PUBLIC HEALTH REPORTS

VOL. 52

MARCH 12, 1937

NO. 11

THE DISTRIBUTION OF BRUCELLA melitensis VARIETY melitensis IN THE UNITED STATES

By ALICE C. EVANS, Senior Bacteriologist, National Institute of Health

REVIEW OF THE LITERATURE

For many years after the recognition of Malta fever in Mediterranean countries, the United States was supposed to be free from that disease, now called brucellosis. The first series of cases in this country was found in Texas.

In 1911 Ferenbaugh reported that he had found 5 cases of Malta fever in Texas. All the patients had been in contact with goats. In the same year Gentry and Ferenbaugh published the results of a more extended investigation in which they diagnosed 7 human cases in addition to the 5 reported by Ferenbaugh. They found that 19.4 percent of the 128 goats which they tested were positive for *Brucella* agglutinins. Their investigations led them to believe that Malta fever had probably been endemic in Texas for at least 25 years.

The next year Yount and Looney reported 5 cases of Malta fever occurring in persons connected with the goat industry in Arizona; and in 1913 Yount reported another case, which was fatal. In that same year Wellman and Eustis diagnosed a case of Malta fever in Louisiana in a patient who had contracted the disease in Texas. They detected Malta fever in this patient by testing for Brucella agglutinins a group of serums which had been found to be negative to the Widal test. Among 46 serums tested, they found the one positive for Brucella agglutinins. It would seem that this one positive result might have suggested that brucellosis was not a rare disease and that further investigations might yield interesting data; nevertheless, no further significant investigations to find cases of brucellosis were carried out until a number of years later, when a definite outbreak occurred. In 1922 Lake, of the Public Health Service, diagnosed 35 cases of Malta fever in Phoenix, Ariz. He traced the source of infection to the milk of infected goats. From information obtained from local health authori-

124378°-37----1

ties and physicians he was convinced that the disease had been present in Arizona at least 14 years.

Thus, during the early part of the present century the recognized cases of brucellosis in this country were associated with goats in the Southwestern States (excepting one isolated case recognized by Craig in Washington, D. C., in 1905). It was believed that the human disease was limited to caprine sources in the Southwestern States.

Recently Meyer and Eddie confirmed the incidence of *Brucella* infections in goats of the Southwest. They studied strains isolated from goats received from Phoenix, Ariz., and Carlsbad, N. Mex., and found that they belonged to the *melitensis* variety.

As the years passed, reports were made of the occasional isolation of the *melitensis* variety in various other parts of the country. Tyndale and Viko reported that the State veterinarian of Utah died in 1923 as a result of handling placental tissue from an infected goat. He had gone to the southern part of the State to investigate the goats following the death of two herders believed to have been infected with *Brucella*. *Brucella* were isolated from the urine of the veterinarian, but the strain was lost without having been classified. The caprine origin of the infection, however, and the severity of the disease, suggest that the strain concerned in these three cases must have been of the *melitensis* variety. Meyer (1936) and his collaborators have worked quite extensively in southern Utah, and they found the goats of that region to be infected with the *melitensis* variety.

In 1925 the writer published the results of a serological classification of *Brucella* strains isolated from man and domestic animals. Many of the cultures were from distant areas in the United States. The collection included cultures of the *melitensis* variety isolated (by the investigators already mentioned) from human cases in Texas and Arizona, one culture from an aborted bovine fetus in Maryland, and another from an aborted equine fetus in Iowa. Later, Huddleson studied these strains in respect to bacteriostatic reactions and confirmed the classification in the *melitensis* variety of all excepting the equine strain, which proved to be atypical in that it agreed with the *abortus* variety in its sensitivity to dyes.

Later, in 1925, after the results of the classification study had been published, the writer received serum from a human case of brucellosis in Rocky Mount, N. C. From this serum the *melitensis* agglutinins were only partially removed by absorption with *abortus* antigen. The reaction indicated that the infection was with the *melitensis* variety. A few years later, Huddleson reported that two strains isolated from bovine sources in Michigan belonged to the *melitensis* variety; and a few years after that Carpenter and Boak reported that they had isolated the *melitensis* variety from cow's milk in three widely separated towns in New York State. These scattered observations suggested In European countries also, cattle have been found to be infected with the *melitensis* variety. Shaw, a member of the British Royal Commission on Mediterranean Fever, reported that he cultivated "Micrococcus melitensis" from the milk of cows in Malta. Descriptions of these strains by which they might have been identified with the melitensis or abortus variety were necessarily lacking. More recently, Taylor, Lisbonne, and Vidal cultivated Brucella organisms from cows' milk in France and identified them with the melitensis variety according to modern methods.

Taylor and his collaborators, and also Gilles, Pérès, and Gulty, have reported that, in the east of France, where human infections are derived from cattle, the *melitensis* variety is responsible for about 95 percent of human infections, although in the cattle of that region the *abortus* variety is the more common cause of infection.

The infection of cattle with the *melitensis* variety is a matter of concern not only on account of the greater number of human cases which an infected cow may cause, but also on account of the greater severity of the human disease caused by the *melitensis* variety. Molinelli made a comparative clinical study of brucellosis indigenous to two sections of Argentina. In the Andes region the human infections are caused by the *melitensis* variety, and the illnesses were found to be much more severe, the mortality higher, the disease lasting longer, and nervous symptoms more common than in the littoral, where the infections are due to the *abortus* variety of the organism.

THE AGGLUTININ ABSORPTION TEST FOR THE DETERMINATION OF THE PREVALENCE OF INFECTIONS WITH BRUCELLA MELITENSIS MELITENSIS

Incidental to the surveys regarding human cases of chronic brucellosis conducted by the United States Public Health Service in several sections of the United States, an opportunity presented itself to investigate the prevalence of *melitensis* infections in man in the surveyed areas by means of the agglutinin absorption test. The use of this test necessitates a discussion of its reliability in identifying the type of infection.

The most satisfactory method for determining the type of infection in any given case of brucellosis is to isolate the organism and study it. Cultures are not readily obtainable from every case, however, and that is particularly true in chronic cases. The agglutinin absorption technique offers another method of obtaining information as to the type of *Brucella* concerned in any given case. If two samples of serum from a brucellosis patient are saturated, the one with the homologous antigen and the other with an antigen of a heterologous variety, and then the remaining agglutinins in each sample are tested against both the homologous and the heterologous antigens, a reaction will occur with the homologous antigen in the sample which was absorbed with the heterologous antigen, but no reaction will occur in the sample absorbed with the homologous antigen. The agglutinin absorption test may be carried out on any serum which has a titer of 1 to 160 or higher.

The advantage of the agglutinin absorption test is that it is applicable to cases from which cultures can not be obtained. On the other hand, there are limitations to the information which may be gained by the use of the agglutinin absorption test. It does not distinguish the *abortus* from the *suis* variety, for they behave alike serologically; further, there is a small percentage of *Brucella* strains which are atypical, with irregular correlations between the serological reactions and other characteristics. Since the usefulness of the agglutinin absorption test in distinguishing the *melitensis* from the *abortus-suis* group depends on the rarity of atypical strains, a knowledge of their prevalence is necessary for correct interpretation of data obtained by absorbing agglutinins from patients' serum.

In 1925, when the writer published the results of the serological classification of 68 *Brucella* strains, there was no other known test by which they could be differentiated. Four years later, when Huddleson devised the bacteriostatic tests which have proved so useful for classifying *Brucella*, 44 strains which had been classified by the writer were sent, at his request, for his study of the correlation of the two systems of classification. He reported irregularities in two strains (4.5 percent). One of the irregular strains was a foreign strain of bovine origin; the other was the equine strain from Iowa which has been mentioned previously.

Recently Veazie and Meyer reported the results of their study of 447 Brucella strains isolated in various parts of the United States and in foreign countries. Among them were 20 strains too rough to be classified. Among the remaining 427 strains only 26, or 5.8 percent, were atypical, in that they failed to conform both serologically and culturally to either the melitensis or abortus-suis type. Only 11 of their irregular strains, all of bovine origin, were isolated in this country. Eight of these 11 strains had been isolated from different cows in a single dairy.

Investigators of other countries have reported results similar to the American findings. Olin and Lindström studied 103 strains, the majority of which were from human cases in Sweden. Two strains (1.9 percent) were atypical. Wilson studied over 300 *Brucella* strains. He divided them into two groups—the main group from many parts of the world, and a smaller group from the northeast, east, and southeast of France. In the main group of 165 strains, only 1 was atypical. In the group of 156 French strains, 41 were too rough to be classified, and a large percentage of the remaining strains gave atypical reactions. Ten subgroups were found. Thus the irregular strains of Wilson's collection were received from one geographical area, where the *Brucella* organism seems to be undergoing a transition.

