

Chemoenzymatic Syntheses and Anti-HIV-1 Activity of Glucose-Nucleoside Conjugates as Prodrugs

Tatiana Rodríguez-Pérez,[†] Susana Fernández,[†] Yogesh S. Sanghvi,[‡] Raymond F. Schinazi,[§] Vicente Gotor^{†,*} and Miguel Ferrero^{†,*}

[†]Departamento de Química Orgánica e Inorgánica and Instituto Universitario de Biotecnología de Asturias, Universidad de Oviedo, 33006-Oviedo (Asturias), Spain

[‡]Rasayan Inc., 2802 Crystal Ridge Road, Encinitas, CA 92024-6615, USA.

[§]Emory University School of Medicine/Veterans Affairs Medical Center, Decatur, Georgia 30033, USA.

ELECTRONIC SUPPLEMENTARY INFORMATION (page 1 of 153 pages)

Contents:

Copies of ¹H, ¹³C, DEPT, and some 2D NMR spectra of compounds **3**, **5**, **7**, **9**, and **10**:

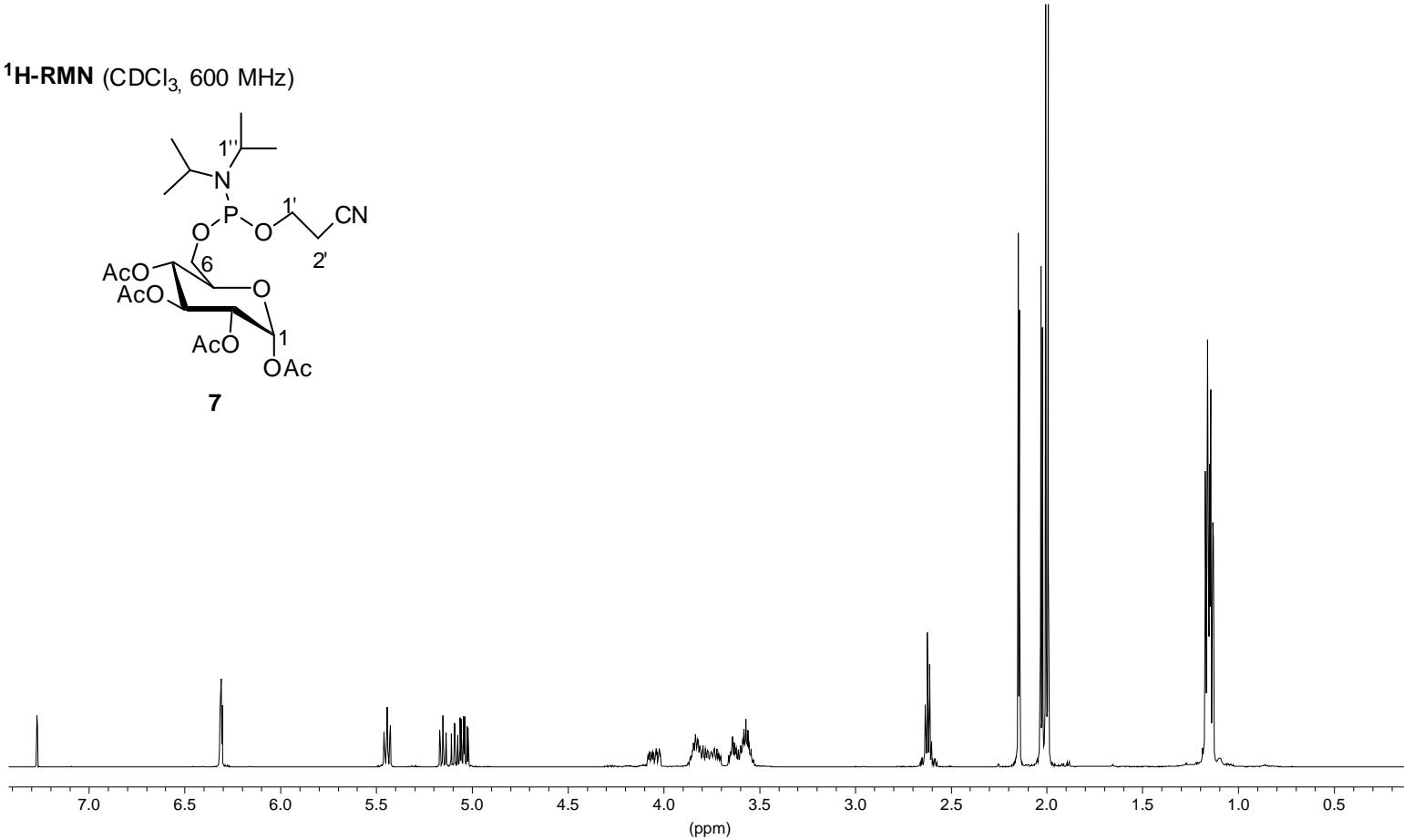
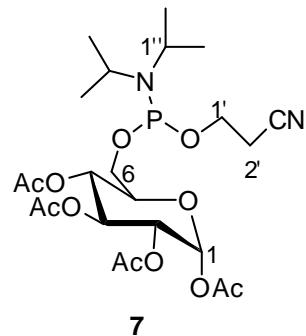
- ¹H NMR of **7** (S4)
- ¹³C NMR of **7** (S5)
- DEPT NMR of **7** (S6)
- ³¹P NMR of **7** (S7)
- COSY of **7** (S8)
- HSQC of **7** (S9)
- HMBC of **7** (S10)
- NOESY of **7** (S11)
- ¹H NMR of **8c** (S12)
- ¹³C NMR of **8c** (S13)
- DEPT NMR of **8c** (S14)
- COSY of **8c** (S15)
- ¹H NMR of **8d** (S16)
- ¹³C NMR of **8d** (S17)
- DEPT NMR of **8d** (S18)
- ¹H NMR of **8f** (S19)
- ¹³C NMR of **8f** (S20)
- DEPT NMR of **8f** (S21)
- ¹H NMR of **8g** (S22)
- ¹³C NMR of **8g** (S23)
- DEPT NMR of **8g** (S24)
- ¹H NMR of **8h** (S25)
- ¹³C NMR of **8h** (S26)
- DEPT NMR of **8h** (S27)

- ^1H NMR of **8i** (S28)
- ^{13}C NMR of **8i** (S29)
- DEPT NMR of **8i** (S30)
- ^1H NMR of **9a** (S31)
- ^{13}C NMR of **9a** (S31)
- DEPT NMR of **9a** (S33)
- ^{31}P NMR of **9a** (S34)
- ^1H NMR of **9b** (S35)
- ^{13}C NMR of **9b** (S36)
- DEPT NMR of **9b** (S37)
- ^{31}P NMR of **9b** (S38)
- COSY of **9b** (S39)
- HSQC of **9b** (S40)
- HMBC of **9b** (S41)
- ^1H NMR of **9c** (S42)
- ^{13}C NMR of **9c** (S43)
- DEPT NMR of **9c** (S44)
- ^{31}P NMR of **9c** (S45)
- COSY of **9c** (S46)
- HSQC of **9c** (S47)
- HMBC of **9c** (S48)
- ^1H NMR of **9d** (S49)
- ^{13}C NMR of **9d** (S50)
- DEPT NMR of **9d** (S51)
- ^{31}P NMR of **9d** (S52)
- COSY of **9d** (S53)
- HSQC of **9d** (S54)
- HMBC of **9d** (S55)
- ^1H NMR of **9e** (S56)
- ^{13}C NMR of **9e** (S57)
- DEPT NMR of **9e** (S58)
- ^{31}P NMR of **9e** (S59)
- COSY of **9e** (S60)
- HSQC of **9e** (S61)
- HMBC of **9e** (S62)
- ^1H NMR of **9f** (S63)
- ^{13}C NMR of **9f** (S64)
- DEPT NMR of **9f** (S65)
- ^{31}P NMR of **9f** (S66)
- ^1H NMR of **9g** (S67)
- ^{13}C NMR of **9g** (S68)
- DEPT NMR of **9g** (S69)
- ^{31}P NMR of **9g** (S70)
- COSY of **9g** (S71)
- HSQC of **9g** (S72)
- HMBC of **9g** (S73)
- ^1H NMR of **9h** (S74)
- ^{13}C NMR of **9h** (S75)
- DEPT NMR of **9h** (S76)
- ^{31}P NMR of **9h** (S77)
- COSY of **9h** (S78)
- HSQC of **9h** (S79)
- HMBC of **9h** (S80)
- ^1H NMR of **9i** (S81)
- ^{13}C NMR of **9i** (S82)
- DEPT NMR of **9i** (S83)
- ^{31}P NMR of **9i** (S84)
- HSQC of **9i** (S85)
- HMBC of **9i** (S86)
- NOESY of **9i** (S87)
- ^1H NMR of **10a** (S88)
- ^{13}C NMR of **10a** (S89)
- DEPT NMR of **10a** (S90)
- ^{31}P NMR of **10a** (S91)
- COSY of **10a** (S92)
- ^1H NMR of **10b** (S93)
- ^{13}C NMR of **10b** (S94)
- DEPT NMR of **10b** (S95)
- ^{31}P NMR of **10b** (S96)
- COSY of **10b** (S97)

- HSQC of **10b** (S98)
- ^1H NMR of **10c** (S99)
- ^{13}C NMR of **10c** (S100)
- DEPT NMR of **10c** (S101)
- ^{31}P NMR of **10c** (S102)
- COSY of **10c** (S103)
- ^1H NMR of **10d** (S104)
- ^{13}C NMR of **10d** (S105)
- DEPT NMR of **10d** (S106)
- ^{31}P NMR of **10d** (S107)
- COSY of **10d** (S108)
- ^1H NMR of **10e** (S109)
- ^{13}C NMR of **10e** (S110)
- DEPT NMR of **10e** (S111)
- ^{31}P NMR of **10e** (S112)
- COSY of **10e** (S113)
- ^1H NMR of **10f** (S114)
- ^{13}C NMR of **10f** (S115)
- DEPT NMR of **10f** (S116)
- ^{31}P NMR of **10e** (S117)
- COSY of **10f** (S118)
- ^1H NMR of **10g** (S119)
- ^{13}C NMR of **10g** (S120)
- DEPT NMR of **10g** (S121)
- ^{31}P NMR of **10g** (S122)
- ^1H NMR of **10h** (S123)
- ^{13}C NMR of **10h** (S124)
- DEPT NMR of **10h** (S125)
- ^{31}P NMR of **10h** (S126)
- COSY of **10h** (S127)
- ^1H NMR of **10i** (S128)
- ^{13}C NMR of **10i** (S129)
- DEPT NMR of **10i** (S130)
- ^{31}P NMR of **10i** (S131)
- COSY of **10i** (S132)
- ^1H NMR of **14** (S133)
- ^{13}C NMR of **14** (S134)
- DEPT NMR of **14** (S135)
- ^1H NMR of **15** (S136)
- ^{13}C NMR of **15** (S137)
- DEPT NMR of **15** (S138)
- ^1H NMR of **17** (S139)
- ^{13}C NMR of **17** (S140)
- DEPT NMR of **17** (S141)
- ^1H NMR of **18** (S142)
- ^{13}C NMR of **18** (S143)
- DEPT NMR of **18** (S144)
- ^1H NMR of **19** (S145)
- ^{13}C NMR of **19** (S146)
- DEPT NMR of **19** (S147)
- ^1H NMR of **21** (S148)
- ^{13}C NMR of **21** (S149)
- DEPT NMR of **21** (S150)
- ^1H NMR of **22** (S151)
- ^{13}C NMR of **22** (S152)
- DEPT NMR of **22** (S153)

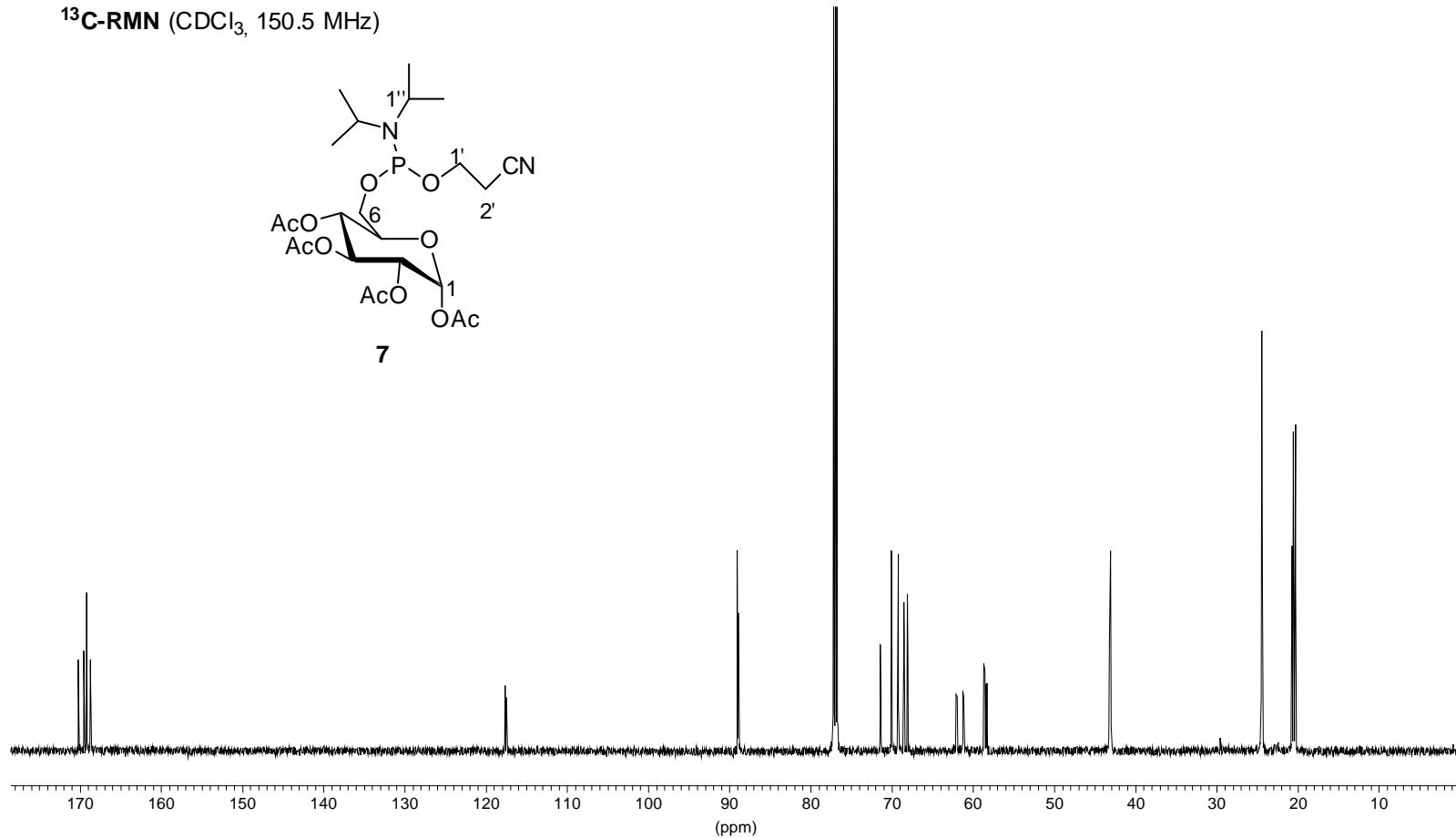
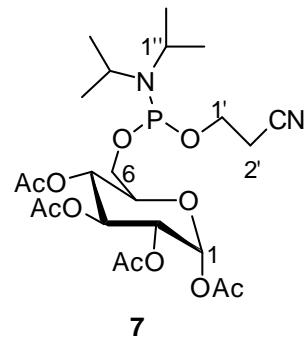
2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)

¹H-RMN (CDCl_3 , 600 MHz)



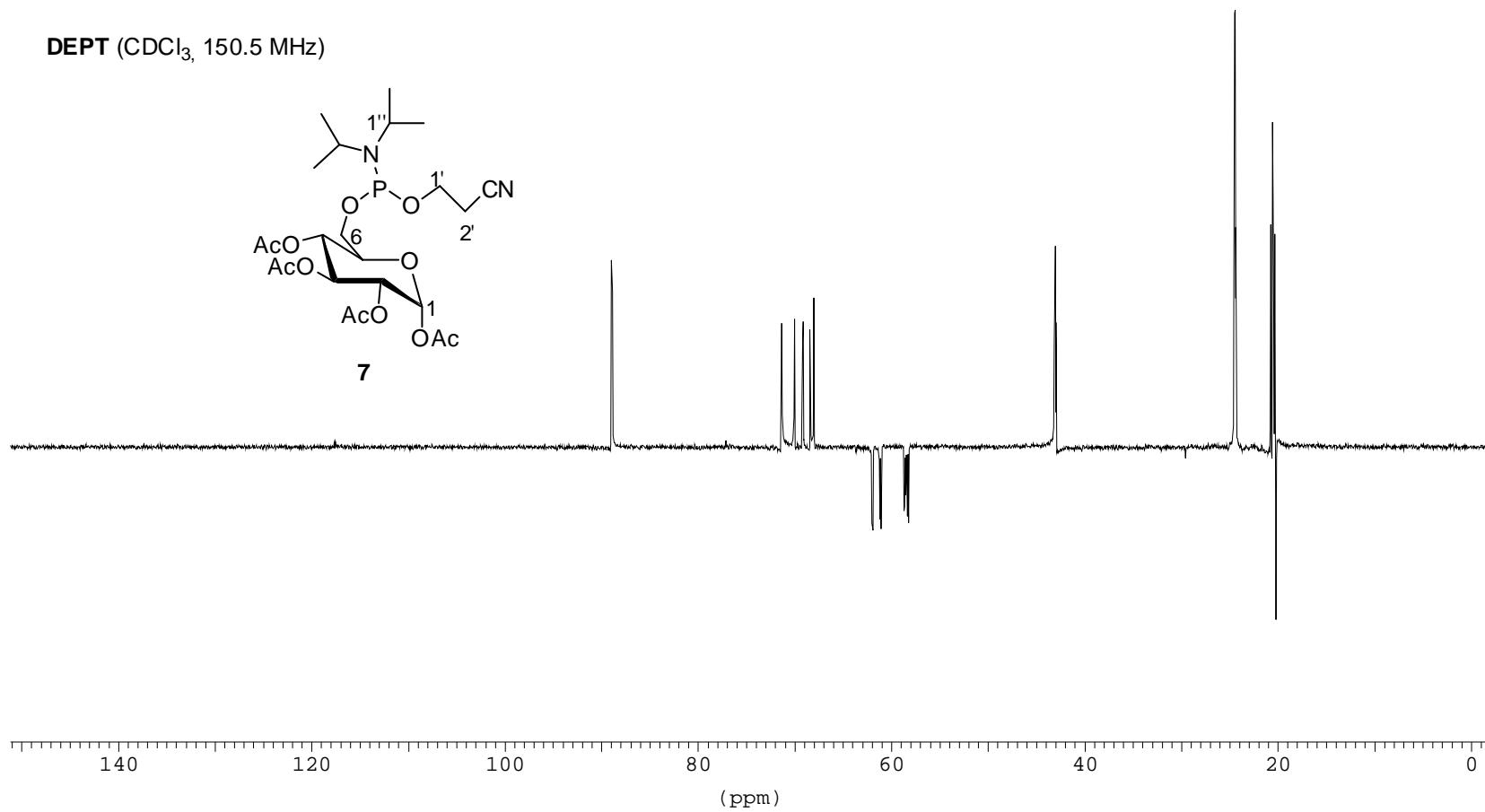
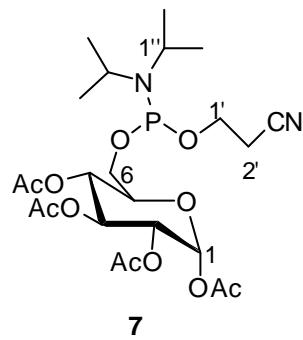
2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)

^{13}C -RMN (CDCl_3 , 150.5 MHz)

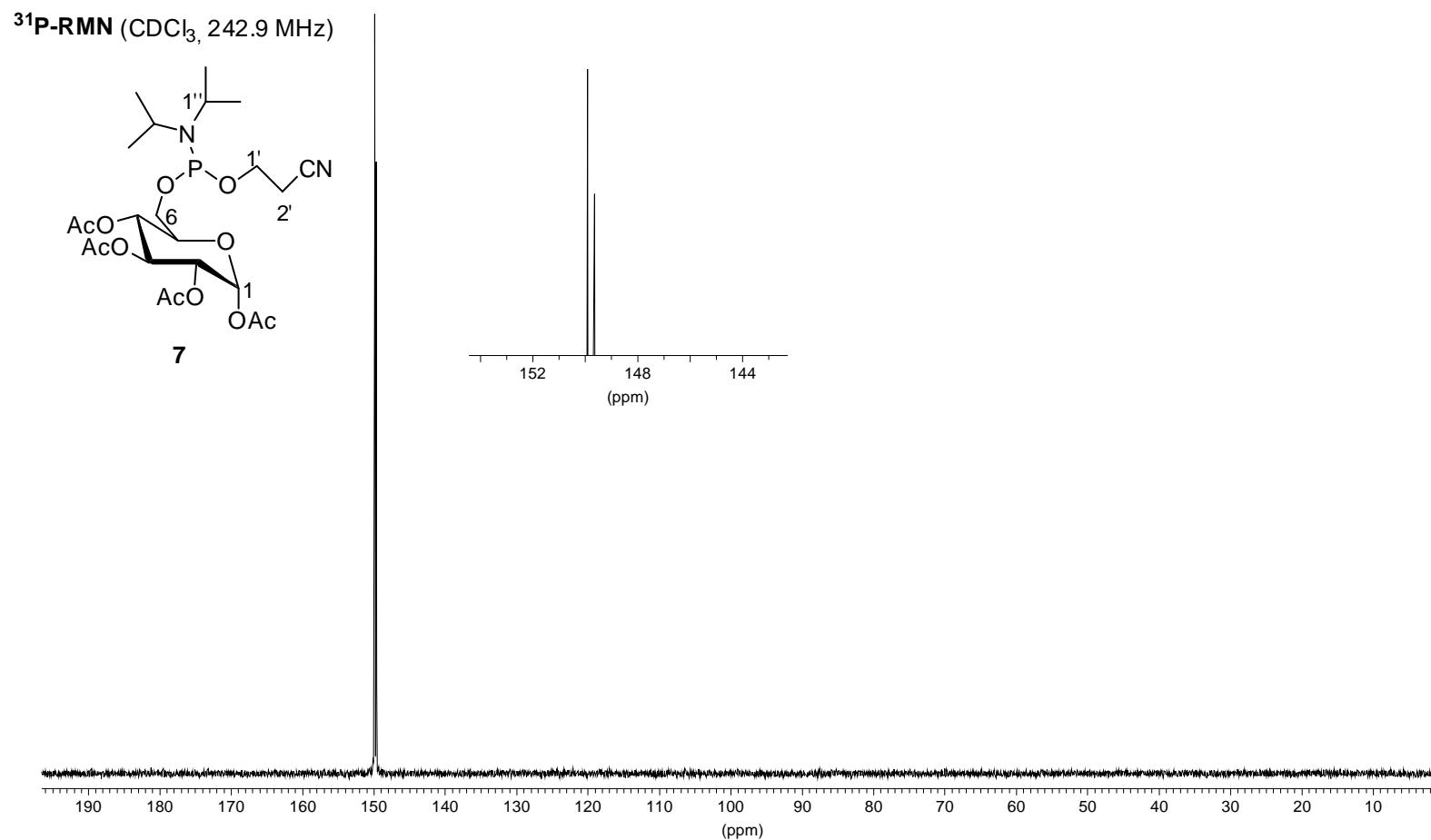


2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)

DEPT (CDCl_3 , 150.5 MHz)

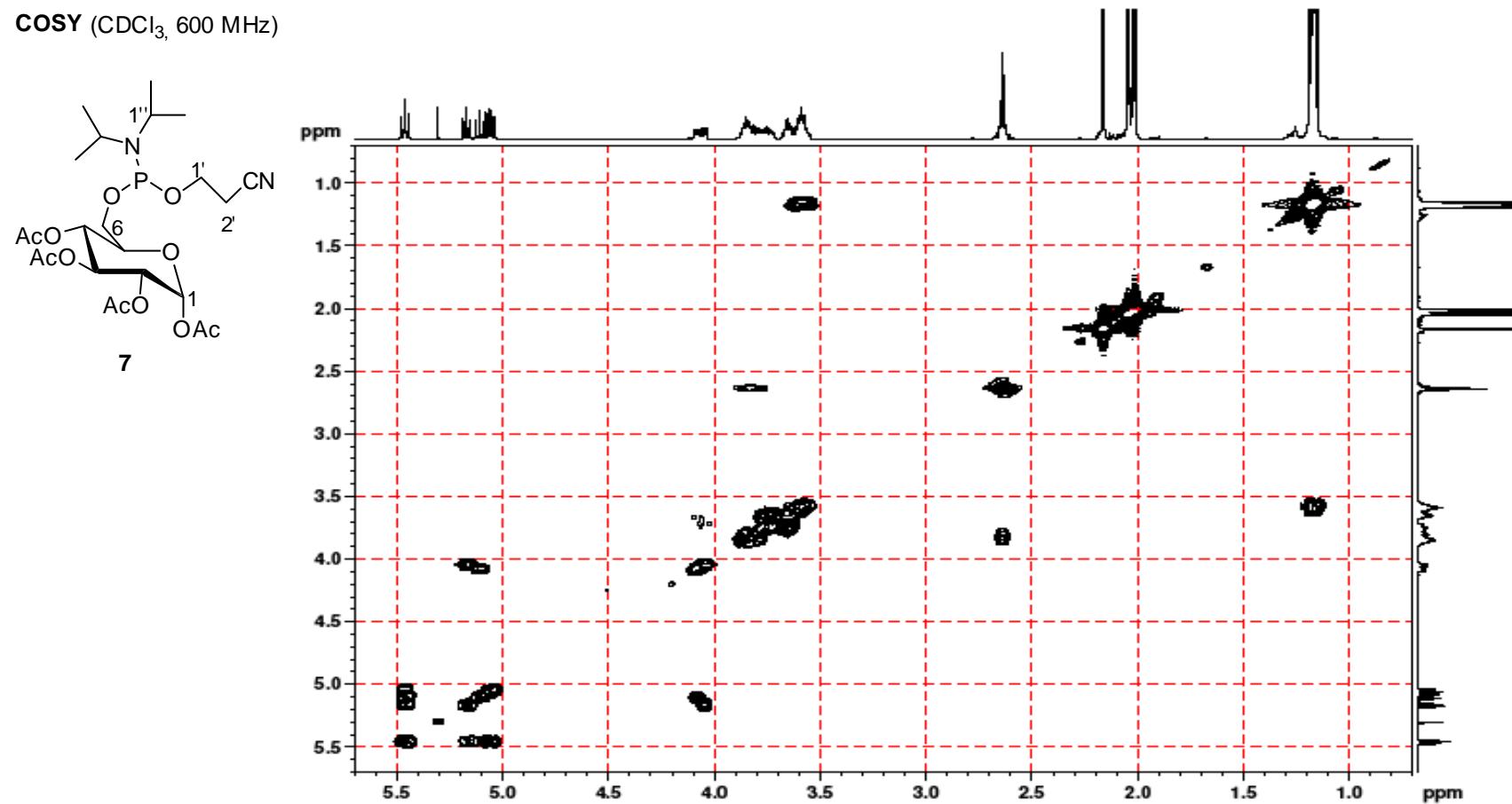


2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)

