

Note

A new method for the synthesis of ureido sugars

Bogusława Piekarska-Bartoszewicz and Andrzej Temeriusz

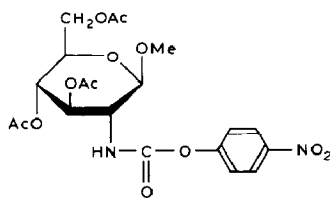
Department of Chemistry, Warsaw University, 02-093 Warsaw (Poland)

(Received September 16th, 1989; accepted for publication, December 8th, 1989)

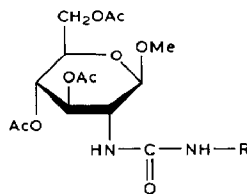
Nitrosoureas are an important class of antitumor agents¹. Analogues of streptozotocin (*N*-nitroso-*N*-methyl urea of 2-amino-2-deoxy-D-glucose)^{2,3} have been described and many other nitrosoureido sugars have been prepared⁴. Nitrosoureido sugars can be obtained easily by aqueous nitrosation of ureido sugars with sodium nitrite³. The most convenient method for the synthesis of ureido sugars involves the reaction of isocyanates with amino sugars⁵, but few isocyanates are available commercially.

Recently, Izdebski and Pawlak⁶ described a convenient new method for the synthesis of *N,N'*-disubstituted ureas and we have adopted this method for the synthesis of ureido sugars.

Bis(4-nitrophenyl) carbonate (1), obtained⁶ by the action of phosgene on sodium 4-nitrophenolate, was reacted in dichloromethane in the presence of 1-hydroxybenzotriazole with methyl 3,4,6-tri-*O*-acetyl-2-amino-2-deoxy-β-D-glucopyranoside⁷ to



2



3 R = Et

9 R = CH₂CH₂CH(CH₃)₂

4 R = Pr

10 R = CH(CH₃)CH₂CH₂CH₂CH(CH₃)₂

5 R = ⁱPr

11 R = cyclohexyl

6 R = Bu

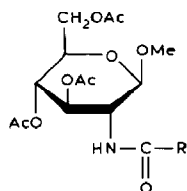
12 R = Bzl

7 R = ⁱBu

13 R = CH₂CH₂Ph

8 R = pentyl

14 R = allyl



15 R = NEt₂

18 R =

16 R = N(C₆H₁₃)₂

19 R =

17 R = N(C₆H₁₁)₂

19 R =

give the 4-nitrophenyl carbamate **2**. Reaction at room temperature was complete within 20 h and 90% of **2** was obtained.

Reaction of **2** in dichloromethane with 1 mol of an alkylamine then gave the alkylureido derivative in good yield. The structures of the ureido sugars **3–19** were confirmed by ^1H - and ^{13}C -n.m.r. spectroscopy.

EXPERIMENTAL

Melting points are uncorrected. Optical rotations were measured on a Perkin-Elmer Model 241 polarimeter. N.m.r. spectra (internal Me_4Si) were recorded with a Jeol FX 90Q (^1H : 90 MHz, ^{13}C : 22.53 MHz) spectrometer. T.l.c. was performed on Silica Gel 60 F₂₅₄ (Merck), using chloroform–acetone (4:1) and detection by u.v. light, or charring with sulfuric acid. Column chromatography was conducted on Silica Gel 60 (Merck 230–400 mesh) in chloroform–acetone (4:1).

