among animals infected by flea bites.

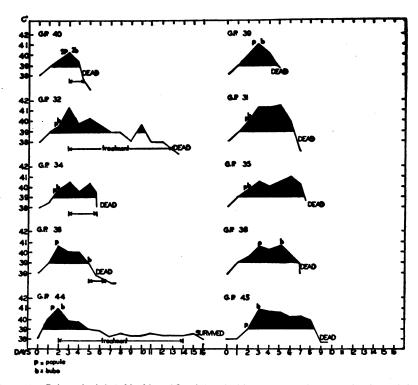


FIGURE 5.—Guinea pigs infected by bites of fleas infected with *P. pestis*, strain B 3035, showing period and range of temperature, survivals, and deaths among those treated with sulfadiazine and sodium bicarbonate and those without treatment.

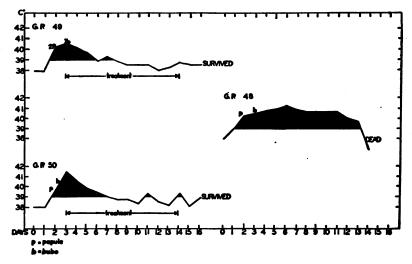


FIGURE 6.—Guinea pigs infected by bites of fleas infected with *P. pestis*, strain B 3035, showing period and range of temperature, survivals, and deaths among those treated with sulfadiazine and sodium bicarbonate and those without treatment.

SUMMARY AND CONCLUSIONS

Fifteen guinea pigs were inoculated intradermally with virulent P. pestis and developed plague, but 13 recovered after treatment with sulfadiazine and showed no evidence of infection at necropsy when killed 21 days after inoculation, and one died with sulfonamide crystals in the kidneys. Thirteen untreated controls died of plague after similar inoculation. One control died of hemorrhage 3 days after inoculation without evidence of plague; one control was killed the thirty-fourth day after inoculation and plague was proved bacteriologically at necropsy.

Eleven guinea pigs infected with plague by flea bites developed the disease, but 7 recovered under treatment with sulfadiazine and showed no evidence of the infection at necropsy 21 days after inoculation. Nine untreated controls which were infected in a similar manner developed the disease and died.

These experiments indicate that the administration of sulfadiazine to guinea pigs in which buboes of plague have been contracted by flea transmission or have been induced by artificial methods simulating flea transmission is of very definite remedial value. The drug treatment should begin as soon as the characteristic buboes and fever have developed and should continue through the febrile period. A blood level of 4 to 7 mg, percent of the drug was usually maintained. but no attempt was made to determine the level required for therapeutic efficiency.

The similarity of the evolution and manifestations of plague induced in guinea pigs and in man by flea transmission lead to the conclusion that this drug may be of great value in the therapy of bubonic plague in man previous to the development of a generalized bacteremia.

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Appendix

Protocols of guinea pigs inoculated with P. pestis or infected by flea bites and treated with sulfadiazine

(First dose 100 mg. sodium sulfadiazine and 100 mg. sulfadiazine. Subsequent doses, 100 mg. sulfadiazine.)

Guines pig No.	Inoculation	Clinit	Olinical course from hour of inoculation	Treatment from hour of inocula-Blood level free sulfadiatine from tion	Blood level free suifediazine from hour of treatment	Results
	1500 <u>+</u> <i>P. petts</i> , strain B 3035, intracutane- ous.	42 hrs. 48 hrs. 66 hrs. 90 hrs. 138 hrs. 162 hrs.	Papule. Bubo. Blood culture negative. Do. Bubo graell, papule healing. Blood culture negative.	48 hrs. First dose. 8 doses dally for 7 days. 2 doses dally for 1 days. 7 total treatment 9 days with 2.400 mg. sulfadiasine, 100 mg. sodium sulfadiasine.	18 hrs. 2 mg. percent. 42 hrs. 5 mg. percent. 90 hrs. 10 mg. percent. 138 hrs. 7 mg. percent.	Survived and killed 21st day. Necropsy, no plague.
(Control)	1500±P. pestis, strain B 3086, intractione- ous.	28 brs. 138 brs. 138 brs. 138 brs.	Papule. Bubo. Blood culture negative. Blood culture positive. Do.	No treatment.		Found deed morning 7th day. Necropsy, plague.
	1800±P. petit, strain B 3035, intracutane- ous.	42 hrs. 66 hrs. 90 hrs. 138 hrs. 162 hrs. 216 hrs. 248 hrs.	Papule. Bubo. Blood culture negative. Do. Papule and bubo healing. Blood culture negative. Animal stek.	72 hrs. First dose. 76 hrs. Second dose. 3 doses daily for 6 days. 2 doses daily for 1 day. Total treatment 8 days, with 2,200 mg, sulfadiazine, 100 mg. sodium sulfadiazine.	18 hrs. 8.5 mg. percent. 66 hrs. 8.0 mg. percent. 114 hrs. 7.0 mg. percent. Post mortem: Free 40.0 mg. percent. Total 64.5 mg. percent.	Found dead morning 11th day no plague. Necropsy, no plague.
(Control)	1500±P. pestis, strain B 3034, intracutane- ous.	42 hrs. 48 hrs. 90 hrs. 112 hrs. 138 hrs. 186 hrs.	Papule. Bubo. Blood culture negative. Blood culture negative. Do.	No treatment		Found deed morning 14th day. Necropsy. plague.
	1500±P. pestis, strain B 3085, intracutane- ous.	22 brs. 188 brs. 188 brs. 188 brs. 216 brs.	Papule. Bubo. Blood culture negative. Do. Papule besling. Papule hesled, very small bubo.	72 hrs. First dose. 3 doses dally for 6 days. 2 doses dally for 1 day. Total treatment 8 days with 2,100 mg. suifadiazine, 100 mg. sodium suifadiazine.	18 hrs. 3.8 mg. percent. 66 hrs. 8.0 mg. percent. 114 hrs. 10.0 mg. percent.	Burvived and killed 21st day. Necropay, no plague.

Found deed morning 18th day. Necropay, plague.	Survived and killed first day. Necropsy, no plague.	Death on 11th day. Necropsy, plague.	Survived and killed žist day. Necropsy, no plague.	Death on 7th day. Necropsy, plague.	Survived and killed zist day. Necropsy, no plague.	Found dead morning 3rd day. Necropey, bemorrhage.	Survived and killed flet day. Necropsy, no plague.
	18 hrs. 4.0 mg. percent. 66 hrs. 8.5 mg. percent. 114 hrs. 7.0 mg. percent.		18 hrs. 3.0 mg. percent. 42 hrs. 2.0 mg. percent. 80 hrs. 9.5 mg. percent. 138 hrs. 6.0 mg. percent.	•	42 hrs. 4.0 mg. percent. 90 hrs. 8.5 mg. percent. 138 hrs. 7.5 mg. percent.		42 hrs. 4.5 mg. percent. 90 hrs. 4.5 mg. percent. 138 hrs. 6.0 mg. percent.
No trestment.	72 hrs. First dose. 3 doses daily for 6 days. 2 doses daily for 1 day. Total trestment 8 days, with 2,100 mg. suifadiscine, 100 mg. sodium suifadiscine, 100	No treatment.	48 hrs. First dose, 3 doses daily for 7 days. 2 doses daily for 1 day. 2 dose daily for 1 day. 7 foral treatment 9 days with 2,400 mg. sulfadiastine, 100 mg. sodium sulfadiastine.	No treatment.	72 hrs. First dose. 3 doses daily for 5 days. 2 doses daily for 2 days. Total treatment 8 days with 2.000 mg. sulfadisatine, 100 mg. sodium sulfadisatine, 4,000 mg.	No treatment.	72 hrs. First dose, 3 doese daily for 5 days. 2 doses daily for 5 days. Total treatment 8 days. with 2,000 mg. sulfadistine, 100 mg. sodium sulfadistine, 4,000 mg. sodium bloarbonate.
Papule. Blood culture negative. Do. Do. Two large buboes, thick inflitration of skin.	Papule. Blood culture negative. Do.	Papule. Blood culture negative. Buod culture negative. Blood culture positive.	Papule and bubo. Blood culture negative. Do. Do. Bubo small. Blood culture negative.	Papule and bubo. Blood culture positive. Do.	Papule. Bubo culture negative. Bubo smaller. Blood culture negative. Bubl. hard bubo.	Papule.	Papule. Bubo culture negative. Bubo smaller. Blood culture negative. Papule healing. Blood culture negative. Papule blood culture negative. Fabule but wall, hard bubo. Guines pig thin, but wall.
42 hrs. 66 hrs. 90 hrs. 138 hrs. 234 hrs.	42 hrs. 90 hrs. 138 hrs. 186 hrs.	42 hrs. 90 hrs. 114 hrs. 138 hrs. 186 hrs.	42 hrs. 66 hrs. 90 hrs. 138 hrs. 162 hrs.	42 hrs. 66 hrs. 90 hrs.	24 hrs. 66 hrs. 114 hrs. 144 hrs. 162 hrs. 210 hrs. 226 hrs.	42 brs.	24 hrs. 42 hrs. 114 hrs. 162 hrs. 210 hrs. 288 hrs.
(Control) B 3086, intracutane- ous.	1800±P. pestis, strain B 3035, intracutane- ous.	1500±P. pestite, strain B 3035, intracutane- ous.	1,600±P. pestu, strain B 3035, intracuta- neous.	1,500±P. pestio, strain B 3035, intracuta- neous.	12.000±P. pesth, strain B 3635, intracuta- neous.	(Control) B 3035, intracutaneous.	12,000±P. pestie, strain B 3036, intra- cutaneous.
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Protocols