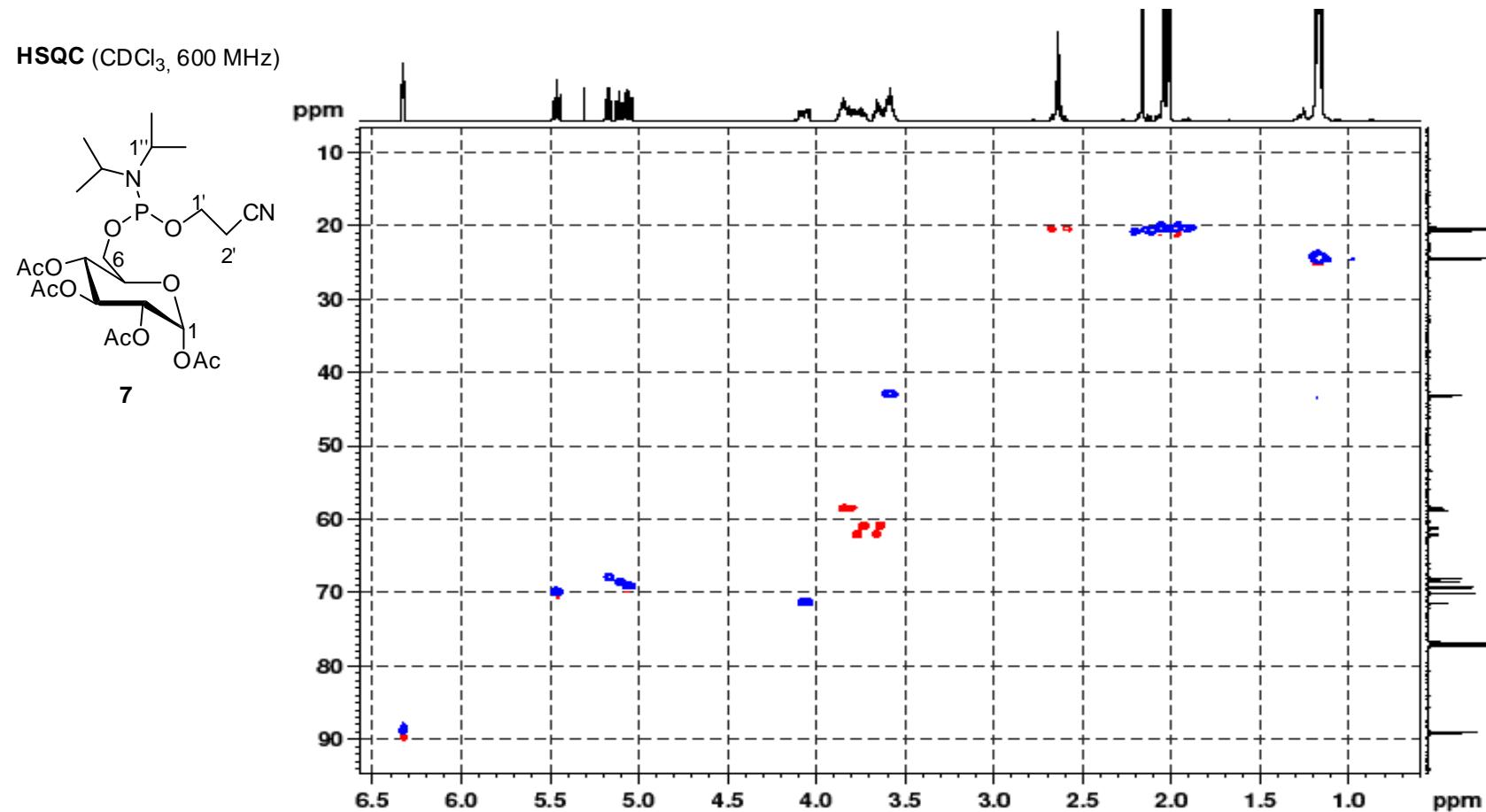


2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)

COSY (CDCl_3 , 600 MHz)

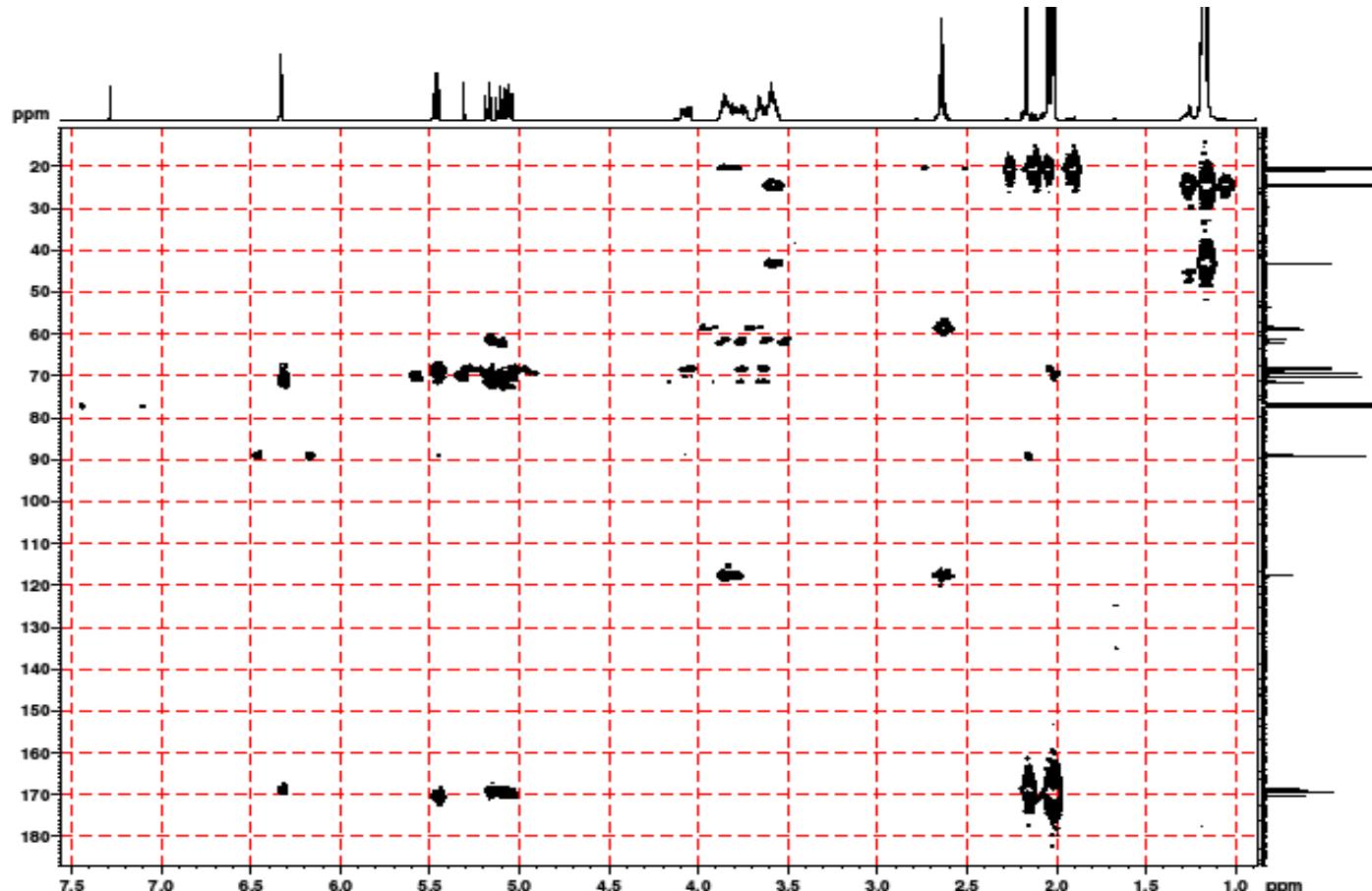
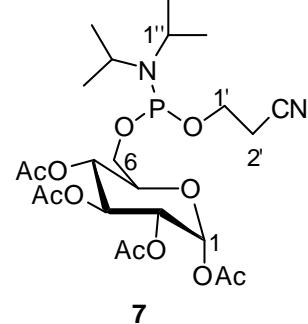


2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)



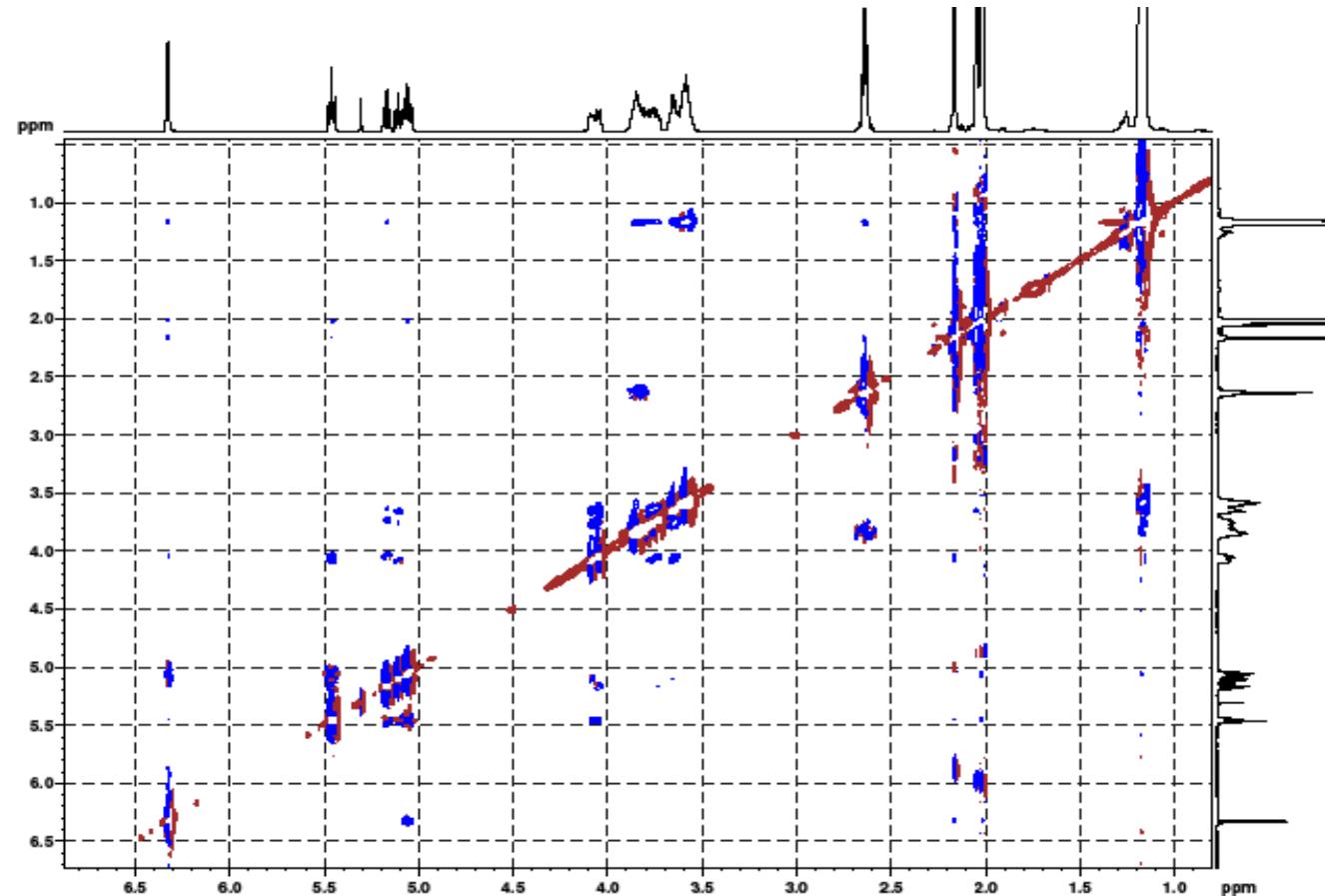
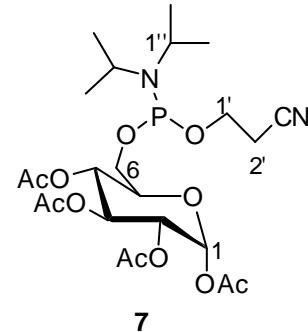
2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)

HMBC (CDCl_3 , 600 MHz)



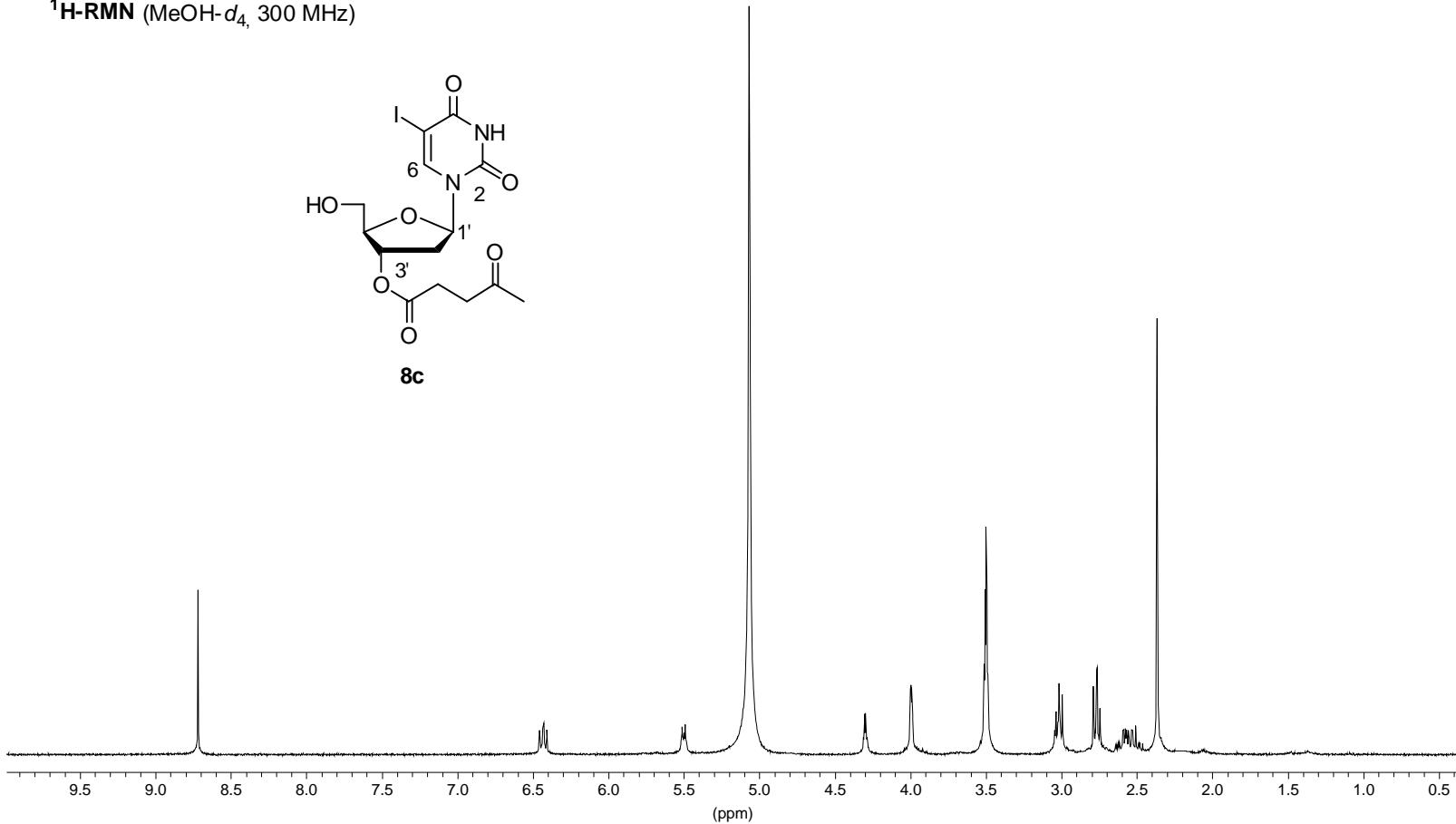
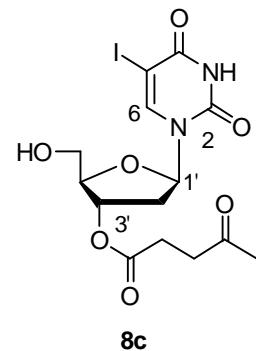
2-Cyanoethyl-*N,N*-diisopropyl-(1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranoside-6-yl)phosphoramidite (7)

NOESY (CDCl_3 , 600 MHz)



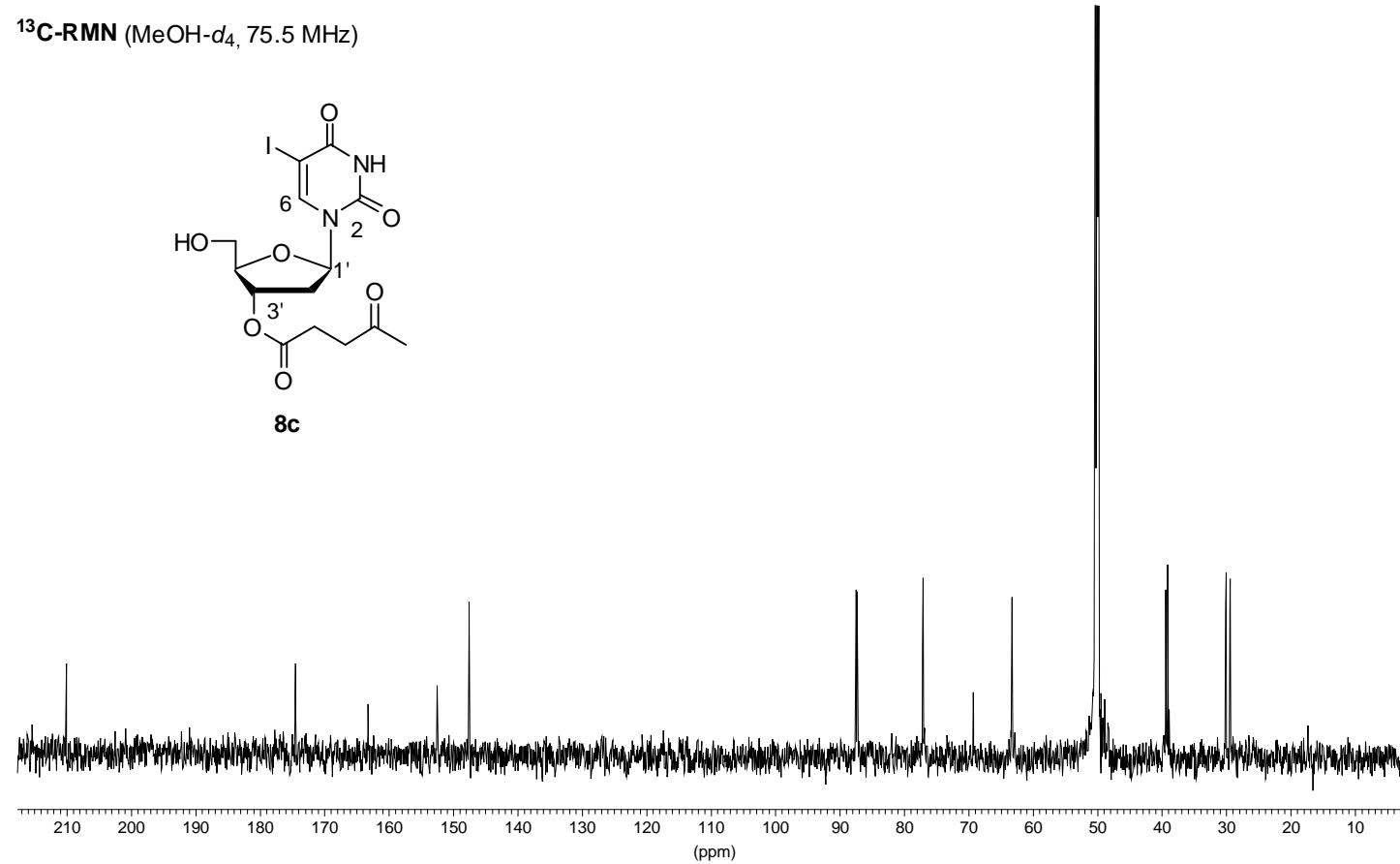
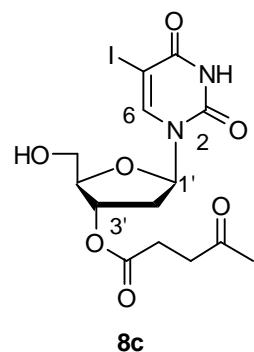
5-Iodo-3'-O-levulinyl-2'-deoxyuridine (8c)

¹H-RMN (MeOH-*d*₄, 300 MHz)



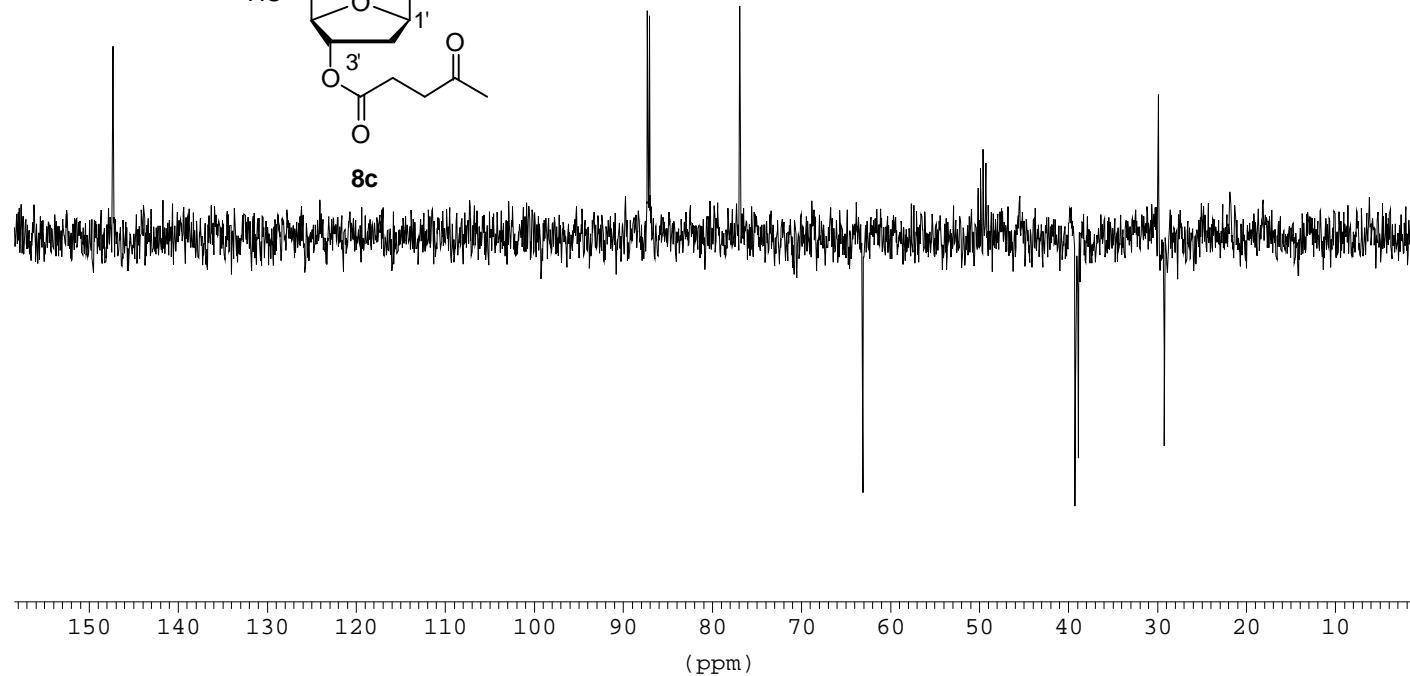
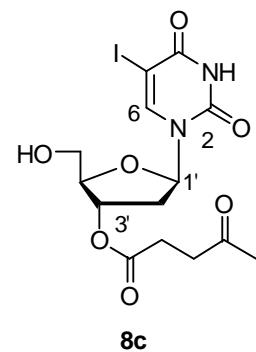
5-Iodo-3'-O-levulinyl-2'-deoxyuridine (8c)

¹³C-RMN (MeOH-d₄, 75.5 MHz)



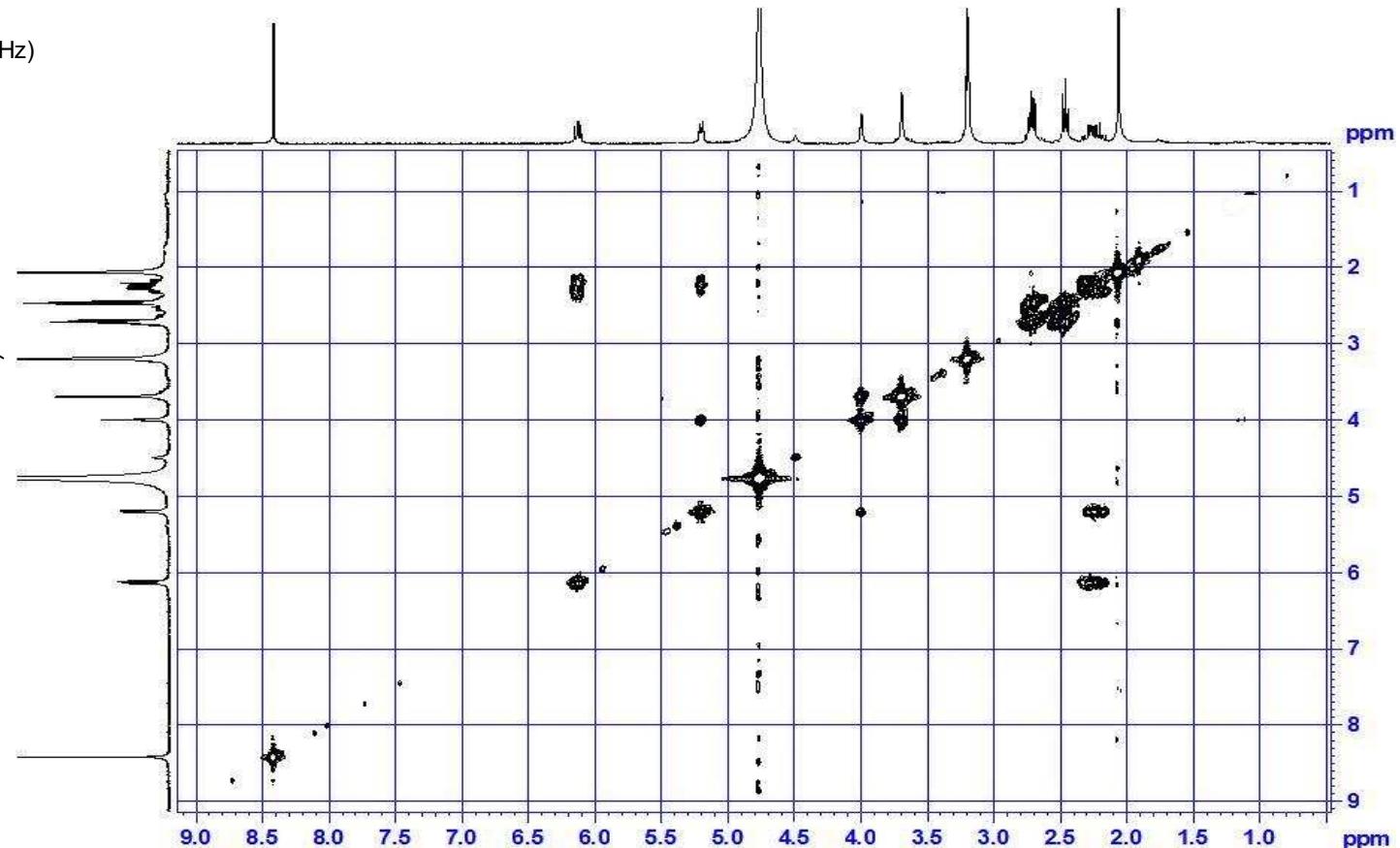
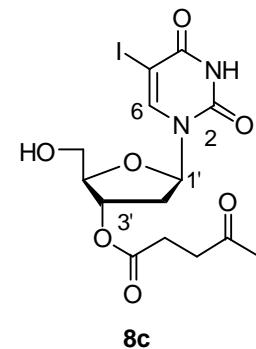
5-Iodo-3'-O-levulinyl-2'-deoxyuridine (8c)

DEPT(MeOH-*d*₄, 75.5 MHz)



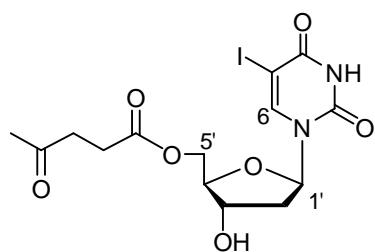
5-Iodo-3'-O-levulinyl-2'-deoxyuridine (8c)

COSY (MeOH-*d*₄, 300 MHz)

