THESIS ON TRYPANOSOMIASIS

By

Alfred C.Sharp, M.B. Ch.B. (Glasg.)

M.D. Degree.

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THESIS ON TRYPANOSOMIASIS.

During the military operations in East Africa in 1918, I had the opportunity of observing cases of trypanosomiasis. Three Europeans were under my care. Owing to the courtesy of Captains Hume. Manson and Brierley. R.A.M.C., I had facilities for observing two Europeans and 14 Africans. These were the only cases reported during the campaign. All the cases, but one, were probably infected with the same type of trypanosome, and were treated on similar lines. From this, it seems possible to draw some conclusions, both regarding the symptomatology and treatment of this deadly disease.

Definition.

tially of Africa, and is the term applied to the condition produced by invasion of the blood of human beings with parasites known as trypanosomes. At first there is an irregular fever and trypanosomes are present in the blood.

Sleeping sickness is the terminal phase of trypano soma infection, when the trypano somes are found in the cerebro-spinal fluid and the chief symptoms are due to the involvement of the nervous system. Great apathy and listlessness with finally a comatose state are features of this stage.

History.

Although its true aetiology was not apprehended until quite recently. sleeping sickness has been known to exist for nearly two centuries. According to Castellani,

the earliest mention of sleeping sockness so far discovered is by John Atkins, in his little book, entitled "The Navy Surgeon" published in 1734, in the appendix to which he describes "the sleeping distemper", common among the negroes on the Guinez Coast, among whom he had travelled in 1731. It was well known to the slave dealers who, on purchasing slaves, always examined the glands at the back of the neck and any slaves whose glands were enlarged were rejected. The first account given of this symptom is by Winterbottom in 1803, who brought out the importance of enlargements of the posterior cervical glands (Winterbottom's sign); the cases he described occured at Sierra Leone.

The occurrence of trypano somes was first found in the blood of animals. In 1879 Lewis, in Calcutta, described the trypano soma lewisi in the rat. In the following year. Evans described a similar parasite.

T.evansi. in the blood of horses in India affected with "surra". This disease the natives had always ascribed to the bite of certain bloodsucking flies; He also found that it infested camels, elephants, buffaloes and dogs.

Fifteen years later, Bruce discovered that "nagana", a fatal disease of cattle in Africa, was due to a trypanosome. T.brucei.

In 1890 Nepveu found trypano somes in the blood of a man in Algeria, but he established no definite relationship between the organisms and the associated morbid conditions, and the discovery did not attract attention.

In 1901 Forde found a parasite in the blood of a European suffering from an irregular non-malarial fever.

in the River Gambia Colony, and in 1902 Dutton recognised the parasite as a trypanosome. Later, Dutton found a similar organism in the blood of a native of the same colony and suggested the name T.gambiense, which the parasite now bears. Subsequently, many cases were described, both in Europeans and natives, and the association of the parasite with a form of irregular fever was established.

In 1902 Castellani discovered trypanosomes in the blood of cases of sleeping sickness and brought about the establishment of the connection between trypanosomiasis and sleeping sickness. Bruce, Nabarro and other investigators confirmed Castellani's findings.

In 1903 Bruce and Nabarro reported that in Uganda the tse-tse fly. Glossina palpalis, was the transmitter of the infection. This conclusion was also reached by Sambon and Brumpt on epidemiological grounds. In 1909 Kleine discovered the tse-tse fly was no mere mechanical transmitter of the trypanosomes, as had been supposed but was a true intermediate host. After feeding G.palpalis on animals infected with T.brucei, he set the same flies to bite fresh animals at varying intervals. Up to sixteen days the flies failed to convey the infection, but from that period onwards to the fortyseventh day, when the experiment concluded, they communicated the trypano some to eight animals. Bruce and others have confirmed Kleine's results and have ascertained that they apply to T.gambiense and other trypano somes.

In 1910 in Rhodesia, Stephens and Fantham established the existence of a more virulent trypano some. T.rhodesiense. They found that it was associated with a

resistant form of the disease, that animals immunized against T. gambiense succumbed to the Rhodesian trypanosome. Later, Kinghorn and Yorke proved that T. rhodesiense was transmitted by Glossina morsitans.

In 1909 Chagas described a form of trypanosomiasis, especially common in children, which is endemic in parts of the state of Minas Geraes in Brazil, and is transmitted to man and the domestic animals by a species of bug, Lamus megistus. The symptoms of this disease are different from the African trypanosomiasis; an acute type usually occurs in children under one year of age, whilst the type of the disease as seen in adults is mainly chronic. (Stitt.)

Geographical Distribution.

The disease is essentially African. According to Castellani, at times, cases were imported to the West Indies from West Africa but the disease soon died out.

No fresh cases occurred as there were no tse-tse flies...

It exists on the west coast of Africa. from Senegal to Mossamedes. It is also present in the Niger and Congo basins, and particularly in Uganda, where it was first noticed in 1900. The more virulent form occurs in Rhodesia.

As regards the East African campaign, the chief areas to be considered are. German East Africa and Portuguese East Africa. In 1902 cases were found in German East Africa. From the writings of German doctors it would appear that the infected areas in German East Africa, at present, are - the southern border of Lake Victoria Nyanza, the vicinity of Lake Tanganyika, and the

neighbourhood of the Rovuma river (which forms the boundary between the German and Portuguese colonies) from Sassawara on the west to Liwale on the east.

Dr Marshall wrote in 1908 that there were two foci of sleeping sickness on Lake Victoria and one on ...
Tanganyika. The disease has been discovered on the islands, Bumbide and Iroba, and on the mainland Ihangiro, opposite to them. Cases of the disease have also been found on Ukerewe Island. Glossina palpalis was found widely distributed on the shore of lake Victoria, and on the rivers in the Shirati district, and sleeping sickness was prevalent there. (The disease was brought along trade routes from West Africa, through the territories watered by the Congo and its tributaries).

In 1910 Wolf met with a case of trypanosomiasis, which came from the Rovuma river area, near its junction with the Sassawara river. It was found that the Glossina morsitans was the transmitter of the Rovuma sleeping sickness and that the trypanosome was identical with T. rhodesiense.

Professor Beck, as a result of his investigations, stated that 10% of the Glossina morsitans in the Rovuma area were infected with trypanosomes.

Previous to the war, no investigation had been made as regards infected areas in Portuguese East Africa, although the Germans considered it most probable that such existed.

In the course of military operations in Portuguese territory, a line of communication was opened up, running from Bandari (on the mainland opposite Port Amelia) through Mahiba (14 miles), Anquabe (44 miles), Medo and

Balama to Lucinje. From Bandari to Anquabe, the road runs through fairly thick bush and is crossed by several small streams, but these dry up in the "dry" seasons. Tse-tse flies were encountered in moderate numbers at Mahiba, and for a distance of seven miles beyond, towards Anquabe. Four of the five cases of trypanosomiasis occurring in Europeans were infected in this area; the fifth case contracted the disease in the Rovuma area. (He was the only one out of 2,000 troops to contract the disease). Careful enquiries were made concerning the native cases and the majority were found to have been in the district adjoining Mahiba, but no definite history could be obtained from the patients themselves.

Another line of communication in Portuguese East

Africa ran from Lumbo (opposite Mosambique) through Nam
pula towards Lake Nyassa - beyond Nampula the veterinary

surgeons reported the prevalence of trypanosomiasis amongst

cattle, but no cases amongst the troops occurred along

the Lumbo line.

carriers, operating in Portuguese East Africa, the disease is not easily contracted in the infected areas, as there were very few cases comparatively - 5 white men (including 2 officers) and 13 natives.

In Africa trypano somiasis is on the increase.

This is probably due to the fact that as the country is being opened up, an increased number of natives are moving from place to place, and larger numbers of Europeans settle in the continent. This has occurred to a much greater extent since war broke out, and one may expect further infected areas.

Etiology.

Stitt states: "The African trypano somiases follow infection with two species of trypano somes; the more virulent type of the disease occurring in South Central Africa being due to trypano some rhodesiense, transmitted by Glossina morsitans, and that of less severe type, but of more general distribution, being due to T.gambiense and transmitted by Glossina palpalis."

Macfie has reported a new trypanosome,

T.nigeriense, from young persons in Nigeria. It is said
to be less virulent than T. gambiense, and to be transmitted by Glossina tachnoides. Bruce considers T.nigeriense as being T.gambiense.

Regarding other forms of trypanosomes in man, Sir P.Manson states: "Seeing that most of the trypanosomes hitherto studied experimentally are capable of living in a variety of vertebrate hosts, it seems probable that other members of this group of parasites, in addition to T.gambiense, T.rhodesiense, and T.cruzi, may find in man at times suitable host. Apparently T.lewisi, T.evansi, and T.brucei do not generally thrive in man, but that circumstance does not warrant the inference that he is completely immune against these and all other species."

The trypanosomiasis parasites have a fusiform body. They vary in size, both in length and breadth.

According to Stitt "the normal type, as found in the blood, varies from 14 to 20 microns, while longer forms, 20 to 24 microns, are growth ones and, in the longest ones (23-33 microns) we have those preparing to divide longitudinally. In width they vary from 1.5 to 2 microns."

The parasite is a motile flagellate organism and

the body (which stains blue), shows two chromative staining masses, the one in the centre being the nucleus, and the one at the end (posterior end), the micronucleus or blepharoplast. Adjoining or surrounding the blepharoplast is a vacuole. Along the whole body runs a wavy membrane - the undulating membrane - which projects beyond the body, at the end away from the blepharoplast, as a flagellum; this end of the body is known as the anterior end.

Fry and Ranken have described the extrusion of granules - sharply defined refractile bodies - and this precedes the disintegration of the trypanosomes, and is especially observable in the more chronic forms of trypanosomiasis of men and animals. If occurring generally, it apparently heralds a disappearance of trypanosomes from the blood. It has been suggested that such granules might be infective, explaining the infectivity of blood from which trypanosomes were absent.

In my cases no granules were observed.

According to Manson in some cases the trypanosomes tend to recur cyclically at intervals of a week or more. There is no uniformity in the number of parasites present, sometimes they are fairly abundant — one or two in each field of the microscope — at other times and in the same patient it may be difficult or impossible, even after prolonged search, to find a single specimen. In my cases the trypanosomes were in comparatively large numbers and fairly easily found. This rather favours the view that the parasite was T.rhodesiense, as usually in early cases of infection, it is difficult to find T.gambiense.

As suggested by Mott, and proved by Greig and others they are generally more easily found in the lymphatic glands, which are often markedly enlarged.