Results	Found deed morning 12th day. Necropsy, plague.	Survived and killed flat day. Necropsy, no plague.	Death on 11th day. Ne- cropsy, plague.	Survived and killed 21st day. Necropsy, no plague.	Found deed morning 12th day. Necropsy, plague.	Survived and killed 21st day. Necropsy, no plague.
Blood level free suffediatine from hour of treatment		48 hrs. 8.0 mg. percent. 96 hrs. 5.5 mg. percent. 144 hrs. 6.6 mg. percent.	•	42 hrs. 4.5 mg. percent 90 hrs. 5.5 mg. percent. 138 hrs. 7.0 mg. percent.		42 hrs. 4.5 mg. percent. 90 hrs. 5.5 mg. percent. 188 hrs. 5.5 mg. percent.
Treatment from hour of inocula- tion	No treatment.	66 hrs. First dose. 8 doses daily for 5 days. 2 doses daily for 2 days. Total treatment 8 days. with 2,000 mg. sulfadiasine, 100 mg. sodium sulfadiasine, 4,000 mg. sodium bicarbo- nate.	No treatment.	72 hrs. First dose. 3 doses dally for 5 days. 2 doses dally for 5 days. 1 rotal treatment 8 days, with 2,000 mg. sulfadiszine, 100 mg. sodium bieschonste.	No treatment.	72 hrs. First dose. 2 doses daily for 5 days. 2 doses daily for 2 days. Total treatment 8 days, with 2,000 mg. salifadisatine, 100 mg. sodium sulfadisatine, 4,000 mg. sodium bloarbonate.
Olinical course from hour of inceniation	Papule. Bubo. Thiok infliretion around large bubo. Blood outlure negative. Blood outlure negative. Guines pig stok.	s. Papule. s. Bubo. s. Blood culture negative. s. Blood culture negative. s. Blood culture negative. Papule healed. s. Blood culture negative.	s. Papule. s. Bubo. s. Blood culture negative. s. Do. s. Blood culture positive.	Bubo. Email and large bubo. Blood culture negative. Blood culture negative. Blood culture negative. Cheeled.	s. Papule. 3. Two buboss. 4. Bood culture negative. 5. Buboes very large with thick inflitration of surrounding skin. Blood culture negative.	Papule. Two bubose. Blood culture negative. Blood culture negative. Blood culture negative. Do. Do. Do. One small bubo.
Inoculation Olini	12,000±P. pestis, 24 hrs. strain B 3036, intra- 114 hrs. 114 hrs. 135 hrs. 135 hrs. 135 hrs. 135 hrs. 135 hrs. 220 hrs. 224 hrs.	12,000±P. pestis, 24 hrs. cutaneous. 114 hrs. 1184 hrs. 1188 hrs. 162 hrs. 230 hrs. 234 hrs.	12,000±P. pestis, 24 hrs. strain B 3035, fatra-114 hrs. 114 hrs. 1162 hrs. 210 hrs.	12,000± P. pestite, strain B 3036, intracutane- 013. 114 hrs. 115 hrs. 210 hrs. 234 hrs.	7	12,000±P. pestis, strain 62 hrs. B 3335, intracuta- 66 hrs. neous. 1144 hrs. 145 hrs. 239 hrs.
Guines pig No.	(Control)	18	(Control)	91	(Control)	90

•			00.			
Found deed morning 11th day Necropsy, plague.	Death on 7th day. Necropsy, plague.	Found dead morning 16th day. Necropsy, plague.	Survived and killed 21st day. Necropay, no plague.	Death on 9th day. Necropey, plagua.	Survived and killed 21st day. Necropsy, no plague.	Death on 8th day. Necropay, plague.
	24 hrs. 8.4 mg. percent.	•	48 hrs. 5.0 mg. percent. 96 hrs. 10.0 mg. percent.		21 hrs. 8.0 mg. percent. 60 hrs. 6.0 mg. percent. 117 hrs. 6.0 mg. percent.	
No treatment	72 hrs. First dose. First Georal dose. 80 hrs. Third dose. 8 doses daily for 3 days. Total treatment 5 days, with 1400 mg. sulfadiasine, 300 mg. sodium bulkalissine, 2,300 mg.	No treatment.	96 hrs. First dose. 100 hrs. Boond dose. 104 hrs. Third dose. 8 doses daily for 4 days. 2 doses daily for 2 days. Total treatment 7 days, with 1500 mg. sulfadiasine, 100 mg. sodium guifadiasine, 3,800 mg. sodium bloarbonate.	No treatment,	78 hrs. First dc ** 3 dosee daily for c.c. 778 2 dosea daily for 2 days. Total treatment 8 days. with 2,000 mg. sulfadissine, 100 mg. sodium sulfadissine, 4,000 mg. sodium blear bonate.	No treatment.
Papule. Bubo. Blood culture positive. Do.	Papule and bubo. Guines pig side. Blood culture positive. Bubo large. Brod culture negative. Guines pig side. Post-mortem blood culture negative.	Papule and bubo. Blood culture negative. Blood culture negative. Large and small bubo. Blood culture negative. Guines pig sick.	Papule. Bubo. Bubo oulture negative. Two small buboes. Blood culture negative. One very small bubo.	Papule and bubo. Two small buboes. Blood culture positive. Do.	Papule and bubo. Guinea pig sid. Guinea pig sid. Blood culture negative. Blood culture negative. Papule open. Bubo smaller. Blood culture negative. Papule	Papule. Bubod culture positive. Two small buboes, cuture positive. Blood culture positive.
48 hrs. 66 hrs. 114 hrs. 162 hrs. 210 hrs.	48 hrs. 72 hrs. 96 hrs. 120 hrs. 144 hrs. 150 hrs.	48 hrs. 96 hrs. 144 hrs. 192 hrs. 246 hrs.	48 hrs. 64 hrs. 144 hrs. 192 hrs. 240 hrs.	48 hrs. 72 hrs. 144 hrs. 192 hrs.	48 brs. 72 brs. 96 brs. 144 brs. 168 brs.	48 hrs. 72 hrs. 96 hrs. 144 hrs. 192 hrs.
12,000±P. pestis, strain B 3036, intracuta- neous.	25,000±P. pestis, strain D 67, intracutane ous.	25,000±P. pestis, strain D 67, intracutane- ous.	28,000± P. pestis, strain D 67, intracutane- ous.	25,000±P. pestis, strain D 67, intracutaneous.	25,000± P. pestis strain D 67, intracutane- ous.	25,000±P. peste, strain D 67, intracutane- ous.
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Results	Found dead morning 7th day. Necropsy, plague.	Death on 13th day. Necropsy, plague.	Survived and killed 21st day. Necropsy, no plague.	Killed on 34th day. Necropsy, plague.	Survived and killed 21st day. Necropsy, no plague.
Blood level free sulfadiazine from hour of treatment	48 hrs. 6.0 mg. percent.		30 brs. 8.0 mg. percent.	•	36 hrs. 7.0 mg. percent. Micro test.
Treatment from hour of inocula- tion hour of treatment	96 hrs. First dose. 100 hrs. Second dose. 104 hrs. Third dose. 3 doses daily for 2 days. Total treatment 3 days, with 800 mr. sulfadiatine, 200 mg. sodium sulfadiazine, 1,800 mg. sodium bicarbonate.	No treatment.	144 hrs. First dose. 2 doses daily for 1 day. 3 doses daily for 1 day. 3 doses daily for 3 days. Total treatment 6 days, with 1,600 mg. sulfadiazine, 100 mg. sodium bulladiazine, 3,400 mg. sodium bloarbonate.	No treatment.	82 hrs. First dose. 2 doses daily for 6 days. 2 doses daily for 6 days. 7 otal frestment, 12 days. with 5.900 mg. sulfadiszine, 200 mg. sodium sulfadiszine, 5,800 mg.
Clinical course from hour of inoculation	s. Papule and bubo. s. Guinea pig thin. rs. Blood culture negative. Small and large bubo. s. Guinea pig stok.	rs. Papule and bubo. rs. Blood culture negative. Small and large bubo. rs. Blood culture negative. rs. Guinea pig sick.	Ta. Papule. Ta. Buboes larger. Ta. Buboes larger. Ta. Blood culture negative. Buboes large. Ta. Blood culture negative.	rs. Papule. rs. Bubo. rs. Bubo large. rs. Bubo large. rs. Do, culture negative. rs. Very large and small bubo. rs. Two very large buboes.	rs. Papule and bubo. rs. Lerge and small bubo. rs. Papule healed, buboes smaller.
Inoculation	25,000± P. pestfs, strain D 67, intracutane- 0us. 150 hrs.	28,000£P. pestiv. strain D 67, intracutane- ous. 192 hrs. 240 hrs.	25,000±P. pestis, 48 hrs. strain D 67, intra- 192 hrs. 122 hrs. 120 hrs. 150 hrs. 168 hrs. 169 hrs. 16	28,000±P pestis, 48 hrs. entaneous. 144 hrs. 192 hrs. 192 hrs. 192 hrs. 240 hrs. 288 hrs. 288 hrs. 288 hrs. 246	Flea bite, strain B 48 hrs. 3035.
Guinea pig No.	82	(Control)	Ŕ	(Control)	\$

