

Fibrinolysis in the central nervous system

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THE FREQUENT RESOLUTION of subarachnoid clots and fibrinous meningitis without the development of meningeal fibrosis or communicating hydrocephalus and the rapid liquefaction of subdural clots suggest that the spinal fluid or the meninges or both possess a competent fibrinolytic mechanism. Although numerous recent investigations¹⁻³ have been performed to study the effects of cerebrospinal fluid on the body's coagulation mechanism, surprisingly few reports are available on the fibrinolytic activity of the spinal fluid and the meninges.

Roberts and Astrup,⁴ in 1957, and Moltke,⁵ in 1958, showed that the dura and pia of various animals contained considerable fibrinolytic activator activity. In 1958, Albrechtsen and associates⁶ found that human cerebrospinal fluid has incomplete fibrinolytic activator activity. The authors are not aware of other reports of fibrinolytic activity in the central nervous system.

A number of drugs capable of enhancing or depressing systemic fibrinolytic activity have become available in recent years. These drugs have been useful in selected patients with systemic fibrinolytic abnormalities, but their effects on central nervous system fibrinolysis have not been clearly demonstrated.

This study was performed to further define the fibrinolytic activity of the spinal fluid and the meninges and to ascertain the possibility of altering this activity by the systemic administration of fibrinolytic enhancing or inhibiting drugs.

MATERIALS AND METHODS

Fibrinolytic activity was assayed by a modification of the fibrin plate method of Astrup and Mullertz.⁷ A homogenous layer of fibrin was formed in a petri dish by the addition of 100 units (National Institutes of Health units) of bovine thrombin to 10 ml. of buffered bovine fibrinogen (0.4% by weight). Heated plates were prepared in the same manner and then heated to 80° C. for fifteen minutes to inactivate any plasminogen accompanying the fibrinogen or thrombin. Streptokinase fibrin plates, heated and nonheated, were prepared by adding 100 units of streptokinase to the fibrinogen solution before thrombin precipitation. The test samples were incubated on both heated and nonheated fibrin plates and streptokinase fibrin plates for eighteen hours at 37° C. The resulting zones of lysis were measured in their two greatest diameters. The product of these measurements represented the fibrinolytic activity of the sample.

Cerebrospinal fluid and peripheral blood samples were obtained from 20 patients during lumbar myelography. Spinal fluid, 0.05 ml. aliquots, was placed on heated, nonheated, and streptokinase fibrin plates and incubated as described above. A euglobulin fraction was prepared from each blood sample by adding

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