In one native case, seen at the Carrier Hospital, Daressalam, which had a cirrhotic liver and developed ascites, the trypanosomes were found in the ascitic fluid.

The parasites are not found in the cerebrospinal fluid during the stage of trypanosomiasis, but are present on the advent of sleeping sickness.

Points in differentiation of T. rhodesiense from T. gambiense:

- "In human blood T. rhodesiense is morphologically indistinguishable from T. gambiense; but if it is passed through the rat a small but variable proportion of the parasites, especially the stumpy forms, will be seen to have their nuclei located posteriorly to the blepharoplast, that is to say, at the non-flagellar end of the organism." (Manson).
- 2. "Whilst both parasites have been cultivated on a special medium, T.rhodesiense is much more difficult to cultivate than T. gambiense." (Manson).
- 3. T. rhodesiense is more virulent for laboratory animals than T. gambiense.
- 4. Animals immunized against T. gambiense succumb to the Rhodesian trypanosome.
- 5. The clinical course of T. rhodesiense in man is more severe, and this type is much more resistant to treatment.
- for the glands are not enlarged to any degree in the Rhodesian infection, whilst they are usually markedly enlarged in cases infected with T. gambiense.
- 7. Two different species of flies transmit the trypano somes, Glossina morsitans transmitting T. rhodesiense and Glossina palpalis transmitting T. gambiense.

As regards the cases amongst the whites under review, Captain Hughes, I.M.S. conducted several experiments in animals, chiefly monkeys and rats.

He injected blood from the patients unto monkeys. Trypanosomes were detected in the animals in six to seven days. One died on the seventh day and two on the twelth day after inoculation. Twice, in the case of the men, a few forms were seen, approaching the so-called posterior nuclear forms, although the nucleus was never located behind the micro-nucleus. In the case of one officer (No.1), death of the monkey occurred on the seventh day, and posterior nuclear forms were obtained, with the nucleus behind the micro-nucleus.

In experiments on rats, infection by inoculating with a patient's blood, was readily induced; the period of incubation averaged 6 days and death took place usually in 12 days. Sub-inoculations from rat to rat, followed much the same course, but posterior nuclear forms were not obtained in the blood of any of the rats.

identical with T. rhodesiense, although the strains varied in virulence in different cases, being very virulent in the case of Officer (No.1), and in Middleton (Case 2 of the men). He based his opinion on the rapid way in which animals could be directly infected from man, its marked virulence in them, and the obtaining of posterior nuclear forms.

The clinical manifestations of the disease were certainly those of T. rhodesiense rather than T. gambiense, considering the non-amenability to treatment with atoxyl,

no marked glandular enlargements. the severity of the symptoms and the death of three of the white men within six months of contracting the disease.

Another important factor, which practically clinches the above opinion, was that G. morsitans was prevalent in the areas from which the cases came, and no G. palpalis was ever found.

The Tse-tse Flies.

Manson describes the Glossinae as "sombrecoloured, narrow bodied flies from about 8 to 12 m.m. long, with a thick proboscis (i.e. proboscis enclosed by the palpi) projecting horizontally in front of the head. The palpi are long, slightly grooved on their inner sides, and closely applied to the proboscis, which they almost entirely conceal, the only uncovered portion being a peculiar large bulb-like expansion at the base. wings are large, of a brownish hue, and present a characteristic venation, the fourth longitudinal vein bending abruptly upwards to meet the short mid-cross vein. the tse-tse is at rest its wings overlap on the back, covering each other like the blades of a pair of scissors; this is a distinctive feature. In the males, beneath the end of the abdomen, the external genitalia form a conspicuous knob-like protuberance, which renders the sexes easily distinguishable."

The genus at present comprises fourteen species and of these G.palpalis and G. morsitans are known to infect man with trypanosomes, although many observers consider that other species of glossinae may be found to convey the disease. Both have a wide distribution.

G. palpalis is present from Senegal to Angola on the west,

and throughout the Lualaba-Congo system to the Victoria Nyanza, Tanganyika, and Upper Nile reaching as far north as Moolo in the Soudan. G. morsitans extends south in Bechuanaland, North Eastern Transvaal, and Zululand. This fly is prevalent in Portuguese East Africa.

Captain McGregor, attached R.A.M.C., investigated the glossinae prevalent along the principal lines of communication in Portuguese East Africa and the southern part of German East Africa, and reported as follows:

- 1. Around Mahiba he found that G. pallidipes and G. morsitans were met with but no G. palpalis; G. pallidipes was the more common variety.
- 2. Between Lumbo and Nampula he found no tse-tse flies, but of the specimens examined which were sent from the district between Nampula and Lake Nyassa, he found G. morsitans, G. Pallidipes and G. fusca, in order of prevalence.
- 2. Captain McGregor also reported on the Lindi line in German East Africa, which ran from Lindi to Nigomago on the Rovuma, and includes a branch road to Tunduru. In the immediate vicinity of Lindi he found no tse-tse; on the roads from Lindi to Negomano and Tunduru he found four species, viz.— G. morsitans, G. pallidipes, G. fusca and G. brevipalpis. G. morsitans was the most common. The percentage of glossinae infected with a species of trypano soma, averaged about 11 per cent.

Habitat of Tse-tse Fly.

The tse-tse fly requires considerable moisture for its existence and is found principally in the trees or bush along the banks of the rivers and shores of the lakes. Removal of the bush would keep down the number of flies, but where there is a supply of water, this is impossible for any length of time, as the bush grows very quickly.

According to Manson, Hodges and his colleagues in

Uganda find that the fly ground proper is always a very narrow strip, not more than 10 to 15 yards, and always along the water's edge; and that the insects very rarely extend their feeding beat beyond 60 yards of this. They may, however, follow with great persistency a man who has just passed through this narrow belt, for several hundred yards, rarely as far as half a mile. The areas where glossinae are found are known as "fly belts" and the natives know the limits of these belts precisely.

They are seldom seen 3000 ft. above sea level.

Reproduction.

The glossinae do not lay eggs, but the eggs hatch and the larvae feed, develop and moult within the body of the parent.

Stitt states: "the female gives birth to a single, yellowish brown, motile larva, which is almost as large as the mother and which, upon reaching the ground, bores its way into a coarse, sandy soil for a depth of about 2 inches and then becomes a pupa. larval stage in the mother lasts about 2 weeks and the pupal stage about a month." "Moisture and sun-light are not favourable for pupal development, the sun being particularly injurious, so that pupae, buried only an inch deep and away from shade, are killed." "Male and female flies bite and transmit the disease. They bite in the day time, usually from 9 a.m. to 4 p.m., and will bite in the sun-light."

On making enquiries from the men who were in the fly areas, this was the common experience, but occasionally they were bitten at night. An entomologist, Mr Pomeroy, who had spent some time in Nigeria, told me he had frequently been bitten during the night time, whilst on trek there.

The Role of the Tse-tse fly in trypano somiasis.

Miss Robertson, in Uganda, found as follows: "In the fly the trypano somes are first established in the posterior parts of the mid-gut. Multiplication occurs and trypano somes of varying sizes are produced. From the loth to the 20th/day long, slender forms develop and pass forward into the proventriculus. These proventricular types reach the salivary glands by way of the hypopharynx. Here further development takes place, multiplication occurs and shorter forms develop, closely resembling the blood type. The development in the salivary glands takes from 2 to 5 days before the forms are infective. The fly is never infective until the glands are invaded."

"Conjugation has not been observed, nevertheless
the fly cycle as a whole has the biological significance
of conjugation."

Stitt states: "High temperatures, 75 to 85 F.

are favourable to development, while low temperatures,

60 to 70 F. are inimical to development, but do not kill

the ingested trypanosomes. This explains the long period

which at times elapses before a fly becomes infective.

Under favourable conditions a fly becomes infective in 20

to 34 days and remains infective the rest of its life, up

to 185 days."

"When tse-tse flies feed on animals infected with trypano somes, only from % to 6% become infective."

Kinghorn and Yorke advanced the view that the

large game of the country is a reservoir of the trypanosomes. This conclusion was supported by the fact that with a view to eradication of the disease certain areas have been depopulated, but upon examining the flies caught in the district a year or more later, infected flies have been obtained — in Uganda, the Government removed the entire population of the Sesse Islands and neighbouring shore of Lake Victoria Nyanza to fly free areas in the interior. Three years afterwards, Bruce ascertained that the local flies could still convey the disease to laboratory animals. This supports the view that the large game act as a reservoir for T. gambiense which is prevalent in Uganda.

Yorke as an experimental sanitary measure has advocated the extermination of the large game.

R.B.Woo snam in an article "The question of the relation of Game animals to Disease in Africa" (Journal of the East Africa and Nat. Hist. Soc., December, 1913), considers that domestic animals (sheep, goats, pigs. dogs, fowls), wild birds, lizards, snakes, frogs and toads may harbour the trypano somes of sleeping sickness and act as carriers of the disease. In which case it would be a hopeless undertaking to attempt to destroy practically all animal life in such extensive areas.

Koch advocated the slaughter of crocodiles.

Bruce and his co-workers have shown that

T. brucei, when injected into rats behaves similarly and cannot be distinguished from T. rhodesiense. From this and other facts these observers suggest that the two trypano somes are the same, that 37% of the wild game in the fly-area harbour pathogenic trypano somes, and that the

water buck, hartebeest, reed buck and duikerbuck are a source of danger to man.

Taute, however, who has carried out research work in Portuguese Nyassaland does not agree with these views. He says that T. brucei can be distinguished from the trypanosomes of sleeping sickness only and solely by the fact of its non-pathogenicity to man. Taute supports this view by certain experiments.

G. morsitans fed on animals infected with T. brucei were subsequently allowed to bite two men; the result was negative, though a number of control animals bitten simultaneously died.

Taute injected himself with 2 c.c. of blood from a trypano some infected dog, the trypano somes being of the Rhodesian type; there was no bad effect. 14 days later he injected 90 c.c. of his own blood into susceptible animals; these animals did not become infected - showing that the human organism has the power to kill this particular trypano some.

As Stitt points out, these experiments are not conclusive. as it is well known that men in good condition are not easily infected by the trypanosomes.

Many think that we may have trypano some carriers and that such persons in the enjoyment of health may act as reservoirs of the virus.

Koch, from observations in Uganda stated that trypanosomes may be communicated by the male during coitus. He noted the infection of 15 women in a fly-free district after the return home of their husbands from fly districts where they had contracted trypanosomiasis. Bernard has noted the same method of infection of prostitutes.