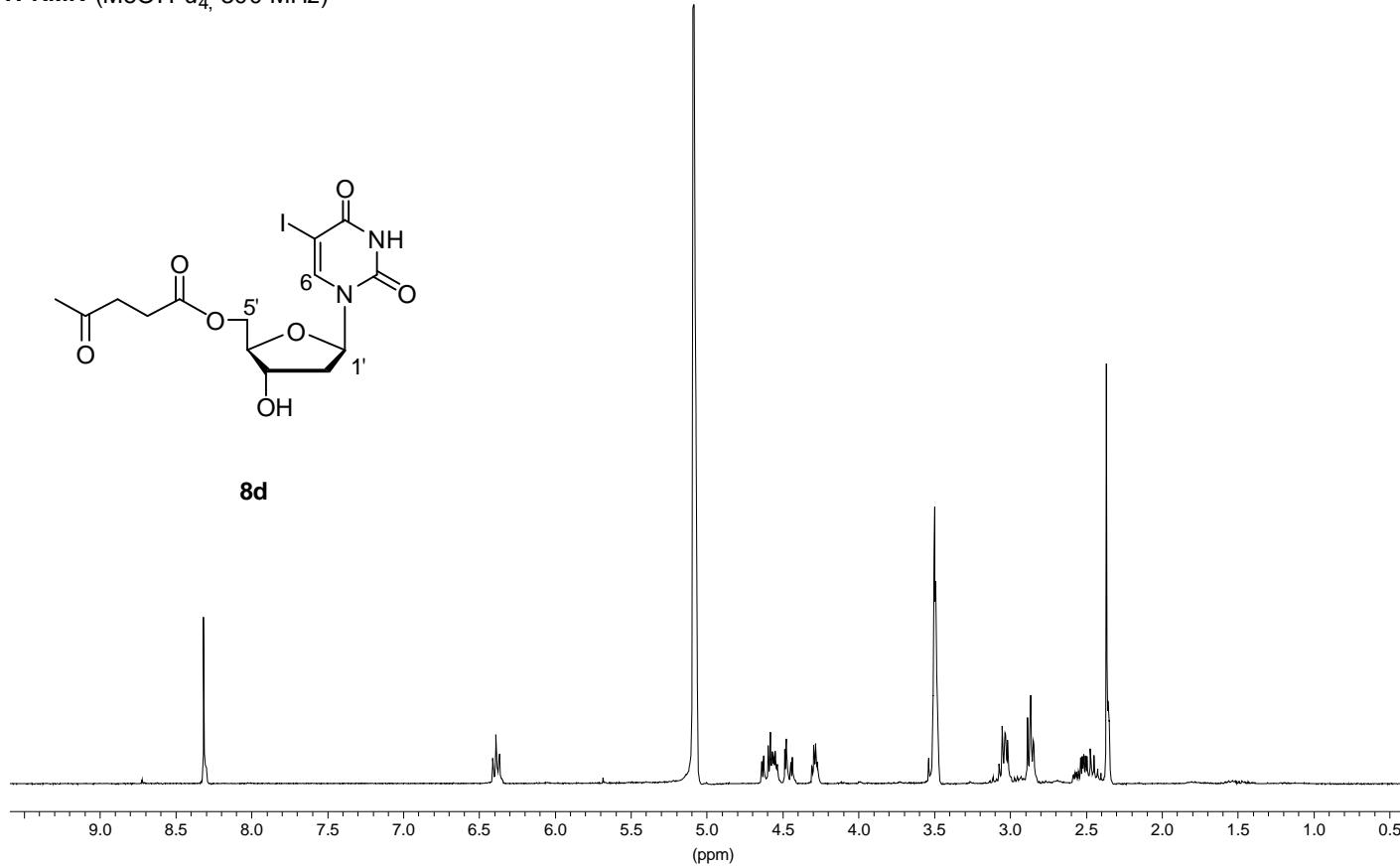


5-Iodo-5'-O-levulinyl-2'-deoxyuridine (8d)

¹H-RMN (MeOH-*d*₄, 300 MHz)

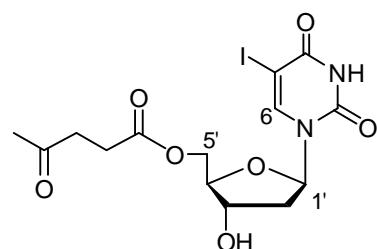


8d

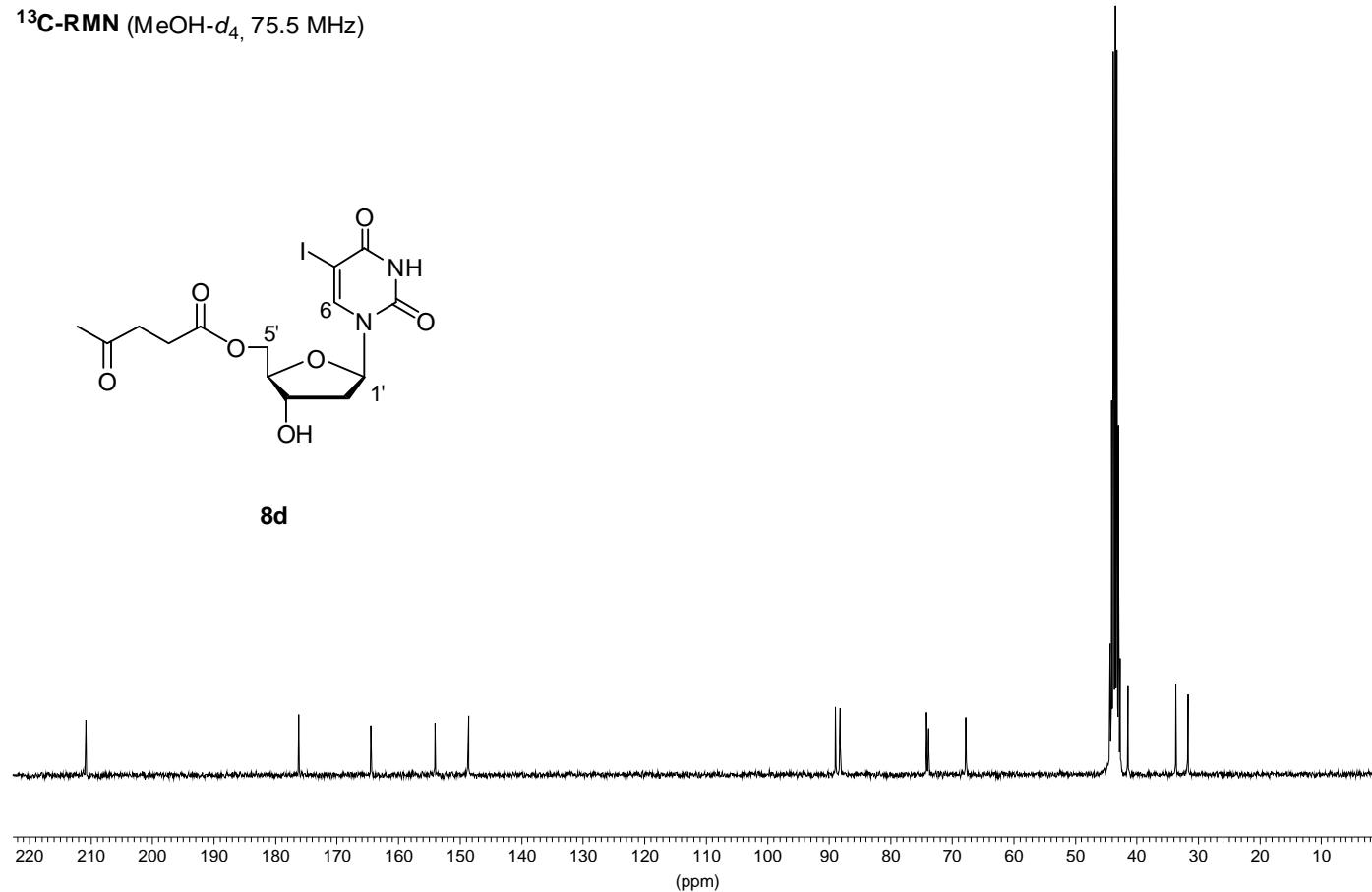


5-Iodo-5'-O-levulinyl-2'-deoxyuridine (8d)

¹³C-RMN (MeOH-*d*₄, 75.5 MHz)

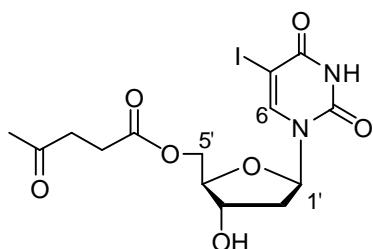


8d

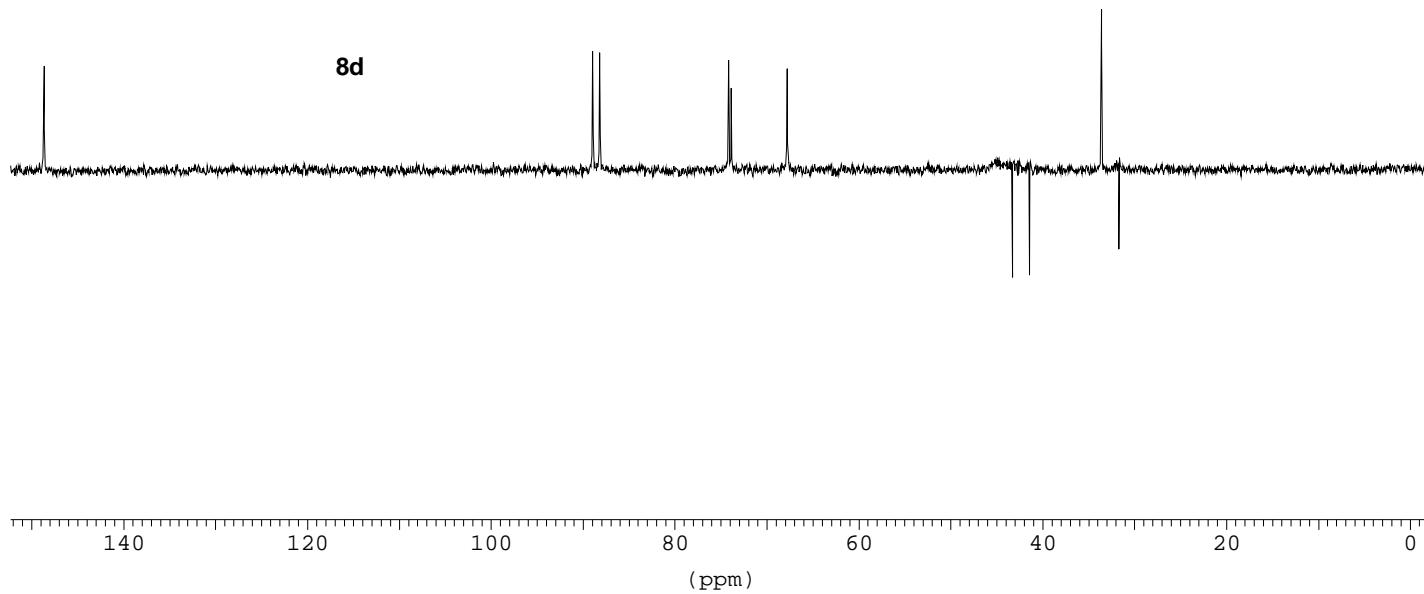


5-Iodo-5'-O-levulinyl-2'-deoxyuridine (8d)

DEPT (MeOH-*d*₄, 75.5 MHz)

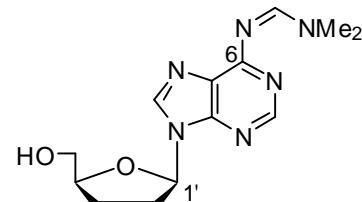


8d

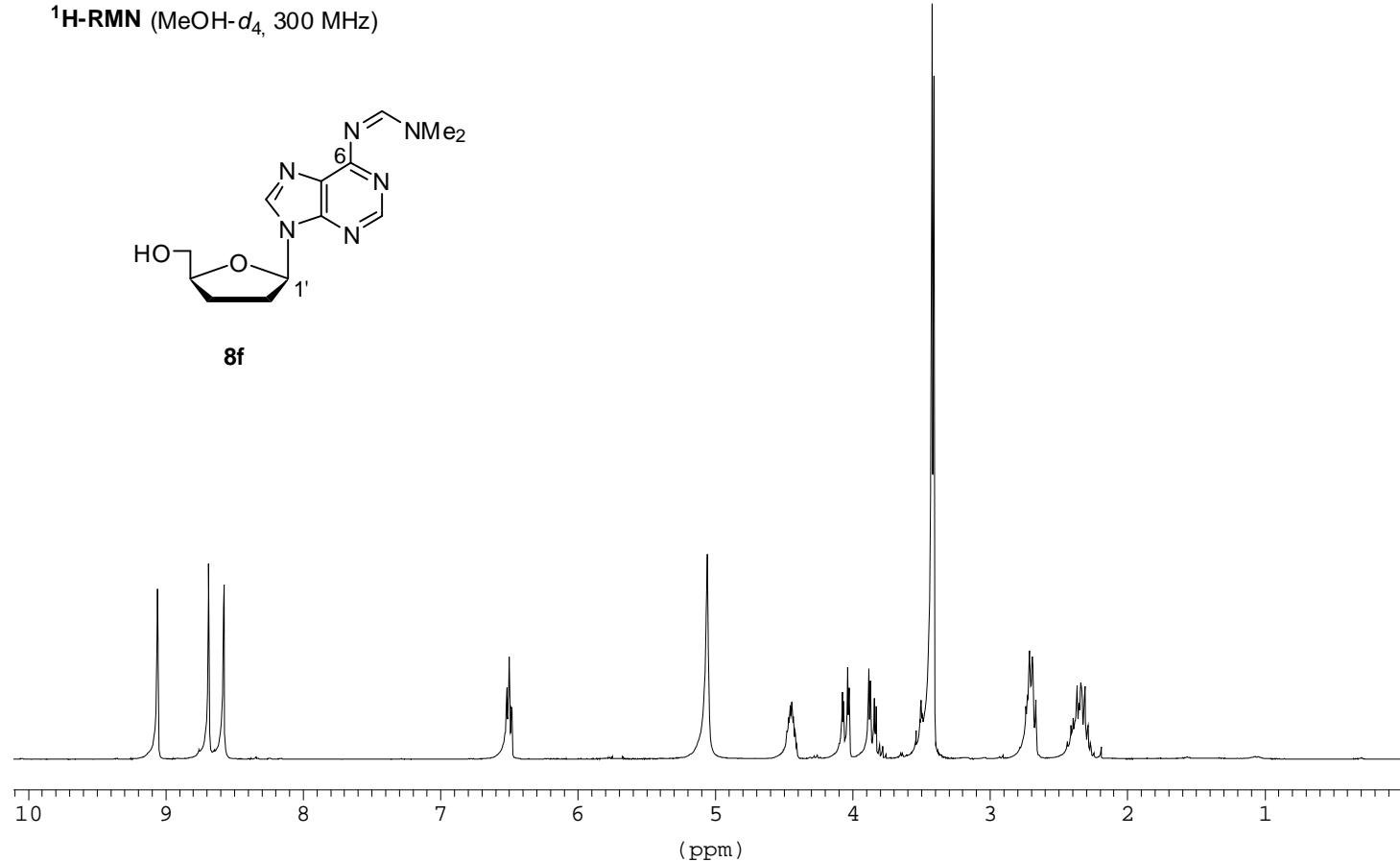


***N*⁶-(Dimethylamino)methylene-2',3'-dideoxyadenosine (8f)**

¹H-RMN (MeOH-*d*₄, 300 MHz)

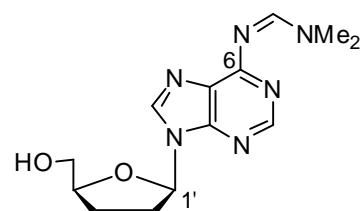


8f

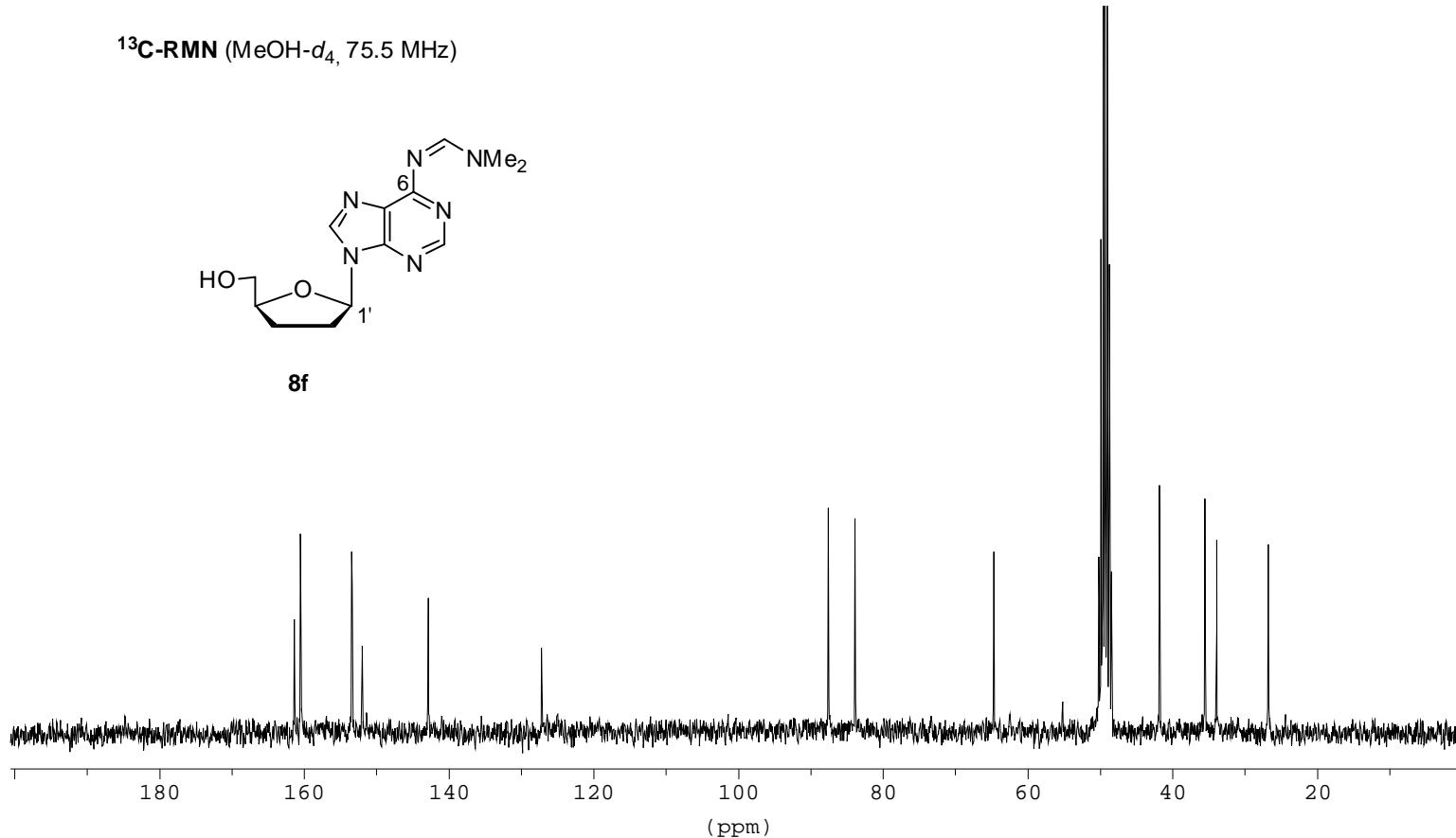


***N*⁶-(Dimethylamino)methylene-2',3'-dideoxyadenosine (8f)**

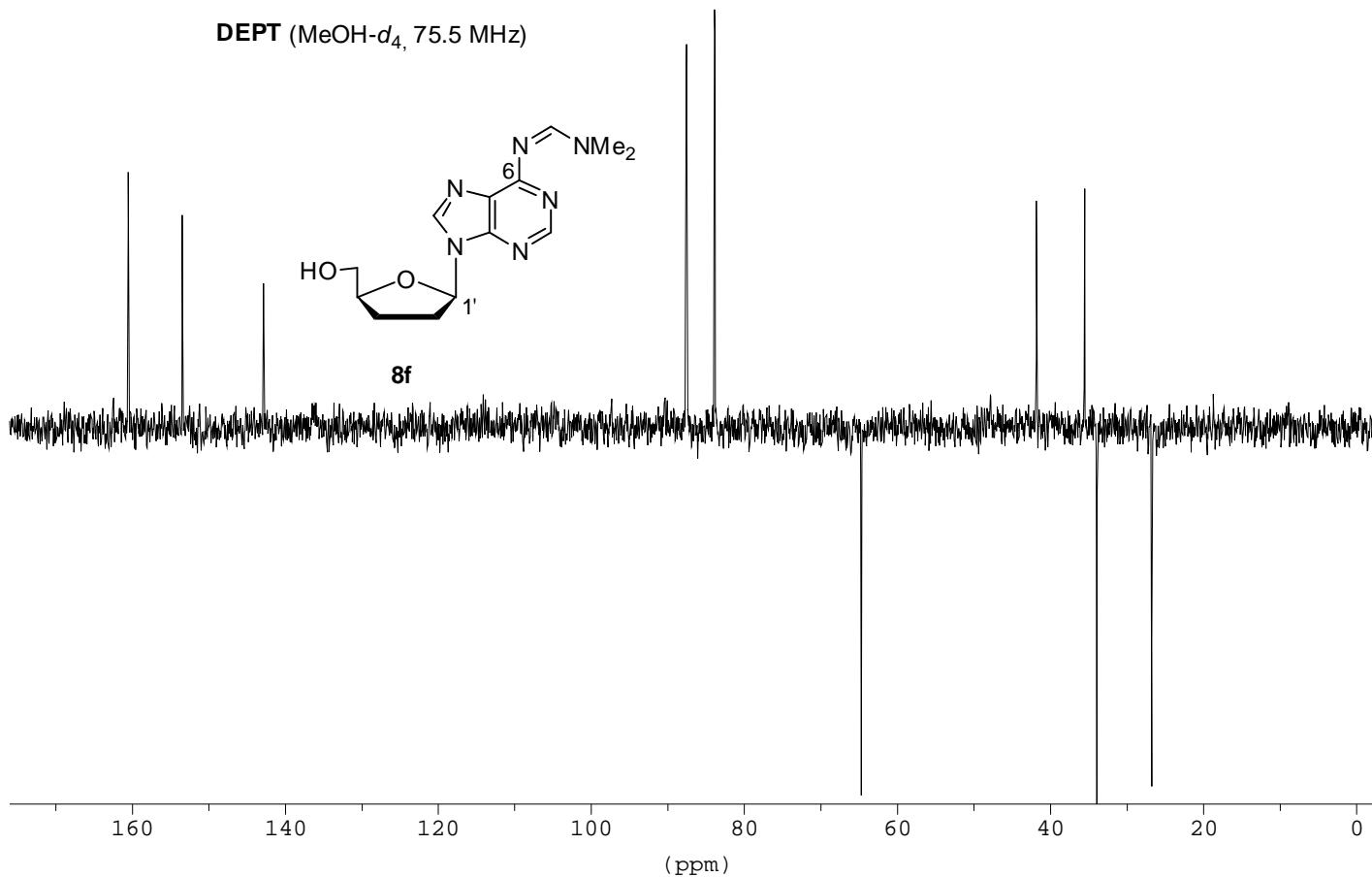
¹³C-RMN (MeOH-*d*₄, 75.5 MHz)



8f

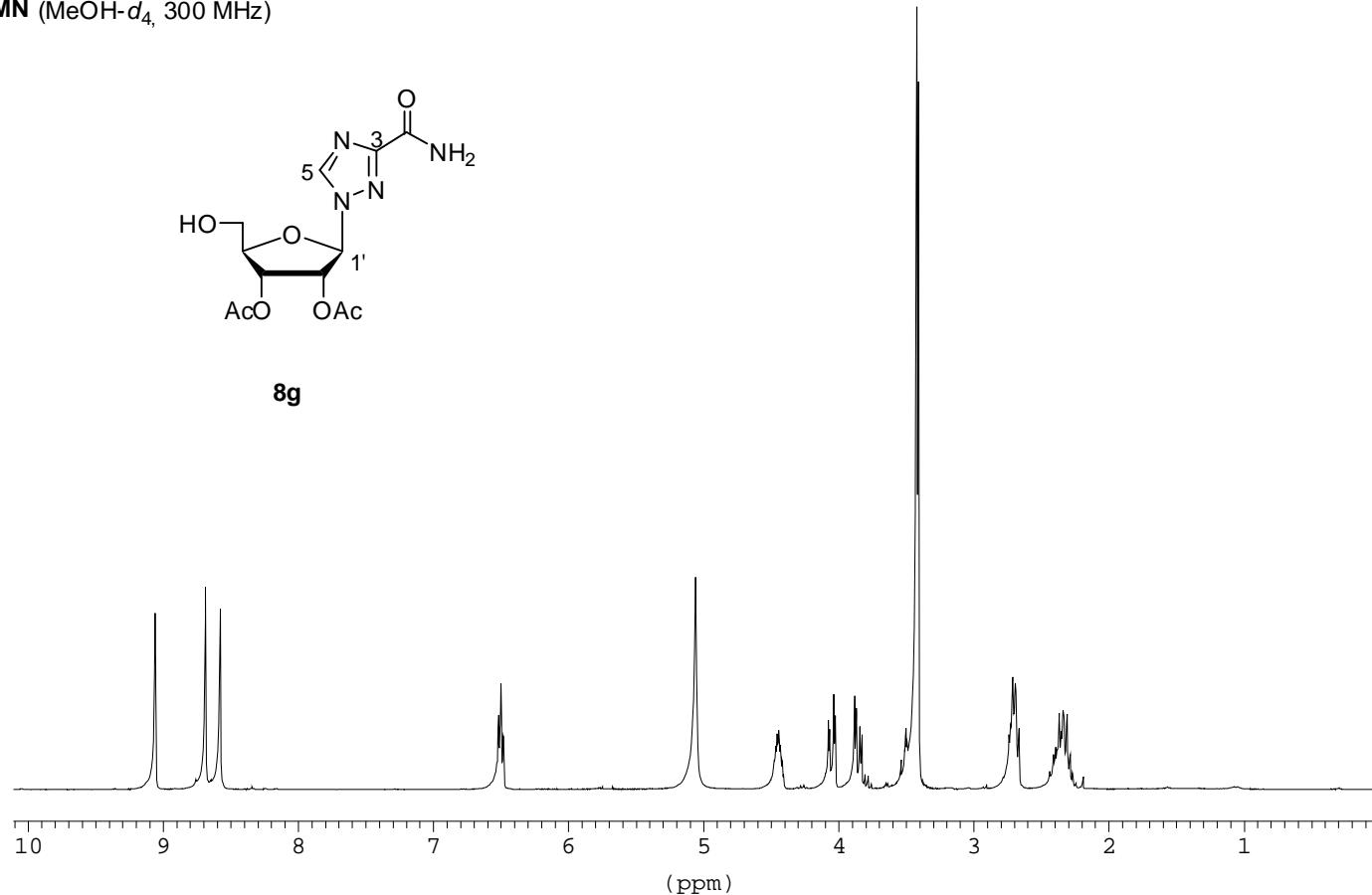
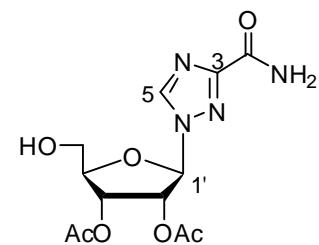


***N*⁶-(Dimethylamino)methylene-2',3'-dideoxyadenosine (8f)**



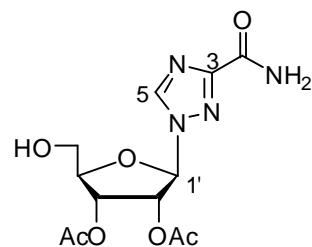
1-(2',3'-Di-*O*-acetyl- β -D-ribofuranosyl)-1*H*-1,2,4-triazole-3-carboxamide (8g)

¹H-RMN (MeOH-*d*₄, 300 MHz)

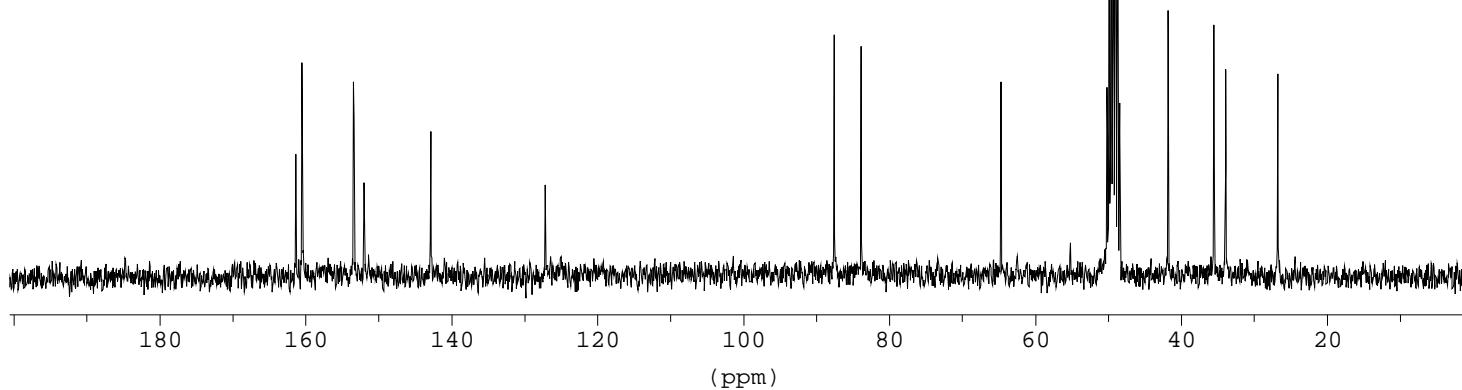


1-(2',3'-Di-*O*-acetyl- β -D-ribofuranosyl)-1*H*-1,2,4-triazole-3-carboxamide (8g)

