

sex of the calves was reported. There was no significant deviation of the sex-rate from that of the corresponding control group.

In another series the centrifuge was run at 1,100 r.p.m., and the rate of flow (16 ml./min.) of the suspension liquid was so high that only one-twentieth to one-fifteenth of the spermatozoa were retained in the centrifuge. They showed very low motility (over-ripeness⁹?). For this reason, and on account of the scantiness of spermatozoa, only 24 inseminations were performed with this material. Of the cows 12 became pregnant, all belonging to different herds. One cow had to be slaughtered later. The 11 calves were all males. In the corresponding control group of 9 calves, 3 were males and 6 females.

The last series of experiments is being repeated.

I am very much indebted to the Artificial Insemination Association of Enköping for allowing me to perform these experiments, and to the chief veterinarian, Dr. T. Svensson, and to Dr. K. Idla for most valuable co-operation. The counter-streaming centrifuge was run by Mr. P.-Å. Lindström. Financial support from Ely Lilly and Co. is gratefully acknowledged.

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Blood-Histamine and Tissue-cell Anoxia in Mental Disease

WORK in our laboratories, *inter alia*, has been concerned for some years with the aberrant morphology¹ and physiological responses²⁻⁴ of the peripheral capillary vascular system of patients with schizophrenia and other mental disturbances. More recently, we have been concerned with the effects of these responses of the minute vasculature on the oxygenation of the adjacent tissue cells. In brief, we have found that, under conditions of increased capillary pressure, lessened local blood-oxygen saturation and capillary corpuscular flow (often with intravascular erythrocytic agglutination), hydration, swelling, vacuolization, granularity and other degenerative changes reminiscent of cellular cloudy swelling occur progressively in the intervascular tissue substance⁵. These changes reverse themselves when the vascular conditions seemingly causing them also suffer a reversal. Although these changes have been observed from time to time in healthy subjects, they are much more characteristically seen, and for much longer periods of time, in psychiatric patients, and especially in psychotics. Also, these vascular and cellular responses seem to coincide in time with significant happenings in the mental state of the patients and the dynamic variations occurring in their state of consciousness (for example, a worsening of the pathophysiology takes place when hallucinations are experienced⁶ or when a myoclonic seizure⁷ occurs in an epileptic, while an improvement in these reactions coincides with recovery or subsidence of these events).

Our interest in the mechanism whereby the cell could conceivably be affected by a diminution in the supply of molecular oxygen in the capillaries led to a study of the histamine metabolism of psychiatric patients. Using the Lowry *et al.*⁸ column

chromatographic microchemical method of estimating blood histamine (modified by trebling the concentration of DNFB per aliquot) and 5-ml. blood samples obtained by venupuncture, we investigated both mentally healthy and mentally sick populations all in the non-fasting state. In the former ($N = 12$) we found a mean blood histamine-level of 88.86 γ per litre ($s = 27.43$), that is, very close to the mean of 82.0 γ per litre for 'true' histamine reported by Lowry *et al.* and the figure of 86 γ per litre obtained by Valentine *et al.*⁹ from 16 non-fasting healthy subjects using Code's¹⁰ modification of the original Barsoum and Gaddum¹¹ bioassay technique. In the latter population the mean levels were invariably greater, those for schizophrenia ($N = 29$) being 106.38 γ ($s = 24.84$) and contrasting with those for epilepsy ($N = 5$) at 139.42 γ per litre ($s = 24.57$), while those for other psychiatric conditions ($N_T = 66$) lay between these two values. An analysis of variance showed the differences between the means to be significant ($P = 0.01$) and no significance could be attached to the possible variables of age and sex.

If these results are confirmed, their repercussions on stress-theory and on the role of histamine metabolism in this theory will be considerable; as they stand they strengthen the growing evidence favouring the relationship between tissue anoxia, mental disease and histamine release. We are currently engaged in extending our experiments in a longitudinal fashion in an endeavour kinetically to correlate the precise vascular and tissue events in circumstances of changing concentrations of blood histamine. The results contained in this communication will be published in greater detail in specialist journals elsewhere.

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Detection of Xanthine Dehydrogenase Activity in Soluble Proteins of Rat Liver separated by Paper Electrophoresis

It has been shown by paper electrophoresis that xanthine dehydrogenase activity is bound to the globulin fractions of rat serum¹. Using differential centrifugation, it was demonstrated that xanthine oxidase and xanthine dehydrogenase activities are present in the supernatant fluid corresponding to the 'cell sap' containing the soluble proteins of the rat liver².

A large number of proteins are present in the 'cell sap' fraction of the rat liver, and they have recently been separated by paper electrophoresis³. We were