

(at C) and equal in intensity to the ethylenic hydrogen absorption (at A).

The NMR spectra of the hydrocarbons which might be designated as cycloheptatriene,⁶ 1,1,4-trimethylcycloheptatriene^{7a} and 1,1,3,4-tetramethylcycloheptatriene^{7b} or the corresponding bicyclic valence tautomers have also been obtained and indicate unequivocally the validity of the cycloheptatriene structures. In each case the ratio of aliphatic to ethylenic hydrogen absorption is that predicted for the monocyclic structure and there is no absorption due to bridge hydrogens.

(6) Kindly provided by Dr. H. L. Dryden, Jr.

(7) Prepared from eucarvone: (a) with sodium borohydride followed by dehydration; (b) with methyl lithium followed by dehydration.

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GONYLEPTIDINE

Sir:

Experiments done by one of us (C. E.) in Montevideo, Uruguay, on the cephalothoracic glands of a South American arachnid of the *Gonyleptidae* family to elucidate the biological significance of the secretion led to the discovery of a volatile antibiotic, named gonyleptidine.

To a drop of protozoan and bacterial culture was added a drop of the substance globally extracted from the secretory pore of the sacciform glands of these gonyleptidae. Cytolysis led us to think of the action being possibly due to proteolytic ferments, but microscopic observation of protozoa and bacteria—in a suspended drop on excavated slides, the bottom of which had a small disc of filter paper soaked in secretion—showed the antimicrobial effect at a distance by emanation. This gave evidence of a volatile antibiotic and not of proteolytic ferments.

Preliminary experiments showed strikingly the action of gonyleptidine on membrane, cytoplasm and nucleus of free cells. Afterwards (Saez and Drets) disturbance was seen in mitosis and meiosis. Isolation of gonyleptidine (M.I.A.) fully confirmed the presence of a volatile antibiotic and allowed quantitative determination of activity. Isolated by distillation from the frozen state, gonyleptidine was characterized as a yellow substance, m.p. 12°, $\lambda_{\text{Water}} 255 \text{ m}\mu$ ($E^{1\%}_{1\text{cm}}$ 1400), which gives color reactions characteristic of quinones. It was found (N.P.B.) effective against at least eighteen genera of bacteria and protozoa, for example, against six strains of *Staphylococcus aureus* at concentrations of 3 to 10 $\gamma/\text{ml.}$, various strains of *B. cereus*, *B. subtilis* or *B. anthracis* (2.6 to 64 $\gamma/\text{ml.}$), *Escherichia coli* strains (3 to 112 $\gamma/\text{ml.}$), *B. tuberculosis* (100 $\gamma/\text{ml.}$), *Trypanosoma cruzi* (100 $\gamma/\text{ml.}$). (C. E., O. Simani, and N. P. B.) Given orally to mice (1 mg./mouse/24 hr.) infected with intestinal parasites, the substance was tolerated perfectly and destroyed the giardias, trichomonas, and hexamites.

While biological studies were continued in Uruguay, a chemical investigation was undertaken at Harvard (L. F. F., M. I. A.). Reduction of

yellow aqueous extract with hydrosulfite, acetylation, and chromatography gave, as a derivative of the major but not the sole quinone component, a substance, m.p. 105–106° (C, 64.69; H, 6.46; acetyl, 38.02; mol. wt., 237), identified by mixed melting point determination as 2,3-dimethylhydroquinone diacetate. Polarographic analysis indicated the presence in gonyleptidine, in addition to a dimethylquinone or quinones ($E^{25\%}_0$ 0.588–590 v.), of a companion quinone of lower potential, and infrared analysis indicated 2,5-dimethyl- and 2,3,5-trimethyl-1,4-quinone to be the probable companions. Hence methods of fractionation were tested on mixtures of the synthetic models. Finally 115 mg. of gonyleptidine was treated at room temperature with 2,3-dimethylbutadiene for selective conversion of 2,3-dimethyl-1,4-quinone to an adduct. When the unreacted quinones were reduced with hydrosulfite and extracted from ether with alkali, the adduct remained in the neutral fraction and was identified as such and as 2,3,6,7-tetramethyl-1,4-naphthoquinone, m.p. 167°. After reoxidation of the hydroquinones, Thiele acetoxylation of the quinone mixture and steam distillation gave a non steam-volatile residue identical with 2,5-dimethyl-1,3,4-triacetoxybenzene (m.p. 107°; mixed m.p.; infrared) derived from 2,5-dimethyl-1,4-quinone, and a steam-volatile quinone identified as 2,3,5-trimethyl-1,4-quinone by ultraviolet and infrared spectra and mixed m.p. of the hydroquinone (m.p. 170°). The amounts of components accounted for (in the order just mentioned) were 71, 11, and 15 mg. Average activities against representative microorganisms in terms of multiples of the activity of gonyleptidine are (same order): Gram positive: -4, +2, +2; Gram negative: +4, +4, +2. Although a great many quinones have been assayed for bacteriostatic activity, these simple benzoquinones have gone neglected. Actually, data to be presented elsewhere show that they are considerably more promising, with respect to potency and retention of activity *in vivo*, than any quinones previously investigated. Acids resulting from addition of thioacetic and β -thio-propionic acid to the methylbenzoquinones and oxidation to the quinones also have interesting bacteriostatic properties.

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SYNTHESIS OF PERCHLOROFULVALENE¹

Sir:

The hydrocarbon fulvalene (I), as yet unsynthesized, has been postulated to be a stable compound possessing resonance energy amounting to about 41 kcal./mole.² 1,2,3,4-Tetraphenylfulvalene (II) was the first fulvalene derivative reported³

(1) The authors wish to express their appreciation to the Hooker Electrochemical Company for financial support of this investigation.

(2) R. D. Brown, *Trans. Faraday Soc.*, **46**, 146 (1950).

(3) E. C. Schreiber and E. I. Becker, *THIS JOURNAL*, **76**, 6125 (1954).