Disagreeing with the results of other investigators is the report of Francis, who studied 23 strains and reported 10 of them (43.5 percent) irregular. His results cannot be explained on the ground that they came from some restricted locality where, as Wilson and also Veazie and Meyer have shown, atypical strains may be widely distributed. Francis' atypical strains were from 5 different localities in the United States, and one group of 3 bovine strains was from Germany. It is not clear why Francis, who studied a relatively few strains, should have obtained results so different from those of other investigators.

If American strains alone are considered, altogether 259 Brucella strains have been studied for correlation between serological and bacteriostatic reactions, with results as shown in table 1. Thus, among the American strains studied, 92.7 percent were typical strains, agreeing in classification when grouped according to bacteriostatic reactions or agglutinin absorption reactions. From this it may be concluded that neither test alone will classify individual strains as belonging to the *abortus-suis* or *melitensis* group with absolute certainty, but that collected agglutinin absorption data will give general information as to the types of infection prevalent in a given locality.

01	Number of	Atypical strains			
Ubserver	studied	Number	Percent		
Evans and Huddleson Francis Veazie and Meyer	29 19 211	1 7 11	3.4 36.8 5.2		
Total	259	19	7.3		

TABLE 1.—Record of atypical Brucella strains isolated in the United States

AGGLUTININ ABSORPTION TECHNIQUE

In the present study the following technique was used to obtain information concerning the variety of the infecting strain in serums of a titer of 1 to 160 or higher:

The cultures used as absorbing antigens were no. 456 of the *abortus* variety and no. 428 of the *melitensis* variety. They were grown on

1 percent glucose agar in Blake bottles. Each bottle was inoculated with the entire growth from 1 agar slant suspended in about 1.5 cc of saline solution. After 48 hours' incubation the growth was washed off with about 15 cc physiological saline solution containing 0.5 percent formalin, by rocking the bottle in the hands. After standing in the refrigerator a few days the dense bacterial suspension was centrifugalized, the clear supernatant fluid was discarded, and saline solution containing 0.5 percent formalin was added to restore the original volume. This suspension was then standardized to a density equivalent to 20,000 parts per million of the silica standard.

In the earlier study (1925) it was found that an antigen of a density of 60,000 p. p. m. would absorb all homologous agglutinins from a serum with a titer of 1 to 640 when the absorption was carried out in a 1 to 5 dilution of the serum; an antigen of half that density would absorb all homologous agglutinins from a serum with a titer half as high; and an antigen of twice that density was required to absorb all homologous agglutinins from a serum with a titer twice as high. In this study an antigen of a density of 60,000 p. p. m. was always used to absorb serums of a titer of 1 to 640; and the density of the antigen was reduced proportionately to absorb serums of lower titer. Serums of higher titer were diluted to a titer of 1 to 640, and the diluted serum was absorbed with an antigen of a density of 60,000 p. p. m. The procedure was as follows:

An equal quantity of glycerine is added to the serum in the field before sending it to the central laboratory. It was found that the test required 1.4 cc of the serum-glycerine mixture to give sufficient absorbed serum of a 1 to 5 dilution for the test. It requires 10.5 cc of the stock antigen to obtain an antigen of a density of 60,000 p. p. m. to absorb the serum in a 1 to 5 dilution. The tube containing 10.5 cc of the stock antigen was centrifugalized, 8.4 cc of clear supernatant fluid was removed, and 1.4 cc of the serum-glycerine mixture was added to the remaining 2.1 cc of antigen. The sediment was emulsified and the tube was then placed in a water bath at 37° C. After 4 hours it was removed to the refrigerator. The next day the tube was again centrifugalized and the agglutinin titer of the clear supernatant fluid was determined by testing with both the abortus and melitensis antigens. For this test 0.5 cc of antigen of a density equivalent to 500 p. p. m. of the silica standard was added to each of the series of tubes containing the diluted serum. Protocols, with the data for two serums, are given in table 2.

Serum	Rerum Treatment of serum				Abortus agglutinins serum diluted 1 to—						Melitensis agglutinins serum diluted 1 to—								
		10	20	40	80	160	320	640	1,280	2,580	10	20	40	80	160	320	640	1,280	2,560
Charlotte, 209.	Not absorbed. Absorbed with abortus. 1 Absorbed with melitensis.	2 0 0	8 0 0	8	8	8	0	0	 		2 4 0	88	4 3	80	80	2	0 		
San Anto- nio, 309.	Not absorbed Absorbed with abortus. Absorbed with melitensis. ³	4	4	4 0 4	40	402	4 0 0	4 0 0	400	8	4	4	400	4 0 0	400	400	4 0 0	40	0

TABLE 2.—Type of Brucella infection as determined by agglutinin absorption tests

1 Results indicate melitensis infection.

* Results indicate abortus infection.

RESULTS OF THE PRES	ENT	STUDY
---------------------	-----	-------

The brucellosis surveys were conducted especially for the purpose of finding chronic cases, which are likely to have serums with a titer of agglutinins too low for the absorption test. Occasionally, however, serums were received at the central laboratory with titers high enough for the test. Table 3 gives the results obtained with the serums from the three survey areas.

 TABLE 3.—Prevalence of Brucella melitensis variety melitensis in 3 survey areas, as indicated by agglutinin absorption tests with patients' serums

Locality	Number of	Abortus	infections	Melitensis	infections	
Locality	tested	Number Percent		Number	Percent	
Charlotte, N. C San Antonio, Tex Kansas City, Kans	7 10 10	2 4 8	28.6 40 80	5 6 2	71. 4 60 20	

Five out of 7, or 71.4 percent, of serums from cases in the Charlotte (N. C.) area; six out of 10, or 60 percent, of serums from cases in the San Antonio (Tex.) area; and 2 out of 10, or 20 percent, of serums from the cases in the Kansas City (Kans.) area gave results indicating infection with the *melitensis* variety. At any rate, the infecting strains in these cases were not of the *abortus-suis* type. They were either *melitensis* infections or infections with atypical strains. Judging from the review of literature, the great majority of these cases must have been infected with the *melitensis* variety. In all of these cases the infection had been contracted in the State where the study was made.

DISCUSSION

It was to be expected that a large percentage of the cases of brucellosis in Texas would be found to be infected with the *melitensis* variety. It was surprising, however, to find that the majority of human cases studied in the North Carolina area and a considerable percentage of the cases in the Kansas area were infected with the *melitensis* variety.

Since comparatively few goats are raised in the United States outside of the Southwestern States, the spread of human brucellosis infections with the *melitensis* variety must depend largely on the susceptibility of cattle to this infection. Hence, the reports of cattle infected with the *melitensis* variety are of great interest.

On account of the much higher virulence of the *melitensis* variety for man, we can expect that whenever it infects cattle in any community in this country our experience will be the same as that in the east of France—the proportion of human infections with the *melitensis* variety to infections with the *abortus* variety will be far greater than the proportion between the two varieties incident in cattle. As in Argentina, we in this country may also expect a greater proportion of severe cases in regions where the *melitensis* variety exists.

Cultural studies are being made on some of the chronic cases in two of the survey areas, and it is hoped that the observations reported here may be extended by the study of strains.

SUMMARY

Human infections with *Brucella melitensis* variety *melitensis* have long been known in southwestern United States. In the literature are found records of occasional human and bovine infections with the *melitensis* variety in various other sections of the United States.

A review is given of the reports in which the grouping of *Brucella* according to serological reactions is correlated with the grouping according to bacteriostatic reactions. Excepting in certain restricted localities, there is a low percentage of atypical strains in which the groupings according to the two systems do not agree. Of 259 American strains which have been studied by various investigators, only 19 (7.3 percent) were atypical. Hence, although agglutinin absorption tests will not classify an individual *Brucella* strain in the *abortus-suis* or *melitensis* group with absolute certainty, collected data will give information as to the types of infections in a given locality.

The results of this study indicate that the percentages of human infections with the *melitensis* variety in the 3 survey areas were as follows: In Charlotte, N. C., 5 out of 7 brucellosis cases, or 71.4 percent; in San Antonio, Tex., 6 out of 10 cases, or 60 percent; in Kansas City, Kans., 2 out of 10 cases, or 20 percent.