^{13}C -RMN (MeOH- d_4 , 75.5 MHz)

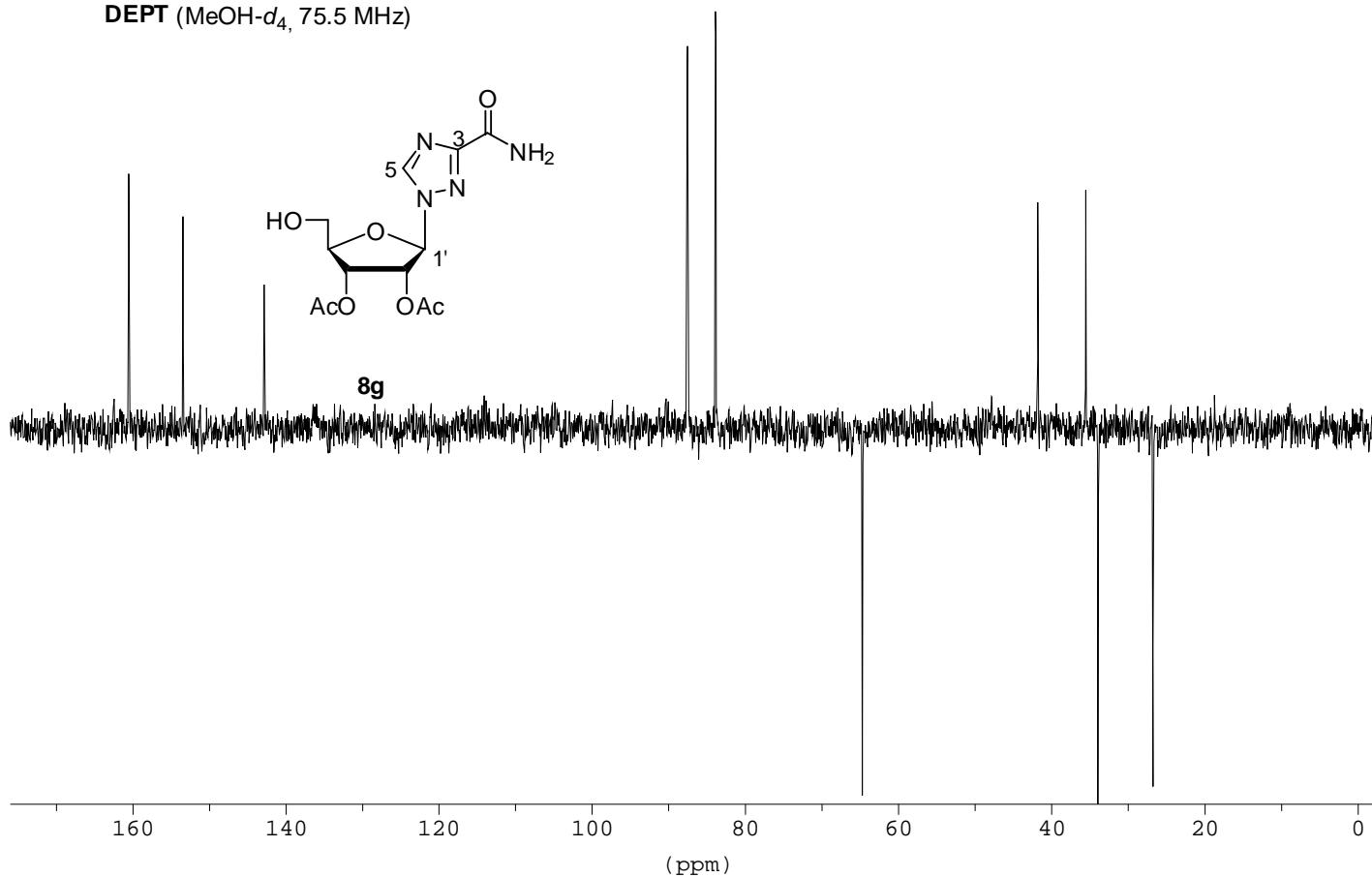


8g



1-(2',3'-Di-*O*-acetyl- β -D-ribofuranosyl)-1*H*-1,2,4-triazole-3-carboxamide (8g)

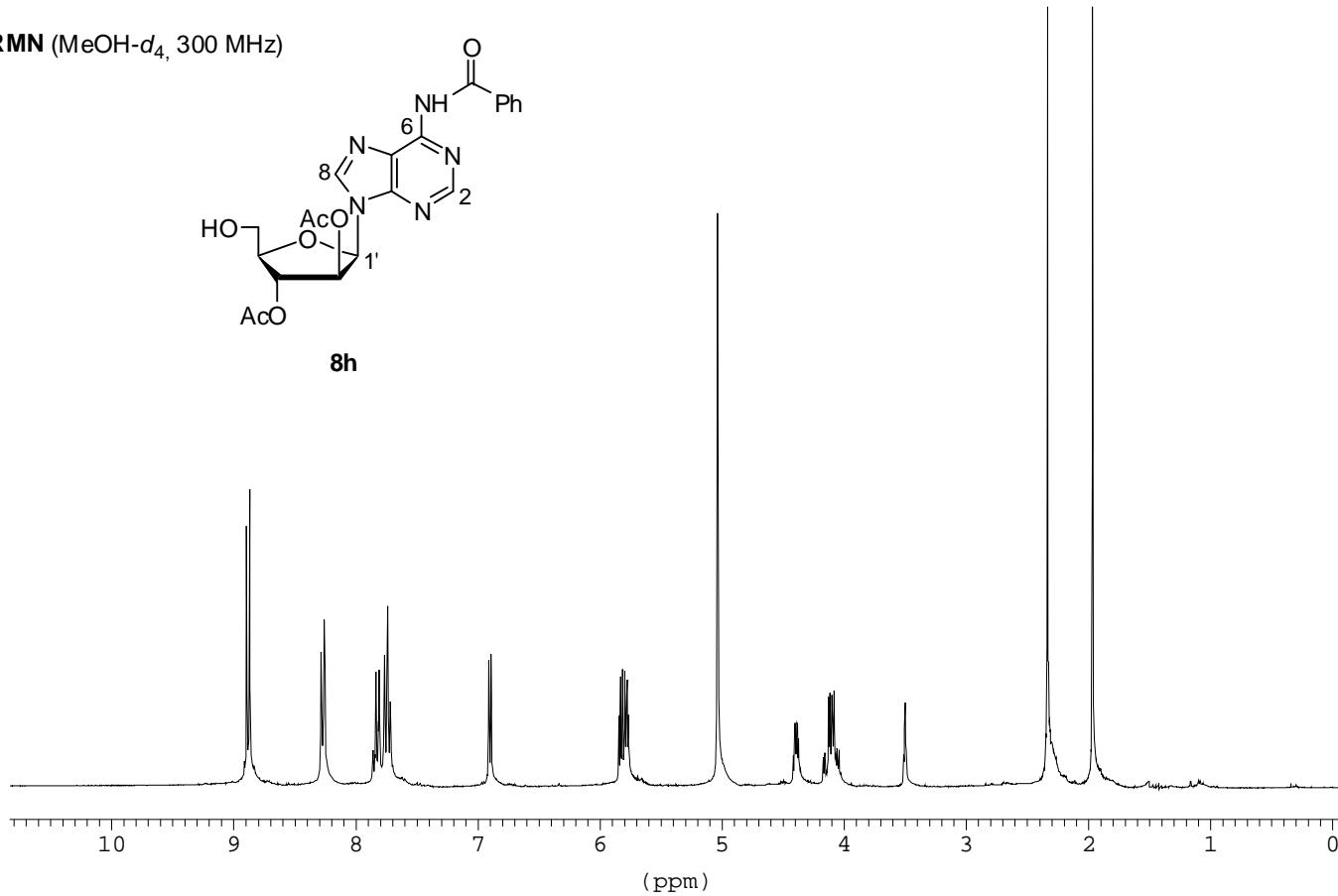
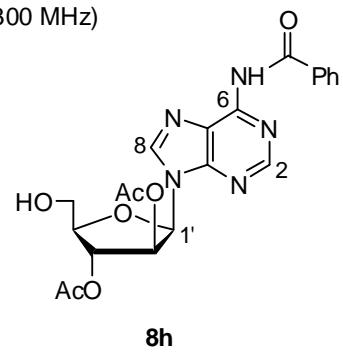
DEPT (MeOH-*d*₄, 75.5 MHz)



***N*⁶-Benzoyl-9-(2',3'-di-*O*-acetyl- β -D-arabinofuranosyl)adenine (8h)**

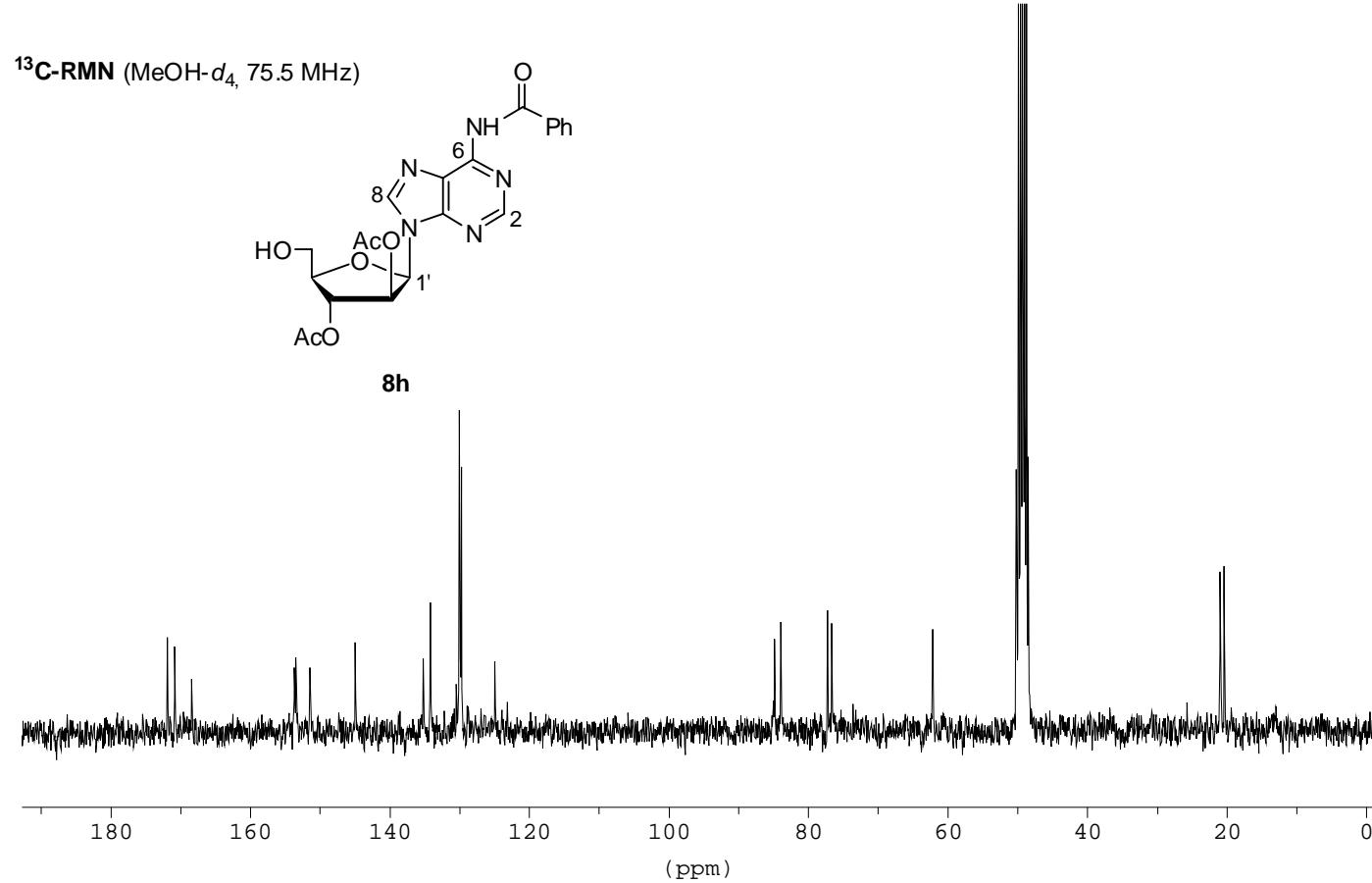
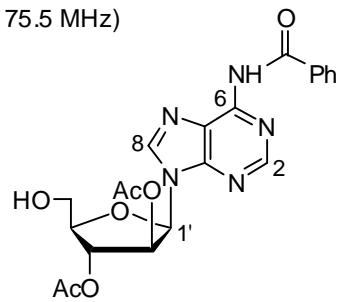
¹H-RMN

(MeOH-*d*₄, 300 MHz)



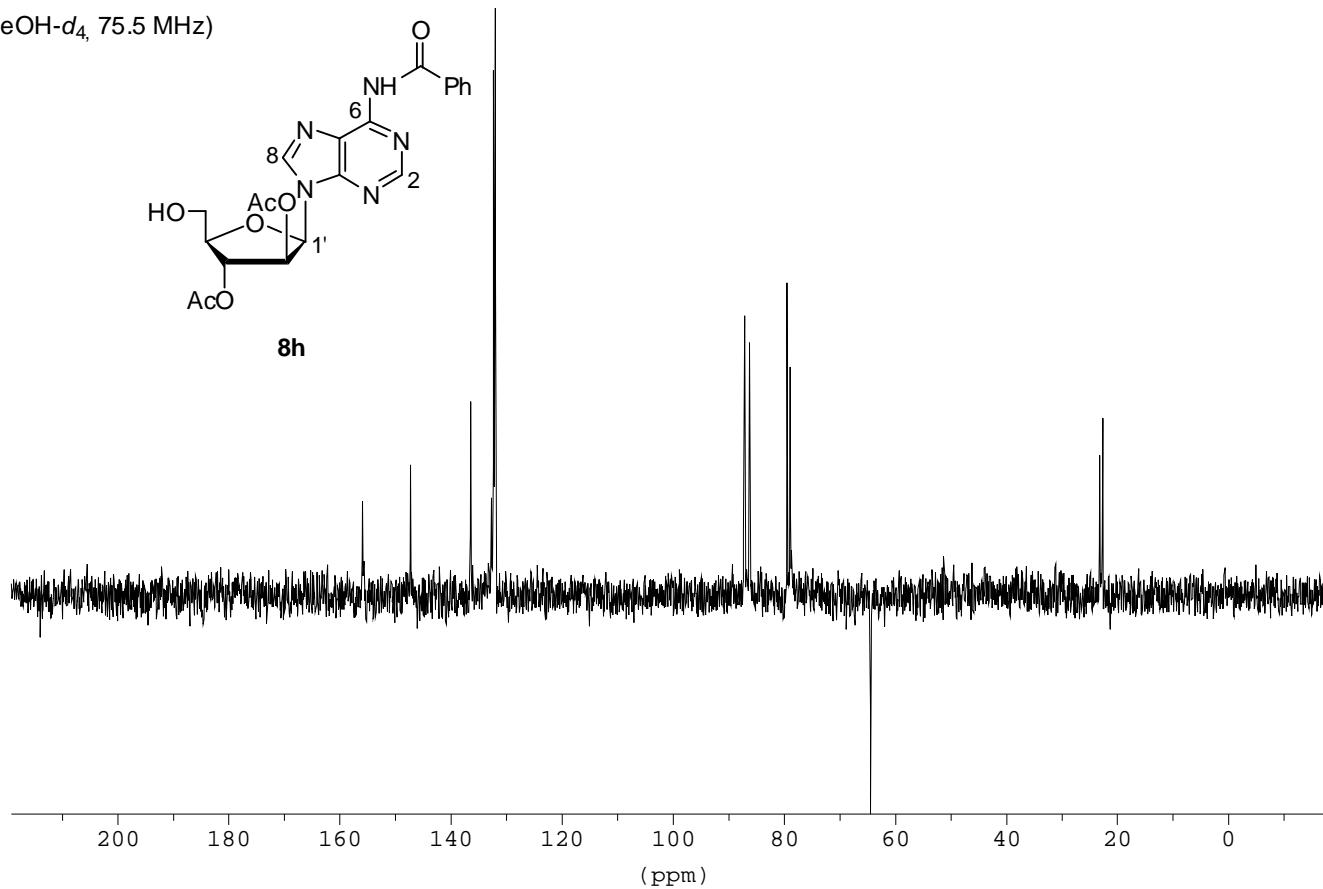
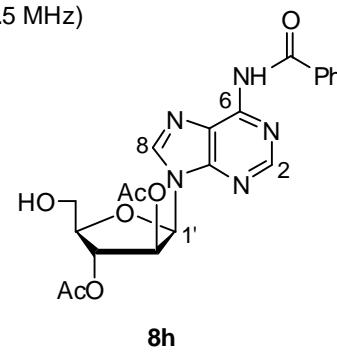
*N*⁶-Benzoyl-9-(2',3'-di-*O*-acetyl- β -D-arabinofuranosyl)adenine (**8h**)

¹³C-RMN (MeOH-*d*₄, 75.5 MHz)



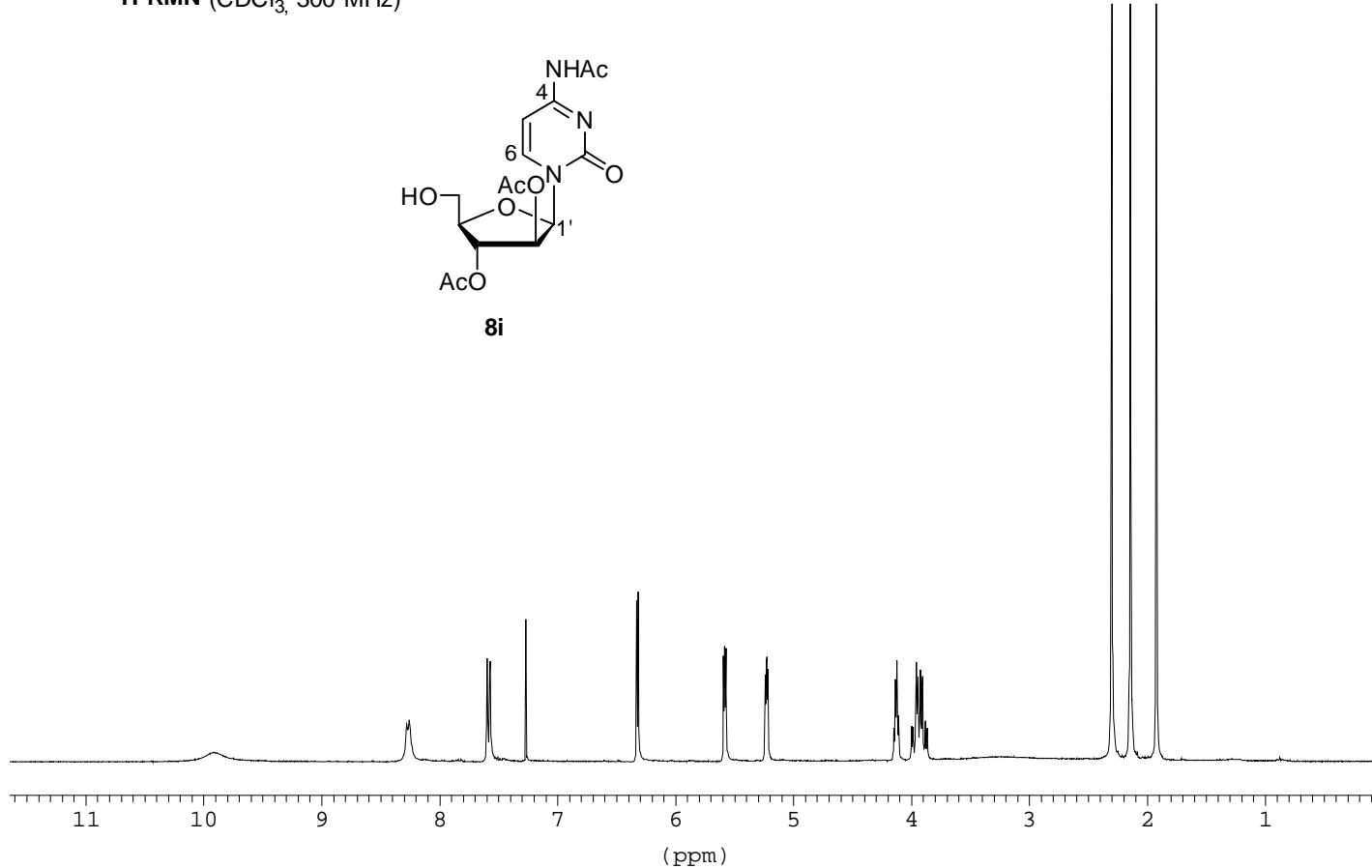
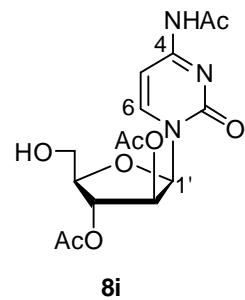
***N*⁶-Benzoyl-9-(2',3'-di-*O*-acetyl- β -D-arabinofuranosyl)adenine (8h)**

DEPT (MeOH-*d*₄, 75.5 MHz)



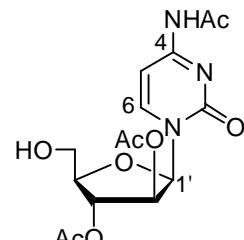
*N*⁴-Acetyl-1-(2',3'-di-*O*-acetyl- β -D-arabinofuranosyl)cytosine (**8i**)

¹H-RMN (CDCl₃, 300 MHz)

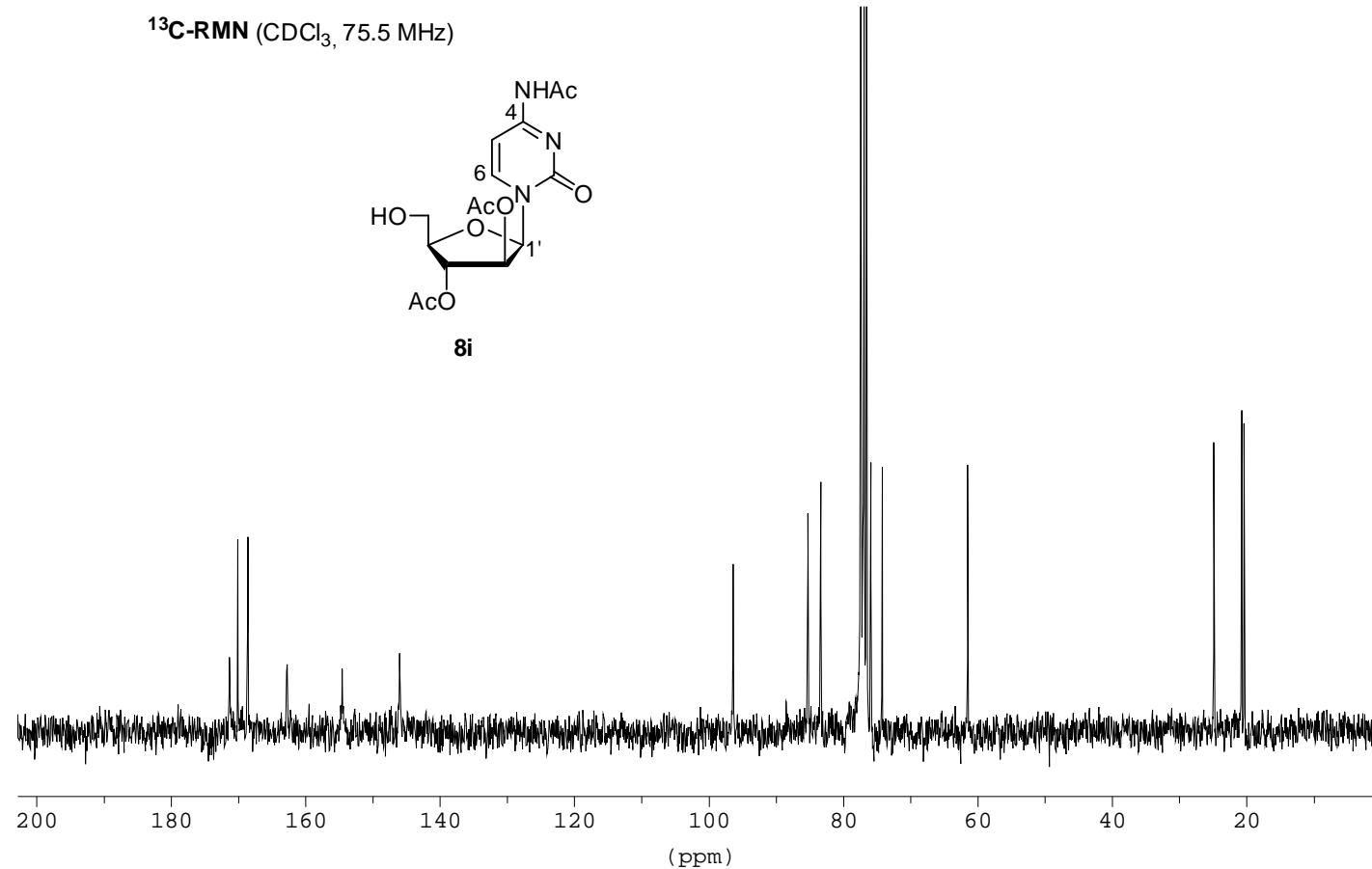


*N*⁴-Acetyl-1-(2',3'-di-*O*-acetyl- β -D-arabinofuranosyl)cytosine (**8i**)

¹³C-RMN (CDCl₃, 75.5 MHz)

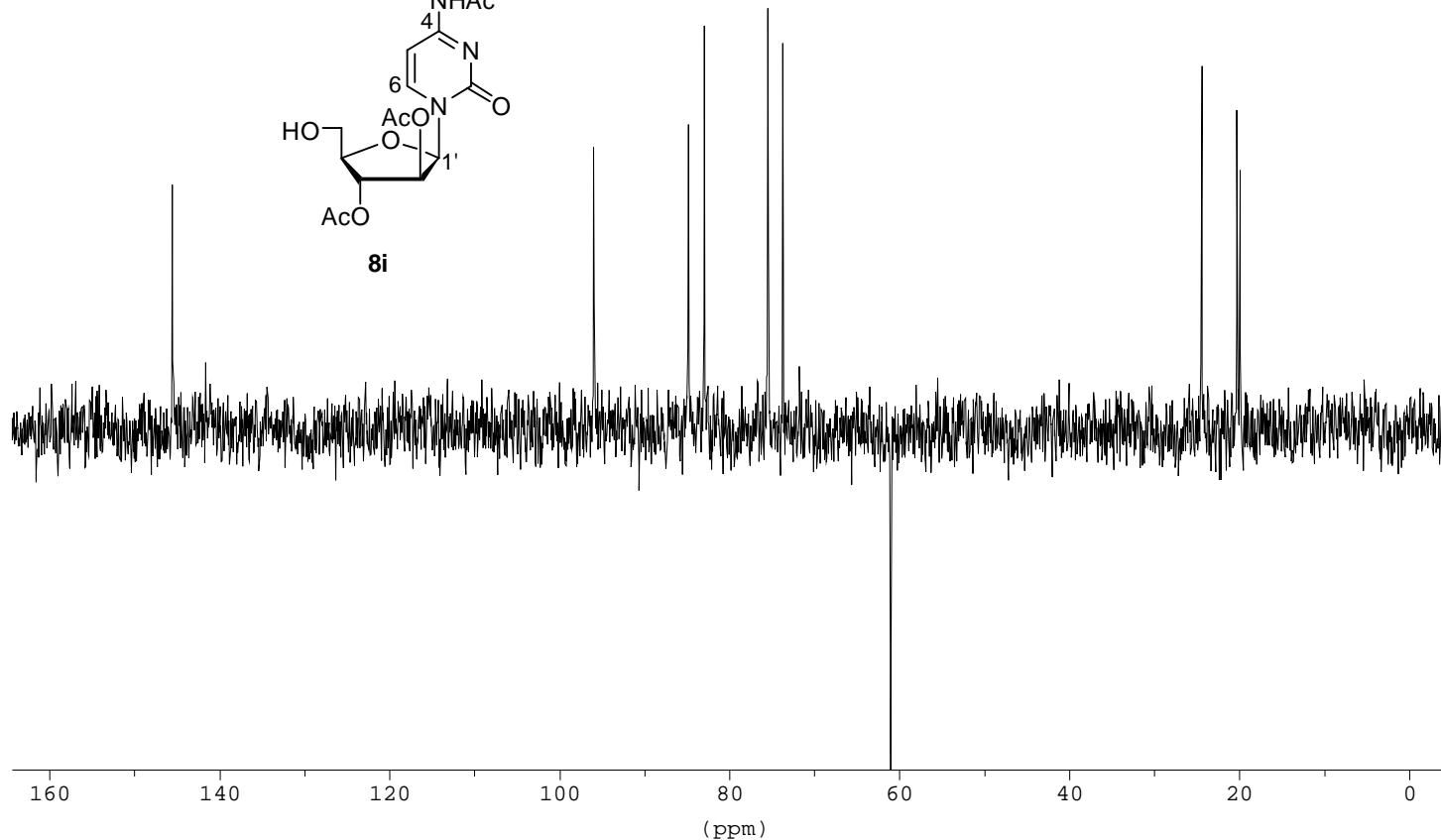
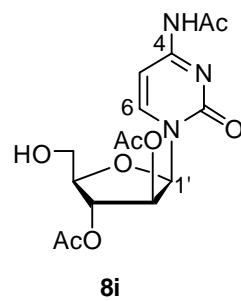


