Isotopic Composition of Boron and its Atomic Weight

AFTER an extensive investigation, we conclude that a boron-11/boron-10 ratio of 4.00 (that is, 80/20) approaches the true natural abundance ratio much more closely than the presently 'accepted' value of 4.31 (81.17/18.83). The latter figure is based on a mass spectrometric investigation of boron trifluoride¹, a notoriously troublesome compound, in view of its adsorption characteristics resulting in instrumental 'memory'. Since publication of that work a number of attempts to verify the abundance ratio from boron trifluoride have resulted in a wide range of values.

On the other hand, abundance ratios calculated by various investigators from mass spectra of boron hydrides have been highly consistent, yielding ratios of approximately $4 \cdot 00$. Lest it be argued that this lower value may be peculiar to boron hydrides, we would like to point out that recent work in this laboratory with other compounds (for example, trimethylboroxine) has yielded the same ratio.

We therefore propose that in the future it would be far more reasonable to use the ratio 4.00 (80/20) to represent the natural isotopic abundance of boron. This would require in addition a revision of the atomic weight from 10.82 to 10.811, a change which seems well within the range of experimental error of earlier investigators².

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¹ Inghram, M., Phys. Rev., 70, 653 (1946).

² German Atomic Weight Commission, Ber. deutsch. chem. Ges., 56A, xvii (1923); 57A, xvi (1924). International Committee on Chemical Elements, J. Amer. Chem. Soc., 47, 597 (1925).

BIOCHEMISTRY

Synthesis of Oxytocin by the Nitrophenyl Ester Method

AFTER the completion of work on the degration of oxytocin¹, this compound was synthesized in this laboratory². Since that time several other synthetic methods leading to biologically active solutions of the crude hormone have been reported³. Recently, an improved synthesis of oxytocin was elaborated⁴ by us. All these syntheses of oxytocin have the common feature of linking peptides together. It is well known, however, that formation of the peptide bond with a protected peptide as the acid component can lead to racemization⁵, the extent of which depends on the components, the conditions and the method. No racemization was observed with the use of the azide method⁶. Nevertheless, the latter procedure can involve the formation of by-products other than diastereoisomers. To avoid racemization it is possible to take advantage of the protecting groups, for example, the carbobenzoxy group, which protects the amino-acid acylated by it not only against unwanted acylation but also against racemization. This protection is effective only if the carbobenzoxy group is joined to the amino-acid, the carboxyl group of which takes part in the peptide bond-forming reaction. Therefore it becomes necessary to start the synthesis with the C-terminal amino-acid, to link to it the next (protected) aminoacid, to remove the protecting group from the resulting dipeptide derivative, to couple to the free amino component thus formed the third (protected) aminoacid, etc. This manner of peptide-chain synthesis is practicable only if the method used for the formation of the peptide bond is unambiguous, leads to only one peptide, and gives high yields. All the mixed anhydride methods were eliminated as they usually