BIBLIOGRAPHY

- Carpenter, C. M., and Boak, R. A.: (1934) Brucella melitensis infection in cattle. J. Bact. 27: 73.
- Craig, Charles F.: (1906) The symptomatology and diagnosis of Malta fever,
- with the report of additional cases. Internat. Clin., 4: 89-115. Evans, Alice C.: (1925) Studies on Brucella melitensis. Hyg. Lab. Bull. No. 143: 1-26.
- Ferenbaugh, Thomas L.: (1911) Endemic Mediterranean fever (Malta fever) in Southwest Texas. J. Am. Med. Assoc., 57: 730-731.
- Francis, Edward: (1931) Agglutinin absorption in undulant fever. Pub. Health Rep., 46: 2416-2437.
- Gentry, Ernest R., and Ferenbaugh, Thomas L.: (1911) Endemic Malta (Mediterranean) fever in Texas. Further notes on its distribution and probable source with report of additional cases. J. Am. Med. Assoc., 57: 1045-1048.
- Gentry, Ernest R., and Ferenbaugh, Thomas L.: (1911) Malta fever in Texas. A report of the serum reaction of one hundred twenty-eight goats in Edwards County. J. Am. Med. Assoc., 57: 1127.
- Gilles, M., Pérès, G., and Gulty: (1932) Fièvre ondulante d'origine bovine à Brucella melitensis. Rev. gen. vet. med., 41: 476-480.
- Huddleson, I. Forrest: (1929) The differentiation of the species of the genus Tech. Bull. No. 100, Michigan State College. 16 pp. Brucella.
- Lake, G. C.: (1922) Malta fever in Southwestern United States. Pub. Health Rep., 37: 2895-2899.
- Meyer, K. F., and Eddie, B.: (1935) The problem of caprine brucella infections in the United States. J. Am. Vet. Med. Assoc., 86: 286-303. Meyer, K. F., Eddie, B., Veazie, L., Stevens, I. M., Stewart, B., and Geiger, J. C.:
- (1934) The heterogenous infection chains as occupational diseases. Bang's disease and Malta fever. Arch. f. Gewerbepath. u. Gewerbehyg., 5: 514-559.
- Meyer, K. F.: (1936) Personal communication.
- Molinelli, Ernesto A.: (1934) Estudio clinico de la fiebre ondulante en la Republica Argentina. El día médico, 7: 207-211.
- Olin, G., and Lindström, B.: (1934) Vergleichende Untersuchungen über verschiedene Brucellatypen, besonders solche schwedischen Ursprungs. f. Bakt. I. Orig., 131: 257-275. Shaw, E. A.: (1906) Mediterranean fever in goats, cows, and other animals. Zent.
- Rept. of Comm. on Medit. Fever, Part IV, pp. 16-26.
- Taylor, R. M., Vidal, L. F., and Roman, G.: (1934) Persistance de Brucella melitensis (variété caprine) chez les vaches naturellment infectées. C. r. Soc. Biol., 116: 132-134. Taylor, R. M., Lisbonne, M., and Vidal, L. F.: (1934) La fièvre ondulante en
- France à la lumière des recherches effectuées par le Centre de Recherches sur la fièvre ondulante 1931-1934. Presse méd., 1882-1883.
- Tyndale, W. R., and Viko, L. E.: (1923). A fatal case of Malta fever. J. Am. Med. Assoc., 81: 1953.
- Veazie, Lyle, and Meyer, K. F.: (1936). The serologic classification of the Brucella group. J. Inf. Dis., 58: 280-287.
 Wellman, Creighton, and Eustis, Allan: (1913). Malta fever in Louisiana. Am. J. Trop. Dis. and Prev. Med., 1: 393-396.
 Wilson, G. S.: (1933). The classification of the Brucella group. A systematic for the brucella group. J. Marca 2019; 516-544.
- study. J. Hyg., 33: 516-541. Yount, C. E.: (1913) Malta fever in the United States. Military Surgeon, 33: 540-545.
- Yount, C. E., and Looney, R. N.: (1912). Malta fever in Arizona, with a pre-liminary report of cases. Southern Cal Pract., 27: 257-261.

PULMONARY TUMORS IN MICE

II. The Influence of Heredity upon Lung Tumors Induced by the Subcutaneous Injection of a Lard-Dibenzanthracene Solution¹

By H. B. ANDERVONT, Biologist, United States Public Health Service

FREQUENCY OF OCCURRENCE

While primary lung tumors are not common in most species of animals, they are known to occur with exceptional frequency in mice. In fact, the prevalence of pulmonary growths in mice is one of the most striking features in the study of malignant tumors in this species. Livingood (12), in 1896, first described a primary lung tumor which arose in a bronchus of an albino mouse and which he diagnosed as an adenocarcinoma. Haaland (7) reported five instances of spontaneous lung tumors in mice, but Tyzzer (21, 22) was the first investigator to give these growths thorough consideration. He observed primary lung nodules in 12 mice ranging in age from 5% months to "very old". among which were 5 white, 3 gray, 2 black, 1 brown, and 1 black and white animals. A detailed and extensive description of each of these tumors was given, and the conclusion was reached that most of them "correspond to a single type, although there are minor variations." Mitosis was observed in only one case, but in two cases the growths extended into bronchi and were undoubtedly malignant. It was found difficult "to decide what name to apply to this type of tumor", but they were finally designated as "papillary cyst-adenoma." A total of 83 spontaneous tumors in 70 mice was found, of which 62 percent were primary lung growths, and the conclusion was reached that, in mice. primary tumors appear to be more frequent in the lungs than in any other organ.

Tyzzer described two types of growth, the first being the "papillary cyst-adenoma", in which the epithelium is arranged in a single layer upon irregular folds of supporting tissue and resembling in certain respects the structure of the lung. The tumor cells are either columnar or cuboidal and have no cilia. He could not decide whether the tumors arose from the epithelium of the bronchi or alveoli, and stated that "in most cases it resembles the bronchial epithelium, but it sometimes resembles the thickened alveolar epithelium." The paucity of mitotic figures was noted, which indicated that the growth rate of such tumors must be very slow. He concluded that some of the tumors "must be considered carcinomata because of the irregular growth of the epithelium." The second type of growth was designated as an "epidermoid carcinoma" of the lung with diffuse growth of epithelium showing a tendency to keratinization. This type of growth is much

¹ From the Office of Cancer Investigations, U. S. Public Health Service, Harvard Medical School, Boston, Mass.

rarer than the papillary cyst-adenoma type. Tyzzer also called attention to the fact that the lungs of mice frequently exhibit areas of chronic inflammatory hyperplasia.

In addition to his descriptions of these growths, to which little has been added by subsequent observers, Tyzzer also discovered the influence of heredity upon their development. Starting with a female mouse with a lung tumor and a male free from tumor, he obtained 62 progeny which reached maturity, and of these 27 percent developed primary lung nodules.

Jobling (10, 11) found primary lung growths to be next in frequency to mammary gland carcinomas in a series of 26 mice exhibiting 41 spontaneous tumors of which 29 arose in the mammary glands and 9 in the lungs. Haaland (8) described 353 primary tumors in mice and, while not giving complete statistics, states that "the adenomatous tumours of the lung vie with the mammary tumours in frequency." His studies led him to conclude that a large proportion of the lung growths are "undoubtedly malignant", but in some cases "their exact nature is uncertain", since many appeared to be hypertrophic changes instead of malignant growths. Nodules occurred only in the lungs of old mice and were often associated with chronic inflammatory processes, which were noted frequently in the lungs of normal mice.

Slye, Holmes, and Wells (17) were the next investigators to comment upon the spontaneous lung tumors of mice. After reviewing the earlier literature, they concluded that these growths are peculiar not only in frequency but also in structure, when contrasted with pulmonary tumors of other species. They observed 160 mice bearing lung nodules in the first 6,000 autopsied mice of Slve's stock: lung tumors constituted one-third of all tumors found in the 6,000 autopsies and were next in frequency to mammary gland tumors. Only those nodules "that seem fairly entitled to be classed as tumors" were included, and in accord with Tyzzer and Haaland they commented upon the "many nodules caused by inflammatory hyperplasia." The authors presented an interesting table of lung tumors in their mice in which the tumors were tabulated as to their growth characteristics. Of the 160 nodules, 20 were classed as "unquestionable carcinomas", 43 as showing "a reasonably sure malignant tendency", 41 as of "doubtful malignancy", and 56 as "benign". The types of growth ranged from those which exhibited active infiltration and regional metastases in the lungs to the "benign" nodules which they regarded as true tumors and not "inflammatory hyperplasia."

The nodules usually appeared in mice over 1 year of age, and sex apparently had little influence on their occurrence. In addition, Slye, Holmes, and Wells were the first to observe the presence of metastases outside the lungs. Four such cases were reported, two exhibiting secondary growths in the mediastinal lymph nodes and two in the mediastinal lymph nodes, chest wall, diaphragm, and kidney. So far as heredity is concerned, the statement was made that "Hereditary influences show a marked relation to the occurrence and character of these lung tumors." Hill (9) raised mice on varied beddings and diets. These factors had little influence on the origin of spontaneous tumors, but in 793 experimental and control mice he found 140 with spontaneous pulmonary growths.