Burvived and killed 21st day. Necropsy, no plagne.	Found dead morning 5th day.	Survived and killed 21st day. Necropsy, no plague.	Death on 6th day. Necropsy, plague.	Survived and killed 21st day. Necropsy, no plagne.	Found dead morning 7th day. Necropsy, plague.	Found deed morning 8th day. Necropsy. plague.
18 hrs. 7.0 mg. percent. 264 hrs. 7.5 mg. percent. Micro tost.		48 hrs. &4 mg. percent.		96 hrs. 6.0 mg. percent. Micro test.		
78 hrs. First dose. 2 doses daily for 6 days. 2 doses daily for 4 days. 7 otal treatment, 11 days, with 8.770 mg. sulfadiasine, 200 mg. sodium sulfadiasine, 5,200 mg.	No treatment.	48 hrs. First dose. 2 doses daily for 7 days. 2 doses daily for 1 day. 1 dose daily for 1 day. 3 doses daily for 2 day. 7 dose daily for 2 days. 7 rotal treatment 13 days. 8 200 mg. sulfadiastne, 200 mg. sodium sulfadiasine, 6,000 mg.	No treatment.	48 hrs. First dose. 52 hrs. Scond dose. 56 hrs. Third dose. 3 doses dally for 9 days. 2 doses 4 doses tally for 9 days. Total treatment 14 days, with 5.700 mg. sulfadisatine, 700 mg. sodium sulfadisatine, 7,600 mg.	No treatment.	72 hrs. First dose. 76 hrs. Recond dose. 80 hrs. Third dose. 18 doses daily for 1 day. Total treatment 2 days, with 600 mg. suifadiatine, 400 mg. so- dium suifadiatine, 1,200 mg. sodium blearbonate.
Papule. Bubo. Papule healed, bubo smaller.	Papule. Bubo. Guines pig sick.	Two papules, one bubo. Blood culture negative. Blood culture negative. Guines pig well. Small residual bubo. Bubo larger. Guines pig not very well.	Papule. Bubo. Blood culture positive.	Two papules, one bubo, Bubo very large. Noisy, difficult breathing. Papules healing. Bubo smaller. Breathing still noisy and uneven. Guines pig ist. Breathing normal. Bubo small.	Papule. Bubo. Two large buboes. Guines pig thin.	Two papules, two buboes.
48 brs. 72 brs. 168 brs.	48 hrs. 72 hrs. 78 hrs.	48 hrs. 96 hrs. 168 hrs. 264 hrs. 342 hrs.	48 hrs. 54 hrs. 96 hrs.	48 hrs. 146 hrs. 212 hrs. 250 hrs.	48 hrs. 50 hrs. 120 hrs.	72 hrs.
	m	м	A	м	A	м
strain	strain				strain	
47 Flee bite, strain B 2005.	(Control) 8035.	Flea bite, strain 3035.	(Control) 3035.	Floa bite, strain 3035.	(Control) 3035.	Figs bite, strain 3036.
	(Control)	5	(Control)	£8	(Control)	40.

Protocols of guinea pigs inoculated with P. pestis or infected by flea bites and treated with sulfadiazine—Continued

Inoculation	lat	<u>8</u>	Citat	Clinical course from hour of inoculation	Treatment from hour of inocula-	Blood level free sulfadiazine from hour of treatment	Results
Flea bite, strain B 72 hrs. 3035.			1	Papule. Bubo.	No treatment.		Found dead morning 6th day. Necropsy, plague.
Flee bite, strain B 66 hrs. 9035. 90 hrs. 114 hrs. 1152 hrs.	m			Papule and bubo. Bubo large. Blood culture positive. Guinea pig weak and sick.	66 hrs. First dose. 70 hrs. Second dose. 74 hrs. Third dose. 2 doses daily for 6 days. 2 doses daily for 6 days.	24 brs. 1.0 mg. percent. 72 brs. 4.0 mg. percent.	Found dead morning 13th day. Necropsy, right inguinal bubo, few P. pestis.
776 hrs. 272 hrs. 276 hrs.	272 hrs 272 hrs 276 hrs	176 hrs 272 hrs 276 hrs		smaller. Guines pig wear and site. Durous smaller. Guines pig fat, buboes small. Throat injured when feeding capsules. Noisy difficult breathing.	Total treatment 10 days, with 2,800 mg, suifadiazine, 800 mg, sodium suifadiazine, 5,400 mg, sodium bicarbonate.		Lung, liver, spleen, no plague. Cervical cellulitis
Fles bite, strain B 48 hrs. 3035.	Ф			Papule and bubo. Blood culture positive. Bubo very large.	No treatment.	•	Found dead morning 7th day. Necropsy, plague.
Flea bite, strain B 48 hrs. 3035.	Д			Papule and bubo. Guines pig sick. Small and large bubo. Edema around large. Blood culture positive.	52 hrs. First dose. 56 hrs. Second dose. 60 hrs. Third dose. 3 doses daily for 1 day. 2 doses daily for 1 day. Total treatment 4 days, with 800 mg. sulffediazine. 200 mg. so- dium sulfediazine, 1,400 mg. sodium bicarbonate.	68 hrs. 4.0 mg. percent.	Death on 5th day. Necropsy, plague.
Flea bite, strain B 48 hrs. 3035.	A			Papule and bubo. Guinea pig sick. Blood culture positive.	No treatment.		Death on 7th day. Necropsy, plague.
Flea bite, strain B 48 hrs. 3038. 120 hrs. 120 hrs. 150 hrs.	м			Papule. Bubo. (Before treatment.) Blood culture positive. Guines pig sick.	120 hrs. First dose. 122 hrs. Second dose. 128 hrs. Third dose. 3 doses dally for 1 day. Total treatment 2 days, with 400 mg. sulfadiazine, 500 mg. so- dium sulfadiazine, 1,200 mg. sodium bloar bonate.		Found dead morning 7th day. Necropsy, plague.
(Control) 3035.				Papule. Bubo. Blood culture positive.	No treatment.		Death on 7th day. Necropsy, plague.