Methyl 3,4,6-tri-O-acetyl-2-deoxy-2-(4-nitrophenoxycarbonylamino)- β -D-glucopyranoside (2). — To a solution of bis(4-nitrophenyl) carbonate⁶ (**1**; 0.91 g, 3 mmol) in dichloromethane (20 mL) were added a solution of methyl 3,4,6-tri-O-acetyl-2-amino-2-deoxy- β -D-glucopyranoside⁷ (0.96 g, 3 mmol) in dichloromethane (20 mL) and 1-hydroxybenzotriazole (0.40 g, 3 mmol). The mixture was stirred for 20 h at room temperature; t.l.c. then revealed no **2**. The mixture was diluted with dichloromethane (50 mL), washed successively with water, M ammonia, water, M hydrochloric acid, and water, then dried, and concentrated. Column chromatography of the residue and recrystallization from chloroform–hexane yielded **2** (1.31 g, 90%), m.p. 178–180°, $[\alpha]_{\text{D}}^{20} + 35^\circ$ (c 1, chloroform). N.m.r. data (CDCl_3): ^1H , δ 8.24 (d, 2 H, J 9 Hz, Ph), 7.32 (d, 2 H, J 10 Hz, Ph), 5.37 (d, 1 H, J 10 Hz, NH), 5.21–5.19 (m, 2 H, H-3,4), 4.68 (d, 1 H, J 8 Hz, H-1), 4.42–4.19 (m, 2 H, H-6a,b), 3.81–3.64 (m, 2 H, H-2,5), 3.56 (s, 3 H, OMe), 2.09, 2.05 (2 s, 9 H, 3 Ac); ^{13}C , δ 170.78, 170.67, 169.42 (CH_3COO), 155.50 (NHCOO) 152.63, 144.93, 125.10, 121.90 (Ph), 101.26 (C-1), 71.84 (C-3,5), 68.43 (C-4), 61.93 (C-6), 57.21 (OMe), 56.67 (C-2), 20.75, 20.64 (CH_3OO).

Anal. Calc. for $\text{C}_{20}\text{H}_{24}\text{N}_2\text{O}_{12}$: C, 49.59; H, 4.99; N, 5.78. Found: C, 49.66; H, 5.06; N, 6.02.

Methyl 3,4,6-tri-O-acetyl-2-(3-alkylureido)-2-deoxy- β -D-glucopyranoside. — To a solution of **2** (1 mmol) in dichloromethane (10 mL) was added a solution of alkylamine (1 mmol) in dichloromethane (10 mL), and the mixture was stored for 5–24 h at room temperature. T.l.c. then revealed no **2**. The mixture was diluted with dichloromethane (20 mL), washed successively with water, M ammonia, water, M hydrochloric acid, and water, then dried, and concentrated. The residue was subjected to column chromatography. The following compounds were prepared in this manner.

Methyl 3,4,6-tri-O-acetyl-2-deoxy-2-(3-ethylureido)- β -D-glucopyranoside (**3**), methyl 3,4,6-tri-O-acetyl-2-deoxy-2-(3-propylureido)- β -D-glucopyranoside (**4**), methyl 3,4,6-tri-O-acetyl-2-deoxy-2-(3-isopropylureido)- β -D-glucopyranoside (**5**), methyl 3,4,6-tri-O-acetyl-2-deoxy-2-(3-butylureido)- β -D-glucopyranoside (**6**), methyl 3,4,6-tri-O-

TABLE I

Physico-chemical and analytical data for 3-19

Compound	Yield (%)	M.p. (degrees)	$[\alpha]_D^{20}$ (c 1, chloro- form) (degrees)	Formula	Analytical data					
					Calc.			Found		
					C	H	N	C	H	N
3	65	116-118	+7.2	$C_{16}H_{28}N_2O_9$	49.23	6.71	7.18	49.14	6.75	7.07
4	70		+5.2	$C_{17}H_{28}N_2O_9$	50.49	6.98	6.93	50.45	7.07	6.88
5	72	152-154	+3.2	$C_{17}H_{28}N_2O_9$	50.49	6.98	6.93	50.50	7.04	7.11
6	75		+5.8	$C_{18}H_{30}N_2O_9$	51.67	7.23	6.69	51.63	7.29	6.65
7	80	123-125	+6.0	$C_{18}H_{30}N_2O_9$	51.67	7.23	6.69	51.61	7.28	6.96
8	89		+5.7	$C_{19}H_{32}N_2O_9$	52.77	7.46	6.48	52.67	7.46	6.37
9	85		+7.9	$C_{19}H_{32}N_2O_9$	52.77	7.46	6.48	52.54	7.58	6.61
10	92		+3.1	$C_{20}H_{32}N_2O_9$	55.68	8.07	5.90	55.76	8.12	6.19
11	86	176-178	+4.5	$C_{20}H_{32}N_2O_9$	54.04	7.26	6.30	54.13	7.24	6.28
12	91		+6.2	$C_{21}H_{34}N_2O_9$	55.75	6.24	6.19	55.52	6.27	6.24
13	82	127-128	+8.2	$C_{22}H_{36}N_2O_9$	56.64	6.48	6.00	56.70	6.45	5.92
14	56		+7.9	$C_{17}H_{28}N_2O_9$	50.74	6.51	6.96	50.79	6.64	7.07
15	83		+10.8	$C_{18}H_{30}N_2O_9$	51.67	7.23	6.69	51.67	7.34	6.80
16	86	102-104	+16.5	$C_{26}H_{46}N_2O_9$	58.85	8.74	5.28	58.79	8.87	5.30
17	90	156-157	+12.8	$C_{26}H_{46}N_2O_9$	59.30	8.04	5.32	59.21	8.02	5.28
18	86	154-155	+16.7	$C_{19}H_{30}N_2O_9$	53.08	7.02	6.51	53.01	6.89	6.49
19	85	179-180	+16.9	$C_{18}H_{28}N_2O_9$	51.92	6.78	6.73	51.99	6.60	6.58