Other investigators have noted the occurrence of primary lung tumors in mice, but it is believed that the reports referred to above are sufficient to establish the fact that in this species pulmonary growths appear with unusual frequency. It is necessary to remember that many of these observations were made in mice which had not been inbred and therefore may be regarded as presenting the incidence of lung tumors in laboratory mice. With the advent of studies pertaining to the effect of inbreeding on the occurrence of cancer in mice, it was only natural that investigators should concentrate on tumors which arose in sites easily accessible to macroscopic examination and the mammary glands were found to meet this requirement. However, the influence of heredity on pulmonary growths in mice has received attention from some investigators.

INFLUENCE OF HEREDITY

Lynch (13, 14, 15) has given this problem extensive consideration by recording the lung tumor incidence in two strains of mice. One of these, strain 1194, was inbred from the sixth to the fifteenth generation by brother to sister matings, "or rarely cousin by cousin", and consisted mostly of black agouti mice, although a few were pinkeyed or brown agouti. Lynch found a lung tumor rate of 6.7 percent in strain 1194 animals in a total of 208 mice that lived longer than 1 year. The youngest age at which a lung tumor was found at autopsy was 18 months. The other strain studied was Lynch's line of the Bagg albinos. In 135 individuals of this strain that lived to be at least 1 year old, Lynch recorded a lung tumor incidence of 37.04 percent. The youngest age at which a lung tumor was found at autopsy was 15 months. Crosses between these two strains produced 14 individuals of the first generation that lived 16 months or longer, and of these, 5, or 31 percent, developed lung tumors. Of the second outcross generation 49 mice lived to be more than 1 year of age, and of these, 11, or 22 percent, developed lung tumors. From these results Lynch suggested that lung tumor susceptibility is inherited as a dominant character.

Lynch also ascertained the incidence of lung nodules in the offspring of mice that had or did not have lung tumors. The results may be briefly summarized as follows: Lung tumors arose in 19 percent of the progeny of mice which were free from lung tumors when autopsied; in 40 percent of the progeny of parents, one of which had lung tumor; and in 48 percent of the progeny of parents, both of which had lung tumors. Since pulmonary growths appeared in the mice whose parents were both free of lung tumor, Lynch suggested that "tumor susceptibility is not only dominant but variable." Other studies led her to conclude that sex had but little influence on the occurrence of lung tumors and that the lung growths were found most frequently in mice coming to autopsy when 2 years of age or older.

Lynch has presented further evidence that lung tumor susceptibility is inherited. A new stock of albino mice, designated as strain D, is introduced which had a lung tumor rate of 34 percent in mice over 6 months of age. A male mouse of strain D was bred to 6 females of strain 1194 and the 45 progeny which lived more than 6 months showed a lung tumor incidence of 24.4 percent. Male mice of the first generation were backcrossed to females of the parent stocks. When backcrossed to females of strain D, the progeny, numbering 404 individuals, gave a lung tumor incidence of 32.2 percent; but when the males were backcrossed to females of strain 1194, the progeny, numbering 218 mice, gave a lung tumor rate of 7.3 percent. Thus it was again indicated that the tendency to develop pulmonary growths is inherited. In other reports Lynch (16) has presented evidence to show that tar-painting may be utilized to demonstrate the inheritance of this organ susceptibility.

A strain of highly inbred mice with a high incidence of spontaneous pulmonary growths has been described by investigators of the Roscoe B. Jackson Memorial Laboratory. These mice designated as strain A have, according to Strong (19, 20), descended from the Bagg albino Bittner (4, 5) states that 55 percent of males living 10 months strain. or longer develop pulmonary growths, and of breeding females which develop mammary gland tumors 36 percent also have lung nodules. In the course of a crossbreeding experiment between this stock and another which does not give rise to lung tumors he found primary lung growths in the hybrid animals. Recently Bittner (6) has published a thorough study of the lung tumor incidence in this interesting strain of mice. Of 123 breeding females coming to autopsy, 26.1 percent had primary lung tumors. Of 126 virgin females, 77 percent developed primary lung growths, the average age at autopsy being 16.6 months. Of 116 breeding males, 71.6 percent exhibited primary pulmonary tumors, the average age at autopsy being 14.8 months. Thus, of 242 virgin females and breeding males of this strain, 181, or 74.7 percent, developed lung tumors spontaneously.

EXPERIMENTAL

Mice of strain A have been used in this laboratory (1, 2) in investigations of the appearance of lung tumors following subcutaneous

injections of a lard solution of 1, 2, 5, 6-dibenzanthracene. It has been found (3) that this carcinogenic compound acts similarly to tar in eliciting lung tumors in mice and that mice of strain A are excellent test animals for such experiments. The idea suggested itself that a cross-breeding experiment between strain A mice and a strain known to exhibit a low incidence, if any, of spontaneous lung growths might be of interest. Mice of the C 57 black strain were chosen as suitable (18) It was decided to inject a lard-dibenzanthracene for the experiment. solution subcutaneously into most of the experimental animals in order to test for susceptibility to lung tumors. This procedure should also yield some information as regards the influence, the age, sex, or color of the progeny might have on their susceptibility to the carcinogenic action of the compound in both the lungs and subcutaneous tissues.

RESULTS OF CROSS-BREEDING

Young adult mice of the pure strains A and C 57 black were mated on July 15, 1935, as follows: 21 females of strain A to males of strain C 57 black and 21 females of strain C 57 black to strain A males. There were 179 black offspring (96 females and 83 males) born between August 15 and October 5, 1935.

On October 30, 1935, the females of the first hybrid generation were mated to their brothers and, as the result of this mating, 665 mice were obtained. The sex and color of the second hybrid generation are summarized below:

	Females	Males	Total
Black Albino Brown	175 86 53	205 86 60	380 172 113
Total	314	351	665

DIBENZANTHRACENE INJECTIONS

A sufficient quantity of a lard-dibenzanthracene solution was made up to last throughout the course of the experiment in order to obviate any difference in results which might be attributed to different solutions. Each cubic centimeter of lard contained 4 milligrams of 1, 2, 5, 6-dibenzanthracene; the procedure for preparing the solution has been described (1) elsewhere.

On November 1, 1935, all living females (38) of both strain A and C 57 black, along with 55 males and 4 females of the first hybrid generation, each received 0.2 cc of the lard-dibenzanthracene solution in the subcutaneous tissue of the right axillary region. The injection was repeated on November 15, 1935.

On January 22, 1936, 92 females of the first hybrid generation which had been used as breeders and 370 mice of the second hybrid generation were available. Of these, 62 mice of the first hybrid and 305 of the second hybrid generations were each given 0.2 cc of the larddibenzanthracene solution in the right axilla. Thirty of the first and 65 of the second hybrid generations and litter mates of the injected mice were set aside as normal controls. The experimental mice received another 0.2 cc injection on February 5, 1936. The color and sex of the injected mice of the second hybrid generation were divided as follows:

	Female	Male	Total
Black Albino Brown	58 50 34	73 51 39	131 101 73
Total	142	163	305

RESULTS IN PURE STRAIN ALBINOS AND C 57 BLACKS

Thirty-eight of these females received the first injection on November 1, 1935. The first tumor was found on February 12, 1936. Between the time of injection and the time of appearance of the first subcutaneous tumor, 15 of the strain A and 8 of the strain C 57 blacks had succumbed to an epidemic of *B. piliformis (23)*. Lung tumors were found in one of the strain A mice; none of the others had tumors when autopsied. Of the 10 remaining C 57 black mice, 9 developed subcutaneous tumors and 1 died without any evidence of tumor. None had macroscopic lung nodules. Of the five strain A mice, four developed both subcutaneous and lung tumors. One died on March 3, 1936, without a subcutaneous tumor but with multiple lung nodules.

RESULTS IN THE FIRST HYBRID GENERATION

These were black mice, 55 males and 4 females, which received an initial lard-dibenzanthracene injection on November 1, 1935. Ten of the males died tumor-free before February 2, 1936, when the first subcutaneous tumor was noted. Of the remaining 49 mice, 47 developed subcutaneous tumors and 42 developed lung tumors; 1 mouse died without a tumor.

There were 62 black mothers of the second hybrid generation which received their first lard-dibenzanthracene injection on January 22, 1936. Only one of these had died when the first subcutaneous tumor appeared on April 23, 1936. Of the remaining 61 animals, 54 developed subcutaneous tumors and 57 developed lung tumors. The last three mice were killed on November 25, 1936. None of these had a subcutaneous tumor, but the lungs of all three contained multiple lung growths. Up to July 23, 1936, 39 of this group developed subcutaneous tumors and had been autopsied; 35 of them exhibited multiple lung nodules. On July 23, 1936, 11 of the uninjected controls were killed and examined for the presence of macroscopic lung nodules; 10 were negative and 1 had a single lung nodule.