			401	
96 hrs. 5.5 mg. percent. Micro Survived and killed 21st test. Necropsy, no plague.	Death on 9th day. Necropsy, plague.	Survived and killed 21st day. Necropsy, no plague.	Survived and killed 21st day, Necropsy, no plague.	Found deed morning 14th day. Necropsy, plague.
Micro				
5.5 mg. percent.		120 hrs. 4.0 mg. percent.	120 hrs. 7.0 mg. percent.	
96 hrs. test.			120 hrs	
48 hrs. First dose. 52 hrs. Second dose. 56 hrs. Third dose. 3 doses daily for 8 days. 2 doses daily for 4 days. Total treatment 13 days. with 3.50 mg. suiladiazine, 600 mg. sodium suiladiazine, 7,000 mg.	No treatment.	72 hrs. First dose. 77 hrs. Second dose. 80 hrs. Third dose. 3 doses daily for 7 days. 2 dose daily for 7 days. 1 dose daily for 1 day. Total treatment 12 days, with 3,000 mg. sulfadlazine, 200 mg. sodium sulladiazine, 5,400 mg. sodium blear bonste.	54 hrs. First dose. 3 doses dally for 7 days. 2 doses dally for 7 days. 1 dose dally for 1 day. 7 rotal treatment 12 days, with 2,800 mg, sulfadiazine, 200 mg. sodium sulfadiazine, 5,600 mg. sodium bicarbonate.	No treatment.
hrs. Papule and bubo. hrs. Bubo large. hrs. Bubo small.	hrs. Papule. hrs. Bubo. hrs. Guines pig sick.	hrs. Two papules. hrs. Two buboos. hrs. Buboes very small.	hrs. Papule. hrs. Bubo. hrs. Massive bubo. hrs. Bubo small.	8 hrs. Pepule. 2 hrs. Bubo.
\$ 52 5	1828	48 72 72 73 74 75	42 48	
train B	strain B	strain B	strain B	strain B
Flee bite. strain B 3036.	Control) 3035.	Flea bite, strain 3035.	Flea bite, strain B 8035.	Flea bite, strain 3085.
	Control)			(Control)

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COMPLEMENT FIXATION IN THE RICKETTSIAL DISEASES— TECHNIQUE OF THE TEST 1

By IDA A. BENGTSON, Senior Bacteriologist, United States Public Health Service

The complement fixation test has been applied in the study of the following rickettsial diseases: endemic typhus (murine typhus), epidemic typhus (European typhus) (1, 2, 3), Rocky Mountain spotted fever (4), Tobia fever of Colombia (probably identical with Rocky Mountain spotted fever), and Q fever of North America and Australia (probably identical) (5).

The test is quantitative and the present report presents its details more fully than has previously been done. For the sake of economy and convenience the various reagents in amounts of 0.2 cc. are employed. The tests are made in tubes measuring $10-11 \times 75$ mm. The reagents, except serum dilutions and antigen, are delivered in the tubes by means of an automatic pipette set at 0.2 cc. for salt solution and complement and 0.4 cc. for the mixture of hemolysin and sheep red corpuscles. Serum dilutions are made by means of a syringe pipette. The total volume in each tube is 1 cc.

REAGENTS

The reagents include the usual hemolytic system, with guinea pig complement, antisheep cells rabbit hemolysin, and sheep red blood corpuscles, the antigen, and standard serum.

THE HEMOLYTIC SYSTEM

- (a) Sheep's red blood cells.—The sheep cells are employed in a 2 percent suspension in 0.85 percent physiological salt solution after being washed at least three times so that the supernatant fluid shows no tinge of red. In the hemolysin titration, 0.2 cc. amounts of the 2 percent suspension are used, while in the test proper 0.4 cc. of the mixture of equal volumes of the 2 percent suspension and the proper hemolysin dilution are used.
- (b) Hemolysin.—The hemolysin is prepared by inoculating rabbits with washed sheep cells according to the method of Kolmer (6) or Kilduffe (7). It is preserved with an equal volume of glycerin.

Titration of hemolysin: The following dilutions are prepared from a 1:100 dilution of hemolysin (2 cc. of a 50 percent glycerinated hemolysin + 98 cc. of 0.85 percent saline): 1:1,000, 1:1,200, 1:1,600, 1:2,000 1:2,400, 1:3,000, 1:4,000, 1:5,000, 1:6,000, 1:8,000, 1:10,000, 1:12,000. These dilutions in 0.2 cc. amounts are transferred to test tubes in duplicate. To each tube is added the following: 0.2 cc. complement diluted 1:40, 0.4 cc. saline, and 0.2 cc. 2 percent sheep red corpuscles.

¹ From the Division of Infectious Diseases, National Institute of Health.

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The tubes are shaken individually after the addition of each reagent and incubated in the 37° water bath for 1 hour. The unit of hemolysin is 0.2 cc. of the highest dilution showing complete hemolysis. Two units of hemolysin are used in the test, and, therefore, the dilution of hemolysin to be used is that one which has twice the concentration of the unit (e. g., if the unit is 0.2 of the 1:3,000 dilution, 2 units are contained in 0.2 cc. of the 1:1,500 dilution).

(c) Complement.—Complement is obtained by bleeding 10 to 15 guinea pigs from the heart and the blood from each guinea pig is collected separately in large test tubes previously rinsed with sterile 0.85 percent salt solution. Care is taken not to cause rupture of the red corpuscles. The needle is removed from the syringe, and the blood allowed to flow slowly into the tube. The tubes are allowed to stand in a refrigerator over-night, and in most of the tubes the clot will have separated from the serum. The serum is removed by means of a Pasteur pipette with a rubber bulb attached. The serums from all the tubes are pooled and then centrifuged to precipitate remaining cells. Tests have been made on various occasions for nonspecific fixability, hemolytic activity, and the presence of natural antisheep hemolysin. Results show that when serums from such a large number of guinea pigs are pooled, these tests are not necessary on individual serums. The complement is preserved by the addition of an amount of saturated NaCl solution equivalent to one-tenth the volume of the serum. The salted complement is kept in the refrigerator and removed only long enough to remove the desired amount of the reagent.

Titration of complement.—The complement is titrated, using a 1:40 dilution prepared as follows: 1 cc. complement, 3 cc. distilled water (to restore tonicity), and 36 cc. 0.85 percent salt solution.

A 0.2 cc. pipette graduated in hundredths is used for measuring the following amounts of diluted complement: 0.08, 0.10, 0.12, 0.14, 0.16, 0.18, 0.20 cc. This is done in duplicate. Sterile 0.85 percent salt solution is added in such amounts that the volume in each tube is 0.4 cc. The antigen in the dilution used in the test is added in amounts of 0.2 cc. The tubes are shaken individually after the addition of each reagent.

The complement mixtures are incubated in the 37° water bath for 1 hour, after which 0.4 cc. of sensitized sheep cells is added (equal parts of 2 percent sheep cells and the dilution of hemolysin which contains 2 units per 0.2 cc., prepared by thorough mixing 10 minutes previously).

Readings are made after a further hour's incubation. The unit is the next to the smallest amount showing complete hemolysis. Two units of complement are used in the test.

ANTIGEN

Antigens are prepared from rickettsiae grown in the yolk sac of fertile hen eggs (method of Cox (8)). Only such yolk sacs are used as contain a large number of rickettsiae. This applies to endemic and epidemic typhus and Q fever. The yield of rickettsiae in Rocky Mountain spotted fever is usually less than in the typhus fevers and Q fever, but even though rickettsiae are not numerous, satisfactory antigens have been prepared.

The infected yolk sacs are weighed and ground in a mortar with alundum or in the Waring Blendor without alundum and prepared as 10 percent suspensions with 0.85 percent salt solution containing 1:10,000 merthiolate. After slow centrifugation to remove coarse particles, the supernatant portion is centrifuged in the angle centrifuge at 4,000 r.p.m. for 1 hour. The precipitate is suspended in 0.85 percent sterile saline containing 1:10,000 merthiolate. After standing in 50 cc. centrifuge tubes in the refrigerator for a week or more a considerable amount of precipitate settles out leaving a somewhat turbid suspension. The supernatant portion is pipetted off and this serves as antigen. The antigens as prepared are stable over a period of several months when stored at ice-box temperature. Epidemic typhus vaccine has been successfully employed as antigen in epidemic typhus. Due to the considerable amount of cross fixation it may also be made use of provisionally in testing suspected endemic typhus semms.

Antigens are titrated against a standard positive serum which has been diluted appropriately, and antigen titers have been found to range from 1/8 to 1/128 for epidemic and endemic typhus and somewhat lower for Q fever. Rocky Mountain spotted fever antigens have been used undiluted and diluted ½ and ¼. The titer or unit of the antigen may be considered to be the smallest amount which gives 4+ fixation with the standard serum. Four units of antigen are used in the test.

The serum selected for titrating antigens may be a serum of moderately high titer (1/64 to 1/128) diluted 1/16 or 1/32. Occasional titrations of antigen against one dilution of serum are desirable, or cross titrations of varying dilutions of antigen against varying dilutions of serum may be carried out.

SERUMS

Serums are inactivated at 56° C. for one-half hour. Serum dilutions are prepared with a syringe pipette. Twofold dilutions are used ranging from 1/4 to 1/512, and higher dilutions are made if the end point is not reached. The 1/4 dilution is prepared by adding 0.3 cc. saline to 0.1 cc. serum. Amounts of 0.2 cc. are carried over to the 0.2 cc. amounts of saline contained in the tubes for the higher dilutions.