TABLE II

¹³C-N.m.r. data for 3-19 (CDCl₃, internal Me₄Si)

Compound	C-1	C-2	C-3	C-4	C-5	C-6	OMe	CH ₃ COO	CH ₃ COO	N-CO-N	Remaining carbon atoms
3	103.21	56.02	71.84	68.97	73.09	62.31	57.00	170.99, 170.67, 169.48	20.75	157.55	35.22, 15.33
4	103.21	56.02	71.68	68.75	72.93	62.20	57.11	170.99, 170.72, 169.48	20.75, 20.64	158.31	42.10, 23.30, 11.27
5	103.21	56.02	71.79	68.86	73.03	62.25	56.99	170.88, 170.61, 169.88	20.70, 20.59	157.56	42.10, 21.24
6	103.16	55.91	71.79	69.08	73.14	62.42	57.00	170.83, 170.61, 169.48	20.70, 20.54	158.42	40.15, 32.24, 20.00, 13.76
7	103.05	55.70	71.68	69.13	73.20	62.47	57.00	170.78, 170.67, 169.53	20.70, 20.59	158.64	47.79, 28.93, 19.99, 18.37
8	103.05	55.53	71.73	69.13	73.30	62.47	57.00	170.78, 170.67, 169.48	20.70	158.80	46.05, 35.32, 26.93, 22.43, 17.07
9	103.10	55.81	71.73	69.19	73.14	62.47	57.00	170.72, 170.61, 169.42	20.60	158.48	39.06, 38.68, 25.79
10	103.05	55.59	71.73	69.35	73.30	62.63	56.94	170.53, 169.61	20.70	157.99	46.00, 39.01, 37.65, 27.96, 23.89, 22.59
11	103.27	56.08	71.79	68.92	73.03	62.36	56.99	170.93, 170.67, 169.48	20.75, 20.64	157.56	48.92, 33.70, 25.68, 24.92
12	103.00	55.53	71.63	69.13	73.14	62.47	56.94	170.67, 169.42	20.64	158.53	139.79, 128.41, 127.00, 126.89, 43.09
13	103.10	56.08	71.73	68.86	72.93	62.25	57.00	170.99, 170.66, 169.42	20.70, 20.59	158.09	139.24, 128.79, 126.35, 41.61, 36.19
14	103.00	55.26	71.68	69.51	73.25	62.74	57.11	170.61, 170.40, 169.58	20.70	158.75	135.94, 114.70, 42.48
15	102.45	55.64	71.89	69.24	73.14	62.52	56.62	170.78, 170.61, 169.37	20.70	156.36	41.34, 13.71
16	102.40	55.91	71.90	69.74	72.81	62.47	56.73	170.61, 169.81	20.64	156.63	47.46, 31.64, 28.50, 26.60, 22.59, 13.98
17	102.24	55.32	71.79	69.30	72.65	62.47	56.78	170.67, 170.61, 170.40	20.70	156.41	56.18, 31.80, 31.47, 26.44, 25.52
18	102.40	55.81	71.79	69.08	73.09	62.47	56.67	170.88, 170.67, 169.42	20.75	156.74	45.13, 25.63, 24.49
19	102.40	55.53	71.79	69.03	73.30	62.42	56.62	170.89, 170.67, 169.42	20.70	156.93	45.57, 25.52