Dourine, a trypano some disease of horses, caused

by T. equiperdum, is transmitted by the sexual act.

According to Griggio, trypanosoma infection is not as a rule transmitted to the foetus; the abortion rate, however, in the infected is increased from the normal 7% in Congo natives to 31.7% and the infant mortality from 29% to 50%.

Incubation Period.

Castellani states - "probably in most cases it does not exceed two or three weeks, and, according to Martin and Lebeeuf's observations in Europeans, it may be even less than 10 days. On the other hand some infected individuals may not show any sign of disease for months, and, it is said, even 5 or 6 years.

Sanderson (Case 1) was the only one to give a definite history re a particular bite and in him symptoms of trypanosomiasis developed as early as 4 or 5 days afterwards. If this was the infecting bite, the incubation period was 4 or 5 days.

Symptoms.

- infected glossina is followed, in a proportion of cases, by a degree of local irritation of greater or less severity." This was present in Case 1 (Sanderson).
- 2. After the invasion period irregular pyrexia

 or period of trypanosome fever ensues. Sir P. Manson states that "irregularity of degree and duration is a feature of the fever."

In the Europeans the irregularity of the

pyrexis was a marked feature, fever being present for varying periods. In Case 1 (Sanderson), after treatment was suspended, there was a rise and fall of temperature, more or less, every day. The chart of Case 3 (Monk), from the date of the first record, for 33 consecutive days shows a temperature above normal. the temperature varying from 990 to 1050, only on 3 days registering 990 or under. Following this, there were 6 days practically of apyrexia, the temperature never reaching above 990. Later, there was a period of 12 days, when the temperature reached 1000 on two occasions only. In Cases beand 2 both of whom stood the treatment badly, there was never a period of apyrexia lasting longer than 2 days. blood examinations made on the Europeans, revealed the presence of trypanosomes usually with pyrexia and absence during the apyrexial period.

of the 15 native carriers, three only were practically apyrexial; attached charts of 9 cases show Cases C and E to be apyrexial. In the others the pyrexia was not so pronounced as in the Europeans, and it was more quickly influenced by treatment. Only occasionally after the temperature was once normal, was there a rise due to trypanosomes being in the blood. Thi was noticed in Cases B. E and F; in B. the rise did not occur until 3 months after the temperature had been normal. Intercurrent disease, however, caused a rise of temperature in many of the cases; the cause was generally malaria, but in one case (Case A.) it was due to relapsing fever.

Stitt states - "although fever may be absent in natives until the onset of the period of sleeping sickness, yet in Europeans there are usually noted febrile

paroxysms, lasting for a few days, and followed by
febrile periods varying from a few days to 2 or 3
weeks. The temperature curve is of a markedly remittent
type, approaching normal in the morning and going up to
103 F. or higher in the evening - a wide daily range."

In the Europeans, the temperature, as a rule, tended to be higher in the evenings, but daily remissions were not a feature in my cases. In the natives with pyrexia the daily remission was a marked feature.

3. Sir P. Manson states - "The heart's action is generally rapid and easily excited."

Stitt states: "Very characteristic is a lowtension rapid pulse." In my cases frequency and low tension of the pulse were a marked feature; the heart's action was easily quickened, being noticed after the least excitement, or exertion. In Case 1, the pulse rate was slow at first and the tension low, later it was quick and the tension still lower. In Case 2, the tension was medium at first, gradually becoming less and when the patient left hospital, it was decidedly low. In Case 3, the tension was low on admission to hospital, but there was a decided improvement in this and also the heart became less excitable, as the treatment began to take effect. Unfortunately, it was impossible to obtain accurate records of blood pressure, as no recording instruments were available.

In the native cases, the low tension and frequency of the pulse were noticed, and also great increase of the pulse rate on the slightest exertion. The pulse rate at first was between 100 and 140. As in Case 3, marked improvement was effected in all the cases

by treatment, the tension being increased and the pulse rate reduced to between 80 and 100.

A slight degree of anaemia was present in all the cases, both European and native.

A common symptom complained of by the Europeans, was of being easily tired. The worst cases amongst the natives, before treatment was instituted, were barely able to crawl about, the others were only able to walk short distances. With treatment a marked improvement was produced.

Symptoms referable to the nervous system:

Head-ache was pronounced in Cases 1 and 2.

usually when the patient was feverish. Headache was not complained of by the natives
and only slightly by Middleton (Case 2.)

5.

- Lack of mental concentration was noticed in the three white men.
 - (c) Insomnia, described in text-books, as being present in the early stage, was only present in Case 1, and in him the sleepless-ness was due to pains in the limbs, resulting from injection of atoxyl. In the native cases insomnia was only present in one case.
- Deep hyperaesthesia or Kerandel's symptom if the patient strikes a limb against any hard object, a degree of disconfort amounting to actual pain is experienced, the sensation being slightly delayed. Kerandel, who suffered from trypano somiasis, noted that the fear of striking against objects, became with him an absolute obsession. None of the natives experienced this, and only in Case 3 of the Europeans was it elicited. although he felt no obsession.
- Other nervous manifestations were present in Case 3 loss of memory for recent events and loss of epicritic sense below the tibial tuberosities. Treatment improved all the above symptoms; four weeks after commencing, Kerendel's symptom was not elicited and the epicritic sense was practically regained. Patient's memory improved, his general appearance became brighter and he was able to concentrate his mind better.

- tremor of the tongue and lips was present to a minor degree at first, but later it was not so marked.
- 7. Glandular enlargements, particularly of the glands in the posterior triangles.

Sir P.Manson states: "that only one gland may be visibly involved or there may be a recognisable general polyadenitis, including the abdominal glands.

The implicated glands may be very prominent or they may not be easily felt."

"No enlargement of the lymphatic glands is said to occur in the victims of infection with T. rhodesiense."

In the native cases no enlarged glands were found. In all the three white men glands were palpable in the posterior triangles of the neck; in Cases 2 and 3 they were palpable in other regions. The glands were usually slightly tender, of medium consistence and varied in size between a coffee bean and an almond, but were never enlarged to the same extent as is seen in cases infected with T. gambiense. As trypanosomes were found easily in the blood, microscopic examination of the gland juice was not done.

8. The Skin.

(a) Erythematous areas may be present. Lieut.Col. Newham, R.A.M.C., (Lecturer at the London School of Tropical Diseases) states: "The rash is characteristic and may appear at any time. This rash appears chiefly on the trunk, particularly on the chest and back.

Sometimes it appears on the forehead but is not usually seen on the limbs. It appears as a definite erythema of

the skin in large rings, perhaps 2 to 3 inches in diameter. The centres of these rings are frequently discoloured, somewhat like a bruise, and the rings are generally incomplete."

In Case 3 (Monk), irregular patches of erythema were seen on the chest about a fortnight after the commencement of symptoms. In one of the Officers (No.1) about the sixth week after the onset of symptoms. I saw the typical rash. This appeared in the front and back of chest in the form of rings; the rings were incomplete, pink in colour and measured from 2 to $3\frac{1}{2}$ inches in diameter The individual rings disappeared in three or four days, fresh ones appearing. The rash was only noticed during a period of ten days.

In the natives no rash was seen.

(b) Localized oedema.

This symptom was a marked feature in the Natives, but was not present in the Europeans. The swelling occurred under the eyelids and in some of the cases the cheeks were puffy.

9. The Spleen.

"The spleen may be enlarged." (Stitt). The spleen was palpable in the three white men; in Cases 1 and 3, half an inch, and in Case 2, three inches below the costal cartilage. In all three cases malaria may have produced the enlargement, and in Case 2, malaria undoubtedly was the chief cause.

The spleen was palpable in the natives, but in no case was it larger than is generally felt.

10. Liver.

Sir P.Manson states: "The liver also may be enlarged." Only Case 2 showed any enlargement and this may have been due to malaria.

cyclitis and choroiditis are stated as sometimes occurring. Sir P.Manson also mentions - "Neuralgic pains, cramps, formication, paraesthesiae, orchitis and painful local inflammatory swellings, which subside without suppuration." Apart from neuralgic pains in the legs, as a result of the administration of atoxyl, none of the above symptoms were noticed in my cases.

Sleeping Sickness Stage.

trypanosomes and toxins is manifested chiefly in the nervous system.

The following is a resume of the symptoms as described in text books:

The tired feeling becomes intensified, the gait is a slow and shuffling one and the patient prefers lying still. He is lethargic and morose and his expression vacant. Later the tongue shows a decided tremor, which may also be present in the lips and hands and more rarely in the legs. The patient gradually sinks into a semi-comatose condition, falling asleep anywhere, even with a morsel of food in his mouth. At this stage he can be aroused and will answer fairly intelligently, but with apathy and retardation. Finally weakness and emaciation increases. Convulsions, local in the limbs, or more general, may occur. Bedsores tend to form. The

coma deepens and the patient dies. Frequently pneumonia or dysentery bring about the end.

The majority of observers consider that the sleeping sickness stage rarely lasts longer than a year without treatment, and with treatment not longer than two years.

in any of the cases under observation. One case amongst the natives, simulated the symptoms on admission. This patient was lethargic and resented being disturbed, his cerebration was very slow, the face was puffy and anaemia was present to a greater degree than in the other cases. No trypano somes were detected in the cerebro-spinal fluid, but on examination of the faeces, ankylostomes were found in great numbers. After treatment with thymol the above symptoms cleared up.

Diagnosis.

Any patient who has resided in tropical Africa recently, and especially if he has been in a tse-tse fly area, must be considered as a possible case of trypanosomiasis.

Apart from irregular pyrexia, all other signs and symptoms may be absent and a definite diagnosis can only be obtained by finding the specific parasite in the blood or glands. Important signs when present are the enlarged glands in the posterior triangles of the neck, and the characteristic rash. The following diseases might be confused with trypanosomiasis: malaria, kala-azar, pellagra and syphilis; of these, no cases of kala-azar or pellagra have been met with in East Africa. The chief disease to exclude is malaria.

In the three white men, malaria was first thought of as the cause of the pyrexia. Case I was admitted to the Base Hospital as a malarial transfer; Case 2 was subject to attacks and benigh tertian parasites were found in his blood after the diagnosis of trypano somiasis had been established; in Case 3, benigh tertian parasites were found in the blood before the trypano somes.

No case showed the tertian periodicity in the temperature chart. On defervescence sweating is seen in both diseases, but it is greater in the benign tertian variety of malaria. As regards symptoms, with a high temperature, in nearly all malarial cases, the patient complains more of pain in the head, body and limbs; in trypanosomiasis the patient is generally sleepy and does not wish to be disturbed, or occasionally he is irritable and has a head-ache, which may be severe. If pyrexia is not influenced by the administration of quinine, malaria can be excluded.