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THE TEST

To the serum dilutions contained in the tubes are added 0.2 cc. of the proper dilution of antigen (4 units) and 0.2 cc. of complement (2 units).

After 1 hour's incubation in the 37° water bath the sensitized sheep cells are added in 0.4 cc. amounts, the hemolysin (2 units per 0.2 cc.) and 2 percent sheep cells having been prepared by thorough mixing 10 minutes previously. After further incubation in the 37° water bath for 1 hour, the test is placed at the cold room temperature and read the following morning. Positive results are recorded as 4, 3, 2, 1+, and trace, and the titer is read as the highest dilution showing 3 or 4+ fixation.

The following controls are set up:

Serum controls. To duplicate tubes of the four lowest dilutions are added the same reagents as for the test except that salt solution is substituted for antigen.

Antigen controls. Antigen controls contain twice the volume of the dilution used in the test, i.e., 0.4 cc., and 0.2 cc. of complement and 0.4 cc. of sensitized cells.

Hemolytic system. The hemolytic system control consists of four tubes containing 0.05, 0.1, 0.15, and 0.2 cc. of the dilution of complement used in the test, these amounts representing ½, 1, 1½, and 2 units. The volume in each tube is made up to 0.6 cc. with sterile saline, and 0.4 cc. sensitized cells added. The tube containing 0.05 should show 1 or 2+ fixation, and the three remaining tubes should be completely hemolyzed.

Standard serum. A standard serum composed of a pooled lot of serums from recovered guinea pigs is titrated with each test, using the same dilutions as for the serums under test.

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THE ALIPHATIC ALCOHOLS: THEIR TOXICITY AND POTENTIAL DANGERS IN RELATION TO THEIR CONSTITUTION AND THEIR FATE IN METABOLISM ¹

A Review

This study covers a review of the literature on the toxicity and potential dangers of monovalent, bivalent, trivalent, and polyvalent alcohols. Each group is followed by a discussion of the relation of the chemical constitution, physical-chemical properties, and metabolic fate of these alcohols to their toxicological action. The study is based on information gathered from approximately 1,200 publications which are quoted in the bibliography.

DEATHS DURING WEEK ENDED MARCH 11, 1944

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

	Week ended Mar. 11, 1944	Corresponding week,
Data for 92 large cities of the United States: Total deaths. Average for 3 prior years. Total deaths, first 10 weeks of year. Deaths under 1 year of age. Average for 3 prior years. Deaths under 1 year of age, first 10 weeks of year. Data from industrial insurance companies: Policies in force. Number of death claims. Death claims per 1,000 policies in force, annual rate. Death claims per 1,000 policies, first 10 weeks of year, annual rate.	9, 526 9, 685 103, 505 597 608 6, 303 66, 357, 378 13, 836 10. 9	10, 181 102, 388 727 7, 284 65, 413, 543 14, 021 11, 2 10, 8

¹ The aliphatic alcohols: Their toxicity and potential dangers in relation to their constitution and their fate in metabolism. By W. F. von Oettingen. Public Health Bulletin No. 281. Government Printing Office, 1943. (Distributed February 1944.) For sale by the Superintendent of Documents, Washington 25, D. C. Price 35 cents.

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 MARCH 18, 1944

Summary

For the second consecutive week the incidence of meningitis for the country as a whole has fallen below that for both the preceding week and the corresponding week of last year. A total of 497 cases was reported, as compared with 517 last week, 614 a year ago, and a 5-year (1939-43) median of 54. The cumulative total for the year to date is 6,087, as compared with 4,659 for the same period last year and 2,689 in 1929, the year of record having the next highest incidence.

Currently, increased incidence was reported in 5 of the 9 geographic divisions. Slight decreases occurred in the Middle Atlantic, the East North Central, and the Mountain areas, and the largest decrease in the East South Central area. Nine States reporting more than 20 cases each (last week's figures in parentheses) are as follows: *Increases*—Massachusetts 25 (8), Ohio 29 (26), Michigan 35 (28), Missouri 26 (20), Virginia 24 (17), California 35 (32); decreases—New York 55 (67), Pennsylvania 39 (40), Illinois 25 (29).

Slight increases were recorded for measles and scarlet fever, with totals of 32,802 and 7,373 cases, respectively, or 46 percent and 67 percent above the respective corresponding 5-year medians. To date, 240,054 cases of measles and 61,731 cases of scarlet fever have been reported, as compared with 5-year medians of 158,612 and 44,084, respectively. The incidence of these diseases to date is above that for any corresponding period since 1938.

A total of 20 cases of poliomyelitis was reported (7 in California), as compared with 19 last week and a 5-year median of 16.

Incidence below corresponding 5-year medians is reported for diphtheria, influenza, smallpox, typhoid fever, and whooping cough. The cumulative figures for all of these diseases except influenza are also below the respective medians.

A total of 9,537 deaths was recorded in 93 large cities of the United States, as compared with 9,548 for the preceding week and a 3-year (1941-43) average of 9,389. The cumulative total to date is 113,209, as compared with 112,524 for the same period last year.

Telegraphic morbidity reports from State health officers for the week ended March 18, 1944, and comparison with corresponding week of 1943 and 5-year median

In these tables a zero indicates a definite report, while leaders imply that, although none was reported, cases may have occurred.

Mar. Mar.		1	Diphth	eria	I	nfluen	za		Measle	8		feningi ningood	
Mar. Mar. 1930 Mar.	Division and State												Me-
Maine		18,	20.	1939-	18.	20.	. 1939- 43	Mar. 18, 1944	20.	1939-	Mar. 18, 1944	20.	1939-
New Hampshire	NEW ENGLAND	1	1		1	ł		1	1	1			
New York	New Hampshire Vermont Massachusetts Rhode Island		0 0 0 0 5 5	0 0	2			17: 70: 37:	3 1: 3 31: 8 1, 39: 6 1:	3 13 0 18 4 830 4 14	25	0 3 34 24	0
New Jersey	MIDDLE ATLANTIC			1	1			l	1	1			
Data Data	New Jersey Pennsylvania	. 8	3	4	13	18	16	1, 360	8 1.46	7 443	15	29	3 1 7
Indiana				١.,		~		2 41	724				
Minnesota	Indiana Illinois Michigan ³ Wisconsin	15 8 9	6 8	7 21 8	9 117 6	13 27	57 41 27	266 1, 280 1, 651	6 403 0 963 1 554	128 648 5 248	9 25 35	18 7	2 1 4 1 1
North Dakota		1				l	١ ـ				_		
SOUTH ATLANTIC Delaware	Iowa Missouri North Dakota South Dakota Nebraska	3 2 2 3	0 6 0	6 1 0	30 3 22 4	1 2 24	28 11 21 2 15	213 474 245 75 70	332 378 3 131 5 52 9 249	196 151 64 27 107	0 26 3 0	0 27 1 0 2	0 0 0 0 1 1
Maryland		1							1	1			
RAST SOUTH CENTRAL	Maryland ² District of Columbia. Virginia. West Virginia. North Carolina. South Carolina. Georgia.	5 0 1 2 7 5	11 0 7 8 8 4	5 10 4 10 3 8	261 40 14 449 24	792 68 77 840 152	792 218 77 774 152	1, 182 129 1, 235 566 2, 106 492 303	73 100 779 36 77 190 187	170 83 376 36 921 190 254	9 1 24 12 11 13 6	25 3 58 3 23 18 9	0 2 1 3 2 1 0 1
Tennessee	EAST SOUTH CENTRAL				- 1								
Arkansas 7 12 7 127 99 291 291 102 102 4 5 0 Louisiana 1 6 6 7 13 27 68 232 154 6 13 1 Oklahoma 6 7 7 155 190 213 88 65 65 4 2 1 Texas 244 34 36 1, 201 1, 543 1, 543 2, 038 1, 160 1, 160 20 28 4 MOUNTAIN Montana 1 2 2 10 24 24 105 343 87 0 0 0 1 Idaho 0 0 0 37 7 72 128 71 0 2 0 0 1 Idaho 1 7 18 36 46 46 350 717 247 0 0 0 0 Wyoming 2 0 0 1 46 5 1116 177 62 1 0 0 0 Colorado 1 7 8 36 46 46 350 717 247 0 0 0 0 New Mexico 1 0 2 4 1 5 140 20 37 0 1 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	Alabama	5 9	7 5	6 6	81	93	161	312	341	165	11 12	8 10	2 2 8 1
Louisiana					1		i					- 1	
Montana 1 2 2 10 24 24 105 343 87 0 0 0 0 10 daho 72 126 71 0 2 0 <td>LouisianaOklahomaTexas</td> <td>1 6</td> <td>6</td> <td>6</td> <td>7 155</td> <td>13 190</td> <td>27 213</td> <td>68 88</td> <td>232 65</td> <td>154 65</td> <td>4</td> <td>13 2</td> <td>0 1 1 4</td>	LouisianaOklahomaTexas	1 6	6	6	7 155	13 190	27 213	68 88	232 65	154 65	4	13 2	0 1 1 4
Idaho													
Washington 2 3 2 1 9 181 947 322 8 9 1	Idano. Wyoming Colorado New Mexico Arizona Utah 3	0 2 1 1 5 0	0 7 0 4 1	0 8 2 2	37 1 36 4 125 158	46 46 1 68 8	5 46 5 209	72 116 350 140 557 31	126 177 717 20 47 466	71 62 247 37 95 155	0 1 0 0 3 1	2 0 0 1 1 7	0 0 0 0 1 0
Washington					i						- 1	- 1	
Oregon 5 4 45 25 28 102 491 421 6 9 0 California 21 23 23 59 74 211 2,094 742 742 35 29 2	Oregon	5		4		25	28	102	491	421	6	9	Ō
Total 243 240 293 3,465 4,536 6,366 32,802 23,150 22,521 497 619 54	Total	243	240	293	3, 465	4, 536	6, 366	32, 802	23, 150	22, 521	497	619	54
11 weeks 32,790 3,191 3,542 314,418 49,953 54,065 4240,054 159,593 158,612 6,087 4,659 587	11 weeks	2, 790	3, 191	3, 542	14, 418 4	9, 953	54, 065	240,054	159, 593	158, 612	6, 087	4, 659	587

