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LETTERS TO THE EDITOR

2,2,6,6-Tetra(hydroxymethyl)cyclohexanol in the Synthesis of New Types of Phospholipids

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Continuing studies on the methods of synthesis of nonglycerol polyol phospholipids, based on trivalent phoshorus compounds and phosphorus(V) cyclic systems [1, 2], we for the first time prepared phos-

pholipids derived from readily available 2,2,6,6-tetra-(hydroxymethyl)cyclohexanol (I). Cyclic polyol I was reacted with acetone in acidic medium to obtain isopropylidene derivatives II and III.

OH
OH
OH
OH
OH
OH
OH
OH
II
III

The yield of compound **II**, mp 92–93°C, R_f 0.48 [Silufol UV-254; hexane–dioxane, 5:1 (system A)], did not exceed 5%; therefore, in further syntheses of phospholipids we used compound **III** only. Protected polyol **III** was purified by triple crystallization from heptane, yield 65%, mp 111–112°C, R_f 0.38 (system A).

Diisopropylidene derivative **III** was phosphorylated with hexaethylphosphorous triamide (**IV**) at 90–100°C with removal of the evolving diethylamine at reduced pressure (380 mm). Crude phosphorodiamidite **V**, $\delta_{\rm p}$ 134.5 ppm, R_f 0.6 [hexane–dioxane, 10:1 (system B)], was converted to thio- and selenoamidophosphates **VI** and **VII** which were purified by crystallization from pentane.

Thioamidophosphate **VI**, yield 60%, mp 91–92°C, δ_P 79.3 ppm, R_f 0.4 (system B).

Selenoamidophosphate **VII**, yield 75%, mp 101–102°C, δ_P 80.2 ppm, R_f 0.4 (system B).

$$\mathbf{III} \xrightarrow{P(\operatorname{NEt}_2)_3} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{[X]} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{[X]} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{OP(\operatorname{NEt}_2)_2} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \\ O \\ O \end{array} \xrightarrow{V} \begin{array}{c} O \xrightarrow{\bullet} O \end{array} \xrightarrow{V$$

X = S (VI), Se (VII).

Phosphoacetals **VI**, **VII** were directly acylated with myristoyl and stearoyl chlorides at 20°C according to [3]. The yield of tetraacylamidophosphates **VIII**–**XI** after purification on a silica gel column (eluent hexane) was 55%.

Tetramyristoylthioamidophosphate VIII, mp 39–

$$R = C_{13}H_{27}, X = S \text{ (VIII)}; R = C_{13}H_{27}, X = Se \text{ (IX)}; R = C_{17}H_{35}, X = S \text{ (X)}; R = C_{17}H_{35}, X = Se \text{ (XI)}.$$

40°C, R_f 0.52 (system B); tetramyristoylselenoamidophosphate **IX**, mp 44–45°C; R_f 0.8 (system A); tetrastearoylselenoamidophosphate **XI**, mp 61–62°C; R_f 0.8 (system A).

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REFERENCES

- 1. Nifant'ev, E.E. and Predvoditelev, D.A., *Usp. Khim.*, 1994, vol. 63, no. 1, pp. 73–92.
- 2. Predvoditelev, D.A. and Nifant'ev, E.E., *Zh. Obshch. Khim.*, 1995, vol. 31, no. 12, pp. 1761–1785.
- 3. Nifant'ev, E.E. and Predvoditelev, D.A., *Usp. Khim.*, 1997, vol. 66, no. 1, pp. 47–56.