To diagnose trypanosomiasis from kala-ezar,
Manson mentions the presence of enlarged glands, local
oedema, and erythema multiforme in trypanosomiasis, and
their absence in kala-azar; also, bacteriological
examination for trypanosomes or Leishman-Donovan bodies if the blood or gland juice prove negative, puncture of
the liver, if enlarged, or spleen, should be done.

In pellagra, there is usually a period of exacerbation of the symptoms in spring; the rash affects symmetrical and sharply delimited patches of skin on exposed parts of the body, and is not ringed or fugitive as that of trypanosomiasis. As regards syphilis, a positive Wasserman is not of much value, as it is often obtained in trypanosomiasis.

The diagnosis finally rests on finding trypano somes in the blood or lymph from an enlarged In all the cases under review, blood lymphatic gland. In the natives, examinations revealed their presence. the parasites were found in the ordinary routine examination of the blood, which was done in every case when admitted to hospital. In the case of the Europeans the parasitas, as a rule, were easily found, one or two trypano somes being seen in nearly every field. best time for taking a specimen of blood, was when the patient had a rise of temperature, but not immediately after the administration of tartar emetic. The following summary indicates the result of 41 blood tests:

- (a) In 24 examinations of the blood with a temperature above 99, trypano somes were present 19 times.
- (b) In 17 examinations with a temperature below 99 trypano somes were only present twice.

In any suspected case, the blood should be examined frequently as the trypanosomes seem to make their appearance in the blood in cycles.

Owing to the difficulty experienced in finding parasites in the peripheral blood, some observers prefer to take 10 to 20, c.c. of blood in about 25 c.c. of citrated salt solution, centrifuging two or three times and examining the sediment of the third centrifugalization. (Nabarro's method). Others prefer, particularly in the earlier stages of the disease, the gland puncture method, using a sterile but dry hypodermic needle. (Grieg-Grey's method).

In the sleeping sickness, stage trypanosomes can usually be found in the cerebro-spinal fluid.

above methods, it is necessary to inoculate susceptible animals, particularly the guinea pig. rat or monkey, with 10 or 20 c.c. of blood or with gland juice from the suspected case, and later examine the blood of these animals for trypanosomes. Apart from diagnosis, such inoculations are of value for testing the virulence of the particular organism and as a test of recovery.

Stitt describes other tests:

- 1. Trypanolysis unheated suspected serum and trypanosomes are incubated together for one hour normal serum may occasionally cause disintegration and treated cases give it in only about 45% of cases. Unfavourable untreated cases give it in about 80% of cases.
- 2. The attachment test is made by making a mixture of inactivated serum, leucocytes and trypano somes and allowing them to be in contact for 20 minutes. A positive test shows attachment of the trypano somes to the leucocytes.

Manson mentions a macroscopic test: "A wet blood preparation exhibits, even to the naked eye, a remarkable clumping of the red corpuscles. Held up to the light, such a preparation has a peculiar granular appearance, produced, as can be seen on microscopical examination, by agglomeration of the corpuscles into heaps and clusters, the usual rouleax arrangement being absent. Such a disposition of corpuscles is significant of, though not peculiar to trypano soma infection."

Castellani states that it is found in cases of filariasis, malaria, syphilis and yaws.

This appearance was noticed in the slides taken from the cases.

Morbid Anatomy.

Manson states: "No gross lesions of the nerve centres, or of any other organ, have been described as being present."

Of the native cases seen postmortem, apart from the local lesion of the disease causing death, macroscopic appearance of the organs was normal, except in one case, which showed signs of irritant poisoning. This patient had been under treatment for 5 months, and apparently he was doing very well, putting on weight and having been free from any rise of temperature for He was given 3 gr. atoxyl and, to begin several days. with, 1 gr. tartar emetic, both twice weekly. As the symptoms were not being controlled, the antimony was increased gradually and for the last 8 injections he had been receiving 3 gr. After the last injection, his temperature began to rise, breathing became rapid, pulse very frequent and weak, and he passed into a comatose condition, dying within 30 hours of the injection. The following is the postmortem report on the cased-

The body is well nourished. Subcutaneous fat is greatly increased.

Brain: Normal.

Heart: Externally the heart is covered with a considerable amount of fat; the cavities are dilated; there is no valvular lesion; the muscle walls are very pale and flabby.

Lungs: Patches of early broncho-pneumonia are

present in both lungs.

Liver: Enlarged and externally fatty.
parenchyma being pale yellow in

colour.

Kidneys: Both are fatty. The parenchyma is very pale and it is difficult to differentiate the cortex from the

medulla.

Intestines: Omentum and appendices epiploices

are very fatty.

Spleen: Slightly enlarged, coft and dark red

Opinion as to the cause of death.

(a) Primary cause: Fatty infiltration and degeneration of the internal organs. The cause of the fatty change is probably due to the cumulative effect of the antimony, but it is possible the atoxyl may have assisted or it may have been the chief cause.

(b) Secondary and immediate cause: Broncho-pneumonia with cardiac failure.

Prognosis.

The progncsis depends on the virulence of the infecting organism, the stage of the disease, and the patient's ability to tolerate the treatment. If the case cannot be removed from the endemic area, the prognosis is bad.

virulence of the infection varies in different parts of Africa. In the case of T. gambiense, the least virulent is the type found in the Congo; the Nigerian type is rather more virulent, but there is usually good hope of recovery; the Uganda type represents the third degree of virulence. The outlook in all the above types is much more favourable than in cases of infection with T. rhodesiense, which is practically always fatal. Lieut.Col. Newham, R.A.M.C., records one case of

rhodesiense infection, which is apparently cured. This patient was treated in London for two years, when all tests for trypano somes proved negative and the patient was able to do his work again.

The cases under review were infected with T. T. rhodesiense and the outlook was very bad. Of the five white men (including two officers) one only (case 3 -Monk) age 20, was able to tolerate the treatment. him, the trypano somes were under control and at the time of writing his chances of recovery were good. The other two men, ages 40 and 52 respectively, were unable to stand the treatment and died from asthenia with oedema within 6 months of the onset of the disease. Neither of the two officers were able to take more than grs. $1\frac{1}{2}$ of antimony, and this often caused a very severe reaction. One officer (No.1) age about 40, died in about 6 months time from asthenia with oedema and towards the end he developed epileptiform convulsions; in the other, age 35, who had only been under treatment for a short time, the trypano somes were not being controlled and also, owing to his anaemic state the veins were collapsed, necessitating dissecting out the vein at each injection, as in Case 2 of the men.

Judging from these cases, it appears that men of middle age, had very little chance of recovery, owing to their inability to take the treatment.

The natives were all able to take the antimony well, without any apparent bad effect. In one case, however, postmortem examination revealed an extensive fatty infiltration throughout the body, with fatty degeneration of the internal organs, and this could only be ascribed to the treatment. (Vide Morbid Anatomy).

Of the other 12 cases, intercurrent disease caused 7 deaths, the causes being:

- 1. Chronic Diarrhoea.
- 2. Relapsing Fever.
- 3. General Debility.
- 4. Pleurisy.
- 5. Pneumonia.
- 6. Pneumonia.
- 7. Pneumonia.

In these cases, there was no other sign of disease in the body. Pneumonia is a likely and dangerous complication in trypanosomiasis; the 3 natives who died from it, were all doing well, when an epidemic of influenza, complicated frequently with pneumonia, broke out in Daressalam. Two of the cases died on the second day after contracting pneumonia. The remaining natives appeared to be doing well, the trypanosomes being kept under control.

Treatment.

At present the chief drugs employed are arsenic and antimony. Ehlrich and Shiga tried colouring compounds belonging to the benze-purpurin group, of which trypan-red is the best known.

Arsenic.

The early observers gave arsenic in the form of liq. arsenicalis by the mouth. This has been superseded by intramuscular injections of a variety of preparations, including atoxyl, soamin, arseno-phenyl glycin, arsacetin, etc., etc. Atoxyl was recommended by Thomas in 1905 and it has been used largely since. Castellani states atoxyl is sodium-p-amino-phenyl arseniate and contains about 23% of arsenic. This preparation was administered in my cases. The atoxyl was dissolved in 10% normal saline; the solution

was made fresh and if at all muddy in appearance,
was not used. It was injected into the gluteal
muscles, at a point 3 inches immediately behind the
anterior superior spine and the dose given was 3 grs.
twice a week. With regard to administration of
atoxyl, Sir P.Manson states: "Unfortunately it has
one serious drawback — in large doses it is apt to
give rise to optic neuritis, and consequent atrophy and
blindness. The drug, therefore, must at once be
stopped on the slightest threatening — dimness of
vision or congestion of the disc — of such a calamity.
Gastro-intestinal irritation and peripheral neuritis
are also indications that it has to be suspended, at
least temporarily, and the subsequent dose reduced."

In Cases 1 and 2 (of the men), the atoxyl had to be reduced to $1\frac{1}{2}$ grs., as both had symptoms of peripheral neuritis. No case of optic neuritis occurred. Case 1 complained of dimness of vision, but this was due to an error in refraction, the retinae being normal.

No reduction of the dose had to be made in any of the natives.

As regards the value of atoxy1, it did not appear to affect the trypanosomes in any of the cases under observation. In Case 1, atoxy1 was given alone at first, but with no appreciable effect. Case A. of the native carriers was tried on it alone, for a few days; the trypanosomes were not controlled and the irregular pyrexia continued until tartar emetic was given.

on the other hand, when atoxyl was not given and tartar emetic was solely used, as in cases C. and D. the trypano somes were controlled just as well, as in those cases where the combined treatment was given.

Apart from the carriers, I saw a case of trypanosomiasis in an askari (native soldier) at the 3rd African Hospital, Daressalam, where atoxyl completely controlled the trypanosomes. The patient was admitted to hospital for a minor ailment, feeling otherwise all right. He presented no symptoms of the disease, apart from the finding of trypanosomes in his blood; these were proved by inoculation tests to be of a much lower virulence than the trypanosomes met with in the case of the Europeans. Captain Hughes, I.M.S., who did the tests, expressed the opinion that the native, who came from West Africa, had probably been infected there. No tartar emetic was given and after the first injection of atoxyl, trypanosomes were not found subsequently in the blood.

It appears that atoxyl does not affect the more virulent strains of trypanosomes, but is efficacious with the milder strains.