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Telegraphic morbidity reports from State health officers for the week ended March 18, 1944, and comparison with corresponding week of 1943 and 5-year median—Con.

	Poi	liomye	litis	8	carlet fe	ver	8	malipo	X	Typh typ	oid and hoid fe	i pera- ver *
Division and State	wend	eek ed—	Me-	Week	ended-	Me-	w	eek ed—	Me-	We		Me-
·	Mar. 18; 1944	Mar. 20, 1943	dian 1939- 43	Mar. 18, 1944	Mar. 20, 1943	dian 1939– 43	Mar. 18, 1944	Mar. 20, 1943	dian 1939– 43	Mar. 18, 1944	Mar. 20, 1943	dian 1939– 43
NEW ENGLAND												
Maine	0 0 2 0 0	0 0 0 1	0 0 0 0	17 447 14	6 8 592 20	3 8 169	000	0 0 0 0	0 0 0 0	1 0 0 2 0	0 1 0	Ó
MIDDLE ATLANTIC New York New Jersey Pennsylvania	0	1 2 2	0	288	183	655 197 342	0 0 0	0	0	5 0 2	8 0 9	5 1 8
Chio	0 0 1 0	0 0 2 0 3	0 0 0 0	507 225 551 264 332	117 224 96	343 191 446 259 186	0 0 1 0	5 1 8 0 1	0 1 8 2 3	1 1 0 2 0	3 1 2 1 0	2 1 2 1 0
WEST NORTH CENTRAL Minnesota	1 0 1 0 0 2 0	0 0 2 0 0	0 0 1 0 0	219 178 173 71 34 86 131	62 94 138 7 15 36 64	88 65 86 21 12 30 67	000000	0 1 0 0 1 0 2	2 1 8 0 1 1	0 0 3 1 0 0	0 1 0 0 0	0 1 2 0 0 0
SOUTH ATLANTIC Delaware	000000000000000000000000000000000000000	0 0 0 1 0 0 0	0 0 0 0 0 0	16 238 222 93 116 41 8 17	6 112 16 53 24 28 8 6 11	16 47 18 36 46 34 5 14 8	000000000000000000000000000000000000000	0 0 0 0 10 0	00000000	0 1 0 3 4 0 1 8	0 7 0 2 2 5 0 2 5	0 1 0 2 2 0 0 3 3
EAST SOUTH CENTRAL KentuckyTennesseeAlabama Mississippi 3	0 0 0	1 0 0 2	0 0 0	77 96 17 4	49 51 18 7	94 75 23 6	0 0 0	0 0 1 0	0 1 1 0	3 0 0 3	0 0 1 0	2 2 2 1
WEST SOUTH CENTRAL Arkansas Louisiana Oklahoma Texas	0 0 0 2	0 0 0 3	0 1 0 1	12 17 16 155	7 11 27 65	7 11 21 65	0 0 0 5	3 1 0 0	3 1 0 2	2 1 2 8	3 1 1 6	3 5 1 6
MOUNTAIN Montana Idaho Wyoming Colorado New Mexico Arizona Utah ¹ Nevada	0 1 0 1 2 0 0	0 0 0 0 0 1	0 0 0 0 0 1 0	53 109 8 71 17 20 109 0	6 12 40 103 1 16 57 6	26 11 11 51 5 7 29 0	0 0 0 1 0 0 0	0 0 0 1 0 0 0	0 0 0 1 0 0	0 0 0 3 3 1 0	0 0 0 0 0	0 1 0 0 0 1 0
PACIFIC Washington Oregon California	0 0 7	1 1 2	0 0 2	360 156 373	46 6 158	46 11 170	0 1 0	0 0 1	0 0 1	1 0 7	1 0 3	1 1 5
Total	20	26	16	7, 373	4, 360	4, 426	8	36	65	70	67	87
11 weeks	263	302	287	61,731	42, 595	44, 084	144	300	521	816	585	834

Telegraphic morbidity reports from State health officers for the week ended March 18, 1944, and comparison with corresponding week of 1943 and 5-year median—Con.

	Wh	ooping	cough			Wee	k ende	d Marc	h 18, 1	944		
Di-later and	Week	ended-	- 1			ysenter	y	En-		Rocky		<u> </u>
Division and State	Mar. 18, 1944	Mar. 20, 1943	Me- dian 1939- 43	An- thrax	Ame- bic	Bacil- lary	Un- speci- fled	ceph- alitis, infec- tious	Lep- rosy	Mt. spot- ted fever	Tula- remia	Ty- phus fever
NEW ENGLAND	36	67	32	0	0	0	0	0	0	0	0	
Maine New Hampshire	0	1 1	5	ĺ	0	0	Ō	0	Ó	Ó	0	0
Vermont Massachusetts	15 94	23 166	35 171	0	0	0 2	0	0	8	0	8	0
Rhode Island	0 38	34 66	28 66	0	0	0	0	0	0	0	0	Ó
Connecticut	300	w		۰	۰	· •	ľ	•	"	"	ا ا	U
New York New Jersey Pennsylvania	121 42 141	423 206 369	423 206 361	0 1 0	3 1 4	7 0 0	0	3 1 2	0	0	0	0 0 0
E. NO. CEN.	80	233	233	0	0		3	0	0	0	0	0
Indiana	16	72	45	0	Ó	Ŏ	0	0	0	0	0	Ó
Illinois Michigan ²	46 120	153 253	124 188	0	0	0	0	0	0	0	0	0
Wisconsin	63	209	182	ŏ	ŏ	Ô	ŏ	ŏ	ŏ	ŏ	ŏ	ŏ
W. NO. CEN.	26	54	41	0	0		0	0	0	0	0	0
Minnesota Iowa	20	18	15	Ŏ	Ō	Ŏ	Ö	Ŏ	Ŏ	ŏ	Ŏ	Ó
Missouri North Dakota	10 2	22 12	31	0	0	0	1 0	1	0	0	0	0
South Dakota	1	1	i	0	0	0	0	0	0	0	0	0
Nebraska Kansas	19 28	61	7 57	0	0	0	0	0	8	0	0	0 1
SOUTH ATLANTIC	_	-	"		•		Ĭ	-	ľ	Ĭ	1	-
Delaware	1	120	4 72	o l	0	0	Ŏ	0	0	0	0	0
Maryland ³ District of Co-	59		'-	0	0	0	. 0	_	0	0	0	
lumbia	3 74	26 55	15 55	0	0	0	0 26	0	0	0	0 2	0
West Virginia	45	55	46	0	0	0	0	0	0	0	0	Ó
North Carolina	115 61	163 43	163 57	0	0 2	0	0	0	0	0	0	0
Georgia	16 14	36 32	36 28	1 0	0	1 1	0	0	0	0	7 0	5
Florida E. 90. CEN.	13	02	20	٠	•	-	١	١	ľ	٠	١	v
Kentucky	61	35	. 51	0	0	0	0	0	0	0	0	0
Tennessee	21 35	122 49	40 31	0	0	0	3	0	8	0	1	0
Mississippi 3				0	0	0	0	0	0	0	0	4
W SO. CEN.	7	42	11	0	1	2			0		1	0
Louisiana	Ó	8	8	O I	2	1	Ŏ	. 0	Ō	Ó	Ō	4
Oklahoma Texas	10 211	23 420	15 208	8	0 11	0 125	8	1 8	0	8	8	0 18
MOUNTAIN				1						- 1		
MontanaIdaho	3 8	12 0	10 3	0	0	0	8	8	0	8	8	0
Wyoming	6	1	3	0	Ó	0	O	Ō	Ŏ.	Ŏ	Ó I	Ó
Colorado New Mexico	24 8	29 41	36 16	0	0	0	0	0	0	8	0	0
Arizona	31	23	20	0	0	0	19	0	0	0	0	0
Utah 2 Nevada	20 0	40 0	87 2	0	0	0	0	8	. 0	8	8	0.
PACIFIC			1			- 1		_ I	1		1	_
Washington	55 36	25 10	61 13	0	8	8	0	0	8	0	8	0
California	106	318	286	ŏ	2		ŏ	0	1	0	0	1
	1, 948	4, 183	4, 024	2	27	152	52	13	1	0	12	34
11 weeks	20, 283	42, 972	43, 609	10 19	276 292	2, 184 2, 268	700 472	115 115	7 5	2 3	113 190	451 548