RESULTS IN THE SECOND HYBRID GENERATION

These animals received their first injection on January 22, 1936, and the first tumor arose on April 23, 1936, just three months later. Prior to April 23, 12 had died without any macroscopic tumor. The animals were examined once each week for the appearance of subcutaneous growths, and all were autopsied for the presence of lung tumors. Up to July 23, 1936, only 10 of these mice died from other causes; and of these, 5 had lung tumors only and 5 were negative in both subcutaneous tissue and lungs.

On July 23, 1936, there were 62 of these mice (17 males and 45 females) alive. All of them and 26 normal controls of the same hybrid generation were killed and autopsied. The findings in these 88 mice are summarized below:

	Injected mice	Control mice
No macroscopic tumor	8	25
Subcutaneous tumor only	1	0
Lung tumors only	43	1
Subcutaneous tumor and lung tumors	10	0

It is seen that 43 of the injected mice had lung tumors only, and it should be mentioned that of the 62 injected mice, 53 exhibited multiple lung tumors, while of the 26 controls only 1 had a single pulmonary growth.

A summary of the findings as regards the appearance of subcutaneous and lung tumors in all the first and second hybrid generations is presented in table 1.

 TABLE 1.—Subcutaneous and lung tumors in first and second hybrid generations following subcutaneous injections of a lard-dibenzanthracene solution

	First hybrid generation	Second hybrid generation	First generation controls killed on 7/23/36	Second generation controls killed on 7/23/36
Died or killed without tumor	12 10	26 60	10	25
Both subcutaneous tumors and lung tumors	8 91	48 171	1	1

An analysis of the results in the second hybrid generation has thus far failed to reveal any influence exerted by color, sex, or pedigree upon the occurrence of the induced lung growths.

THE INFLUENCE OF SEX ON THE APPEARANCE OF SUBCUTANEOUS TUMORS

As stated previously, the mice were examined each week for the presence of subcutaneous tumors; and as the tumors arose, the mice were placed in other cages for observation or were killed and autopsied. As the experiment progressed, this procedure drew attention to the fact that cages containing male mice of the second hybrid generation were emptied earlier than those in which the females were kept. At the conclusion of the experiment the time of appearance of the subcutaneous tumors in both the first and second hybrid generations was tabulated, according to sex. The findings are summarized in table Attention is directed to the first column of figures in the table, 2. which indicates the number of mice alive at the time the first subcutaneous tumor arose and not the total number of mice injected. In the table the numbers of new tumors discovered each week are listed according to the sex of the animal in which they arose. Just below these figures and in heavy face type is the weekly percentage of the total number of mice of each sex developing subcutaneous tumors. It is seen that the males of both generations responded to the carcinogenic agent by developing tumors earlier than did the females.

	Total or killed umber of without tumors subcutane- ous tumor	43 1220 1220 1320 133
	8- -	
\$		
37		
36		
34	ith t	
32	ice w	
31	E Jo	-8-6
8	Bgos	8-6-8-6
8	oent,	
36	l per	22 - 22 - 22 - 22 - 22 - 22 - 22 - 22
37	tota	2832381
3	q (þ	~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~
ส	rs en	
3	ouin	801937740 8019377940
8	lew t	8-355-3
19	a of n	80828 ⁷ 828
18	aber	~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~
11	Nun	**************************************
9	(B)	•844 <u>8</u> 844
15		N0004
1		0 0 0 0
	Number of injected mice alive when first tumor arose	45 (6) 65 (6) 66 (6) 147 (6) 146 (6) 146 (6)
	Bear	Male Female Male Female
Time in weeks.	Hybrid generation	First. Do

 $T_{\Delta BLB}$ 2.—Time in weeks of the appearance of dibenzanthracene tumors

SUMMARY AND CONCLUSIONS

In this experiment 21 females of strain A were bred to males of strain C 57 black, and 21 females of strain C 57 black were bred to males of strain A. The mice of strain A are known to be susceptible to both spontaneous and induced lung tumors, while the mice of strain C 57 black are known to be very resistant to the development of all spontaneous growths, and in this laboratory no lung tumors have thus far been induced in them by the subcutaneous injection of a larddibenzanthracene solution. The progeny of this mating has been designated as the first hybrid generation. Females of the first hybrid generation were mated to their brothers to procure mice designated as of the second hybrid generation.

Most of the first hybrid generation were injected subcutaneously with a lard-dibenzanthracene solution. Out of 121 of the injected animals, 11 died without tumor before the appearance of the first subcutaneous tumor at the injection site. Practically all the remaining 110 mice were kept under observation for the occurrence of subcutaneous or lung tumors; the last 3 were killed 11 months after the initial subcutaneous injection. Of the 110 animals, 101, or 91.9 percent, developed a subcutaneous tumor and 99, or 90 percent, developed lung nodules.

Of the second hybrid generation, 305 mice of black, white, or brown coat color were injected subcutaneously with a lard-dibenzanthracene solution. Twelve of these died tumor-free before the appearance of the first subcutaneous tumor. The remaining 293 mice were kept under observation for 6 months, during which time 231 developed a subcutaneous tumor or died from other causes. At the end of the 6month period the remaining 62 mice were killed and autopsied for subcutaneous and lung tumors. It was found that up to 6 months after the initial injection of these 293 mice, 231, or 78.8 percent, had developed a subcutaneous tumor, and 219, or 74.7 percent, had developed lung nodules.

The occurrence of all subcutaneous tumors in the two generations cannot be compared, because the first generation mice were kept for 11 months and the second generation mice kept for 6 months after the initial lard-dibenzanthracene injection. However, table 2 reveals that 88, or 80 percent, of the first generation had developed subcutaneous tumors 6 months after the first injection.

The first generation mice were hybrids from reciprocal crosses between a strain of mice in which the females exhibit a high incidence of spontaneous mammary tumors (strain A) and a line in which the females show a low incidence of mammary tumors (strain C 57 black). Investigators (18) at the Roscoe B. Jackson Memorial Laboratory have found that such reciprocal crosses reveal an extra-chromosomal influence in the occurrence of spontaneous mammary gland tumors, for hybrids derived from females which belong to high tumor lines develop more mammary tumors than those derived from females belonging to low tumor lines. The findings in the first hybrid generation mice of this experiment indicate that the progeny from either outcross were equally susceptible to the carcinogenic action of dibenzanthracene. This was to be expected, for previous investigations (1) have shown that both high and low spontaneous mammary tumor lines are susceptible to the development of induced dibenzanthracene tumors.

So far as the occurrence of lung tumors is concerned, 90 percent of the first hybrid generation mice and 74.7 percent of the second hybrid generation developed these growths. The fact that 62 of the second generation mice were killed 6 months after injection, and before a subcutaneous tumor was present in most of them, had little influence on the percentage developing lung nodules, for it will be recalled that 53 of the 62 mice exhibited lung nodules when examined at autopsy.

The absence of lung growths in a high proportion of uninjected controls of both the first and second hybrid generations shows that the lung nodules in the injected mice did not arise spontaneously and, in addition, shows that the lungs of the 62 second generation mice killed on July 23, 1936, were more responsive to the induction of tumors than the subcutaneous tissues, for 43 had multiple lung tumors without any evidence of a subcutaneous growth. This finding is similar to the results obtained in other experiments (3) with strain A mice.

The presence of lung tumors in a high percentage of animals of both the first and second hybrid generations shows that the susceptibility of this organ to tumor formation induced by the subcutaneous injections of a lard-dibenzanthracene solution is inherited in a dominant manner.

In both hybrid generations the carcinogenic compound produced subcutaneous tumors earlier in the male mice. It is essential to note that the females of the first hybrid generation were injected approximately 3 months after their brothers had received their initial injection. Furthermore, practically all the females had raised a litter before their injections began. Hence, the age at the time of injection or the influence of breeding may have been of some significance in the later appearance of tumors in the females of this generation. These factors, however, cannot be held responsible for the difference which occurred in the second hybrid generation, for all of these females were virgins and all were injected at the same time as their male litter mates. Previous experiments in this laboratory have not revealed any such difference in susceptibility between the sexes. The reason for the results obtained with these outcross animals remains obscure.