In 1907 Ehlrich and his pupils. Franke and Roehl, discovered that trypanosomes may, after a time, become atoxyl - resistant. Sir P.Manson states: "Unless contra-indicated, atoxyl and antimony treatment should be combined or alternated. One or both should be instituted immediately the diagnosis of trypanosomiasis is established."

"As pointed out by Ehlrich, certain strains of trypanosomes are arsenic fast," that is they resist the action of the drug - a property which becomes hereditary and is continued

in sub-inoculations into the lower animals. This undoubtedly accounts for the failure of arsenical treatment in a large proportion of cases. Presumably there are strains having a similar resisting power to antimony. As the atoxyl-fast trypanosomes may respond to antimony, and vice versa, both drugs should be employed in every case."

"Intravenous injection of arsenophenylglycin, in doses
of about 1 gramm (15 grs.) intravenously has been highly
recommended. Recent reports from German East Africa
state that of 35 treated with this drug, 6 died of the
effects of the drug. Salvarsan and neosalvarsan have
been used but apparently without particular success."

"Very favourable reports have recently been made from the use of galyl and ludyl, arsenical compounds."

Antimony.

Plimmer and Thomas introduced tartar emetic.

Regarding the giving of antimony Sir P. Manson states: "In two cases of trypanosomiasis in man in which I used this drug intramuscularly the consequent local pain and irritation (culminating in tissue necrosis) were so excessive that in this form the treatment had to be abandoned. One of the patients, however, continued the drug by the mouth to the extent of $1\frac{1}{2}$ to 2 gr. daily, very highly diluted, a little being given in all his food and drink, and along with atoxyl injections, for over two years. He appears to be quite well."

"It can be given, freely diluted in normal saline, per rectum; but the efficacy of this method has not been proved. Though troublesome and not free from

risk, the intravenous method is the best and the one now generally adopted. The dose is $\frac{1}{2}$ gr., gradually increased to $1\frac{1}{2}$ gr., in 6 oz. or thereabouts of sterile normal saline."

"As with atoxyl, there are differences in the methods of employing antimony injections. Some give them daily in courses of 15 days at intervals of 15 days; others, once a week; others again, according to the indications supplied by temperature and the microscopical examination of the blood."

Stitt states: "Probably the best treatment is one in which three doses of atoxyl are followed by from 10 to 15 daily injections of 0.1 gram. $(1\frac{1}{2} \text{ gr.})$ of tartar emetic. The course is repeated after an interval of 3 weeks. It is advisable to give a hypodermic of caffein a few minutes before the tarter emetic to lessen depression."

In the cases under observation, tartar emetic was never tried by the mouth or intramuscularly. In Case 2, as no prominent vein could be obtained, the drug was administered for 2 weeks per rectum. It is doubtful if any of the solution was absorbed, as the trypano somes were certainly not controlled and there was no constitutional reaction afterwards, as one had with the same dose given intravenously.

In the Europeans, the tartar emetic, dissolved in normal saline, was given intravenously, using a funnel, tube and needle, in the same way as salvarsan is administered; in all, about 8 oz. of saline were given at each injection. On two occasions, following the injection, pain and stiffness at the site of the injection, was

complained of. This was probably due to a slight leakage of the drug into the cellular tissues.

In the natives, a 10 c.c. syringe was used and the solution of tartar emetic was injected direct into the vein. This method was quite efficacious and no local trouble followed any injection.

The important points to bear in mind, in administering the drug, are:-

- That there should be no leakage into the surrounding tissues.
- 2. That the solution should be well diluted.
- 3. That the solution should enter the vein at blood heat.
- 4. That the drug should be administered slowly.

The general effects produced by injection of tartar emetic.

The effects on the temperature (see Charts) in the Europeans, were noticed in relation to the presence or absence of trypano somes in the blood - blood slides were taken just previous to the injection - with no trypano somes in the blood, there was generally little or no reaction; with trypano somes present, there was always a rise of temperature, which was greater when the parasites were plentiful, than when scanty. The rise was often preceded by a rigor. During the first injections or after any increase in the dose of the drug, the rise of temperature was marked, shooting up to 104° or 105°; later. as seen in Case 3, the rises were not so pronounced and definite rigors were absent. In the majority of the native cases, the injections at first caused a sharp rise of temperature, which was down on the following day. Preceding the rise, the patient experienced usually a

slight chilly sensation. but no rigor. This occurred in different cases, for varying periods, from a few days to 8 weeks following the first injection; afterwards, there was rarely any effect on the temperature.

Other effects in the white men.

In Case 1, at first, construction of the chest with a short cough occurred immediately after the injection, followed in 1 to 12 hours by a rigor; later, for a period, there was only a slight reaction, until the patient had a very severe one, with gastro-intestinal irritation and profound collapse. No further injections were given.

In Case 2, the first injection caused severe gastro-intestinal irritation with collapse. After subsequent injections, a rigor generally occurred with pain in the bottom of the back. Later, the dose was increased to $1\frac{1}{2}$ gr. but the reaction was too great and intravenous injections were discontinued.

In Case 3, injections of 1 gr. caused only slight constitutional symptoms, usually a slight rigor, occurring about 1 hour afterwards, with a rise of temperature, which was usually down by the next day. With doses of 2 and $2\frac{1}{2}$ gr. immediately following the injection patient had a slight spasmodic cough, lasting for 2 or 3 minutes, and later, usually a severe rigor with a sharp rise of temperature, which dropped to normal by the next day.

The native cases were very little effected apart from the rise of temperature, which was frequently preceded by a chilly feeling. With doses of 2, $2\frac{1}{2}$ and 3 gr.

a short cough occurred as in case 3.

with reference to the cause of these effects. the most probable explantion, is the trypanolytic action of the drug, with liberation of endotoxin.

Dose of tartar emetic used and the patient's progress:-

In all cases, 1 gr. twice a week, was given at the commencement. In Cases 1 and 2, the reaction was so severe, that no increase in the dose could be made, and the trypanosomes soon became tolerant to it. Case 1 was a bad patient to treat, finally refusing to have any further injections. The tartar emetic ameliorated his symptoms at first, and trypanosomes were not found in the blood so frequently as after the injections had been stopped, when the parasites were generally present and in greater numbers, and the patient became progressively weaker and died.

case 2 was unfortunate in having no prominent veins and the dissection of individual veins with slow healing of the skin, added to the difficulties of his treatment. The small dose, whilst affecting the trypanosomes, did not control them and there was no improvement in the patient's general condition. Before transferring him to a hospital ship, treatment with a solution of antimonium oxide, given intramuscular, was substituted for the tartar emetic, and this was persevered with, but the patient became weaker and died 3 months later in South Africa.

In Case 3 the reaction was never severe and after 1 gr. twice a week, for 30 days, the dose was increased to 2 gr; this was continued for 17 days, when $2\frac{1}{2}$ gr. was given. An increase in dose was made, when it appeared

the trypanosomes were not being controlled; this being judged both by their continued presence in the blood and by the temperature chart. The outlook in this case was decidedly good, as both the patient's general condition improved and the clinical signs of the disease abated.

(Since the paper was written, Case 3 has died from pneumonia following influenza, five months after contracting trypanosomiasis. Up to this time he was doing apparently very well, taking the treatment well and the trypanosomes were under control)

The contrast between the natives and Europeans was marked as regards the amount of tartar emetic which could be tolerated; in the natives, the dose was quickly increased by $\frac{1}{2}$ gr. a time, until 3 and $3\frac{1}{2}$ gr. were given, twice weekly. The natives responded quickly to the treatment, as evidenced by the general improvement in their condition and by the early disappearance of trypanosomes from the blood, slides being examined every day, but it is questionable, judging from the case reported under morbid anatomy, whether they will be able to stand taking such high doses as $3\frac{1}{2}$ gr. over a prolonged period, without serious effects on the internal organs.

Length of Treatment necessary.

The absence of trypano somes in the patient's blood, may give one a false sense of security. This was exemplified in some of the native cases, trypano somes reappearing in the blood after being absent for some time:

In case B for 4 months m m E m 3 m and m m F m 1 m

According to Castellani, treatment should be continued for at least 2 years and then inoculation tests must be negative before stopping treatment. Lieut.Col. Newham recommends inoculating a monkey, 3 or 4 times, with blood from the patient; if the monkey shows no signs of the presence of the trypanosomiasis parasite, the patient may be considered free. Apart from bacteriological findings, a good test is the patient's capacity for physical endurance.

Prophylaxis.

Segregation of the infected cases and diminishing the number of tse-tse flies in any area are the chief
measures.

Where you have a restricted area, sanitary measures can exterminate the disease, as was accomplished in the island of Principe, in Portuguese West Africa.

Here, according to an article published in a White Paper (published Ap. 22, 1914) on the question of native labour, the workers carried cloths covered with glue or other sticky substance on their backs when working in fly areas, and a great number of flies were caught in this way.

Other measures in addition were taken - jungle clearing, drainage, blood examination, segregation of infected cases and killing any possible animal reservoir.

To exterminate the disease by sanitary measures alone, in such widely infected areas as met with in Africa, is impossible at present, but it would be a great advance if by vaccine or serum therapy immunity could be produced in any exposed people.

Prophylaxis as carried out in East African Campaign.

The ideal plan would have been to keep all troops out of infected areas. This was impossible, but instructions were issued that any troops passing through the fly belt zones, should proceed as quickly as possible, and if it was necessary to pitch a camp, the bush had to be cleared for a number of yards around. No troops were allowed to wear "shorts" (trousers leaving the knees bare), and a protection for the face and neck was given to each man, made from green netting with a fine mesh, similar to mosquito netting. Also, if possible, gloves had to be worn or otherwise bamber oil was is sued for smearing any exposed parts. All cases of sickness, coming from the area, had to have repeated microscopic examinations of the blood done, to exclude trypanosomiasis, and were kept under observation for one month.

Conclusions.

- to man. Large numbers of troops and carriers were in the infected districts for considerable periods. In one instance, a body of 2,000 troops marched through a fly stricken area and only one case occurred. The blood of the 1,999 others was examined with negative results.
- The Europeans were debilitated and their resisting power was probably lowered.
- 3. It seems clear that the incubation period may be as short as 5 days. (Vide Case I).

- The only symptom constantly present. in all the cases under my observation was irregular pyrexia, and the diagnosis depended entirely on the finding of the parasite.
- Atoxyl had no apparent effect on the trypano somes.

 Antimony seems to kill the parasites, but in the severer infections with T. rhodesiense, the trypano somes seem to become resistant.