8i

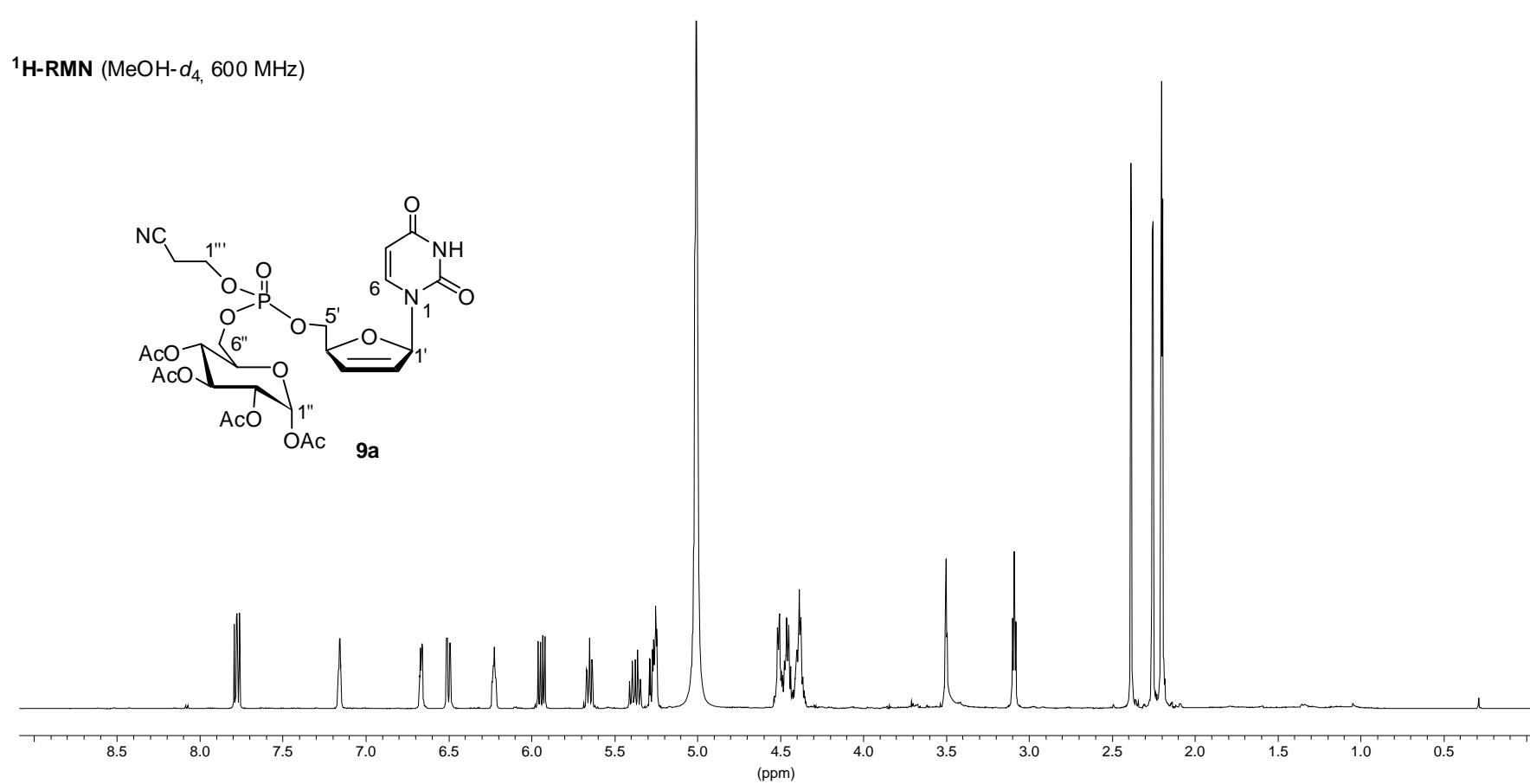


*N*⁴-Acetyl-1-(2',3'-di-*O*-acetyl- β -D-arabinofuranosyl)cytosine (**8i**)

DEPT (CDCl₃, 75.5 MHz)

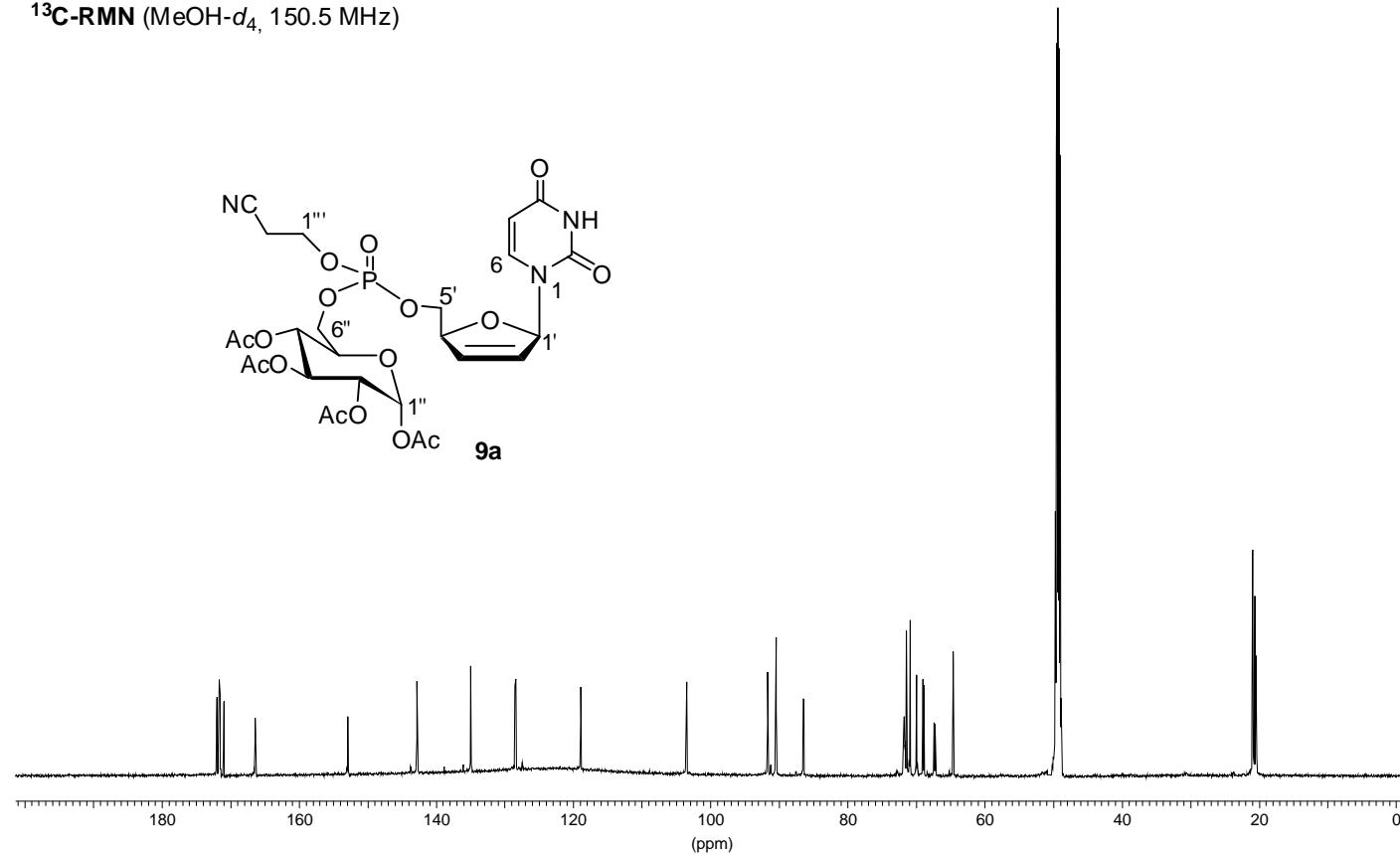
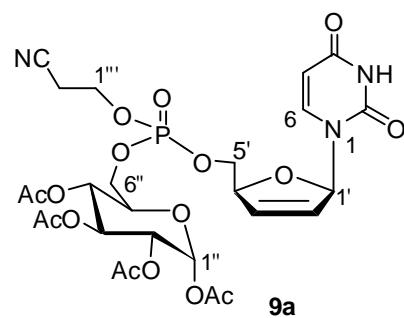


2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxyuridin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9a)



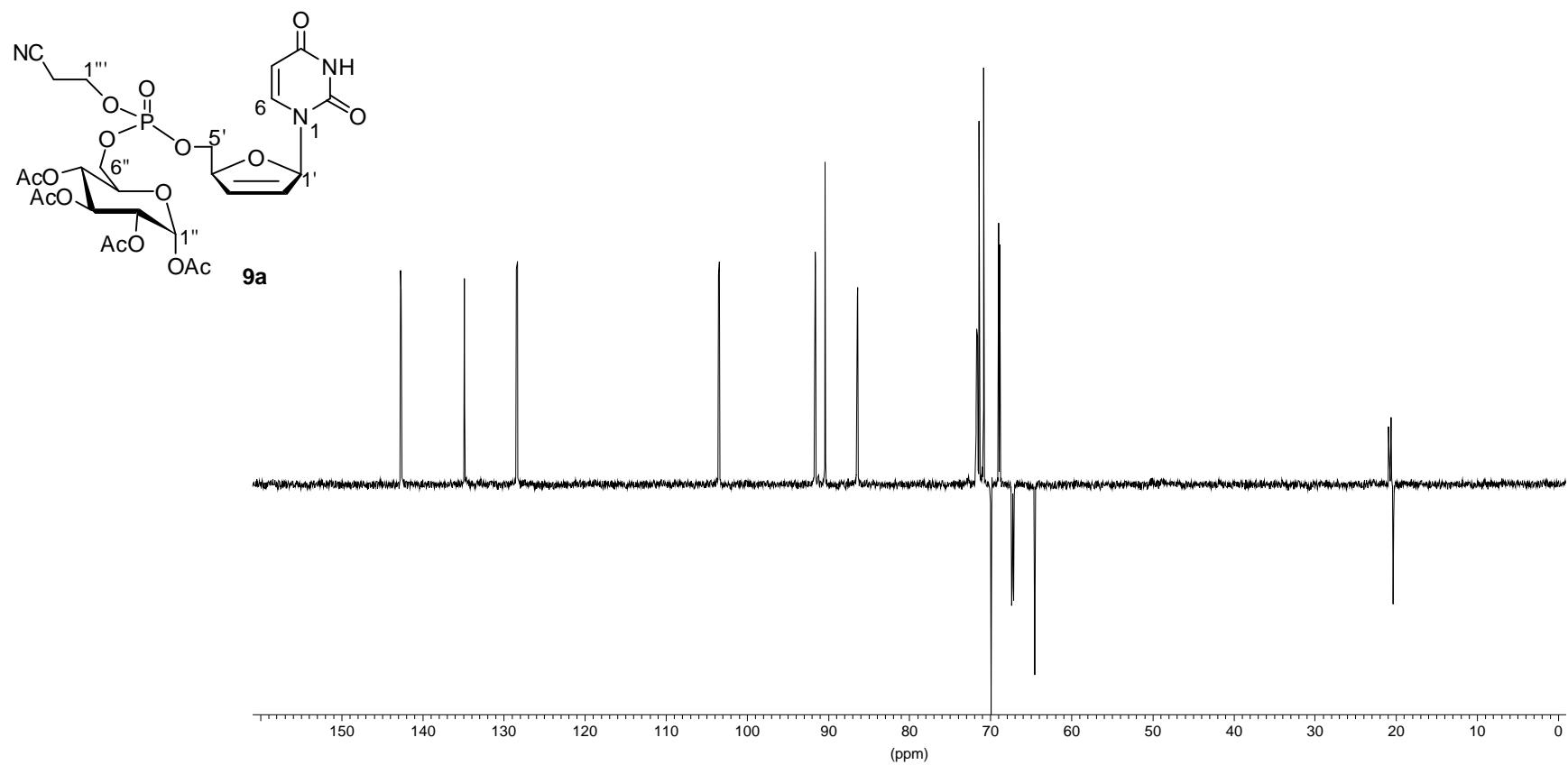
2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxyuridin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9a)

¹³C-RMN (MeOH-*d*₄, 150.5 MHz)

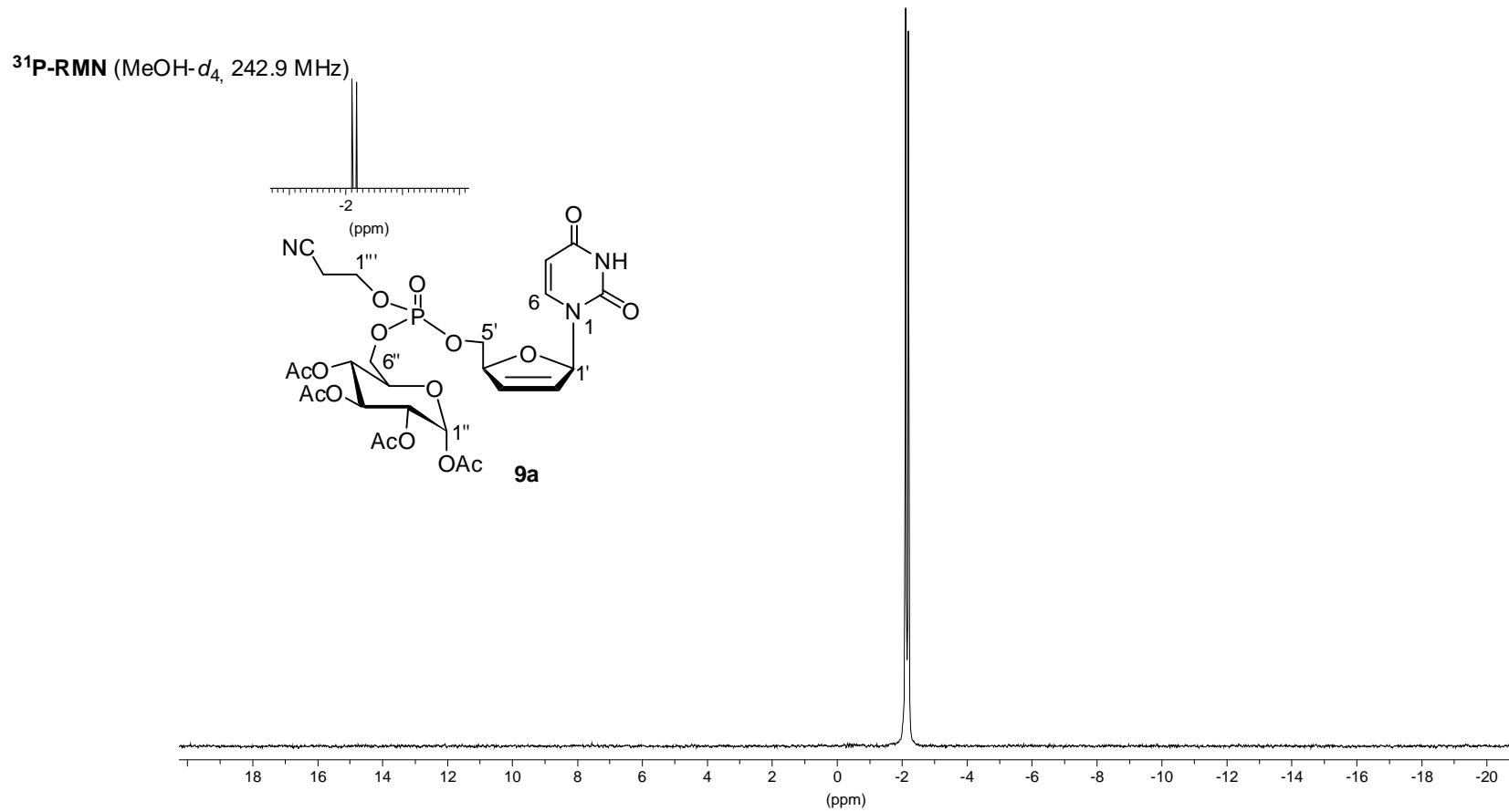


2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxyuridin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9a)

DEPT (MeOH-*d*₄, 150.5 MHz)

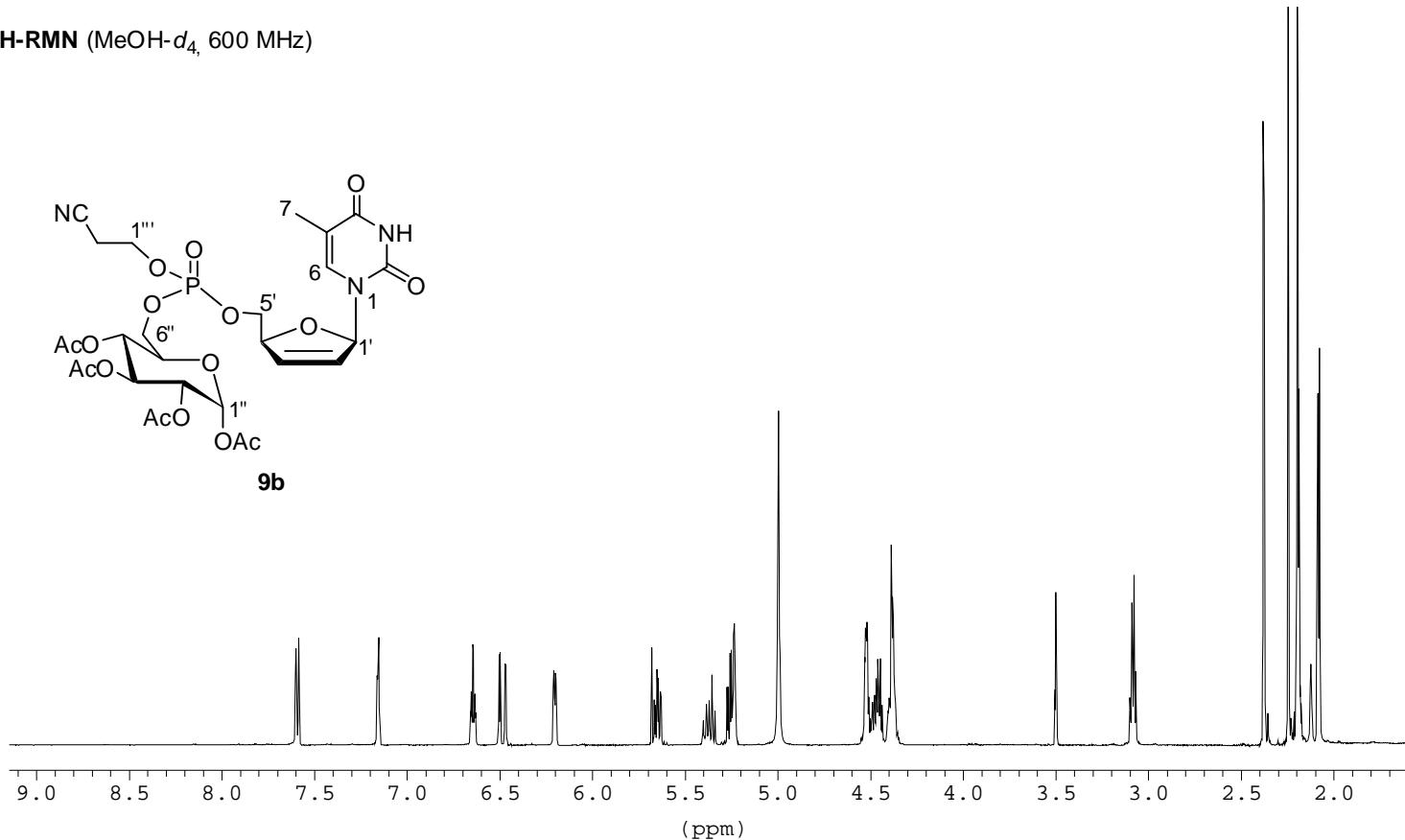
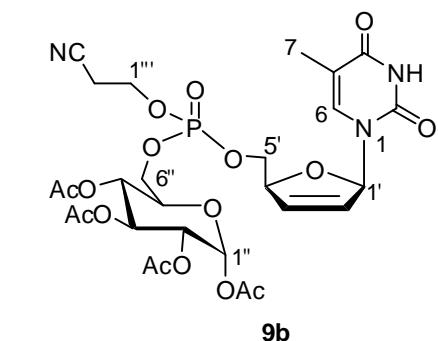


2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxyuridin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9a)



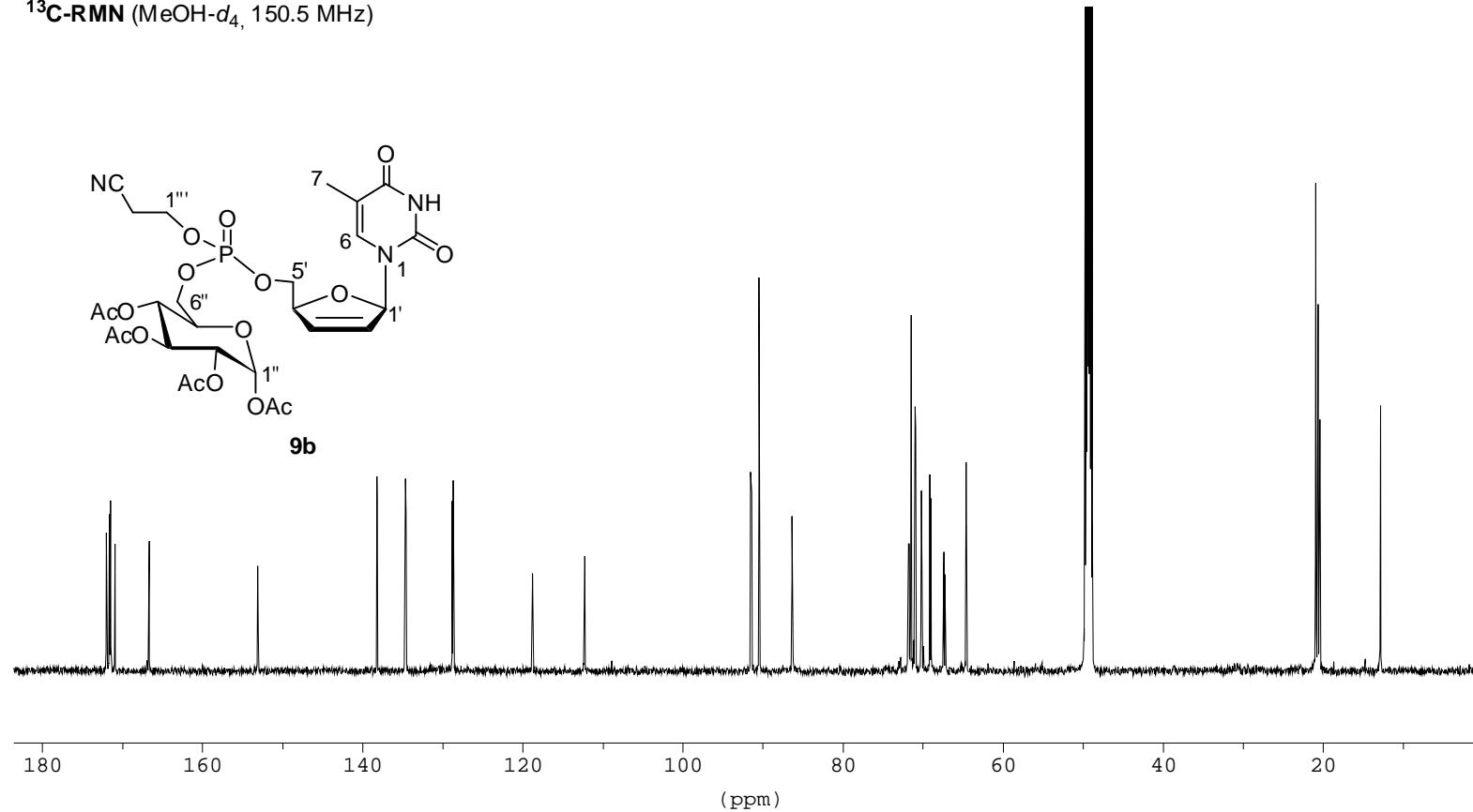
2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxythymidin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9b)

¹H-RMN (MeOH-*d*₄, 600 MHz)



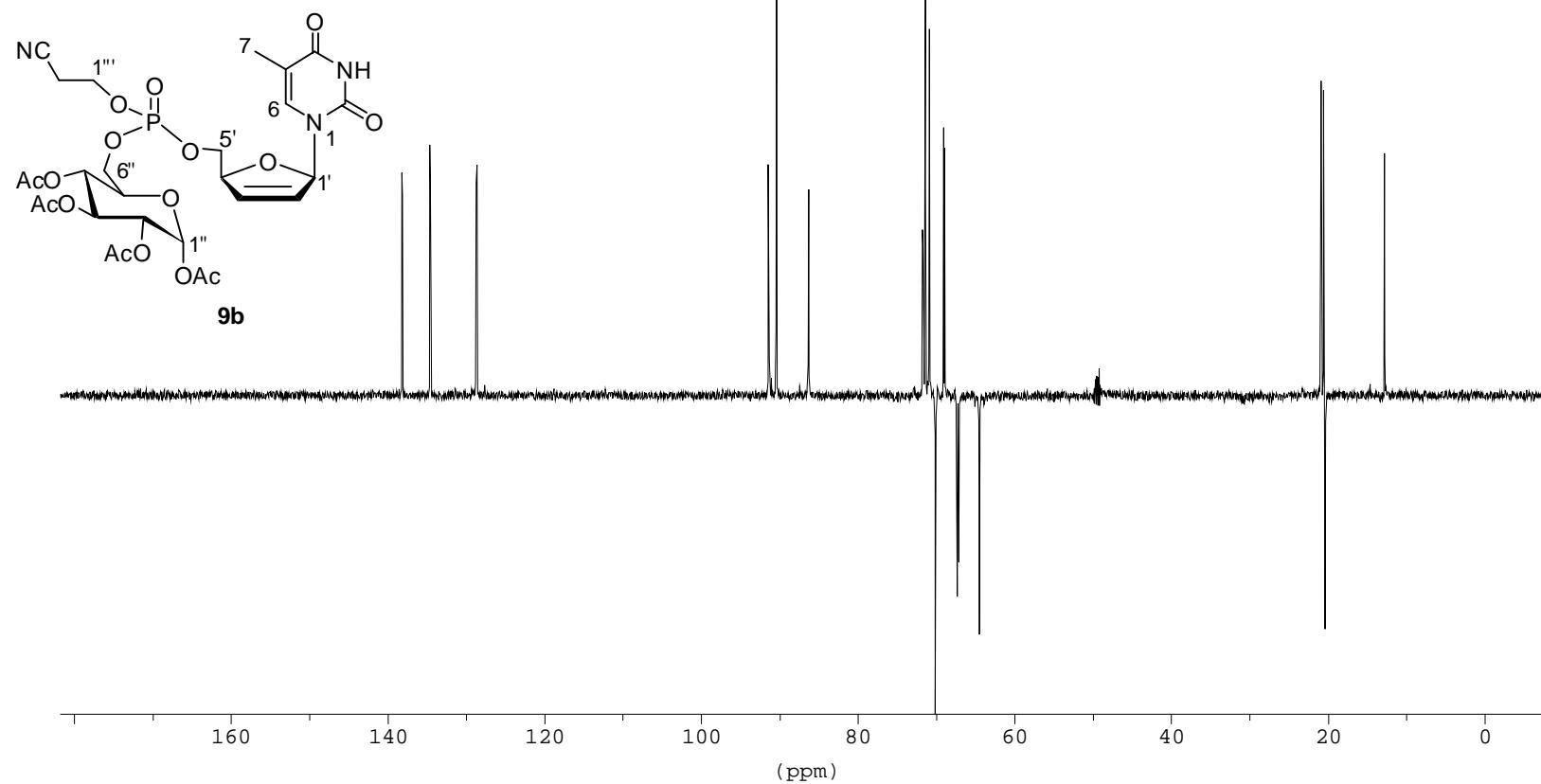
2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxythymidin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9b)