¹ New York City only.
² Period ended earlier than Saturday.
³ Later information shows 62 cases of diphtheria in Texas for the week ended March 11, instead of 26 as previously reported.
¹ The following are corrected cumulative figures for measles, respectively, for the first 7, 8, and 9 weeks of the year: 114,762, 141,835, and 176,073.
¹ Including paratyphoid fever cases reported separately as follows: Indiana, 1; Virginia, 1; Georgia, 1; Texas, 1; Washington, 1; California, 3.

WEEKLY REPORTS FROM CITIES

City reports for week ended March 4, 1944

This table lists the reports from 85 cities of more than 10,000 population distributed throughout the United States, and represents a cross section of the current urban incidence of the diseases included in the table.

		fifeo	Influ	enza		menin-	atha	8	2800	_	De.re.	cough
	Diphtheria cases	Encephalitis, in tious, cases	Cases	Deaths	Measles cases	Meningitis, m. gococcus, cas	Pneumonia desths	Poliomyelitis	Scarlet fever c	Smallpox cases	Typhoid and I	Whooping co
NEW ENGLAND								l				
Maine: Portland	0	0		0	24	0	5	0	17	0	0	1
New Hampshire: Concord Vermont:	0	0		0	0	0	2	0	2	0	0	0
Barre	0	0		0	0	0	0	0	1	0	0	. 0
Boston Fall River Springfield Worcester	5 0 0	0 0 0		0 0 0	90 22 54 0	10 0 2 0	10 2 0 9	000	86 3 31 47	000	0	16 1 2 2
Providence	1	1		0	259	6	6	0	8	0	0	9
Connecticut: Bridgeport	0	0	2 1	0	30 2 147	2 1 8	0	0	4 8 6	0 0 0	0	0 0 6
MIDDLE ATLANTIC												
New York: Buffalo New York Rochester Syracuse New Jersey:	0 12 0 0	0 0 0	10	1 1 0 0	1, 658 1 1	2 37 3 0	9 84 3 2	0 0 0	22 298 7 3	0 0 0	0 0 0	0 38 2 11
New Jersey: Camden Newark Trenton Pennsylvania:	0 0 0	0	i	0 1 0	1 59 6	0 10 1	2 4 7	0	31 17 8	0 0 0	0 0 0	0 9 0
Pennsylvania: Philadelphia Pittsburgh Reading	2 0 0	0	5 5	8 5 1	53 222 5	12 8 0	34 25 3	0	86 23 2	0	0	15 5 1
BAST NORTH CENTRAL												
Ohio: Cincinnati Cleveland Columbus Indiana:	2 0 0	0 0 0	1 2	0 0 2	44 1, 088 191	4 9 2	3 5 1	0	43 61 10	0	0	2 18 3
Fort WayneIndianapolisSouth BendTerre Haute	0 1 0 1	. 0		0 1 0 0	0 22 2 0	0 8 0 0	1 7 0 8	0	2 54 4 0	0	2 0 0 0	0 10 0 0
Illinois: Chicago Springfield	2	0	. 3	5	83 42	29	28	o l	190	8	0	9 1
Michigan: Detroit	5	0	3	1 0	75 18	12	13	0	90 2	8	1 0	7
Grand Rapids Wisconsin: Kanosha	ŏ	0		ŏ	230	ŏ	i	ŏ	10	ŏ	ŏ	Ö 1
Milwaukee Racine Superior	0	0	2	0	52 13 3	6 0	13 1 0	0	94 3 24	0	0	23 7 0
WEST NORTH CENTRAL												
Minnesota: Duluth Minneapolis St. Paul	0 1 0	0		0 1 0	9 820 572	0 1 1	0 7 5	0	20 58 83	0	0	12 3 2

See footnotes at end of table.

City reports for week ended March 4, 1944—Continued

									,			
	88	firfec	Influ	enza		menin-	eths	Calles	99968	_	Dara- cases	cough
	Diphtheria ca	Encephalitis, in tions, cases	Cases	Deaths	Measles cases	Meningitis, m gococcus, car	Pneumonia desths	Poliomyelitis	Scarlet fever o	Smallpox cases	Typhoid and typhoid fever	Whooping of
WEST NORTH CENTRAL— continued				,								
Missouri: Kansas City St. Joseph St. Louis Nebrasks:	1 0 0	1 0 0	10	3 0 1	31 0 228	3 4 20	15 0 14	0	37 5 29	0	0	1 0 4
Omaha Kansas: Topeka Wichita	0 0	0	i	0 1 0	9 243	2 1 1	11 3 6	0	29 4 1	0	0	1 8 0
SOUTH ATLANTIC	Ĭ		-	Ĭ			Ĭ		-	J		
Delaware: Wilmington Maryland:	0	0		0	6	1	4	0	1	0	0	. 0
Baltimore Cumberland Frederick	9 0	0 0 0	8	1 0 0	706 0 0	5 0 0	19 1 0	0 0 0	64 0 1	0	.0 0	12 0 0
District of Columbia: Washington Virginia:	0	0	2	0	136 13	2 0	9	0	232 0	0	0	3 1
Lynchburg Richmond Roanoke	ő	0	5 2	0 3 0	188 126	4 0	· 2 2 1	ő	8 2	. 0	0	0
West Virginia: Charleston	0	0		0	0 2	0	0	8	9 20	0	0	0
Winston-Salem South Carolina: Charleston	0	0	1 35	0	87 61	0	1 2	0	3 0	0	0	8
Georgia: Atlanta Brunswick	0	0	17	0	50 57	1 5	3	0	6	0	0	0
Savannah Florida: Tampa	0	0	3	0	4 9	0	0	0	. 1	0	0	0 1
EAST SOUTH CENTRAL						į			Ì			
Tennessee: Memphis Nash ville	8	0	4 0	1 2	12 3	15 5	2 3	0	14 9	0	0	11 0
Mobile	0	0	Ż	0	16	1	3	0	0	0	0	2
Arkansas: Little Rock	0	0	2	0	39		1	0	1	٥		2
Louisiana: New Orleans	3 1	1 0	8	3 1	35 0	8	6 7	0	5	0	1 0	0
Texas: Dallas Galveston Houston San Antonio	2 0 6 0	0 0 0	16 2	2 0 0 2	85 4 49 12	3 0 2 1	8 1 10 10	0 0 0 0	8 1 2 0	0 0 0	0 0 1 0	1 0 0 0
MOUNTAIN									,			
Montana: Billings Great Falls Helena Missoula Idaho:	0 2 0 0	0 0 0	9	0 0 0	3 8 1 0	0	1 1 0 1	0	5 5 0 5	0	0 0 0	0 0 0
Boise	0	0	l _.	0	11	0	0	0	10	0	0	0

See footnotes at end of table.

City reports for week ended March 4, 1944-Continued

		infeo	Infi	ents.		mentn-	ą	1	1		2 8 2 8	danoo
	Diphtheria on	Encephalitis, i	Canada	Deaths	Measles cases	Meningitis, ma	Pneumonia de	Poliomyelitis	Souriet fever o	Smallpox case	Typhoid and I	Whooping oo
Colorado: Pueblo	0	0		0	14	0	2	0	2 35	0	0	3
PACIFIC Washington: Sectile	8	000		0	35 46	2 0	5	8	20	. 0	. 0	ű
Spokane	32 7 1	0	25	300	172 9 38	5 0 8	14 0 5	1 0	15 78 41 3 43		1 0	11 2 ,1 5 6
Total	100	3	191	48	8, 460	274	486	ì	2, 222	0	7	296
Corresponding week, 1943 Average, 1939–43	55 73	2	181 616	37 1 58	4, 510 24,309	167	532 1 518	4	1, 398 1, 517	0 11	8 18	1, 151 1, 035

¹ 3-year average, 1941–43. ² 5-year median.