TABLE III

¹H-N.m.r. data (δ in p.p.m.)^a for 3-19

3	5.97–5.78 (m, 2 H, 2 NH), 5.01 (t, 1 H, $J_{2,3} = J_{3,4} = 10$ Hz, H-3), 4.81 (dd, 1 H, $J_{3,4} 10$, $J_{4,5} 9.6$ Hz, H-4), 4.52 (d, 1 H, $J_{1,2} 8$ Hz, H-1), 4.18–4.08 (m, 2 H, H-6a,6b), 3.72–3.55 (m, 2 H, H-2,5), 3.33 (s, 3 H, OMe), 2.96–2.70 (m, 2 H, CH ₂), 2.01, 1.96, 1.90 (3 s, 9 H, 3 Ac), 0.96 (t, 3 H, $J 8$ Hz, CH ₃).
4	5.92 (d, 1 H, $J 12.7$ Hz, NH), 5.76 (m, 1 H, NH), 5.09 (t, 1 H, $J_{2,3} = J_{3,4} = 10$ Hz, H-3), 4.80 (dd, 1 H, $J_{3,4} 10$, $J_{4,5} 9.4$ Hz, H-4), 4.52 (d, 1 H, $J_{1,2} 8$ Hz, H-1), 4.17–4.06 (m, 2 H, H-6a,6b), 3.97–3.75 (m, 2 H, H-2,5), 3.38 (s, 3 H, OMe), 2.93–2.75 (m, 2 H, NCH ₂), 2.03, 1.97, 1.90 (3 s, 9 H, 3 Ac), 1.39–1.19 (m, 2 H, CH ₂), 0.81 (t, 3 H, $J 7$ Hz, CH ₃).
5	5.79 (d, 1 H, $J 8$ Hz, NH), 5.69 (d, 1 H, $J 8.4$ Hz, NH), 5.10 (dd, 1 H, $J_{2,3} = J_{3,4} = 10.5$ Hz, H-3), 4.82 (t, 1 H, $J_{3,4} 10.5$, $J_{4,5} 9.5$ Hz, H-4), 4.52 (d, 1 H, $J_{1,2} 8.4$ Hz, H-1), 4.37–4.06 (m, 2 H, H-6a,6b), 3.93–3.57 (m, 2 H, H-2,5), 3.34 (s, 3 H, OMe), 2.95 (m, 1 H, CH), 2.02, 1.96, 1.89 (3 s, 9 H, 3 Ac), 0.99 (d, 6 H, $J 6.6$, 2 CH ₃).
6	5.99–5.92 (m, 1 H, NH), 5.82 (d, 1 H, $J 8.6$ Hz, NH), 5.14 (t, 1 H, $J 9.4$, $J 9.8$ Hz, H-3), 4.84 (t, 1 H, $J 9.4$ and 9.8 Hz, H-4), 4.55 (d, 1 H, $J_{1,2} 8.4$ Hz, H-1), 4.20–4.08 (m, 2 H, H-6a,6b), 3.95–3.78 (m, 2 H, H-2,5), 3.34 (s, 3 H, OMe), 2.92 (m, 2 H, NCH ₂), 2.02, 1.96, 1.84 (3 s, 9 H, 3 Ac), 1.35–1.24 (m, 4 H, 2 CH ₂), 0.86 (t, 3 H, $J 5.6$ Hz, CH ₃).
7	5.91 (m, 1 H, NH), 5.75 (d, 1 H, $J 8.2$ Hz, NH), 5.11 (dd, 1 H, $J_{2,3} 10$, $J_{3,4} 9.5$ Hz, H-3), 4.80 (t, 1 H, $J_{3,4} = J_{4,5} = 9.5$ Hz, H-4), 4.50 (d, 1 H, $J 8.5$ Hz, H-1), 4.22–4.10 (m, 2 H, H-6a,6b), 4.00–3.47 (m, 2 H, H-2,5), 3.34 (s, 3 H, OMe), 2.85–2.52 (m, 2 H, CH ₂), 2.02, 1.96, 1.88 (3 s, 9 H, 3 Ac), 1.71–1.42 (m, 1 H, CH), 0.79 (d, 6 H, $J 6.6$ Hz, 2 CH ₃).
8	5.80 (m, 1 H, NH), 5.75 (d, 1 H, $J 8.5$ Hz, NH), 5.10 (t, 1 H, $J_{2,3} = J_{3,4} = 10$ Hz, H-3), 4.80 (dd, 1 H, $J_{3,4} 9.4$, $J_{4,5} 10$ Hz, H-4), 4.51 (d, 1 H, $J_{1,2} 8$ Hz, H-1), 4.17–4.06 (m, 2 H, H-6a,6b), 3.88–3.51 (m, 2 H, H-2,5), 3.34 (s, 3 H, OMe), 3.10–2.67 (m, 2 H, CH ₂), 2.01, 1.96, 1.89 (3 s, 9 H, 3 Ac), 1.37–1.15 (m, 4 H, 2 CH ₂), 0.89–0.80 (m, 5 H, CH ₂ , CH ₃).
9	5.09–5.72 (m, 1 H, NH), 5.76 (d, 1 H, $J 9.4$ Hz, NH), 5.09 (dd, 1 H, $J_{2,3} 10$, $J_{3,4} 9.4$ Hz, H-3), 4.80 (t, 1 H, $J_{3,4} = J_{4,5} = 9.4$ Hz, H-4), 4.51 (d, 1 H, $J_{1,2} 8.5$ Hz, H-1), 4.23–4.05 (m, 2 H, H-6a,6b), 3.95–3.45 (m, 2 H, H-2,5), 3.34 (s, 3 H, OMe), 3.03–2.81 (m, 2 H, NCH ₂), 2.01, 1.96, 1.89 (3 s, 9 H, 3 Ac), 1.70–1.20 (m, 5 H, CH, CH ₂), 0.84 (d, 6 H, $J 6$ Hz, 2 CH ₃).