In view of the fact, that inhabitants of an area infected with trypanosomiasis, seem to acquire a natural immunity to this disease, and that the small African humped oxen are known to be immune to Nagana, it seems possible that in vaccine or serum therapy lies our chief hope for both prevention and cure.

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DRIVER SANDERSON, G.W.

S.A.S.C.

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DRIVER SANDERSON G.W.

SOUTH AFRICAN SERVICE CORPS, M.T.

TRYPANOSOMIASIS

918. ch 14

The patient is 40 years of age and a Driver in the Motor Transport Service.

He served in the Boer War, having Enteric Fever and later settled in South Africa, where he enjoyed good health.

Patient commenced service in East Africa in July, 1916. From September 1916 to July 1917 he was on duty in the Rufigi area, where he was often bitten by the Tsetse fly but the bites never troubled him.

Previous to the present illness he has had several attacks of Malaria, being seven times in hospital; also he has had two attacks of Dysentery and after the last attack at the end of December 1917 he was sent South for recuperation.

On his return to East Africa at the beginning of February 1918, he was sent to Port Amelia and from February 7th. to the 20th.

was on convoy duty between Maheba and Ankuabe, about 35 miles inland. During this time he was frequently bitten by the Tsetse fly, causing a sharp sting, but the effects passed off in a few minutes. During the middle of the day on Feb. 21st. or 22nd., whilst drawing water at the river near Maheba, patient was bitten behind the left shoulder; this was more painful than any previous bite and later, irritable, so in the evening he had the part painted with Iodine. Next day he felt no irritation or pain.

On February 26th. patient felt seedy, suffering from severe pain all over the head; down back of neck and in the loin; he was feverish: through the night he perspired and felt much better on the following morning, but he had no desire for food.

Patient, himself, thought it was an attack of Malaria, but Col. Newham, R.A.M.C. dated the commencement of Trypanosomiasis from this time, and if so, and the severe bite was the infecting bite, the incubation period was 4 or 5 days.

On February 27th. he was admitted to the Camp Hospital at Maheba. Just before mid-night, he woke up with dreadful pains through the

head, chiefly at the back and radiating down the back of the neck to the 7th. Cervical Spine; also he had severe pains in the body, which were most severe in the loin; he experienced no rigor but felt feverish.

On February 28th. at 6-0 a.m., Temperature rose to 101°; he felt seedy all day and did not perspire; he had a good night and felt better the following morning, March 1st, when he was transferred to the hospital at Port Amelia. For the next few days patient felt quite well, except when he in the sun's rays when he had headaches, slight fainting attacks, and was easily tired. On March 8th. he was transferred as a Malarial case per hospital ship to Dar-es-Salaam, being admitted to No.2.South African General Hospital on March 10th. 1918. On examination nothing abnormal was detected and patient said he felt alright, apart from feeling weak and being easily tired.

Temperature normal. Pulse 90. Blood slide was negative to Malarial parasites.

Treatment. Quinine Sulphate grs.10. ter in die and ordinary diet. Patient was allowed up between 4 and 6pm.

Until March 14th. he was comfortable, when he complained of feeling tired, head-ache and pains all over body. Temperature at 7pm. was 101.

- h 15. Patient was depressed and drowsy. Temperature 101.2 in morning and at 10 pm. 99.
 - 16. Temperature remained high, the evening temp. reached 105.

 Through the night he perspired freely.
 - 17. Major Semple, R.A.M.C., O.U., Bacteriological Laboratory, reported that Trypanosomes had been found in the blood slide specimen, which had been taken on March 16th.

 Other clinical signs were then noticed:General condition: Patient is emaciated, physique poor. Slight enlargement, about the size of an almond, and tenderness of one gland in the posterior triangle on the right side, just behind the Sterno-Mastoid; no other glands were enlarged. There was no rash and no localised oedema; no ocular changes (Retinae normal); no

deep hyperaesthesia; Knee jerks were normal; there was no Babinski

and no Ankle clonus; epicritic sense and sense to heat and cold were normal; patient appreciated two points of needles at a distance of 1 inch in the transverse direction and from 1\frac{1}{6} to 2 inches in the long axis of the limb.

HEART: No enlargement; no adventitious sounds.

PULSE: 80 per minute, was of low tension.

LUNGS: Normal.

SPLEEN: Palpable & no tenderness.

LIVER: Not palpable.

Since admission patient's mental attitude has been variable, sometimes being bright and cheerful, and at other times, usually when his temperature was above normal, he was irritable or apathetic and listless. His appetite was good except when he had a temperature. Bowels constipated.

Absolute rest in bed was ordered and as much nutriment to be taken as possible; the Quinine was stopped. Liquid Cascara was given for the constipation and this had to be supplemented sometimes with Calomel and Salts.

ch 18. Atoxyl grs.3. dissolved in Normal Saline, was injected into the Gluteal muscle at & point 3ins. directly behind the anterior superior spine. No after effects noticed.

Injection was repeated on the 20th., 22nd. and 25th.

- 22. Bacteriological Report: Trypanosomes not found.
- 26. do. do. Trypanosomes found in blood.
- 27. Tartar Emetic, grs.1. dissolved in 2 ozs. Normal Saline was injected intravenously at the elbow: A solution of Normal Saline alone was first run through funnel, tube and needle into vein, to make sure there was no escape of any fluid into the cellular tissue; the solutions were kept at blood heat, the bottles being immersed in warm water. At the end of the injection, which lasted about 4 minutes, patient experienced a sharp and constricting pain over the lower part and front of chest; this lasted about 1 minute and was accompanied with a short cough. Pulse in the evening was 108. Temperature 100.
- 28. Patient complained of acute pain over right side of chest.
- 29. Comfortable.

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- h 30. In morning pulse and temperature were normal. At 2pm.

 Tartar Emetic, grs.l. was given; immediately afterwards patient complained of constricting pain at the lower part and front of chest and he had a short irritable cough. At 3-30pm. he had a rigor, temperature rising to 104 by 6pm.
 - 31. Patient comfortable.
- 11 1. do. do. Atoxyl grs.2 injected.
 - 2. do. do.

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- 3. Bact. Report: Trypanosomes were present in blood.

 He complained of pains in limbs, coming and going. He was given a mixture containing Sod. Salicyl grs.10. and Potass. Tedide grs.5. and as he had been sleeping badly he was given at bed-time a mixture containing Potass. Bromide and Chloral Hydrate as grs.10.
- 4. Temperature at 2pm. was 98. At 2-15pm. Tartar Emetic, grs.l. was injected; constricting pain in chest and irritable cough followed immediately after injection. One hour afterwards patient had a rigor and Temp. at 6pm. was 104, Pulse 116. Through the night her perspired freely.
- 5. At 6am. Temperature was normal. Pulse 92.
- 6. Patient comfortable. Atoxyl grs. 21 injected.
- 8. Temperature at 2pm. was normal. Tartar Emetic grs. 1. was injected constricting pain was only slightly felt and there was no cough or rigor afterwards. At 6pm. Temp. was 101. Pulse 99.
- 9. Patient comfortable. Atoxyl grs.22 injected.
- 11. He complained of aching pains and tenderness in legs and thighs, chiefly gelt at night. At 2pm. temp. was 99. At 2-30pm. Tartar Emetic, grs.l.was injected, slight pain and cough followed and at 3-15pm., rigor: at 6pm. Temp. was 102 and Pulse 130 pulse weak and very low tension: at 9pm. temp. was 100: through the night he slept well.
- 12. Patient still complained of pains in limbs: Knee jerks were absent.

 Atoxyl was injected, but reduced to grs.12.
- 13_ Patient comfortable.
- 14. Towards evening he complained of pains in the limbs.

 Bact. Report: No trypanosomes found.
- 15. Bact. Report: Trypanosomes were present in blood.

Patient felt rather seedy: Temp. 100 at 10am. Pulse 90. Tartar Emetic injected at 2pm. No rigor followed: At 6pm. Temperature as 101. Through the night he perspired.

- 16. Atoxyl grsz lt injected.
- 17. Patient falt better.

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- 18. Tartar Emetic, grs.l.was injected at 3pm., the Temp. being 99.2; there was no pain, cough or rigor afterwards: Temp. at 6pm. was 101.4: Pulse 108.
- 19. Atoxyl grs.la injected. Bact. Report: No trypanosomes found in blood.

Differential count of blood was as follws:-

Polymorph 63% Large mononuclear 9% Lymphosytes 27% Eosinophils 1%

Several pigmented mononuclear seen.

Patient having complained of dimness of vision, Oculist reported on condition of eyes as follows:- "The fundus of each eye "appears to be quite normal. The discs are a good colour and the dimness of vision is due to an error of refraction".

- 20. Patient comforable and eating well.
- 21. Felt sleepy and tired.
- 22. He did not feel well, but complained of nothing definite, only wanting to be left alone. Tartar Emetic, grs.l. was injected at 2pm. Temp. was 100: at 6pm. Temp. was 103: Pulse 96.

 Patient had a restless night.
- 23. Patient was easier in the morning. Atoxyl grs. 12 injected.
- 24. Patient was much better.

Physical emaminations-

HEART: No adventitious sounds. Right border 1 to right of Mid-Sternum. Left border in line with left nipple.

PULSE: 94, of low tension.

LUNGS: Normal.

LIVER: Normal.

SPLEEN: Palpable 1 inch.

GLANDS: One gland palpable and slightly tender behind each Sterno-Mastoid; no enlarged gland in axilla or groin. GENERAL CONDITION: poor: decided loss of weight as evidenced by wasting of muscles and subcutaneous tissue in arms, legs and back.

No rash present: No localised oedema:

ABDOMEN: showed slight gaseous distension: no fluid present.

NERVOUS SYSTEM: Tactile sense - good: Hot and cold sense good.

No deep hyeraesthesia present: No pains complained of

in calves of legs or thighs: Knee jerks absent:
No Ankle clonus: No Babinski: Eye reflexes normal:

Urine - no albumen, no sugar.

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of bile.

- 25. At 10am. patient had a rigor, Temp. reaching 101: Tartar Emetic grs.1. was given at 2pm.
 - Bact. Report: Trypanosomes were present in blood.
 - 26. Patient comfortable. Atoxyl grs.lt injected.
 - 27. Patient complained of pain in left ear. No swelling was present in front or behind the ear and there was no discharge.
- -28. He felt seedy and still complained of pain in the left ear- on examination nothing abnormal was detected.
- Patient had a severe rigor \(\frac{1}{12} \) hour afterwards and at \(\frac{12}{12} \) me. temp. was 104,8: Pulse 130 and weak: In the afternoon he was retching and vomiting the vomit was acid, slightly bilious but there was no blood: At llpm. patient was very collapsed, pulse thready and uncountable; Strychnia and Digitalis were given hypodermically, and alcohol in the form of Champagne and Brandy by the mouth.
- 30. Stimulants had to be continued and intravenous saline was given in the morning and in the afternoon rectal saline with Brandy.