REFERENCES

- (1) Andervont, H. B.: Pub. Health Rep., 49: 620 (1934).
- 2) Andervont, H. B.: Pub. Health Rep., 50: 1211 (1935).
- (3) Andervont, H. B.: Pub. Health Rep., 52: 212-221.
- (4) Bittner, J. J.: Am. J. Cancer, 25: 113 (1935)
- (5) Bittner, J. J.: Proc. Soc. Exp. Biol. and Med., 34: 42 (1936).
- (6) Bittner, J. J.: Am. J. Cancer, 27: 519 (1936).
 (7) Haaland, M.: Ann. de l'Inst. Pasteur, 9: 165 (1905).
- (8) Haaland, M.: Fourth Scientific Report of the Imperial Cancer Research Fund, London, 1911.
- (9) Hill, L.: Lancet, 1: 966 (1931).
- 10) Jobling, J. W.: Proc. Soc. Exp. Biol. and Med., 6: 10 (1908). (11) Jobling, J. W.: Monograph Series, Rockefeller Institute for Medical Research, New York, 1910.
- (12) Livingood, L. E.: Johns Hopkins Hosp., Bull., 7: 177 (1896).
- (13) Lynch, C. J.: J. Exp. Med., 43: 339 (1926).
- (14) Lynch, C. J.: J. Exp. Med., 54: 747 (1931). (15) Lynch, C. J.: Am. J. Clin. Path., 6: 293 (1936).
- (16) Lynch, C. J.: J. Exp. Med., 46: 917 (1927).
- (17) Slye, M., Holmes, H. F., and Wells, H. G.: J. Med. Res., 30: 417 (1914). (18) Staff of the Roscoe B. Jackson Memorial Laboratory, C. C. Little, Di-
- rector: Am. J. Cancer, 27: 551 (1936). (19) Strong, L. C.: J. Hered., 25: 119 (1934). (20) Strong, L. C.: J. Hered., 27: 21 (1936).

- (21) Tyzzer, E. E.: Fourth Report of the Caroline Brewer Croft Fund Cancer Commission, Boston, Mass., 1907.
- Tyzzer, E. E.: J. Med. Res., 21: 479 (1909). (22)
- (23) Tyzzer, E. E.: J. Med. Res., 37: 307 (1917).

DEATHS DURING WEEK ENDED FEBRUARY 20, 1937

(From the Weekly Health Index, issued by the Bureau of the Census, Department of Commerce)

	Week ended Feb. 20, 1937	Correspond- ing week, 1936
Data from 85 large cities of the United States: Total deaths	10, 403 9, 475 74, 766 586 4, 504 69, 207, 100 16, 541 12. 5 11. 6	10, 445 67, 022 617 3, 965 67, 958, 356 14, 938 11, 5 10, 8

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

Cases of certain communicable diseases reported by telegraph by State health officers for weeks ended Feb. 27, 1937, and Feb. 29, 1938

	Diphtheria		Infi	Influenza Mer		Measles		gococcus ingitis
Division and State	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1966	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936
New England States:								
Maine	1	1	212	5		255	6	1 0
New Hempshire	·i			l i	78	13	i ă	i ă
Vermont	-		1	-		406	Ň	l X
Maggachmatte					768	016		12
Rhode Island	l i	Ĭ			201	43	2	
Connectiont	2	4	169	17	474	91	2	. 5
Middle Atlantic States:		1 ⁻	100				~	-
New York	31	51	1.45	178	430	2 636	11	97
New Jersey	ő	18	134	62	1 190	150		
Pennsylvania	45	41		-	219	797	ž	Å
East North Central States:								Ű
Ohio	38	35	447	127	99	421	13	12
Indiana	9	27	133	48	ii	40		- ī
Minois	37	89	162	42	36	28	7	18
Michigan	16	7	4	10	52	ü	2	
Wisconsin	ĩ	2	220	64	14	84	ī	3
West North Central States:	-	-					-	Ŭ
Minnesota	6	3	1	2	32	289	2	1
Iowa	2	15	8	6	2	- 4	ō	3
Missouri	19	19	944	650	8	20	8	12
North Dakota	1	5	7	12	2		ŏ	ō
South Dakota	ī	8	9	2			i	ž
Nebraska	5	9	30		13	29	ō	õ
Kansas	9	15		32	6	12	ŏ	å
South Atlantic States:								
Delaware	1		7		76	66	0	1
Maryland ¹	8	9	372	72	554	146	2	11
District of Columbia	10	22	28	2	75	25	1	7
Virginia	14	11			269	86	14	48
West Virginia	14	12	1, 252	218	1	21	10	9
North Carolina	25	16	173	482	64	55	8	8
South Carolina ¹		3	1, 346	1, 509	54	12	2	16
Georgia 3	8	6	1, 262	1, 819			8	9
Florida	9	8	35	33	5	6	0	2
East South Central States:								
Kentucky	13	16	493	80	243	73	17	42
Tennessee	11	14	844	338	10	52	10	0
Alabama 3	37	28	1, 546	2, 383	26	5	8	1
Mississippi 2	71	4		l			2	3

See footnotes at end of table.

Cases	of certain	. communica b	le diseases	reported b	y telegraph	by State	health	officers
	for	weeks ended F	Peb. 27, 193	87, and Feb	. 29, 1936–	-Continue	ed be	-

	Diphtheria		Infi	10128	Measles		Menin	Meningocoocus meningitis	
Division and State	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	
West South Central States:	 			140					
Louisiana. Okiahoma ⁴ Texas	11 12 7 29	15 10 15	980 389 974 3, 480	140 78 256 655	8 12 310	51 13 418	8 2 7 14	3 3 10 5	
Mountain States: Montana Idaho Wyogning	2 2	1	132 67 50	45 3	2 34 2	16 15	0	1	
Colorado New Mexico Arizona	6 4 5	4 9 5	167 269	8 304	6 62 239	9 18 37	1 0 2	21	
Pacific States: Washington Oregon	4	1 1 2	5 196	4 267	20 43 7	261 733	0 1	1	
California	21	33	1, 915	1,661	110	1,890	13	11	
First 8 weeks of year ⁴	4, 584	5, 218	209, 415	48, 179	37, 714	10, 390 54, 856	1, 247	1, 645	
Christian Francisco F	Polion	yelitis	Scarle	t fever	Sma	lipox	Typho	id fever	
Division and State	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	
New England States:									
New Hampshire Vermont Massachusetts Rhode Island Connecticut	00000	1 0 0 0 1	15 23 10 255 56 88	14 3 19 285 35 89	000000	000000000000000000000000000000000000000	0 0 1 0 2	0 1 3 0 1	
Middle Atlantic States: New York New Jersey Pennsylvania	0 0 0	1 1 1	953 174 561	1, 277 520 512	4 0 0	0 0 0	2 1 3	6 3 1	
East North Central States: Ohio Indiana Illinois Michigan Wicenstn	1 1 1 0	00000	493 216 582 771	491 344 969 297	7 8 40 3	0 1 6 0	3 0 4 1	5 8 2 1	
West North Central States: Minnesota Iowa Missouri North Dakota South Dakota Nebraska	020000	000000000000000000000000000000000000000	169 351 292 47 73 106	372 106 219 124 62 238	2 35 46 22 5 0	3 20 17 17 31 23	1 0 1 1 0	1 3 2 0 1	
Kansas South Atlantic States: Delaware Maryland ¹ District of Columbia Virginia West Virginia North Carolina South Carolina ¹	0 0 0 0 0 1 0	0 0 1 0 1 0	378 4 51 21 35 55 33 6	325 9 98 30 57 44 34 5	22 0 0 0 0 0 0 0	47 0 0 0 0 1 0 0	1 1 1 3 8 3	1 8 1 2 1 0 0	
Georgia ³ Florida	02	Ŏ	14	24 4	Ō	1	2	1 6	

See footnotes at end of table.

March 12, 1937

318

	Polion	nyelitis	Scarle	t fever	Sma	llpox	Typhoid fever	
Division and State	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936	Week ended Feb. 27, 1937	Week ended Feb. 29, 1936
East South Central States:								
Kentucky	0	1	65	76	0	0	2	(
Tennessee	0	0	18	24	0	0	4	
Alabama		a a a a a a a a a a a a a a a a a a a		30	0	0		
Mississippi	0	0	12	14	0	U	8	1 1
west South Central States:								
Arkausas	l i			10	ő	2		
Oblehome 4		i i		20	6	3		
Tores	5	1	77	26	2	Š		
Mountain States	-				-	v	20	
Montene		•	47	127	20		0	1
Ideho	Ň	ň	25	107	1	4	ŏ	1
Wyoming	ň	ŏ	36	127	3	i	ŏ	ៅ
Colorado	ŏ	ŏ	45	154	ĭ	ŝ	ŏ	
New Mexico	ň	ŏ	35	112	ô	ŏ	ă	
Arizona	ŏ	ŏ	16	34	ŏ	ž	ĭ	2
Utah ?	ŏ	ŏ	23	143	ŏ	ī	ō	ō
Pacific States:		-			Ť	-	-	
Washington	0	0	63	81	5	13	2	1
Oregon	1	ī	23	34	24	1	Ō	ō
California	Ō	8	219	410	9	ī	i	4
Total	16	17	6, 969	8, 777	283	233	82	88
First 8 weeks of year *	181	141	50, 571	60, 128	2, 364	1, 688	885	786

Cases of certain communicable diseases reported by telegraph by State health officers for weeks ended Feb. 27, 1937, and Feb. 29, 1936—Continued

New York City only.
 Week ended earlier than Saturday.
 Typhus fever, week ended Feb. 27, 1937, 16 cases, as follows: South Carolina, 3; Georgia, 9; Alabama, 4.
 Exclusive of Oklahoma City and Tulsa.
 Figures for 8 weeks ended Feb. 27, 1937, include delayed reports.