10	5.67 (d, 1 H, $J 9.4$ Hz, NH), 5.09 (dd, 1 H, $J_{2,3} = J_{3,4} = 10$ Hz, H-3), 4.80 (t, 1 H, $J_{3,4} = J_{4,5} = 10$ Hz, H-4), 4.52 (d, 1 H, $J_{1,2} 8.5$ Hz, H-1), 4.51 (d, 1 H, $J 8.6$ Hz, NH), 4.50–4.06 (m, 2 H, H-6a,6b), 3.78–3.64 (m, 3 H, H-2,5, NCH), 3.35 (s, 3 H, OMe), 2.02, 1.97, 1.90 (3 s, 9 H, 3 Ac), 1.51–1.11 (m, 7 H, CH, 3 CH ₂), 0.97 (d, 3 H, $J 6.6$ Hz, CH ₃), 0.85 (d, 6 H, $J 6.6$ Hz, 2 CH ₃).
11	5.83 (d, 1 H, $J 10$ Hz, NH), 5.72 (d, 1 H, $J 10$ Hz, NH), 5.11 (dd, 1 H, $J_{2,3} 10$, $J_{3,4} 9.4$ Hz, H-3), 4.82 (dd, 1 H, $J_{3,4} 9.4$, $J_{4,5} 10.3$ Hz, H-4), 4.52 (d, 1 H, $J_{1,2} 8.5$ Hz, H-1), 4.26–4.07 (m, 2 H, H-6a,6b), 3.97–3.48 (m, 3 H, H-2,5, CH), 3.36 (s, 3 H, OMe), 2.04, 1.98, 1.91 (3 s, 9 H, 3 Ac), 1.72–1.17 (m, 10 H, 5 CH ₂).
12 ^b	7.25 (m, 5 H, Ph), 5.18–4.94 (m, 3 H, H-3,4, NH), 4.40 (d, 1 H, $J_{1,2} 8.5$ Hz, H-1), 4.37–4.16 (m, 5 H, H-6a,6b, CH ₂ , NH), 3.46 (s, 3 H, OMe), 2.07, 2.00, 1.98 (3 s, 9 H, 3 Ac).
13	7.28–7.22 (m, 5 H, Ph), 5.97–5.88 (m, 2 H, 2 NH), 5.11 (dd, 1 H, $J_{2,3} 9.9$, $J_{3,4} = 9.4$ Hz, H-3), 4.88 (t, 1 H, $J_{2,3} = J_{4,5} = 9.4$ Hz, H-4), 4.52 (d, 1 H, $J_{1,2} 8$ Hz, H-1), 4.29–4.08 (m, 2 H, H-6a,6b), 4.29–4.09 (m, 2 H, H-2,5), 3.35 (s, 3 H, OMe), 3.30–3.15 (m, 2 H, NCH ₂), 2.72–2.58 (t, 2 H, $J 7$ Hz, CH ₂), 2.02, 1.98, 1.90 (3 s, 9 H, 3 Ac).
14	6.09–5.63 (m, 3 H, 2 NH, =CH), 5.12 (dd, 1 H, $J 10$ and 9.4 Hz, H-3), 5.03, 5.01 (2 d, 2 H, $J 8.5$ Hz, =CH ₂), 4.82 (dd, 1 H, $J 10$ and 9.4 Hz, H-4), 4.54 (d, 1 H, $J_{1,2} 8.5$ Hz, H-1), 4.31–4.07 (m, 2 H, H-6a,6b), 3.97–3.50 (m, 4 H, H-2,5, CH ₂), 3.37 (s, 3 H, OMe), 2.04, 1.99, 1.92 (3 s, 9 H, 3 Ac).
15	6.15 (d, 1 H, $J 9$ Hz, NH), 5.19 (t, 1 H, $J_{2,3} = J_{3,4} = 10$ Hz, H-3), 4.82 (dd, 1 H, $J_{3,4} 10$, $J_{4,5} 9.4$ Hz, H-4), 4.62 (d, 1 H, $J_{1,2} 8$ Hz, H-1), 4.26–4.07 (m, 2 H, H-6a,6b), 3.85–3.62 (m, 2 H, H-2,5), 3.35 (s, 3 H, OMe), 2.03, 1.97, 1.88 (3 s, 9 H, 3 Ac), 0.98 (2 t, 6 H, $J 7.0$ Hz, 2 CH ₃).
16 ^b	5.42 (dd, 1 H, $J 8.4$ and 8.9 Hz, H-3), 5.05 (dd, 1 H, $J 8.4$ and 8.9 Hz, H-4), 4.71 (d, 1 H, $J_{1,2} 8.5$ Hz, H-1), 4.41 (d, 1 H, $J 8$ Hz, NH), 4.26–4.17 (m, 2 H, H-6a,6b), 3.80–3.55 (m, 2 H, H-2,5), 3.52 (s, 3 H, OMe), 2.08, 2.00 (2 s, 9 H, 3 Ac), 1.76–1.02 (m, 16 H, 8 CH ₂), 0.89 (t, 6 H, $J 4.7$ Hz, 2 CH ₃).

