Tetrahedron Letters No.9, pp. 985-988, 1966. Pergamon Press Ltd. Printed in Great Britain.

> THE NATURE OF CORPAVERINE Tetsuji Kametani and Kazumi Ohkubo Pharmaceutical Institute, School of Medicine Tohoku University No. 85, Kitayobancho, Sendai, Japan Richard H. F. Manske Dominion Rubber Co. Ltd., Research Laboratories, Guelph, Ontario, Canada (Received 14 December 1966)

In the previous papers(1,2) it has been reported that corpaverine consisted of two components, (-)-capaurine (I) and sendaverine (II).



The purpose of the present investigation was to study the relation between (-)-capaurine and sendaverine, eventually showing that corpaverine is a molecular compound (1:1) of both substances as shown by thermal analysis.

A mixture (ca. 10 mg.) of (-)-capaurine, m.p. 162<sup>o</sup>, and sendaverine, m.p. 134.5 - 135.5<sup>o</sup> (3), was used for thermal analysis

985

of two components system. Sintering and melting points were determined in a sulphuric acid-bath (Table 1). The samples we prepared as follows. After both specimens had been dissolved a small amount of chloroform, the solvent was distilled, and t resultant residue was dried at  $40^{\circ}$  for 24 hr. under reduced pressure.

## Table 1

## Sendaverine %

100 85.5 80 73 68 66 60 50 45 43 30 22 10 Sintering point 134.5° 123° 124° 126° 125° 127° 127° 134° 134° 134° 133° 133° 147° Melting point 135.5° 129° 127° 130° 132° 133° 132° 133° 136.5° 136° 138° 144° 158°

> The first transition level was observed at  $123 - 124^{\circ}$  and second one at  $133^{\circ}$ . Three melting point curves intersect with each other at one eutectic point and one transition point. T: intersection of the two sintering point curves occurs in the sition which corresponds to a molar equivalent (1:1) of (-)capaurine and sendaverine as is shown in Fig. 1. Furthermore recrystallization of a mixture (1:1) of (I) and (II) from met afforded corpaverine as colourless needles, m.p.  $136 - 138^{\circ}$  ( The optical rotation of corpaverine agrees satisfactorily wit that of the caluculated value determined as follows;

986



Fig. 1. - Sendaverine % against (-)-capaurine

130° 120° 30 20 100 90 80 70 60 50 40 10 0 371.42 299.36 1 capaurine Molecular ratio sendaverine

Caluculated  $\left[\alpha'_{1}\right]_{0}$  of corpaverine =  $-271^{\circ}$  x  $\frac{1}{1+0.806}$  +  $0^{\circ}$ 

= 
$$-150.1^{\circ}$$
 [Found:  $[\alpha]_D^{20} - 154.2^{\circ} (CHCl_3) (4)$ ]

These facts show that corpaverine is a molecular compound of (-)-capaurine and sendaverine, reminiscent of the somewhat analogous tetrahydroshobakunine (5).

\* (-)-capaurine: 
$$[\alpha]_{D}^{24}$$
 -271° (CHCl<sub>3</sub>) (6).

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- 3. This melting point was determined in a sulphuric acid-bath and uncorrected. When this sample was measured on a hotstage apparutus, it melted at  $139 - 140^{\circ}$ .
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