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- cus menin- gitis	Diph- theria	Influ- enza	Mala- ria	Mea- sles	Pel- lagra	Polio- mye- litis	Scarlet føver	Small- pox	Ty- phoid fever
January 1857 Arizona California Kansas Mississippi Montana North Dakota North Dakota North Dakota Vermont Wisconsin	10 80 8 9 6 8 7	16 147 48 81 12 8 8 8	3, 333 19, 447 14, 943 13, 872 10, 600 956 43 215 6, 110	8 8 1, 605	602 \$56 35 986 10 139 603 18 84	 1 144 	1 14 2 1 0 0 0 0 2	94 1, 375 1, 346 48 921 960 231 39 1, 205	0 40 120 61 145 0 86	3 25 3 21 5 1 1 8 4

319

Summary of monthly reports from States-Continued

January 1937		January 1937-Continue	d	January 1937-Continue	bđ
-	Cases		Cases		Cases
Chicken pox:		Hookworm disease:		Septic sore throat-Contd.	
Arizona	263	Mississippi	844	Rhode Island	1
California	8. 415	Impetigo contagiosa:		Wisconsin	18
Kanaas	466	Kansas	1	Tetanus:	
Mississippi	791	Montana		California	2
Montana	236	Jaundice, enidemic:	-	Rhode Island	ĭ
North Dakota	163	California	2	Trachoma:	-
Rhode Jelend	238	Kanees	ĩ	Arizona	21
Vermont	453	TANKORY'	-	California	8
Wienonein	2 282	California	2	Mississinni	ă
Toroma.	a, 00a	Mumpe	-	Montene	1
Miadariani	1	A sigona	119	Thereamie	•
Desenter:	-	Callornia	9 663	California	1
Dysencery.		Kanasa	720	Vanana	;
Arizous		Mindasimmi	109	Wieconsin	1
California (amoebic)	10	Mastana	040	W BOULISIII	1
Camornia (Dacmary)	19	Mioneth Dabata	200	Typuus lever.	
Kansas (Dacinary)		North Dakota	201		1
Mississippi (amoedic)	.09		12	Undulant lever:	•
Mississippi (bacillary) -	179	Vermont	134		ž
Montana (amoebic)	1	Wisconsin	730	California	5
Encephalitis, epidemic or		Ophthaimia neonatorum:		Kansas	0
lethargic:		California	2	Mississippi	1
California	4	Mississippi	5	Vermont	- 4
Montana	2	Paratyphoid fever:		Wisconsin	1
North Dakota	1	California	1	Vincent's infection:	
Wisconsin	1	Puerperal septicemia:		Kansas	5
Food poisoning:		Mississippi	20	North Dakota	5
California	41	Rabies in animals:		Whooping cough:	
German measles:		California	106	Arizona	26
Arizona	21	Mississippi	30	California	1, 278
California	89	Scables:		Kansas	107
Kansas	6	Kansas	4	Mississippi	269
Montana	13	Montana	2	Montana	19
Rhode Island	1	Septic sore throat:		North Dakota	4
Vermont	12	Arizona	1	Rhode Island	132
Wisconsin	3	California	18	Vermont	115
Granuloma, coccidioidal:	-	Kansas	3	Wisconsin	439
California	4	Montana	17		
~ withus the					

WEEKLY REPORTS FROM CITIES

City reports for week ended Feb. 20, 1937

This table summarizes the reports received weekly from a selected list of 140 cities for the purpose of showing a cross section of the current urban incidence of the communicable diseases listed in the table. Weekly reports are received from about 700 cities, from which the data are tabulated and filed for reference.

Otata and aller	Diph-	Inf	luenza	Mea-	Mea- Pneu-		Small-	Tuber-	Ty-	Whooping	Deaths,
State and city	Cases	Cases	Deaths	Cases	deaths	fever cases	Cases	deaths	fever cases	cough cases	all Causes
Maine:		9			•	1			•		
New Hempshire	U U		v	v		-	ľ	1 1	v	°	
Concord	6		3	0	4	2	0	6	0	6	14
Manchester	ŏ		Ă	ŏ	7	ō	ŏ	ŏ	ŏ	ŏ	20
Nashua	ĬŎ			Ŏ	7	Ō	Ō		Ŏ	Ŏ	
Vermont:	-			-	-	-	-		-		
Barre	0		1	0	0	1	0	1	0	6	2
Burlington	0		0	0	0	6	0	0	0	0	7
Rutland	0		0	0	3	1	. 0	0	0	0	11
Massachusetts:											
Boston	8		5	10	60	52	0	9	1	146	812
Fall River	2		0	10	8	3	0	1	0	2	43
Springfield	0		0	27	2	1	0	2	0	10	43
Worcester	0		0	187	11	6	0	3	0	36	66
Rhode Island:											
Pawtucket											
Providence	U	14	1	203	13		U	4	U	10	80
Connecticut:				42		10					40
Bridgeport	Ň		ź	50	10	10	Ň		Ň		90
Maruoru		32	, v	4	14	10			Ň		01
New Haven	-	- 11	-	• •	•			۷ V			
New York:											
Buffalo	0		2	42	20	25	11	7	0	40	201
New York	48	74	14	151	168	457	ō	9 a l	Ă	112	1.617
Rochester	õ		8	- õ	- 8	- 8	ő	Ĩ	ő	-11	94
Syracuse	ŏ		i	32	15	70	ŏ	il	ŏ	22	71

	Diph	_ Inf	luenza	Mea-	Pneu-	Scar-	Small-	Tuber-	Ty-	Whoop-	Deaths,
State and city	theris cases	Com	Deaths	sles cases	deaths	fever	pox cases	culosis deaths	fever	cough	all
		Cases	Deatins			Cases			Cases	Cases	
New Jersey:											
Camden Newark	1	;-	2	480	4	3	0	0	0	13	35
Trenton	ŏ	6	2	2	6	6	Ŏ	4	ŏ	ĩõ	54
Philadelphia	7	35	20	12	60	291	0	26	0	101	630
Pittsburgh Reading	2	14			29 5	50 3		10	0	36 22	178
Scranton	ĩ			Õ		19	Ŏ		Ŏ	3	
Ohio:		.								_	
Cleveland	32	149	11	33	18 34	21 76	Ö	10	0	7 57	183 240
Columbus Toledo	2	119	13	35	12	11		10	8 8	0 35	128
Indiana:						10					10
Fort Wayne	ŏ		Ō	ő	6	12	Ő	5	ŏ	2	10 28
South Bend	0		5 1	10	28	45 6	0	5		22	119 26
Terre Haute	1		0	Ó	Ó	3	Ō	Ō	Ō	Ō	38
Alton	0		0	0	1	4	0	1	0	0	10
Elgin	13 0	29	4	14	82	243	Ő	33	2	83 13	800 10
Moline Springfield	0		0	0	3	0	0	2	0	12	12
Michigan:			-		20	-	ů	10	Ň		
Flint	1		1	• 	3	22	ŏ	10	ŏ	3	32
Wisconsin:	0		1	8	2	15	0	1	0	17	48
Kenosha Madison	0		0	0	0	4	0	1	0	3	4
Milwaukee	ŏ	3	3	2	15	72	ŏ	4	ŏ	21	138
Superior	Ö		1	0	1	3	ő	ő	8	0 5	12 14
Minnesota:											
Duluth	0		2	0	20	23	0	0	<u>o</u>	3	30
St. Paul	1		ŏ	î	7	15	ŏ	2	ŏ	42	61
Cedar Rapids	0			0		2	0		0	0	
Davenport Des Moines	0			8		3 41	0		0	2 -	30
Sioux City	ō			Ŏ		27	ŏ.		ŏ	2 -	
Missouri:	, i			Ů		20			"		
Kansas City St. Joseph	1	7	4	4	67	84 10	24	12	8	4	101 56
St. Louis	9		- 4	i	18	50	2	5	ŏ	69	228
Fargo	0		0	1	0	8	0	0	0	0	3
Minot	ő		<u>0</u>	0	0	ő	1	0	Ö	8-	9
South Dakota:	0			0	1	5	0		0		
Nebraska:											
Kansas:					11	•				•	8/
Lawrence Topeka	0		0		2 11	0 12	0	8	8	8	5 27
Wichita	0	2	2	2	7	10	Ó	1	Ō	ŏ	33
Delaware:											
Maryland:			Z	32	8	•	٥	Ů	"	2	40
Baltimore Cumberland	8	35 1	4	362	51 3	17	8	15	0	63	273 14
Frederick	ŏ.		ō	ŏ	2	ŏ	ŏ	ŏ	ŏ	ŏ	4
Washington	5	27	10	63	34	* 23	0	14	1	18	220
Lynchburg	1		1	3	3	0	0	2	0	2	15
Norfolk	0	23	0 R	0	9	5	<u>8</u>	0	8	Õ	34
Roanoke	ĭ٢		ŏl	72	3	il	ŏI	2	ŏl	6	19