- 17^b** 5.54 (dd, 1 H, J 8.9 and 10.3 Hz, H-3), 5.05 (dd, 1 H, J 8.9 and 10.3 Hz, H-4), 4.86 (d, 1 H, $J_{1,2}$ 8 Hz, H-1), 4.37 (d, 1 H, J 8 Hz, NH), 4.27–4.08 (m, 2 H, H-6a,6b), 3.82–3.62 (m, 2 H, H-2,5), 3.52 (s, 3 H, OMe), 3.40–3.25 (m, 2 H, 2 CH₂), 2.08, 2.01 (2 s, 9 H, 3 Ac), 1.81–1.68 (m, 20 H, 10 CH₂).
- 18^b** 5.37 (dd, 1 H, $J_{2,3}$ 10, $J_{3,4}$ 9.4 Hz, H-3), 5.07 (t, 1 H, $J_{3,4} = J_{4,5} = 10$ Hz, H-4), 4.77 (d, 1 H, J 9 Hz, NH), 4.68 (d, 1 H, $J_{1,2}$ 8 Hz, H-1), 4.27–4.17 (m, 2 H, H-6a,6b), 3.81–3.63 (m, 2 H, H-2,5), 3.51 (s, 3 H, OMe), 3.33 (m, 4 H, 2 CH₂), 2.05, 2.02 (2 s, 9 H, 3 Ac), 1.56 (m, 6 H, 3 CH₂).
- 19^b** 5.33 (dd, 1 H, $J_{2,3}$ 10, $J_{3,4}$ 9.4 Hz, H-3), 5.07 (t, 1 H, $J_{3,4} = J_{4,5} = 9.4$ Hz, H-4), 4.39 (d, 1 H, J 10 Hz, NH), 4.65 (d, 1 H, $J_{1,2}$ 8 Hz, H-1), 4.26–4.17 (m, 2 H, H-6a,6b), 3.90–3.64 (m, 2 H, H-2,5), 3.51 (s, 3 H, OMe), 3.36–3.22 (m, 4 H, 2 CH₂), 2.09, 1.95 (2 s, 9 H, 3 Ac), 1.92–1.82 (m, 4 H, 2 CH₂).

