thesized from the glucose (evidence obtained from chemical analyses of glycogen, total carbohydrate and total fat content); (d) that the cell volume, measured by a centrifuge method, increases with time.<sup>5</sup> The optical density of the suspending medium did not change during 6 hours at  $25^{\circ}$  C. The evidence obtained indicates, therefore, that the increase in optical density of the suspension is primarily due to progressive changes in optical characteristics of the cells and that these changes accompany the formation and storage of anabolic products within the cells.

On the basis of the above results it is proposed that the changes in optical characteristics of veast cells during assimilation might be used as indicators of anabolic activity. Thus, the rate of increase in optical density of the cell suspension might be conveniently employed as a measure of the "rate of assimilation." A rapid, optical method for use in exploratory experiments dealing with the dynamic aspects of the assimilation process would obviously reduce the total time and effort demanded by a kinetic type of attack on the mechanism of carbohydrate and fat assimilation. The long, laborious chemical methods of analysis would still be employed, but only for each series of check experiments. Our present densitometer can be employed for this purpose within certain limits. It has been sufficiently reliable for the qualitative detection of whether or not assimilation is taking place in a given yeast suspension in which the total number of cells remains constant with time; it also has given a linear relation between the opacity of the suspension and the per cent. increase in dry weight of the cells during the early stages of carbohydrate assimilation (i.e., during about the first 2 hours at 25° C.). We hope to develop a more satisfactory method of measuring specifically the change in cellular opacity in order to follow with greater accuracy the entire course of carbohydrate and fat storage.

Details of the experiments reported in this paper will be published shortly.

THEODORE J. B. STIER M. ISABEL NEWTON HERBERT SPRINCE

BIOLOGICAL LABORATORIES, HARVARD UNIVERSITY

## THE USE OF PURIFIED THROMBIN AS AN HEMOSTATIC AGENT<sup>1</sup>

RECENTLY we have described a thrombin preparation which is far more powerful than any previously

 $^5$  Cf. O. W. Richards and T. L. Jahn, Jour. Bact., 26: 385–391, 1933; and L. G. Longsworth, *ibid.*, for discussions of the effect of an increase in cellular opacity and volume during growth by cell division on measurements of population density by the photoelectric method.

<sup>1</sup> Aided by a grant from The John and Mary R. Markle Foundation. Funds for a technical assistant were supplied by the Graduate College, State University of Iowa. reported.<sup>2</sup> In a 1 per cent. solution, 1 cc will clot 1 cc of blood within 2 seconds. Animal studies show that such a solution is highly effective when applied locally to check hemorrhage. It has been found to be non-toxic when used to control the oozing of blood from operative surfaces. For this purpose it is applied by means of an atomizer. Under these circumstances a thin film of blood forms almost instantly, and this seals the finer vessels. The larger vessels are controlled by ligature as a preliminary measure.

With the thrombin spray, profuse bleeding from incised liver tissue can be checked completely in 5 seconds or less. Prolonged bleeding from bone is often very troublesome to the surgeon at the time of operation. With the application of thrombin we are able to stop this bleeding within 5 to 10 seconds. This obviates the need of bone wax, which is a foreign body. With the use of thrombin, we have been able to resect portions of brain and to check with ease the hemorrhage which is otherwise so difficult to control.

The same beneficial effects have been obtained in operations on dogs with a prothrombin deficiency which can be produced by chronic chloroform administration.<sup>3</sup> In such dogs, bleeding is very profuse at operation and may be fatal unless thrombin is applied.

The important applications to human surgery and to human bleeders await further study regarding the sterility of the thrombin preparations.

> W. H. SEEGERS E. D. WARNER

K. M. BRINKHOUS

H. P. SMITH

11. F. SMIT

DEPARTMENT OF PATHOLOGY, STATE UNIVERSITY OF IOWA

## ISOLATION OF CRYSTALLINE HETERO-TRYPSIN FROM BEEF PANCREAS

NORTHROP and Kunitz have prepared from beef pancreas the two crystalline enzymes, trypsin and chymotrypsin.<sup>1</sup> For both of these enzymes, specific substrates have been synthesized in this laboratory. Trypsin was shown to hydrolyze benzoyl arginine amide, while chymotrypsin was found to split benzoyl tyrosyl glycine amide. Another synthetic substrate, benzoyl glycyl lysine amide, was found to be hydrolyzed by crude pancreatic extracts, but not by trypsin or chymotrypsin.<sup>2</sup> Therefore, the hydrolysis of this lysine compound was attributed to an unknown enzyme

<sup>2</sup> W. H. Seegers, H. P. Smith, E. D. Warner and K. M. Brinkhous, *Jour. Biol. Chem.*, 123: 751, 1938; *ibid.*, 126: 91, 1938.

91, 1938. <sup>3</sup> H. P. Smith, E. D. Warner and K. M. Brinkhous, *Jour. Exp. Med.*, 66: 801, 1937.

<sup>1</sup> M. Kunitz and J. H. Northrop, Jour. Gen. Physiol., 19: 991, 1936.

<sup>2</sup> M. Bergmann and W. F. Ross, *Jour. Am. Chem. Soc.*, 58: 1503, 1936; M. Bergmann, J. S. Fruton and H. Pollok, SCIENCE, 85: 410, 1937.