Dysentery, amedic.—Cases: New York, 2; Philadelphia, 5; St. Louis, 1; Baltimore, 1; Los Angeles, 1.
Dysentery, bacillary.—Cases: Providence, 1; Buffalo, 8; New York, 1; Philadelphia, 2; Detroit, 1; St. Louis, 1; Richmond, 2; Charieston, 8. C., 3; Los Angeles, 1.
Dysentery, unspecified.—Cases: Baltimore, 1; San Antonio, 4.
Tuleremia.—Cases: St. Louis, 1.
Typhus fever.—Cases: Charleston, S. C., 1; Mobile, 1; Houston, 1.

Rates (annual basis) per 100,000 population, by geographic groups, for the 85 cities in the preceding table (estimated population, 1942, 34,022,600)

	case , infec- rates	influenz		ienza	rates	menin- case	death	CBS6	case	rates	Case	qSnoo
	Diphtheria ca rates	Encephalitis, i tious, case re	Case rates	Death rates	Measles case r	Meningitis, me gococcus, rates	Pneumonia de rates	Poliomyelitis crates	Scarlet fever c rates	8	Typhoid and paratyphoid fever case rates	20 8
New England Middle Atlantic East North Central West North Central South Atlantic East South Central West South Central West South Central Mountain Pacific	14. 9 6. 3 6. 4 7. 9 17. 4 0. 0 35. 3 33. 0 71. 9	2.5 0.0 0.0 2.0 0.0 0.0 3.0 0.0	7. 5 9. 8 6. 4 21. 8 128. 8 53. 8 88. 2 148. 4 43. 8	0.0 5.4 6.4 11.9 7.0 26.9 23.5 16.5	1, 565 899 1, 092 3, 792 2, 514 278 659 643 536		84. 7 77. 4 49. 2 121. 1 80. 0 71. 7 126. 5 148. 4 49. 1	0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0	531 222 347 528 604 206 529 1, 022 358	0.0 0.0 0.0 0.0 0.0 0.0	0.0 0.0 1.8 0.0 1.7 0.0 5.9 0.0 1.8	92 36 48 52 40 117 9
Total	15. 4	0.5	29. 4	7.4	1, 300	42.1	74.7	0. 2	341	0.0	1.1	45

TERRITORIES AND POSSESSIONS

Hawaii Territory

Plague (rodent).—Rats proved positive for lague have been found in Hamakua District, Island of Hawaii, T. H., as follows: Paauhau-January 24, 1944, 1 rat, January 27, 1 rat, February 4, 1 rat; Kapulena-February 2, 1 rat, Februar / 9, 1 rat; Kukuihaele-February 2, 1 rat.

Panama Canal Zone

Notifiable diseases—January 1944.—During the month of January 1944, certain notifiable diseases were reported in the Pan ma Canal Zone and terminal cities as follows:

Disease	Pa	nama	c	olon	Can	al Zone	Zone	side the and ter- al cities	Т	otal
	Cases	Deaths	Cases	Deaths	Cases	Deaths	Cases	Deaths	Cases	Deaths
Chickenpox Diphtheria Dysentery (amebic) Dysentery (bacillary) Leprosy Malaria Messies Meningitis, meningococcus Mumps Paratyphoid fever Pneumonia Relapsing fever Tuberculosis Typhoid fever Whooping cough	10 6 2 13 1 1 10	7	11 1 2 2 2	3	2 118 66 49 3 20 3	1	1 1 1 2 56 1 1	1 1 1 1 2 9	26 8 1 6 189 70 1 68 3 220 2 2 2 3 1	2 1 1 1 1 1 1 1 2 39

¹ 62 recurrent cases.
² Reported in the Canal Zone only.

FOREIGN REPORTS

CANADA

Provinces—Communicable diseases—Week ended February 19, 1944.— During the week ended February 19, 1944, cases of certain communicable diseases were reported by the Dominion Bureau of Statistics of Canada as follows:

Disease	Prince Edward Island	Nova Scotia	New Bruns- wick	Que- bec	On- tario	Mani- toba	Sas- katch- ewan	Al- berta	British Colum- bia	Total
Chickenpox Diphtheria Encephalitis, infectious		9 7	6	215 22	343 1	58 2 1	50	113	281	1,069 38 1
German measles Influenza Measles	1	5 65 29	1	50 662	29 94 568	6 1 63	15 76	7 241	30 43 17	142 204 1, 657
Meningitis, meningococ- cus	1	29 12	1	1 83 71	4 172 191	79 57	8 22	2 55 87	3 56 83	10 483 524
Tuberculosis (all forms) Typhoid and paraty- phoid fever Undulant fever.		1	1	87	69 1 2	14		14	20 1	206 10 2
Whooping cough		7		58	64	14	2	11	25	181

JAMAICA

Notifiable diseases—4 weeks ended February 12, 1944.—During the 4 weeks ended February 12, 1944, cases of certain notifiable diseases were reported in Kingston, Jamaica, and in the island outside of Kingston, as follows:

Disease	Kingston	Other localities	Disease	Kingston	Other localities
Cerebrospinal meningitis Chickenpox Diphtheria Dysentery Erysipelas	7 2 3	2 35 3 2 1	Leprosy Poliomyelitis Puerperal sepsis Tuberculosis Typhoid fever	25 11	8 2 1 61 54

SWEDEN

Notifiable diseases—November 1943.—During the month of November 1943, cases of certain notifiable diseases were reported in Sweden as follows:

Disease	Cases	Disease	Cases
Cerebrospinal meningitis Diphtheria Dysentery Encephalitis, epidemic Gonorrhea Hepatitis, epidemic Paratyphoid fever	9 280 98 4 1, 773 943 14	Poliomyelitis Scarlet fever Syphilis Typhoid fever Undulant fever Weil's disease	246 3, 033 141 8 4

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

NOTE.—Except in cases of unusual incidence, only those places are included which had not previously reported any of the above-named diseases, except yellow fever, during the current year. All reports of yellow fever are published currently.

A table showing the accumulated figures for these diseases for the year to date is published in the PUBLIC HEALTH REPORTS for the last Friday of each month.

(Few reports are available from the invaded countries of Europe and other nations in war zones.)

Plague

Egypt—Suez.—During the week ended February 26, 1944, 3 cases of plague with 2 deaths were reported in Suez, Egypt.

Smallpox

Egypt—Suez.—During the week ended February 26, 1944, 18 cases of smallpox with 2 deaths were reported in Suez, Egypt.

India—Bombay.—Smallpox continues in epidemic form at Bombay, India. According to official reports for the week ended February 12, 1944, 223 new cases with 102 deaths occurred as compared with 192 cases and 55 deaths reported for the preceding week.

Indochina (French).—For the period January 21-31, 1944, 48 cases of smallpox were reported in French Indochina.

Sudan (French).—For the period January 11-20, 1944, 165 cases of smallpox with 14 deaths were reported in French Sudan.

Turkey.—For the month of December 1943, 1,488 cases of smallpox were reported in Turkey.

Typhus Fever

Bulgaria.—For the period January 6-19, 1944, 80 cases of typhus fever were reported in Bulgaria.

Guatemala.—For the month of January 1944, 155 cases of typhus fever with 27 deaths were reported in Guatemala.

Hungary.—For the 2 weeks ended February 19, 1944, 121 cases of typhus fever were reported in Hungary.

Netherlands.—For the 3 weeks ended January 22, 1944, 5 cases of typhus fever (including 3 cases in Amsterdam) were reported in the Netherlands. For the week ended January 29, 2 cases of typhus fever were reported in Amsterdam.

Union of South Africa—Cape Province.—Information dated March 6, 1944, states that official reports indicate a wide prevalence of typhus fever in Transkei region, Cape Province, practically all among the native population, where 282 cases were reported in one week. The principal outbreaks have occurred more than 300 miles from Port Elizabeth. Precautionary measures are being taken by the Government.

Yellow Fever

Brazil.—Deaths from yellow fever have been reported in Brazil as follows: Amazonas State, Benjamin Constant, December 21, 1943, 1 death; Matto Grosso State—Coronel Ponce, January 15, 1944, 1 death; Cuiaba, January 2, 1 death; January 19, 1 death.