 Throughout the day vomiting continued, the vomit consisting chiefly
- Hiccough commenced in the early morning and continued throughout the day; during the morning patient vomited once: tongue was thickly coated with fur; motions were loose and there was incontinence of the bowels: otherwise his general appearance was better: At 6pm. Temp. was 98 and Pulse 100: Pain was still complained of in the left ear: Mixture of Bismuth and Salol was given.

- 2. Hiccough stopped: patient felt sickly but otherwise better:

 Tongue was very furred: Bowels were loose and incontinent.

 Atoxyl grs.li injected. Bact. Report:- Trypanosomes were present in blood in large numbers.
- 3. Bowels still loose and incontinent: Stomach felt uncomfortable, patient feeling sickly but he did not womit.
- 4. Bact. Report: Trypanosomes found in blood. Patient was rather drowsy: No sickly feeling: In each groin three glands were palpable and slightly tender.
- 5. Atoxyl grs.2.injected. Patient was drowsy and not eating well.
- 6. Still drowsy and no desire for food.
- 7. It was proposed to patient to administer Tartar Emetic again; this had been previously hinted at, but on both occasions patient refused to have it.
- 8. Atoxyl grs.2.injected.

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9. Tartar Emetic grs.22 given per Rectum; this was only retained half an hour.

Patient had been apathetic and listless ever since the big reaction to Tartar Emetic, but on being informed that he was to be shortly transferred to South Africa where his home was, he immediately seemed to take on a new lease of life, being brighter and more cheerful and was quite confident he would be alright once he arrived there.

- 10. Oedematous swelling about the ankles was noticed and fluid was present in the abdominal cavity: A mixture containing Digitalis and Diuretic was given.
- 11. Bact. Report: Trypanosomes were found in the blood.
- 12 & 13. Patient is bright and more interested in things going on in the Ward.
- 14 & 16. Atoxyl grs.2.injected.
 - 17. Oedema about ankles is a little better: no appreciable alteration in amount of fluid in abdomen.
 - Patient was transferred to Hospital Ship for transfer to S.A.

 Patient died two months later in South Africa, the chief cause being cardiac failure, as evidenced by increase of oedema.

eg. 10/1918. Patient died.

PULSE PER MINUTE MINUTE ... Day of Disease, Dates of Observation. Temperature, Fabrenheit. Corps. Motions. 1000 1040 1010 1020 1030 1050 1060 Disgase rypanotomicadio S Time, Time, Time. 5 Time. Time. or Time. Time. Time. 16 Date of Admission 10-3-18 Date of Discharge Time. Rank and Name & Sanderson GW. (To be attached to the case sheet,) CLINICAL CHART. Time. Time. Time. 19 Time. • Result Time. 20 Time. 68 Hospital Station 48 Time. 2 641 0 Pime. Time. 28 A. F. B181 15-2-10. Gratis.

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S.A. Road Corps.

Previous to 1896 patient was a Sailor, when he had occasional attacks of Malaria, and once a venereal sore, which soon healed and no internal treatment was necessary.

colony or the Transvaal but he was never in Rhodesia. During this time he kept quite well. Patient served in the German West Campaign.

and proceeded to Port Amelia in the middle of March, going inland to Maheba. Early in April he had a slight attack of Malaria and he was treated with Quinine.

but did not feel any bite and does not know the date. His present illness dates from May 6. when he began to feel seedy, developing severe pains in all his joints and being feverish; he thought it was an attack of Malaria, but oh the whole the pains were more severe this time than he had experienced previously on May 10th. he was admitted to hospital at Port Amblia.

Whilst here patient remembers a Tsetse fly settling on his face.

12 & 13. Blood examined microscopically for parasites, proved Negative.

Trypanosomes found in blood in large numbers.

He was given Liq. Arsenicalis, m, iii, three times a day after food.

Patient was transferred to Base Hospital, Dar-es-Salaam per
Hospital Ship and was admitted to 84 British General Hospital on
25th. May, 1918.

Examination on Admission.

Patient states he feels alright, except that he has no energy and could not do any work, being easily tired.

General conditions Rather stout, debilitated and anaemic.

Tongue: Furred at back and on protuding a fine tremor is

present.

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15.

Skin: No rash is present on body: no local oedema:

Glands: about the size of an almond are felt in the posterior

triangle of the neck on the left side; no tenderness is complained

of.

10. Smaller glands are palpable in the right posterior triangle. The left epitrochlear and left inguinal glands are enlarged. The glands were of medium 200 consistence. dia i Nervous System: Memory is poor for recent events: Pupil reflexes 18 Normal: Knee jerks Normal: No ankle clonus: No Babinski; No deep hyperaesthesia: epicritic sense and sense to hot and cold are normal. · 」、1996年,1998年(1997年)(宋成城家)等數數(Heart: Fatty, sounds being soft and muffled: Arteries sclerosed. Pulse 100, medium tension. 20. Spleen: Palpable 3", no tenderness. Palpable 1 tender : The out of the back Liver: Urine: No Albumen, No sugar. Treatment and Effects. Lass lap normal size 24. M. 8. J. J. Atoxyl gr.3. was given intramuscularly in gluteal region. 8 An attempt was made to give Tartar Emetic gr.1. dissolved in normal Saline by direct intravenous injection, but no prominent vein could be obtained and there was no certainty the needle was in the vein, so the solution was given per rectum. I lease sais 11. This was continued twice weekly until June 11., and was then 8 discentinued as the Trypanosomes were not being controlled. 18. A vein was dissected out at the wrist and Tartar Emetic, gr.1. dissolved in Normal Saline was given intravenously; 15 minutes afterwards patient had a rigor and complained of severe pain in the number region; } hour later he vomited and 4 hours after the injection he was very collapsed. 13. rationt had rallied somewhat but was still sickly; evidence of 13 stomatitis was present, the tongue being swollen and thickly 29. coated with fur, and swelling of the lip was present. Atoxyl grs. 3. wes injected. The obligation so the effect. 1 Bacteriological Report: No trypanosomes found in blood. He was still sickly but he felt better in himself: Stomatitis 14. rather worse. 15. Still sickly and no desire for food; Tartar Emetic gr.1. was give intravenously - no pain or collapse afterwards.

- 16. Patient for 2 days had complained of pains in limbs; the knee jerks were found to be absent so Atoxyl was reduced to grs.l.
- 17. Still sickly and stomatitis troublesome. The same sectoriological Report: Trypanosomes present.
- 18. Patient not so well; sickly; Pulse 110; tension low. Tension
- 19. Mouth rather better although lip was swollen and sore.

 Tartar Emetic, grs.l. given intravenously there was a slight riger one hour afterwards, also pain in bottom of back was complained of, but this soon passed off.
- Patient comfortable; eating better. Atoxyl grs.lf injected
 † hour afterwards slight rigor with retching and slight pain in
 back. 2 hours afterwards he was comfortable.

 Bact. Report: Trypanosomes present.
- 21. Fairly comfortable. Complained of pains in legs. Lip normal size and mouth alright.
- 22. Tartar Emetic, gr.1. injected; followed by a slight rigor 20 minutes afterwards, retching and slight pain in back; in 2 hours after injection patient was comfortable.
- 23. Patient comfortable and eating well. Atoxyl grs.l injected.

 Bact. Report: To Trypanesomes present.
- 24 & 25. Keeping quite well.

d

- 26. Tartar Emetic, gr.l.injected rigor & hour after intravenous injection, but only slight pain in back complained of and there was no Gastro-intestinal irritation.
- Patient is comfortable and eating well. Atoxyl grs.l injected.

 Bact: Report: No Trypanosomes present, but Benign Tertian

 parasites found. Quinine Hydrochlor. grs.5 given every 3 hours.

 Comfortable.
- 2 cc. of solution of Antimon Oxide was injected subcutaneously a hours afterwards felt nausea, but otherwise no ill effect.
- Bact: Report: Trypanosomes and Benign Tertian parasites present in blood.

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Case 3.

549802. SAPPER R. MONK.

R.E. Telegraphs.

No.549802. Sapper R. M O N K., R.E. Telegraphs.

Age. 20. Service 9 months.

Disease. TRYPANOSOMIASIS.

en.Hosp. essalam. 13/18. Patient landed in East Africa on October 1st., 1917 and was stationed at Daressalam until March 21st. when he proceeded to Port Amelia. On April 1st. he went to Ankuabe, about 40 miles inland.

Patient did not experience any sickness until the end of April. Whilst im Camp at Meza, during the first night he was unable to use his mosquito net properly, being obliged to lie in the open on the ground. The following day he had no appetite and felt feverish; the next day he was better: on the third day he suffered from severe frontal headache and giddiness, he felt feverish and on reporting sick he was admitted to Hospital. He was treated as Malaria and discharged in the second week of May.

Patient was sent on duty to a station 10 miles beyond Meza - a small stream and a pond were about 250 yards from the Camp.

Plenty of flies were present during the day, but he never saw a Tsetse fly and was never bitten, it was cold at night and there were few mosquitoes. For 100 yards around, the Camp had been cleared of bush.

After his discharge from hospital patient kept well for five weeks, except for one night when he was a little feverish and had pains in head; he took Aspirin, grs.5. and on the following morning felt alright.

Whilst here patient ran alternate days on a motor bicycle to nkuabe, 25 miles distant; he never saw any flies.

On one journey in the middle of June, he had to stop and repair the mathine and in doing so made a skin deep cut on the first finger of the beft hand. At the time he paid no attention to this but in the evening felt stiff in the elbow and armpit.

On the following day he reported sick at the Camp Hospital and the arm was treated with fomentations. He was detained four days at the Camp Hospital; at night he was feverish and unable to sleep, patient felt a burning sensation all over body, had pains in the left arm and severe pain in the

head.

13.

- 22. He was transferred to Ankuabe where fomentations were continued. Whilst at Ankuabe patient ate solid food well and does not remember being feverish.
- 27. Patient was transferred to 19th. Stationary Hospital, Port

 Amelia chart shows subsequent course of Pyrexia.

 Blood examination negative.
- 28. " Benign Tertian.

Patient was treated with Quinine - no effect on Temperature and temperature chart did not show tertian periodicity.