¹³C-RMN (MeOH-*d*₄, 150.5 MHz)



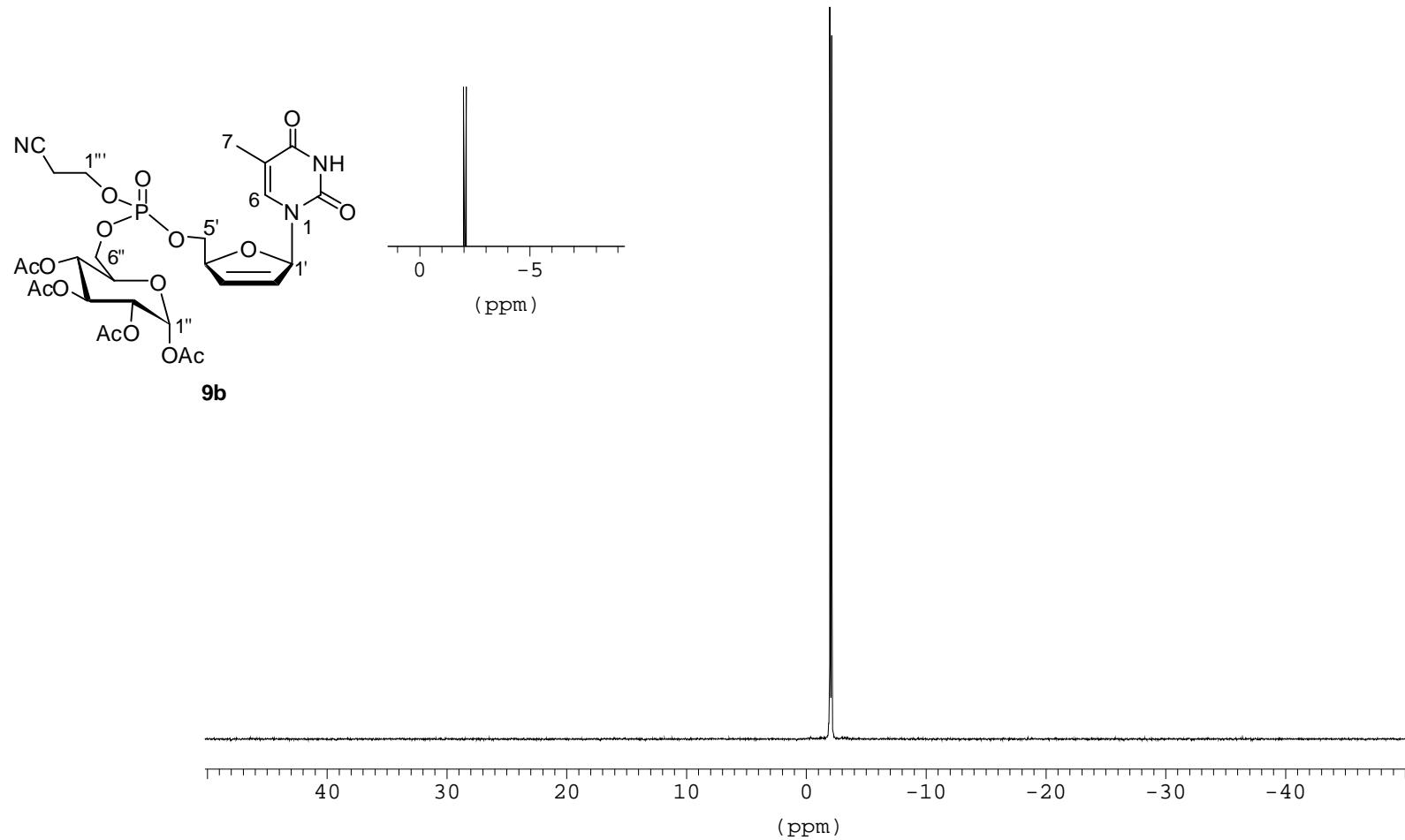
2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxythymidin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9b)

DEPT (MeOH-*d*₄, 150.5 MHz)

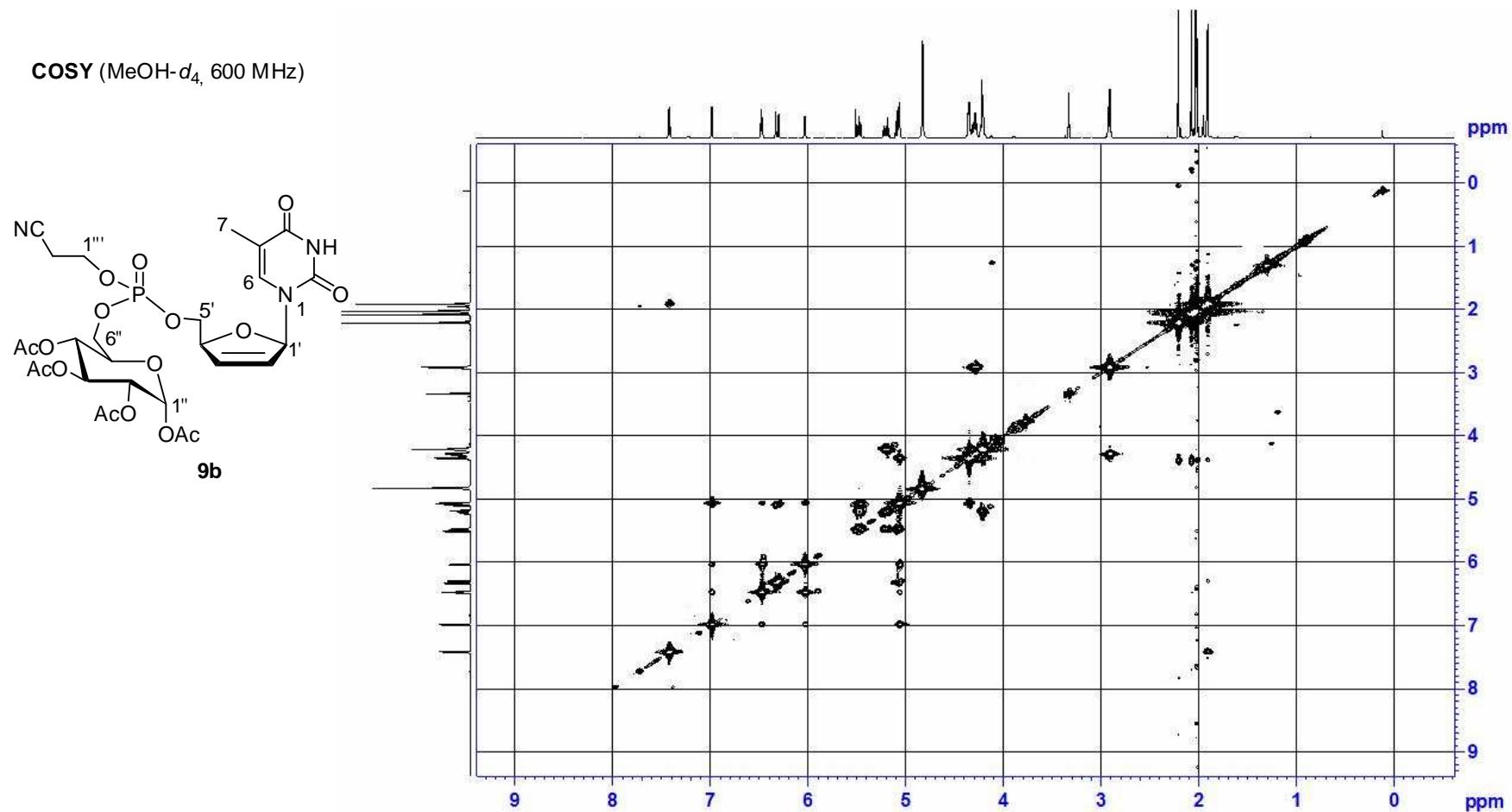


2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxythymidin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9b)

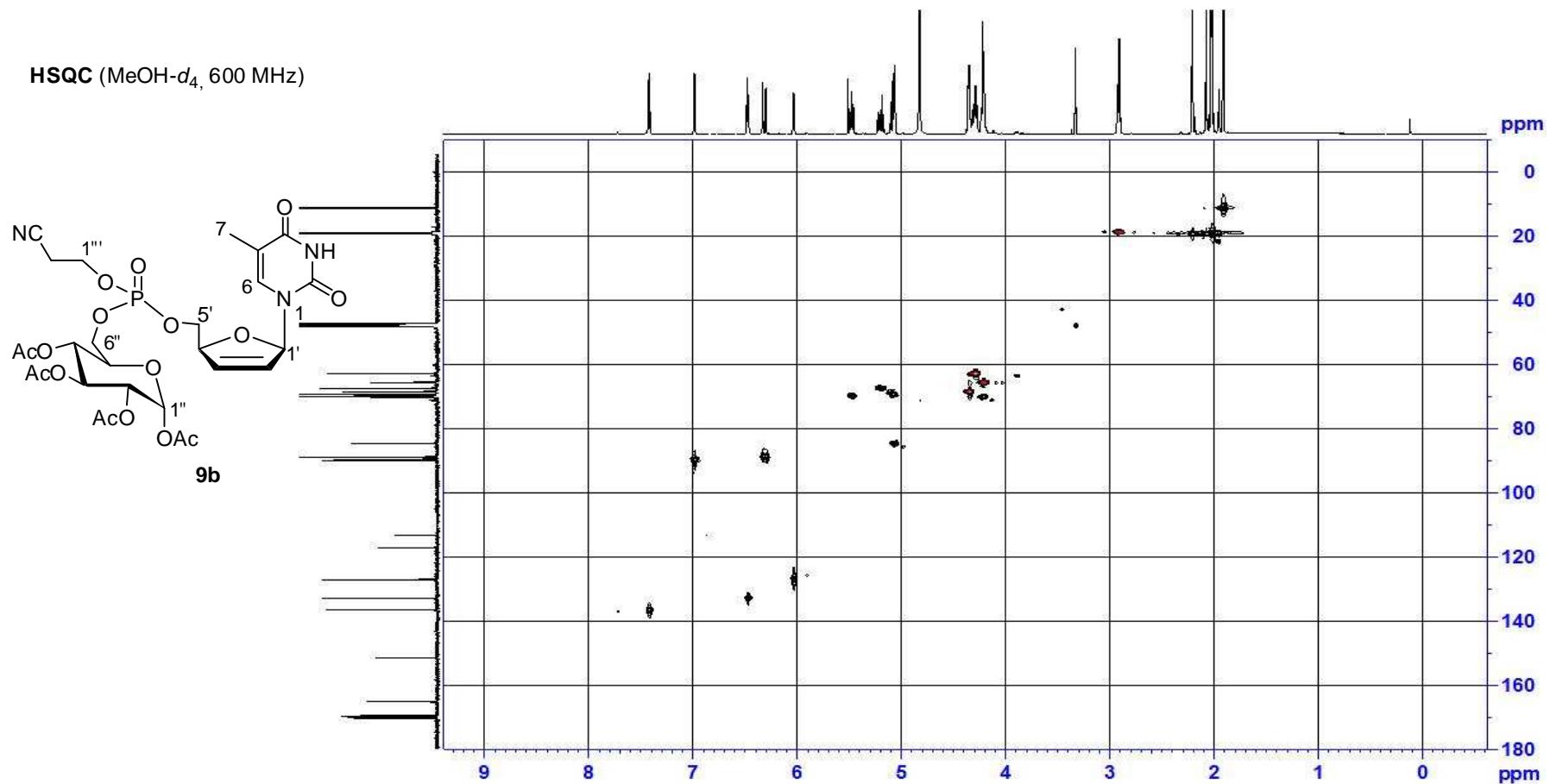
^{31}P -RMN (MeOH- d_4 , 242.9 MHz)



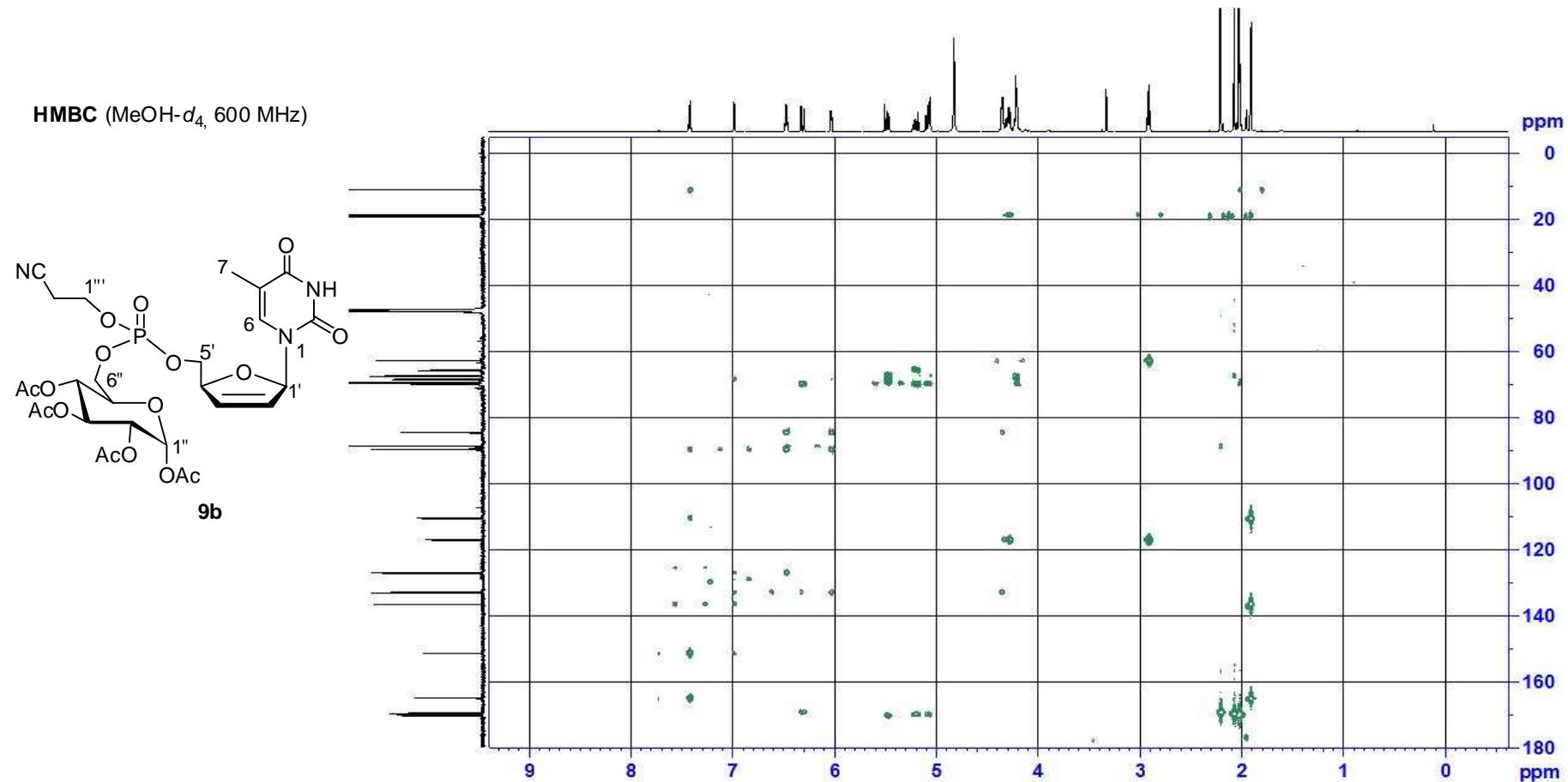
2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxythymidin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9b)



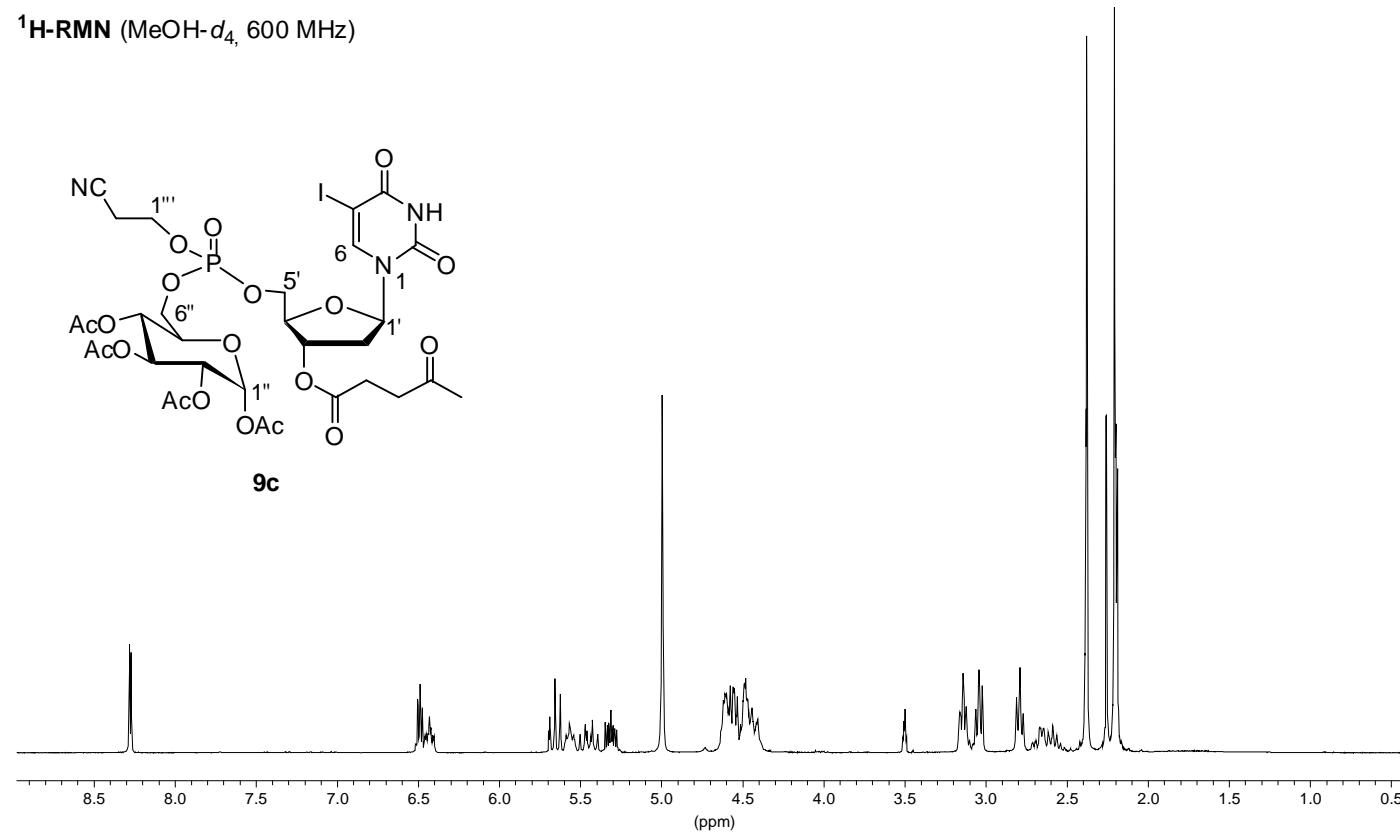
2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxythymidin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9b)



2-Cyanoethyl-(2',3'-didehydro-2',3'-dideoxythymidin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9b)

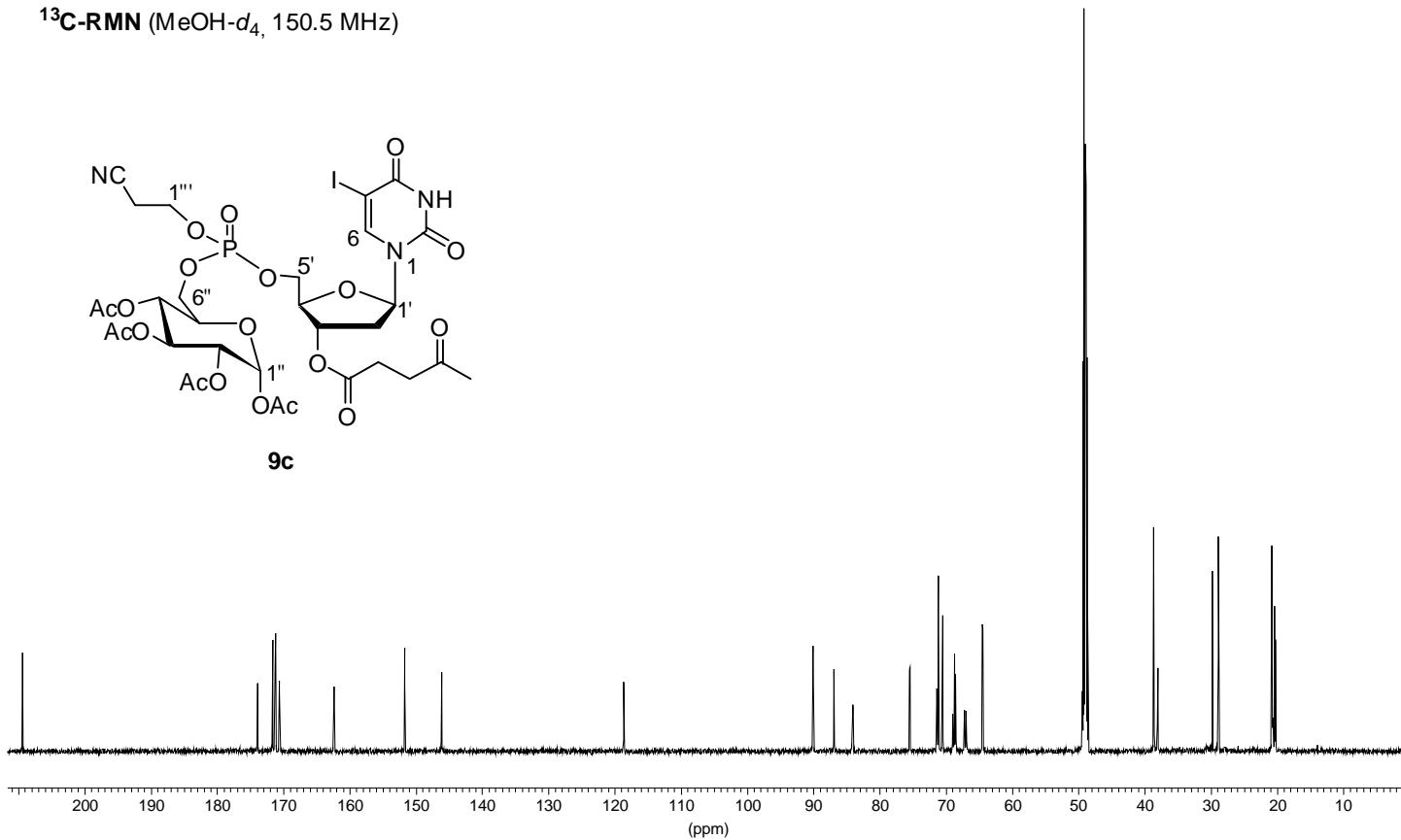
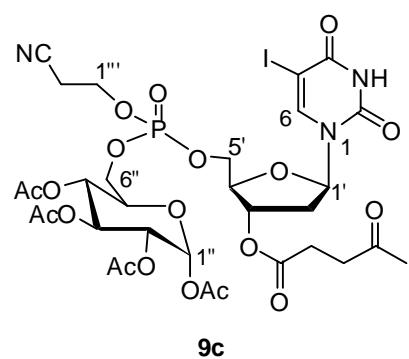


2-Cyanoethyl-(3'-*O*-levulinyl-5-iodo-2'-deoxyuridin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9c)



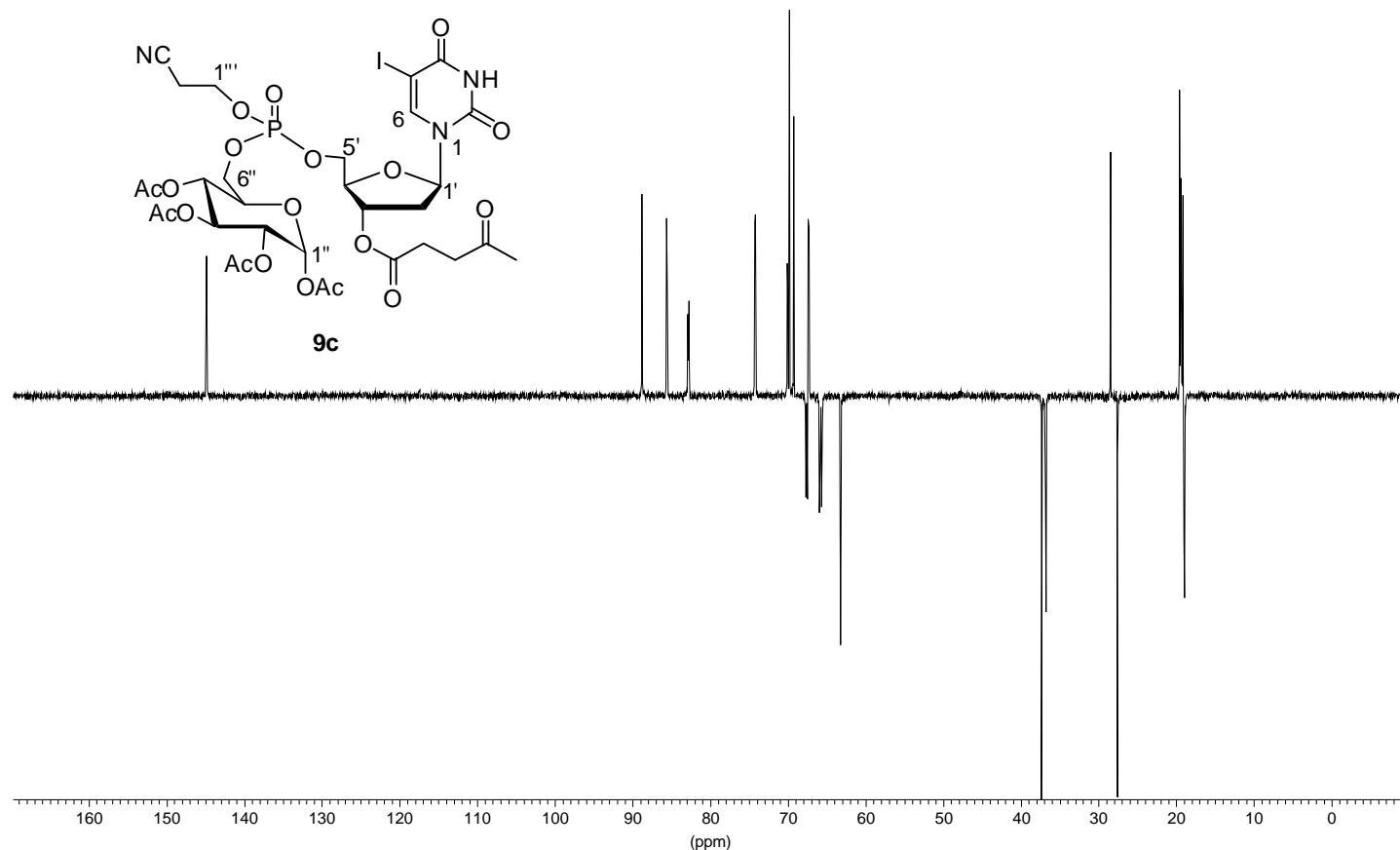
2-Cyanoethyl-(3'-*O*-levulinyl-5-iodo-2'-deoxyuridin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9c)

¹³C-RMN (MeOH-*d*₄, 150.5 MHz)