City reports for week ended Feb. 20, 1937-Continued

T

-

T

-

City reports	fo r week	ended	Feb. 20,	1937Continued	
• •	•		•		

State and city	Diph-	Inf	luenza	Mea-	Pneu- moria	Scar- let	Scar- let pox		Ty- phoid	Whoop- ing	Deaths,
	Cases	Cases	Deaths	08.965	deaths	Dever Crises	Cases	deaths	lever Cases	cases	CBUSES
West Virginia: Charleston Huntingdon Wheeling	0	9	0	0	8	1 2	0	1	0	0	25
North Carolina: Gastonia	0	1	0	0	0	0	0	0	0	0	01
Wilmington Winston-Salem.	1 0	1	Ŏ	0 1	1 5	08	Ő	12	0 0	0 1	21 7 20
Columbia Florence	1 0 0	174	400	000	448	2 0 0	0000	0 1 0	2 0 0	0 0 0	31 26 17
Georgia: Atlanta Brunswick	1	443 1	9	0	12 0	1	0	80	0	2	98 5
Florida: Miami Tampa	1	120 12 2	1	2	1	2	0	2	0	• • 1 0	40 89 22
Kentucky: Ashland	o	30	0	0	8	0	0	ņ	0	0	11
Lexington Tennessee:	0	4	4	12	7	1	Ö	2	0	ů	18 26
Memphis Nashville	000		20 4	3 1 0	29 7	0 8	0	7 8	0	34 4	37 146 65
Birmingham Mobile Montgomery	2 0 1	201 15 6	9 5	0 0 0	7 9 	5 0 1	0 0	4 3	0 1 0	5 0 0	75 39
Arkansas: Fort Smith Little Rock	0		i	0	14	4 1	0	8	0	0	25
Lake Charles New Orleans Shreveport	1 9 0	75	0 10 2	0 0 0	2 35 10	0 2 0	0000	0 12 2	0 0 1	0 0 0	192 43
Oklahoma: Muskogee Tulsa	0			0		2 5	0		0	0	
Texas: Dallas Fort Worth	4	24	24 3	8 78	19 10	13 1	0	3	0	11	115 • 54
Houston San Antonio	2 0	•	8 15	0 7	26 17	1 0	0	4 9	1 0	0 1	33 98 100
Montana: Billings Great Falls	0		03	0	1	1	0	0	0	0	7 11
Helena Missoula Idaho:	0		0	17 0	8 2 1	18	0	0	0	0	58
Colorado: O o l o r a d o Springs	0		1	0	2	12	0	1	0	0	10
Denver Pueblo New Mexico:	Ŏ 1		8 0	6 1	7 2	11 1	Ŏ	8 0	Ŭ O	45 0	93 7
Albuquerque Utah: Salt Lake City.	0 2	38	0	0 8	0 6	6 10	0	3 0	0	9 8	9 41
Nevada: Reno											
Washington: Seattle Spokane Tacoma	0 0 1	4	4 4 2	1 0 0	12 8 4	5 3 6	0 0 0	5 0 0	0 0 0	3 4 0	111 45 43
Oregon: Portland Oalifornia:	0	6	4	2	8	7	8	0	0	2	85
Los Angeles Sacramento San Francisco	18 1 8	155 134 87	18 3 10	24 1 1	75 7 14	85 5 81	0 0 0	29 3 9	0 0 0	65 1 20	490 41 210

322

State and city	Meningococcus meningitis		Polio- mye-	State and city	Menin men	gococcus ingitis	Polio- mye- litis
	Cases	Deaths	Cases		Cases	Deaths	Cases
Massachusetts: Boston	8 1 9 1 1 1 2 2 1 2	2 0 5 0 1 1 0 1 0	0 0 0 0 0 0 0 0 0 0 0 0 0	Maryland: Baltimore District of Columbia: Washington Georgia: Atlanta Kentucky: Ashland Louisville Alabama: Birmingham Alabama: Birmingham Alabama: Birmingham Alabama: Birmingham Galveston Galveston	4 2 1 1 4 1 0 1 2 1	2 1 0 5 0 2 0 1 0	0 0 0 0 0 0 0 0 0
Detroit Flint Minnesota:	1 1	0 1	0	Houston Montana: Billings	3 1	Ŭ 0	ŏ
Minneapolis St. Paul Missouri:	1	1 0	000	Colorado: Denver California:	2	1	0
Kansas: Wichita	2	1	0	Los Angeles Sacramento San Francisco	2 1 2	5 0 0	1 0 0

City reports for week ended Feb. 20, 1937-Continued

Encephalitis, epidemic or lethargic.—Cases: New York, 1; Philadelphia, 1; Flint, 1; Washington, D. C., 1; Louisville, 1. Pellagra.—Cases: Baltimore, 1; Winston-Salem, 2; Charleston, S. C., 1; Savannah, 2; Birmingham, 1; Mobile, 1; Los Angeles, 1; San Francisco, 1. Rables in man.—Deaths: Atlanta, 1. Typhus fever.—Cases: New York, 2; Savannah, 3; Miami, 2.

FOREIGN AND INSULAR

AUSTRIA

Vital statistics-1935.—The following table shows the number of marriages, births, and deaths in Austria for the year 1935:

Population	6, 760, 631	Deaths from—Continued:	
Marriages	45, 375	Influenza	1, 404
Births	91, 111	Malaria	6
Total deaths	92, 390	Measles	163
Deaths from:		Scarlet fever	103
Accidents	2, 313	Sepsis	913
Cancer and other malignant tumors	11,854	Suicide	2.506
Congenital debility	1, 807	Syphilis	406
Diabetes	783	Tuberculosis	7.343
Diphtheria	992	Typhoid fever and paratyphoid faver.	121
Heart disease	15, 365	Whooping cough	209

CUBA

Habana—Communicable diseases—4 weeks ended February 13, 1937.—During the 4 weeks ended February 13, 1937, certain communicable diseases were reported in Habana, Cuba, as follows:

Disease	Cases	Deaths	Disease	Cases	Deaths
Diphtheria Leprosy Malaria Measles	17 1 184	1 	Pcliomyelitis Scarlet fever Tuberculosis Typhoid fever	¹ 2 1 14 140	34

¹ Includes imported cases.

GERMANY

Vital statistics—Third quarter 1936.—Following are vital statistics for Germany for the third quarter of 1936:

and the second se	Number of marriages.	147, 055	Number of deaths	172, 394
	Number of marriages per 1,000 population	8. 7	Number of deaths per 1,000 population	10. 2
	Number of births	812, 162	Deaths under 1 year of age	18, 824
	Number of births per 1,000 population	18. 5	Deaths under 1 year of age per 100 live births.	5. 9
1	Number of stillbirths	7, 739		0.0

ITALY

Vital statistics—1936.—Following are vital statistics for Italy for the year 1936:

 Number of marriages
 810, 822 | Number of live births per 1,000 population.
 22. 2

 Number of marriages per 1,000 population.
 7. 2 | Total deaths.
 582, 612

 Number of live births
 955, 189 | Deaths per 1,000 population.
 13. 5

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

NOTE.—A table giving current information of the world prevalence of quarantinable diseases appeared in the PUBLIC HEALTH REPORTS for February 26, 1937, pages 255-267. A similar cumulative table will appear in the PUBLIC HEALTH REPORTS to be issued March 26, 1937, and thereafter, at least for the time being, in the issue published on the last Friday of each month.

Cholera

India—Bassein.—During the week ended February 20, 1937, 4 cases of cholera were reported in Bassein, India.

Plague

Hawaii Territory—Island of Hawaii—Hamakua District—Paauhau Sector.—A rat found March 1, 1937, in Paauhau Sector, Hamakua District, Island of Hawaii, Hawaii Territory, has been proved plague infected.

Smallpox

Siam—Tak Province.—During the week ended February 20, 1937, 26 cases of smallpox were reported in Tak Province, Siam.

Typhus Fever

Trans-Jordan-Kerak District.-During the week ended February 20, 1937, 3 cases of typhus fever were reported in Kerak District, Trans-Jordan.

Yellow Fever

Gold Coast—Accra.—On February 10, 1937, 2 cases of yellow fever were reported at Accra, Gold Coast.

Ivory Coast.—During the week ended February 20, 1937, yellow fever was reported in Ivory Coast as follows: 3 cases in Adzope, Agneby Circle, and 1 suspected case in Nzimcomoe Circle.