- Blood examination Trypanosomes present.
 - "Has small palpable glands in triangles of neck, in both axillae" and in groins".
- 5. "Small irregular patches of superficial erythema were visible on "chest wall".

DIET.		TREATMENT.	1994(2)
Boof Toa.	July 4-7.	Q. Hydrochl. grs.30.	i.
Jelly.	* 8.	Q.H.grs.30. Tabs. Soamin gr.1	•
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Patient vomited occasionally - generally after a dose of Quinine or Mag. Sulphate.

Apart from being feverish he felt alright whilst at Port Amelia.

- He was transferred to H.M.H.S.*Dongola* and disembarked at an aressalam on July 13th., being admitted to 84th. British General Hospital.
 - On admission patient complains of no pain and is comfortable.

 Apatite variable, sometimes good and at other times poor.

 He says he has lost weight lately.
 - General Condition. Poor physique. Anaemia slightly: No localized oedema: No rash detected: body fairly well nourished: Muscles flabby. In both posterier triangles of neck enlarged glands are detected; the glands are about the size of a coffee bean, of medium consistence and are not tender, although the patient says they

axilla, but a gland the size of an almond is present in the left axilla - this gland is tender. Glands are enlarged in both groins in the Transverse group - those in the left are not tender, but

tenderness is present in glands of right grain.

have been tender. No gland is present in the right

Epitrochlear glands are not palpable.

Memory rather poor lately.

Heart No adventitious sounds: nor enlargement.

Pulse 96: low tension.

Lungs. Normal.

Spleen Palpable hlaf inch: tender.

Liver Normal.

Nervous system. Patient has noticed his memory has been poor lately. Deep hyperaesthesia is a marked symptom - tapping him over the shin bones causing him to wince; this is present also in the upper limbs, but to a lesser degree.

Knee reflexes - active.

Plantar Reflex - slightly more active than normal:
no Babinski present.

Ankle clonus - not present.

Testing tactile sense as judged by stroking the skin with a piece of cotton wool - this was absent below the Tibial Tuberosities, but over the rest of the body it was present and in some places the was sensation produced/excessive, causing local twitching of the muscles.

Hot and cold test - normal.

Testing with pricking with 2 needles held apart.

Sensation was less in the lower limbs - taking the axis transversely to the limbs he appreciated two points at a distance of $1\frac{1}{4}$ and in the length of the limb it varied from 2 to $2\frac{1}{4}$.

Eye reflexes - normal. No nystagmus present.

- Ty 13. Patient was comfortable and had no complaint.

 Bacteriological Report: Trypanosomes present in blood: No
 - Malarial parasites.
 - 14. Comfortable.
 - 15. Tartar Emetic grs.l. injected intravenously at 9-30p.m. no immediate effects apart from complaining of pain in arm and headache. Temperature rose from 101 to 104: towards evening he was quite comfortable.
 - 16. Atoxyl grs.3. injected intramuscular.

 Bacteriological Report: No Trypanosomes.
 - Comfortable during the day but rather drowsy.
 - 17. Patient complains of feeling a little dizzy and is drawsy.

 18. Tartar emetic gr.l. given intravenous rigor # hour after injection
 - Temperature rose from 99 to 103: no complaint of pain afterwards.
 - 19. Atoxyl grs.3.injected intramuscular: Still drowsy.

 Batt. report: Trypanosomes present.
 - 20. Patient did not sleep very well on account of a burning sensation through body.
 - 21. Febrile: thirsty: pulse 112 per minute, weak and low tension:

 Memory bad: drowsy:
 - 22. Tartar Emeritic grs.l.given slight rigor 20 minutes afterwards.
 otherwise comfortable.
 - 23. Atoxyl grs.3.injected. Comfortable but slight numbness in legs complained of. Bact. report: Trypanosomes present.
 - 24. Knee jerks are slightly increased: Plantar reflexes are very active: Deep hyperaesthesia is not so marked: No rash detected: Patient is very emaciated: otherwise is comfortable and has no pain.
 - 25. Tartar Emetic grs.l.given rigor 3 hours after injection and patient complained of pain in shoulder and arm.
 - 26. Atoxyl grs.3.injected left arm is stiff at the elbow but there is no local swelling. Bact. Report: Trypanosomes present.
 - 428. Patient comfortable.
 - Tartar Emetic grs.l.administered 4 rigor 1 hour afterwards.

 He complained of no pain and felt comfortable apart from feverish-
 - Atoxyl grs.3.injected.
 - Bact. Report: Trypanosomes not present.
 - Patient comfortable.

- 1. Tartar Emetic grs.1. given- no rigor or discomfort afterwards.
- 2. Atoxyl grs.3.injected. Patient feels well and is eating well:

 Kerendels symptom less marked: Tactile sense improved: Knee

 jerks very active: Plantar reflex flexor and active: No rash.

 Bact. Report: Trypanosomes not found.
- & 4. Patient comfortable and taking food well.
 - 5. Tartar Emetic grs.1.given while giving the injection he complained of pain in the same shoulder: 20 minutes afterwards had rigor, temperature reaching 103.
 - 6. Atoxyl grs.3.given. Patient comfortable, pulse 90, tension low.

 Bact. Report: Trypanosomes present, but scanty.
 - 7. Tongue clean: General appearance good: Pulse 84: Heart Normal:

 Lungs Normal. He complains of a feeling of deadness in both feetfeet are normal in appearance: Plantar reflex active: deep
 hyperaesthesia is not elicited. Knee jerks active: No rash.
 - 8. Tartar Emetic grs.l.given no rigor afterwards and no pain.

 Seen by Col. Newham, R.A.M.C. who recommended increasing Tartar

 Emetic to grs.2.
 - 9. Atoxyl grs.3.injected. Patient comfortable.

 Bact: report: Trypanosomes not frank present.
- 10 &11.Patient comfortable.
 - 12. Tartar Emetic grs.2.given immediately afterwards patient had a slight irritating cough: 15 minutes later rigor; Temperature rising to 104.8: Vomited soon after rigor; later complained of frontal headache.
 - 13. Atoxyl grs.3.injected. Bact. Report: Trypanosomes present.
 - 14. Pulse 99: tension improving, not quite so low as when admitted:
 No pain: eating well.
 - 15. Tartar Emetic grs.2. given severe rigor 2 hours afterwards:

 Temperature rose to 105; patient complained of pain in front

 of head.
 - Atoxyl grs.3.injected. Bact. Report: Trypanosomes present.
 - 17. Knee reflexes active: Plantar reflex flexor: Epicritic sense normal in legs: No deep hyperasthesia.
 - 18. Patient comfortable: Pulse 96, moderate tension.
 - 19. Tartar Emetic grs.2.given immediately after injection a short irritable cough commenced and lasted 2 minutes: No rigor.

- 20. Atoxyl grs.3.injected. Patient comfortable.

 Bact. Report:- No Trypanosomes found.
- 21. Patient comfortable.

& 25.Patient comfortable.

- 22. Tartar Emetic grs.2. given Cough commenced immediately following injection and continued 5 minutes: No rigor.
- 23. Diarrhoea probably due to eating fruit.

 Atoxyl grs.3. injected. Bact. Report: No trypanosomes found.
- 26. Tartar Emetic grs.2.injected No rigor after injection.
- 27. Atoxyl grs.3.given. Bact. Report: Trypanosomes present.
- 28. Patient complained of pain in head. Temperature rose in the evening to 103.2. Glands palpable in right and left posterior triangles and in right and left groin; they vary in size from a pea to an almond and are slightly tender: gland in left axilla is barely palpable and is not tender: no epitrochlear glands felt: Spleen is not palpable. Bact. Report: Trypanosomes present.
- Patient more comfortable, no headache, Temperature at 10a.m. 100.

 Tartar Emetic grs.2 given A slight cough after injection:

 rigor 1 hour afterwards: he did not complain of any pain.
- 50. Atoxyl grs.3.injected.

Nervous system - physical examination.

No deep hyperaesthesia: Epicritic sense not quite as good as on Aug 17/18, but much better than when admitted. Hot and cold test good. Tests of appreciation of two sharp points similar result as when admitted: Knee jerks active: No ankle clonus: Plantar reflex - flexor.

- on front and top of head. Temperature rose to 101.
- 1. Patient comfortable. Bact. Report: No Trypanosomes found.
- 2. Tartar Emetic grs.2 given Severe fit of coughing immediately followed injection: No other effect.
- 3. Atoxyl grs.3. injected.
- 4. Patient comfortable. Bact. Report: No Trypanosomes found.
- Tartar Emetic grs.22 given No rigor, but temperature rose to 102. Other manifestations in connection with the nervous system, e.g. deep hyperaesthesia and epicritic sense &c.,

were practically normal.

- 6. Atoxyl grs.3.injected. Bact. Report: Many Trypanosomes present
- 7. Patient comfortable and eating well.
- 8. Tartar Emetic grs.2 injected: No rigor but temperature rose to 102.5. and patient complained of frontal headache.

Blood film taken before injection showed Trypanosomes present.

Patient was evacuated per Hospital Ship for transferrence to England. His condition was greatly improved as compared with his condition on admission, July 13/18. This improvement was most marked as regards his mental outlook, the patient being brighter and more interested in himself and his surroundings. Other manifestations in connection with the nervous system, e.g. deep hyperaesthesia and epicritic sense &c., were practically normal. His general appearance was much better, being fuller in the face and body, and he had certainly gained weight. The tension of the pulse was improved as compared with the tension on admission. No cedema was present and no rash.

ADDENDA.

Monk was transferred to the Seamen's Hospital, London. He was doing all right until he contracted influenza with pneumonia in the beginning of November, 1918 and he died in 4 days time.

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TRYPANOSONIASIS

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TRYPANOSOMIASIS.

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TRYPANOSONIASIS.

CHART OF CASE D. (NATIVE).

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PULSE PER MINUTE RESPIRATIONS PER MINUTE ... Day of Disease, Dates of Observation. Fahrenheit. Corps -1030 Motions 1040 970 100 101 1050 1060 1070 1020 Time. Time. Time. Time. Time. Time. Time. Time. Date of Admission 56/1 Date of Discharge Rank and Name (To be attached to the case sheet.) CLINICAL CHART, Time. Time. 6 Time. Time. Result Time. Time. 0 6 Пше. Hospital Station 0 Time. Time. ادو Time. Time. Time. A. F. P. 1814.

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CHART OF CASE E. (NATIVE).

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TRYPANOSOMIASIS.

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CHART OF CASE F. (NATIVE).

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