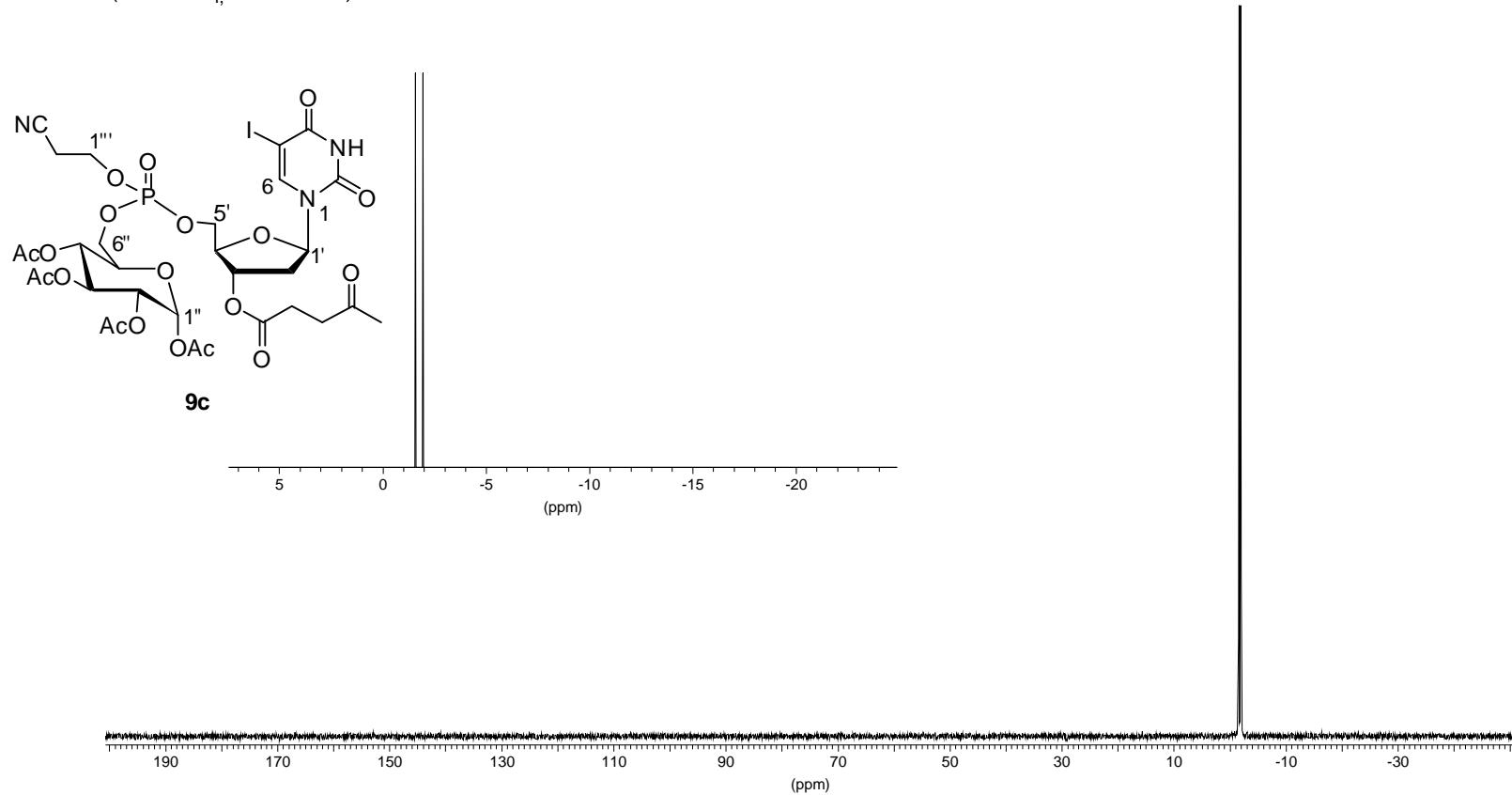
2-Cyanoethyl-(3'-*O*-levulinyl-5-iodo-2'-deoxyuridin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9c)

DEPT (MeOH-*d*₄, 150.5 MHz)

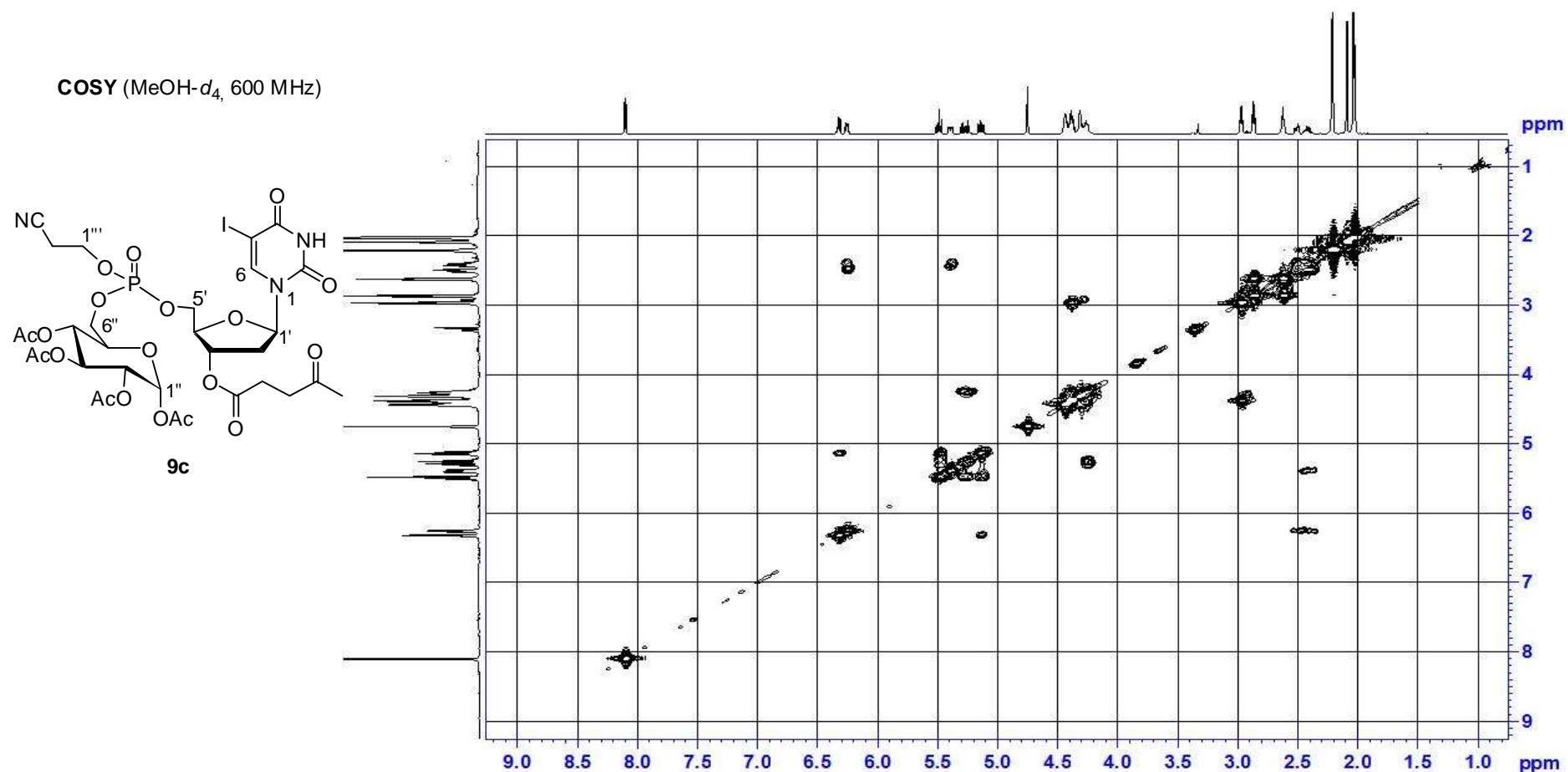


2-Cyanoethyl-(3'-*O*-levulinyl-5-iodo-2'-deoxyuridin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9c)

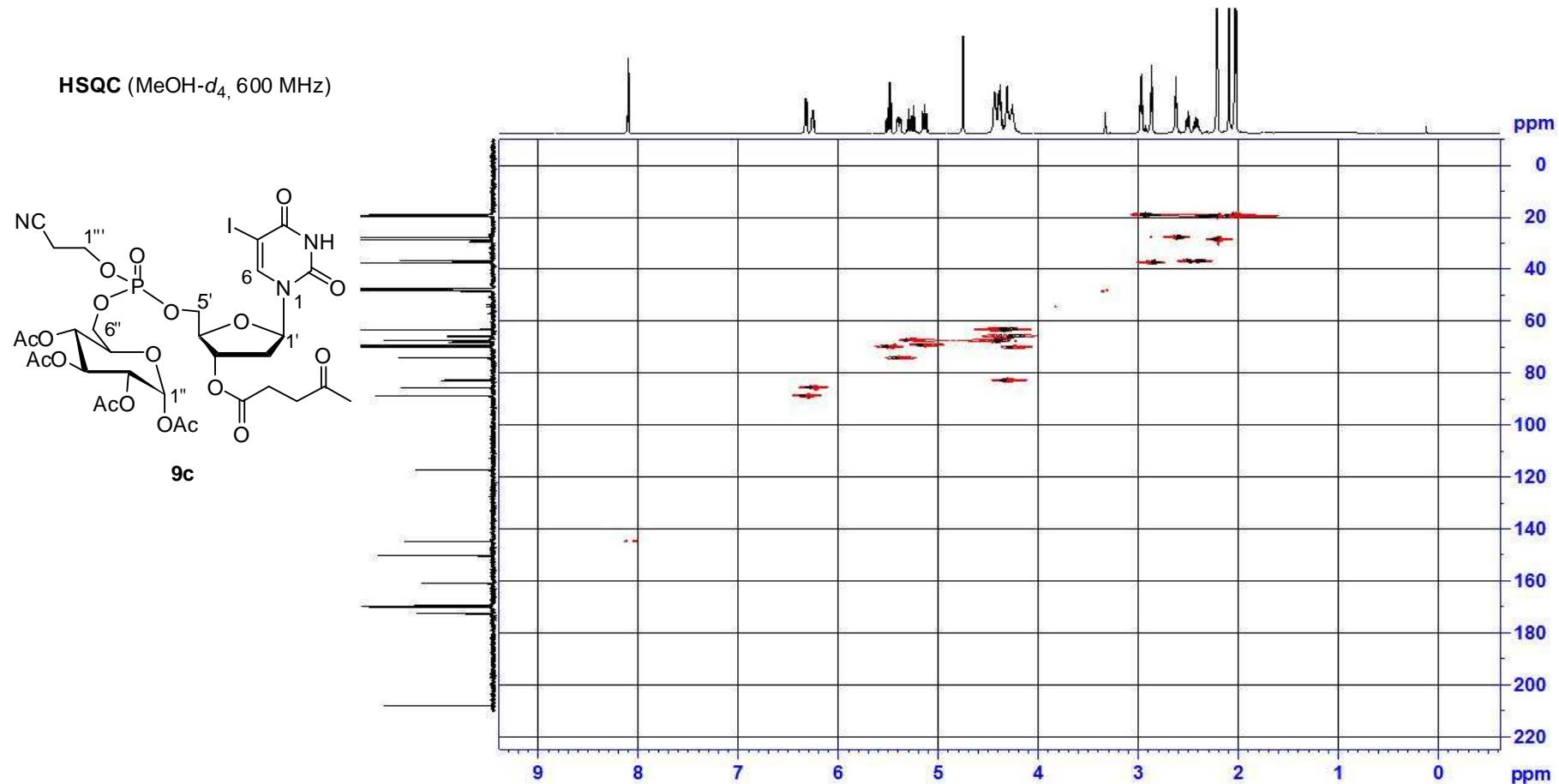
$^{31}\text{P-RMN}$ ($\text{MeOH-}d_4$, 242.9 MHz)



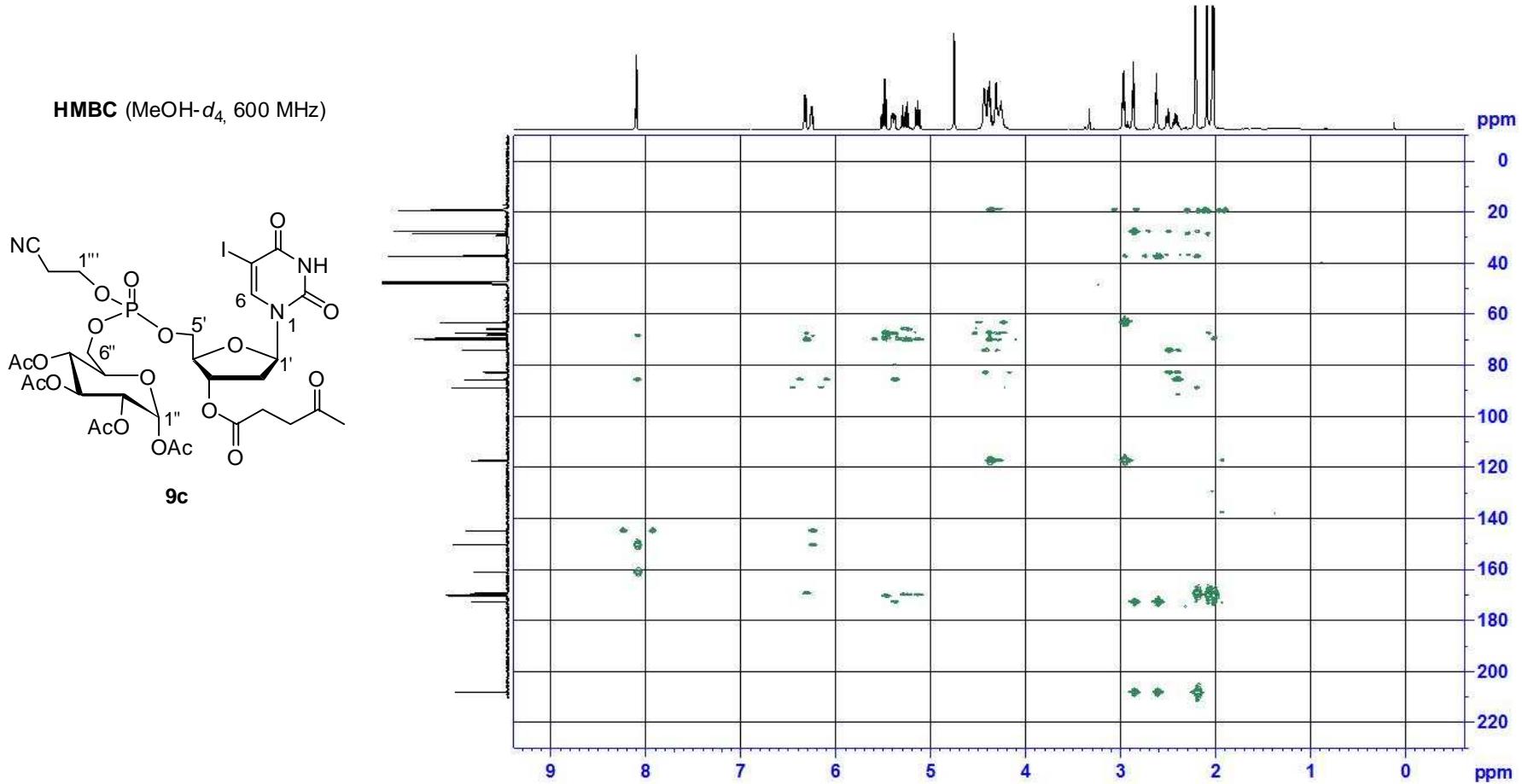
-Cyanoethyl-(3'-*O*-levulinyl-5-iodo-2'-deoxyuridin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9c)



*-Cyanoethyl-(3'-*O*-levulinyl-5-iodo-2'-deoxyuridin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9c)*

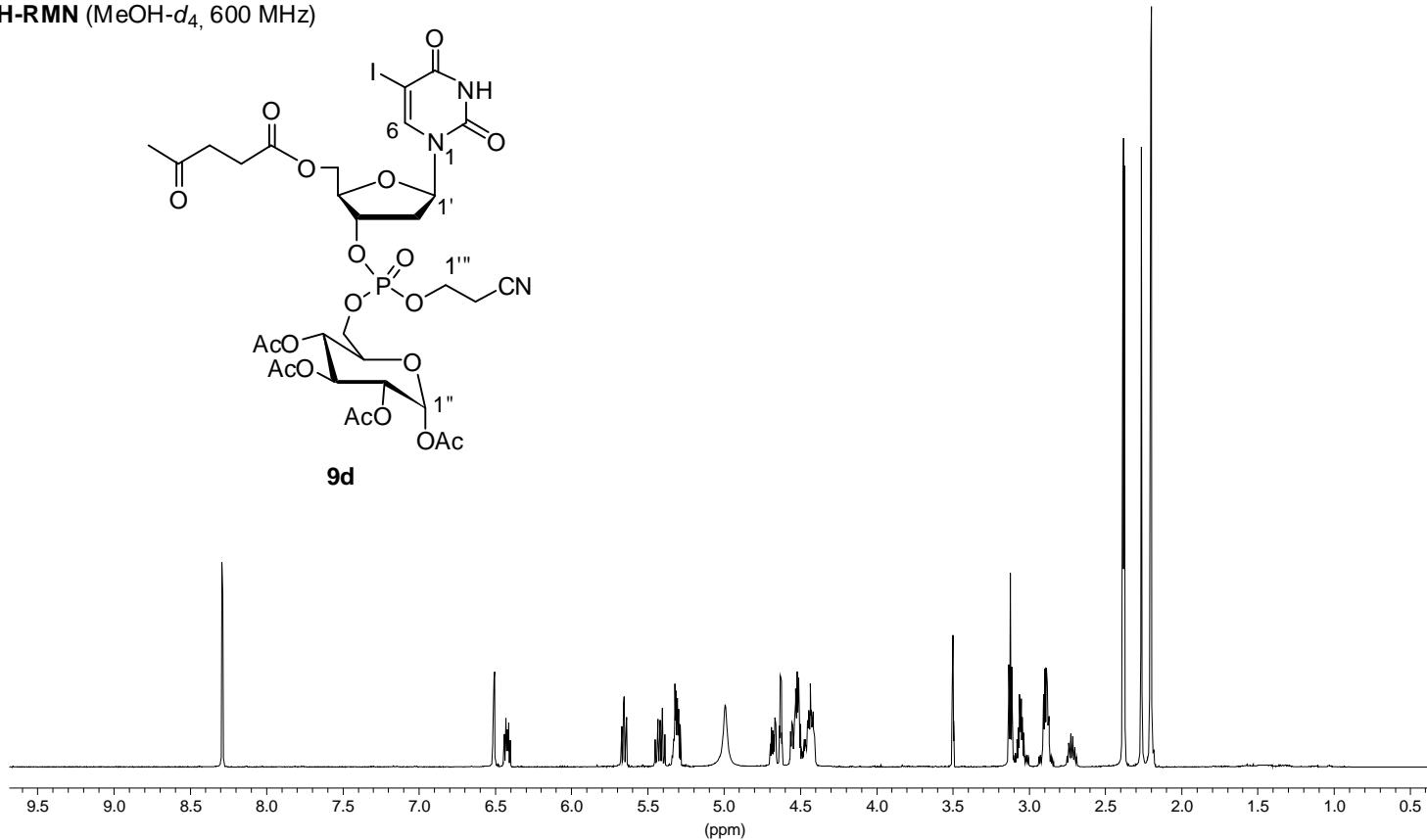
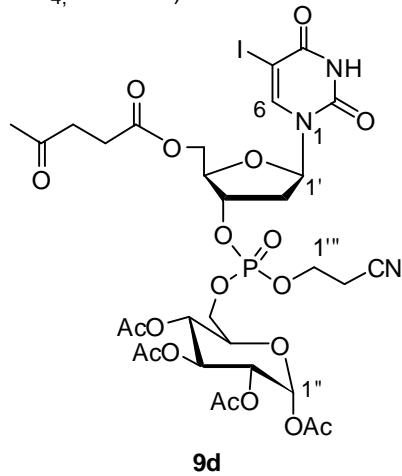


-Cyanoethyl-(3'-*O*-levulinyl-5-iodo-2'-deoxyuridin-5'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9c)



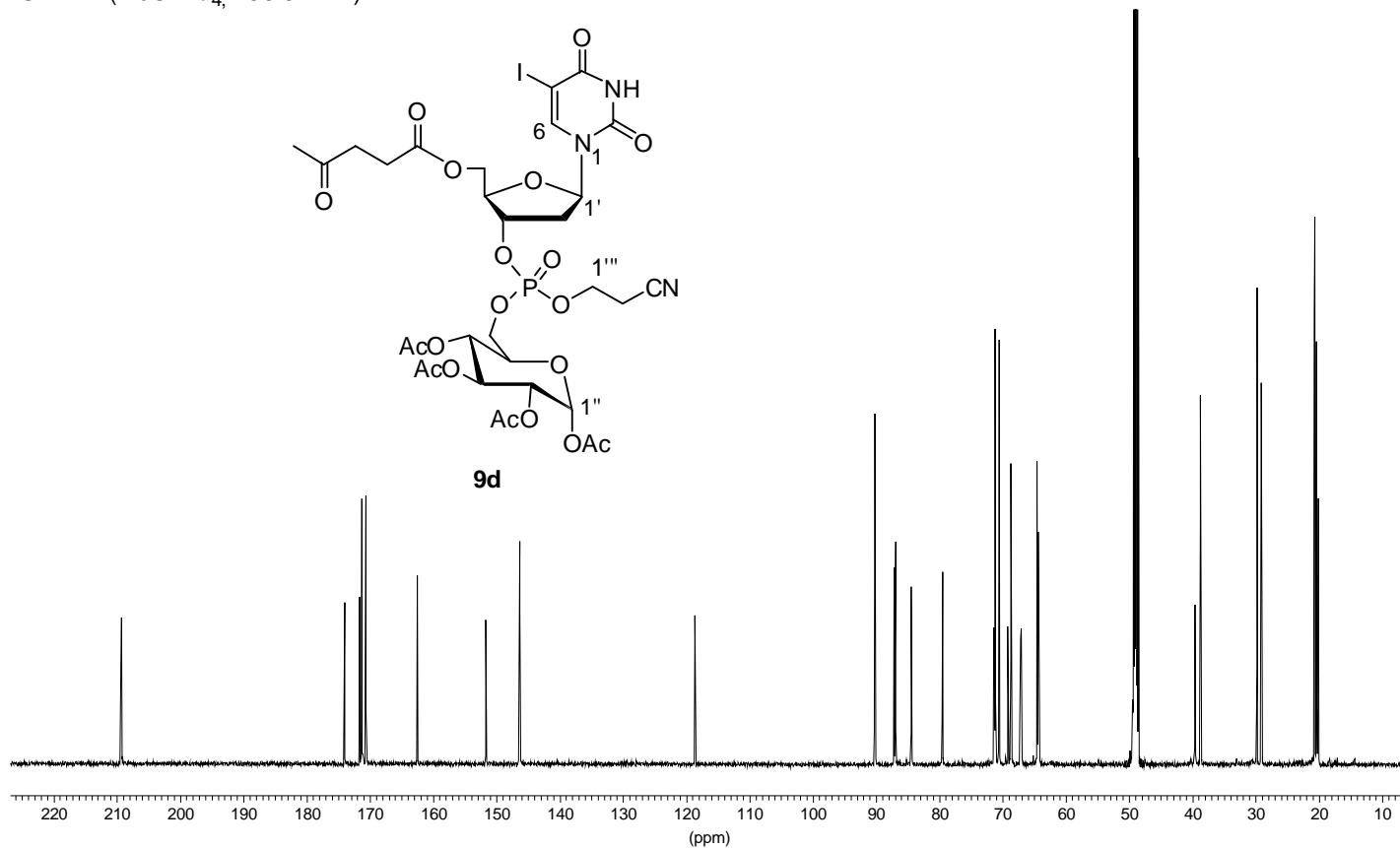
2-Cyanoethyl-(5'-*O*-levulinyl-5-iodo-2'-deoxyuridin-3'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9d)

¹H-RMN (MeOH-*d*₄, 600 MHz)

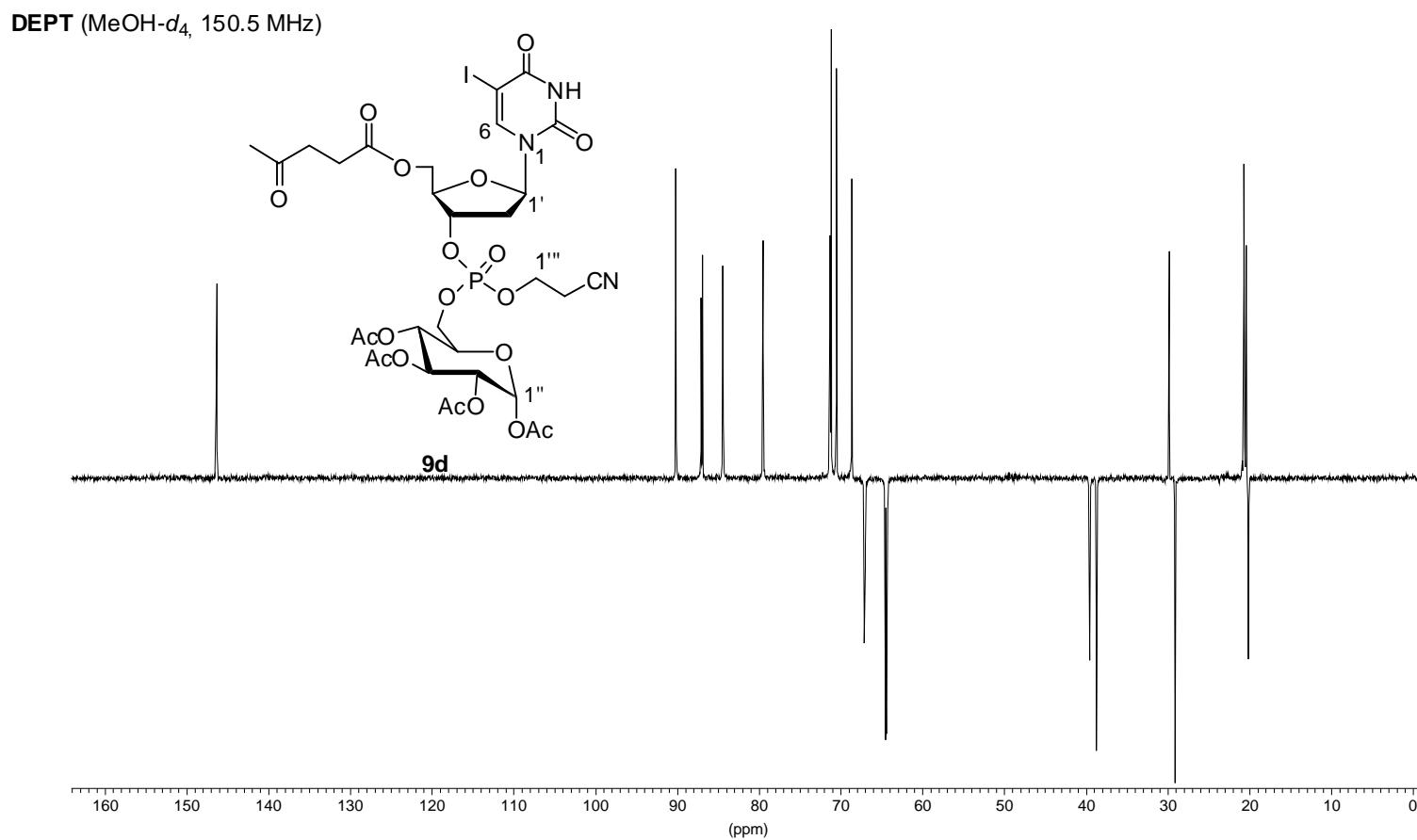


2-Cyanoethyl-(5'-*O*-levulinyl-5-iodo-2'-deoxyuridin-3'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9d)

¹³C-RMN (MeOH-*d*₄, 150.5 MHz)