^a For a solution in (CD₃)₂SO (internal Me₄Si). ^bFor a solution in CDCl₃.

acetyl-2-deoxy-2-(3-isobutylureido)- β -D-glucopyranoside (**7**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3-pentylureido)- β -D-glucopyranoside (**8**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3-isopentylureido)- β -D-glucopyranoside (**9**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-[3-(1,5-dimethylhexyl)ureido]- β -D-glucopyranoside (**10**), methyl 3,4,6-tri-*O*-acetyl-2-(3-cyclohexylureido)-2-deoxy- β -D-glucopyranoside (**11**), methyl 3,4,6-tri-*O*-acetyl-2-(3-benzylureido)-2-deoxy- β -D-glucopyranoside (**12**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3-phenethylureido)- β -D-glucopyranoside (**13**), methyl 3,4,6-tri-*O*-acetyl-2-(3-allylureido)-2-deoxy- β -D-glucopyranoside (**14**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3,3-diethylureido)- β -D-glucopyranoside (**15**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3,3-dihexylureido)- β -D-glucopyranoside (**16**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3,3-dicyclohexylureido)- β -D-glucopyranoside (**17**), methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3-piperidinoureido)- β -D-glucopyranoside (**18**), and methyl 3,4,6-tri-*O*-acetyl-2-deoxy-2-(3-pyrrolidinoureido)- β -D-glucopyranoside (**19**). The yields and structural data for these compounds are shown in Tables I–III.

REFERENCES

- 1 R. J. Weinkam and H. S. Lin, *Adv. Pharmacol. Chemother.*, **19** (1982) 1–33.
- 2 J. J. Vavra, C. DeBoer, A. Dietz, L. J. Hanka, and W. T. Sokolski, *Antibiot. Annu.*, 1959/1960, (1960) 230–235; R. R. Herr, T. E. Eble, M. E. Bergy, and H. K. Jahnke, *ibid.*, 1959/1960, (1960) 236–240; R. R. Herr, H. K. Jahnke, and A. D. Argoudelis, *J. Am. Chem. Soc.*, **89** (1967) 4808–4809; E. Hardegger, A. Meier, and A. Stoss, *Helv. Chim. Acta*, **52** (1969) 2555–2560.
- 3 E. J. Hessler and H. K. Jahnke, *J. Org. Chem.*, **32** (1970) 245–246.
- 4 P. Roger, C. Monneret, J. P. Fournier, P. Choay, R. Gagnet, C. Gosse, Y. Letourneux, G. Atassi, and A. Gouyette, *J. Med. Chem.*, **32** (1989) 16–23, and references therein.
- 5 T. Suami and T. Machinami, *Bull. Chem. Soc. Jpn.*, **43** (1970) 3013–3015.
- 6 J. Izdebski and D. Pawlak, *Synthesis*, (1989) 423–425.
- 7 J. C. Irvine, D. McNicoll, and A. Hynd, *J. Chem. Soc.*, **99** (1911) 250–261.