Cbz -Leu-O ϕ NO ₂ (2) Cbz -Pro-O ϕ NO ₂
H-Gly-OEt Cbz-Leu-Gly-OEt
(1) $HBr/AcOH$
$\mathbf{M}_{3}/\mathbf{M}_{6}\mathbf{OH} \qquad (2) \ \mathbf{U}_{5}\mathbf{Z}\cdot\mathbf{S}\cdot\mathbf{B}\mathbf{Z}\cdot\mathbf{U}_{9}\mathbf{S}\cdot\mathbf{O}\phi\mathbf{N}\mathbf{O}_{2}$
Cbz-Pro-Leu-Gly-OEt
(1) HB/ACOH (2) CONTACT (2)
Chz-S-Bz-Cys-Pro-Leu-Gly(NH)
(1) HBr/AcOH
(2) Cbz -Glu(NH ₂)O φ NO ₂
$Cbz-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \longrightarrow$
Bz
(1) $HBr/AcOH$
(2) Cbz-fleu-O ϕ NO ₂
$Coz-Glu(NH_2)$ -Asp (NH_2) -Cys-Pro-Leu-Gly (NH_2)
P-
132
(1) HBr/AcOH (2) Chz-O-Bz-Tyr-OcNO
$Cbz-Ileu-Glu(NH_{\circ})-Asp(NH_{\circ})-CvS-Pro-Leu-Glv(NH_{\circ}) \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-O\phi NO_{2} \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-OADA (2) Cbz-O-Bz-Tyr-OADA (2) Cbz-O-Bz-Tyr-OADA (2) Cbz-O-Bz-Tyr-OADA (2) Cbz-O-Bz-Tyr-O-BZ-DA (2) Cbz-O-Bz-Tyr-O-BZ-DA (2) Cbz-O-BZ-DA (2) Cbz-O-BZ-DA (2) Cbz-DA (2) Cbz-DA (2) Cbz-DA (2) C$
$Cbz - Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(2) Cbz-O-Bz-Tyr-O\varphi NO_2} \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(2) Cbz-O-Bz-Tyr-O\varphi NO_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O-Bz-Tyr-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O-Bz-Tyr-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-D_2} \xrightarrow{(2) Cbz-D_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-D_2} (2) Cbz-D_$
$Cbz-Ileu-Glu(\mathbf{NH}_2)-\mathbf{Asp}(\mathbf{NH}_2)-CyS-\mathbf{Pro-Leu-Gly}(\mathbf{NH}_2) \xrightarrow{(1) \ \mathbf{HBr}/AcOH}_{(2) \ Cbz-O-Bz-Tyr-O\phiNO_2}$
$Cbz-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(2) Cbz-O-Bz-Tyr-O\varphi NO_2} \xrightarrow{(1) HBr/AcOH^*} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(2) Cbz-O-Bz-Tyr-O\varphi NO_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O-Bz-Tyr-O\varphi NO_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-O-Bz-Tyr-O_2} \xrightarrow{(2) Cbz-D_2} $
$Cbz-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(2) Cbz-O-Bz-Tyr-O\varphi NO_2} (1) HBr/AcOH^* (2) Cbz-S-Bz-Cys-O\varphi NO_2 \xrightarrow{(2) Cbz-S-Bz-Cys-O\varphi NO_2} \xrightarrow{(2) Cbz-S-Bz-Cys-O\varphi ND_2} \xrightarrow{(2) Cbz-S-Bz-Cys-Az} \xrightarrow{(2) Cbz-S-Bz-Cys-Az} \xrightarrow{(2) Cbz-S-Bz-Cys-Az} \xrightarrow{(2) Cbz-Az} (2$
$Cbz-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(2) Cbz-O-Bz-Tyr-O\varphi NO_2} \xrightarrow{(1) HBr/AcOH^*} (2) Cbz-S-Bz-Cys-O\varphi NO_2 \xrightarrow{(2) Cbz-S-Bz-Cys-O\varphi NO_2} \xrightarrow{(2) Cbz-S-Bz-Cys-O\varphi ND_2} \xrightarrow{(2) Cbz-S-Bz-Cys-Dy ND_2} \xrightarrow{(2) Cbz-S-Bz-Cys-Dy ND_2} (2)$
$Cbz-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \xrightarrow{(1) HBr/AcOH} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(2) Cbz-O-Bz-Tyr-O\varphi NO_2} (2) Cbz-O-Bz-Tyr-O\varphi NO_2 \xrightarrow{(1) HBr/AcOH^*} (2) Cbz-S-Bz-Cys-O\varphi NO_2 \xrightarrow{(2) Cbz-S-Bz-Cys-O\varphi NO_2} \xrightarrow{(2) Cbz-S-Bz-Cys-O\varphi ND_2} \xrightarrow{(2) Cbz-S-Bz-Cys-O\varphi ND_2} \xrightarrow{(2) Cbz-S-Bz-Cys-Oy-S-Bz-Cys-Dy ND_2} \xrightarrow{(2) Cbz-S-Bz-Cys-Oy-S-Bz-Cys-Dy ND_2} (2) Cbz-S-$
$\begin{array}{c} (1) & \text{HBr/AcOH} \\ (2) & \text{Cbz-O-Bz-Tyr-O\phiNO}_2 \\ & & \\ Bz \\ \\ Cbz-Tyr-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \\ & & \\ Bz \\ \\ & & \\ Bz \\ \\ & & \\ Bz \end{array} \xrightarrow{(1) & \text{HBr/AcOH}^* \\ (2) & \text{Cbz-S-Bz-Cys-O\phiNO}_2 \\ \hline \\ & & \\ (2) & \text{Cbz-S-Bz-Cys-O\phiNO}_2 \\ \hline \\ & & \\$
$\begin{array}{c} \begin{array}{c} (1) & \text{HBr/AcOH} \\ (2) & \text{Cbz-O-Bz-Tyr-O\phiNO_2} \end{array} \\ \\ (3) & \text{Cbz-O-Bz-Tyr-O\phiNO_2} \end{array} \\ \\ (4) & \text{Cbz-Tyr-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2)} \end{array} \\ \\ (5) & \text{Cbz-Tyr-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2)} \end{array} \\ \\ \\ \begin{array}{c} (1) & \text{HBr/AcOH}^{\bullet} \\ (2) & \text{Cbz-O-Bz-Tyr-O\phiNO_2} \end{array} \\ \\ (2) & \text{Cbz-S-Bz-Cys-O\phiNO_2} \end{array} \\ \\ \\ \end{array} \\ \\ \begin{array}{c} (1) & \text{HBr/AcOH}^{\bullet} \\ (2) & \text{Cbz-S-Bz-Cys-O\phiNO_2} \end{array} \\ \\ \\ \end{array} \\ \\ \end{array} \\ \\ \end{array} \\ \\ \end{array} \\ \\ \begin{array}{c} (1) & \text{HBr/AcOH}^{\bullet} \\ (2) & \text{Cbz-S-Bz-Cys-O\phiNO_2} \end{array} \\ \\ \\ \end{array} \\ \\ \end{array} \\ \\ \end{array} \\ \\ \begin{array}{c} (1) & \text{HBr/AcOH}^{\bullet} \\ (2) & \text{Cbz-S-Bz-Cys-O\phiNO_2} \end{array} \\ \\ \\ \end{array} \\ \\ \end{array} \\ \\ \end{array} \\ \\ \end{array} $
$\begin{array}{c} (1) \ \text{HBr/AcOH} \\ (2) \ \text{Cbz-O-Bz-Tyr-O\phiNO}_2 \\ \\ (3) \ \text{Cbz-O-Bz-Tyr-O\phiNO}_2 \\ \\ (4) \ \text{Cbz-O-Bz-Tyr-O\phiNO}_2 \\ \\ (5) \ \text{Cbz-O-Bz-Tyr-O\phiNO}_2 \\ \\ (6) \ \text{Cbz-CyS-Tyr-Ileu-Glu(NH_2)-CyS-Pro-Leu-Gly(NH_2)} \\ \\ (6) \ \text{Cbz-CyS-Tyr-Ileu-Glu(NH_a)-Asp(NH_a)-CyS-Pro-Leu-Gly(NH_a)} \\ \end{array}$
$\begin{array}{c} (1) \ \text{HBr/AcOH} \\ (2) \ \text{Cbz-O-Bz-Tyr-O\phiNO}_2 \\ \\ Bz \\ \\ Cbz-Tyr-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \\ \\ Bz \\ \\ \\ Cbz-CyS-Tyr-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2) \\ \\ \\ (2) \ Cbz-S-Bz-Cys-O\phiNO_2 \\ \\ (2) \ Cbz-S-Bz-Cys-O\phiNO_2 \\ \\ \\ \\ (2) \ O_2 \ (air) \\ \\ \\ \\ (2) \ O_2 \ (air) \\ \\ \\ \\ \\ \\ \\ (2) \ O_2 \ (air) \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\$
$\begin{array}{c} \begin{array}{c} (1) & \text{HBr/AcOH} \\ (2) & \text{Cbz-O-Bz-Tyr-O\phiNO}_2 \end{array} \\ \\ (3) & \text{Cbz-Cys-S-Bz-Cys-O\phiNO}_2 \end{array} \\ \\ (3) & \text{Cbz-Cys-S-Bz-Cys-O\phiNO}_2 \end{array} \\ \\ (4) & \text{Cbz-Cys-S-Bz-Cys-O\phiNO}_2 \end{array} \\ \\ (5) & \text{Cbz-Cys-S-Bz-Cys-O\phiNO}_2 \end{array} \\ \\ \\ \\ (5) & \text{Cbz-Cys-Tyr-Ileu-Glu(NH_2)-Asp(NH_2)-CyS-Pro-Leu-Gly(NH_2)} \end{array} \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ \\ $

(1) HBr/AcOH

Fig. 1