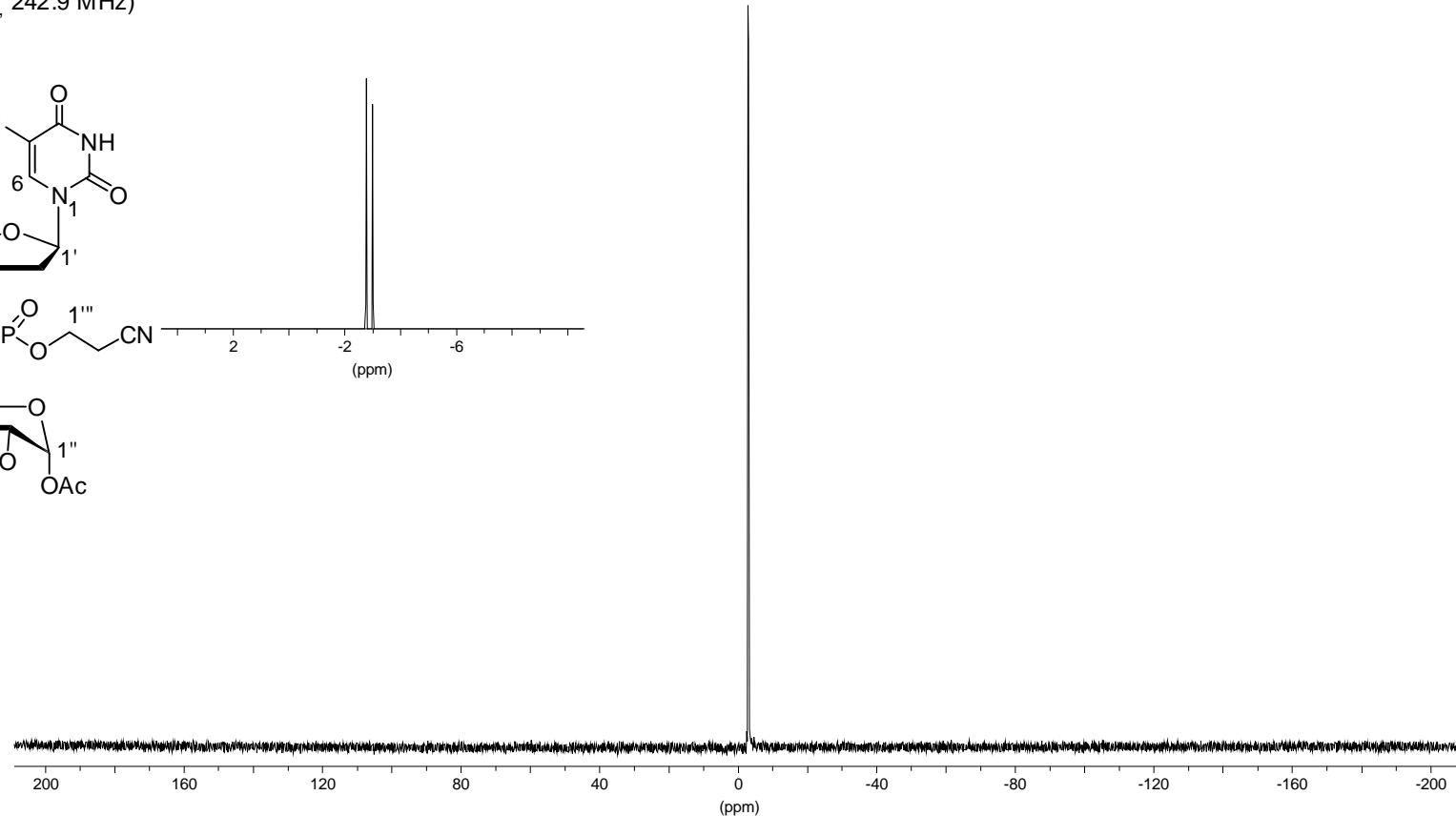
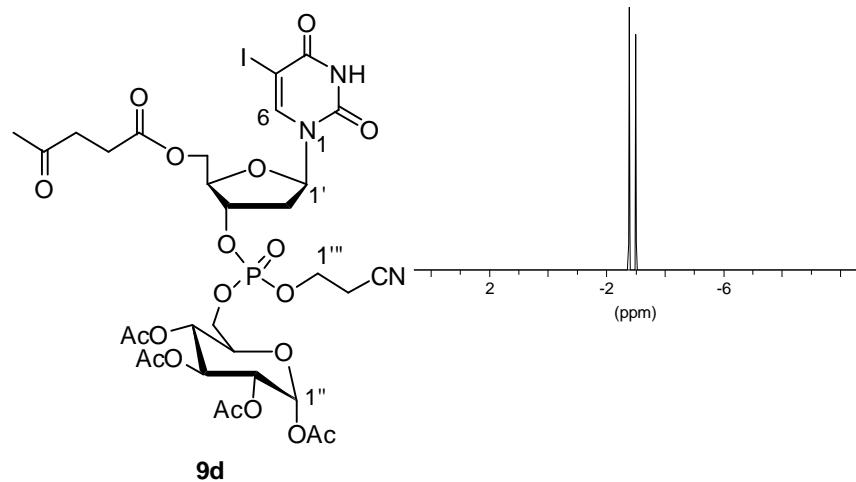


2-Cyanoethyl-(5'-*O*-levulinyl-5-iodo-2'-deoxyuridin-3'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9d)

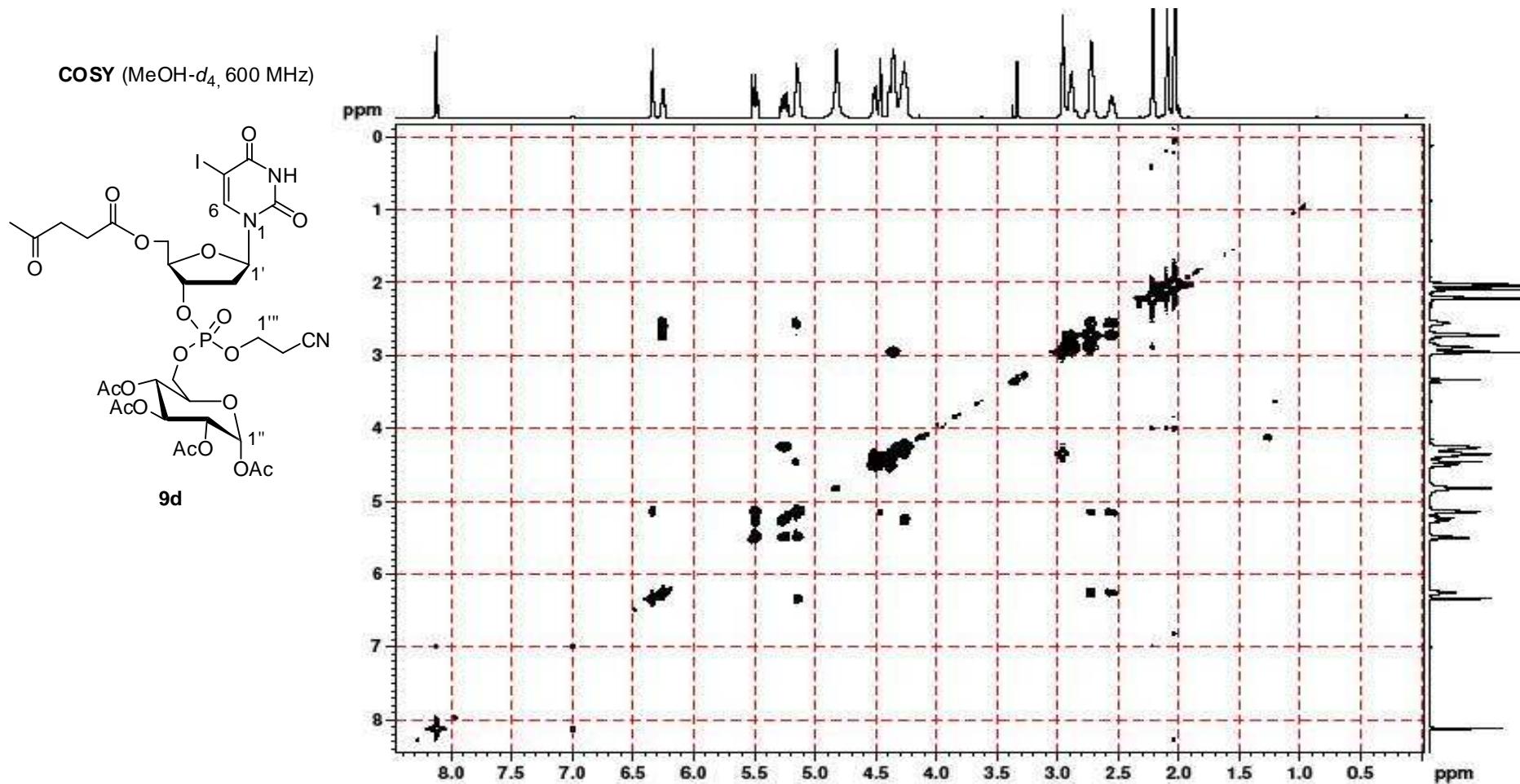


2-Cyanoethyl-(5'-*O*-levulinyl-5-iodo-2'-deoxyuridin-3'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9d)

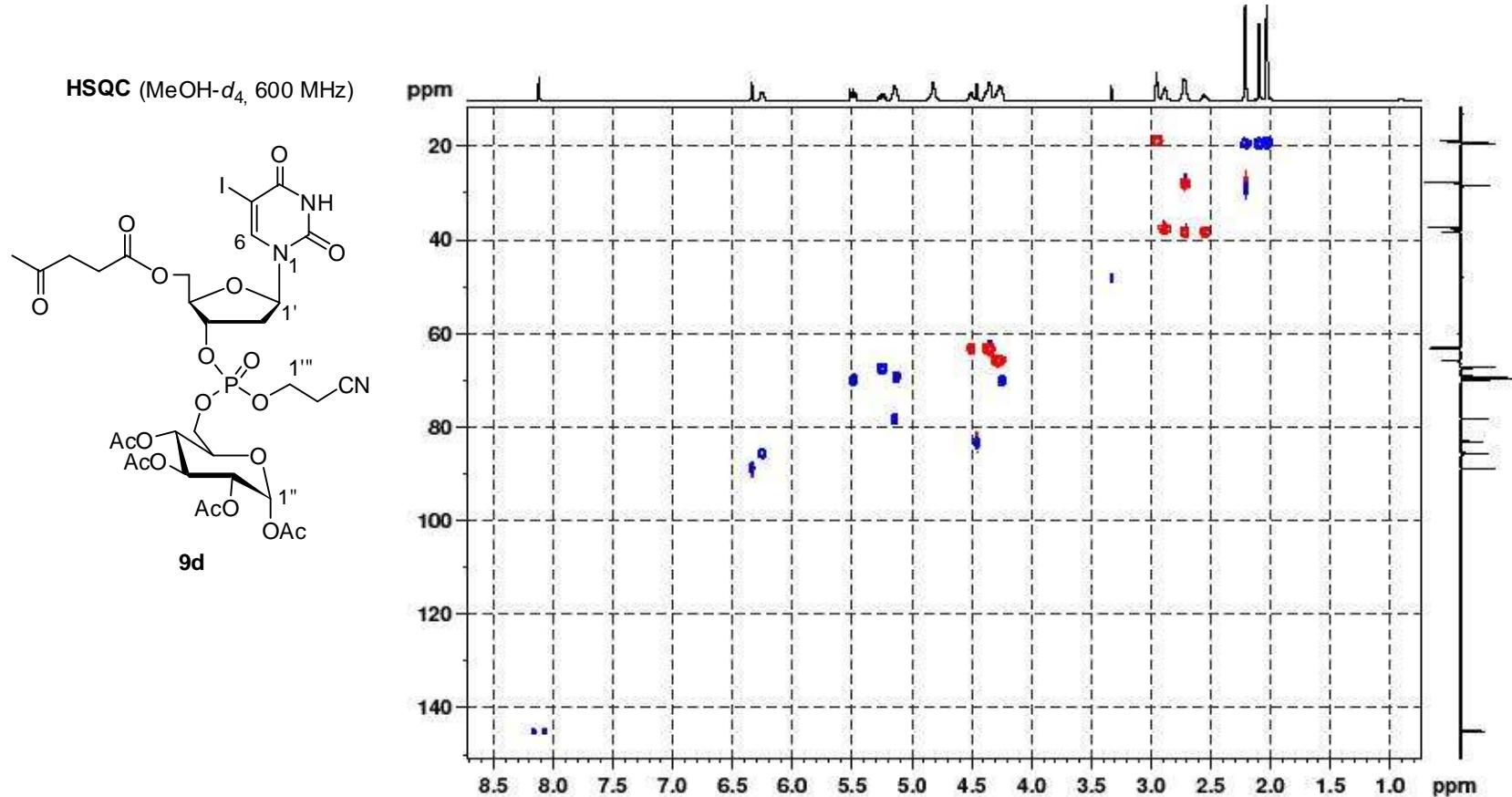
^{31}P -RMN (MeOH- d_4 , 242.9 MHz)



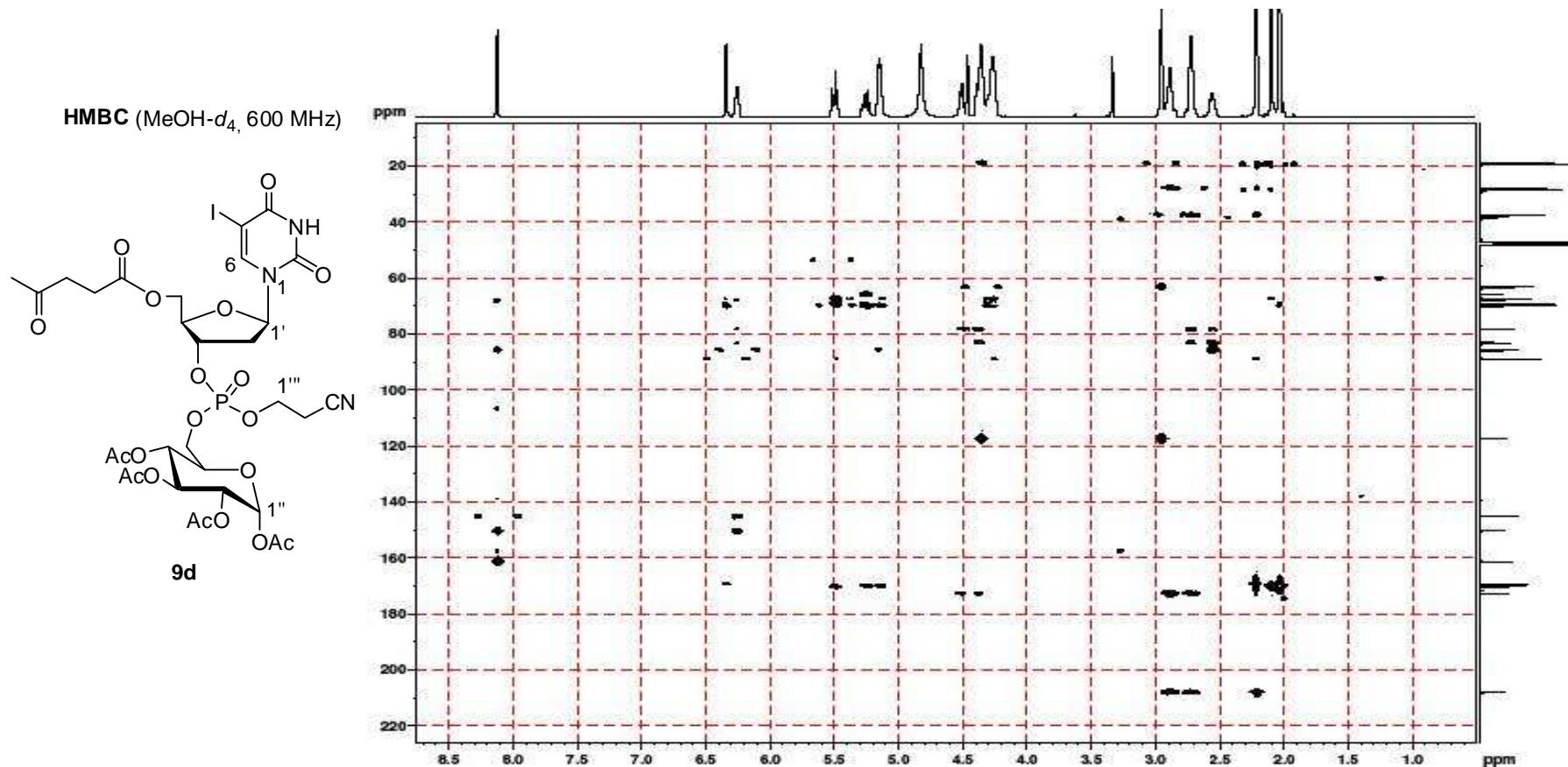
2-Cyanoethyl-(5'-*O*-levulinyl-5-iodo-2'-deoxyuridin-3'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9d)



2-Cyanoethyl-(5'-*O*-levulinyl-5-iodo-2'-deoxyuridin-3'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9d)

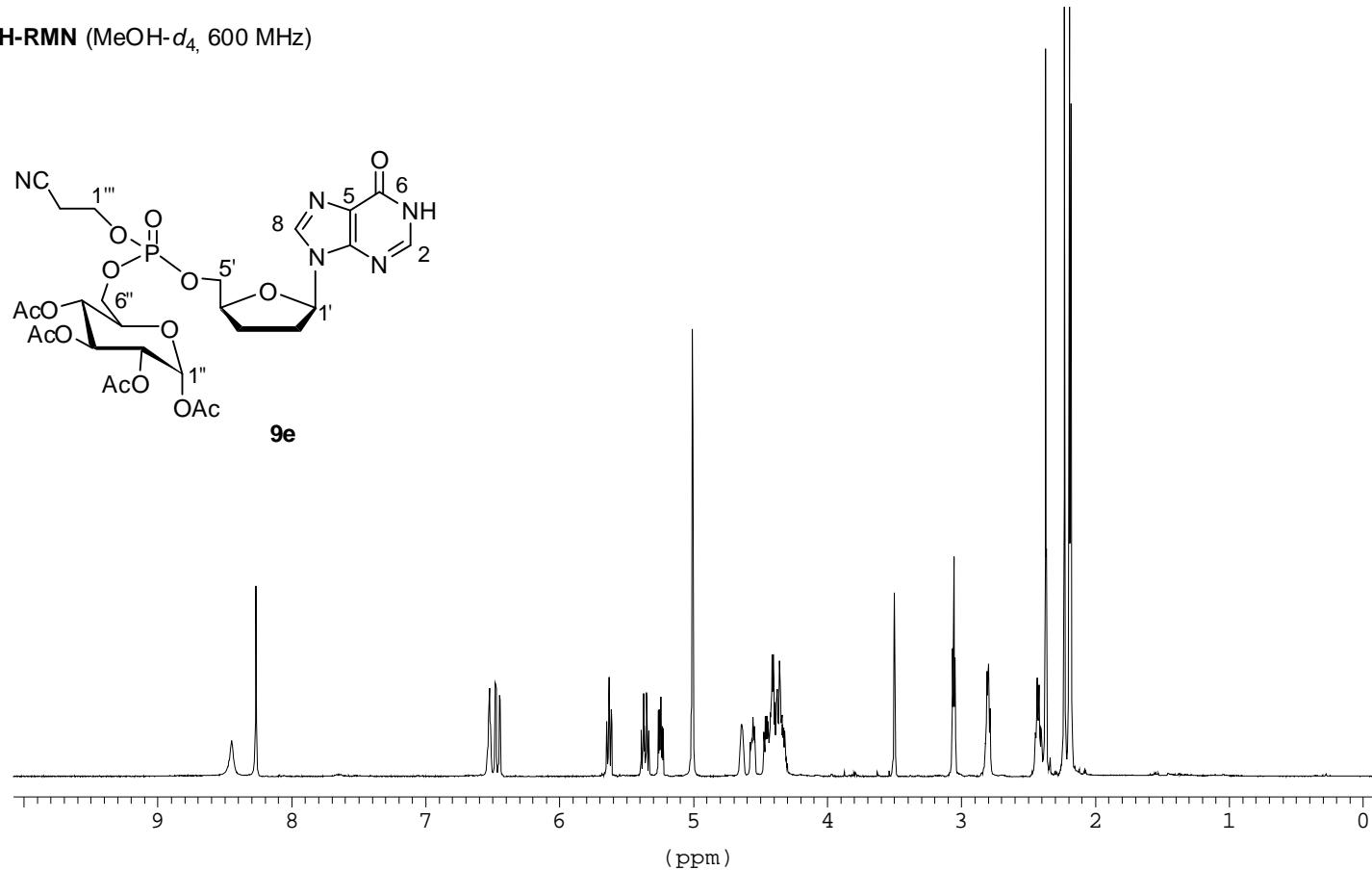
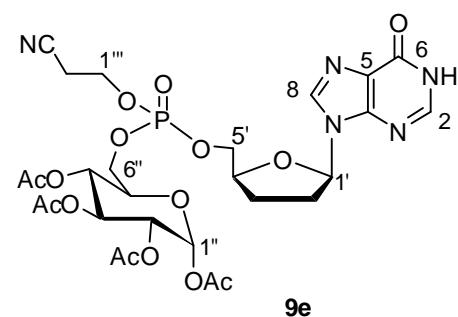


2-Cyanoethyl-(5'-*O*-levulinyl-5-iodo-2'-deoxyuridin-3'-yl)-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9d)



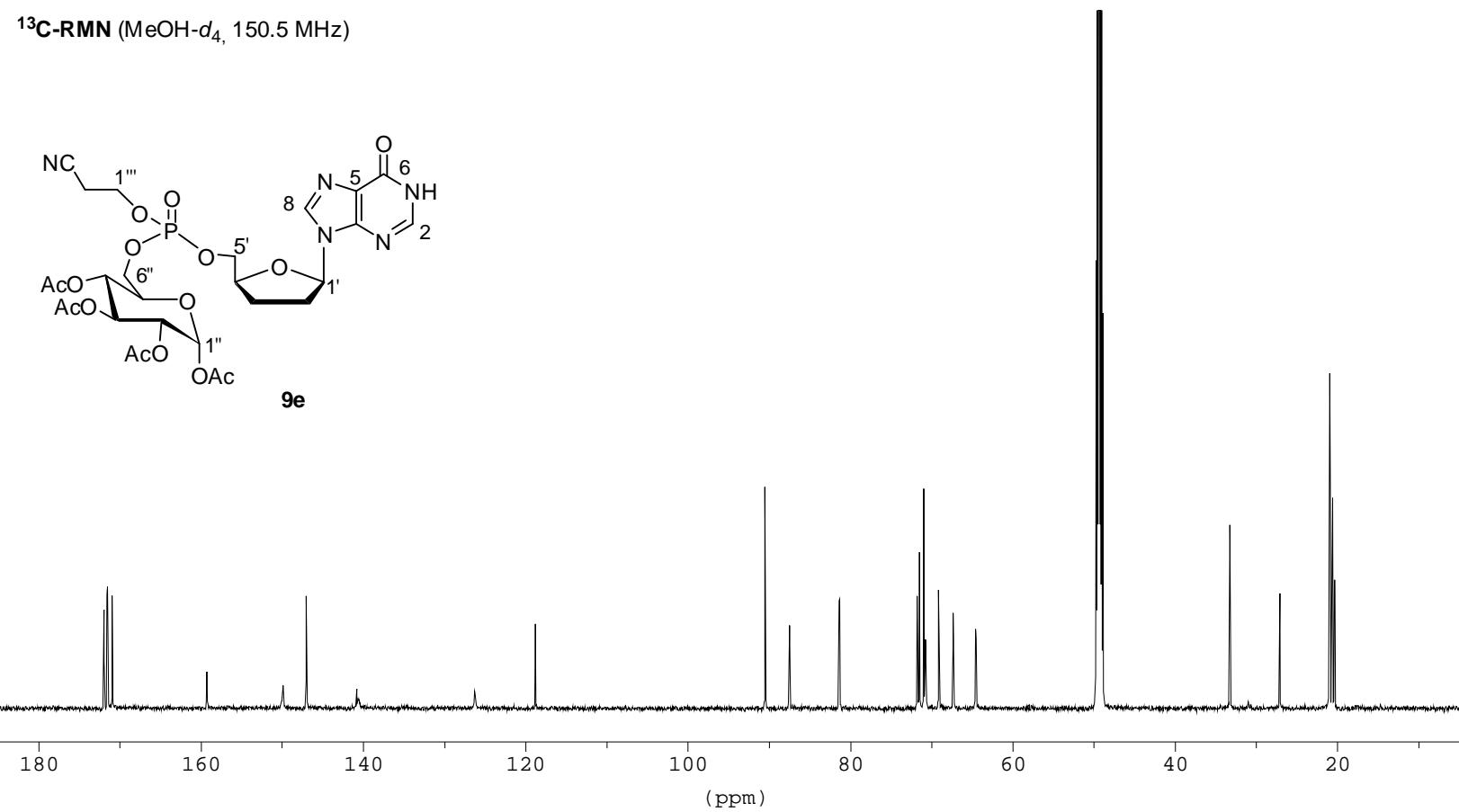
2-Cyanoethyl-(2',3'-dideoxyinosin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9e)

$^1\text{H-RMN}$ ($\text{MeOH-}d_4$, 600 MHz)



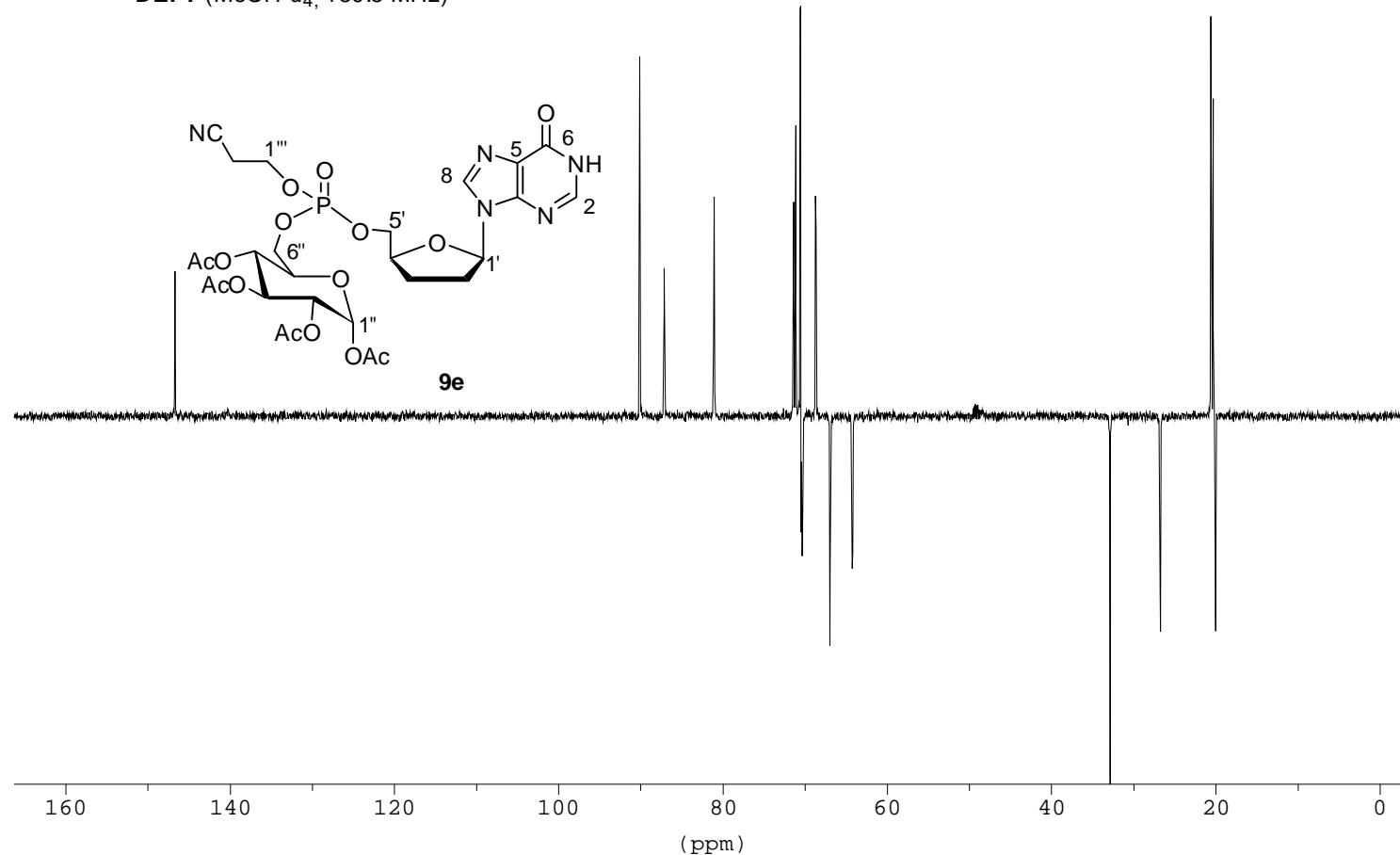
2-Cyanoethyl-(2',3'-dideoxyinosin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9e)

^{13}C -RMN (MeOH- d_4 , 150.5 MHz)



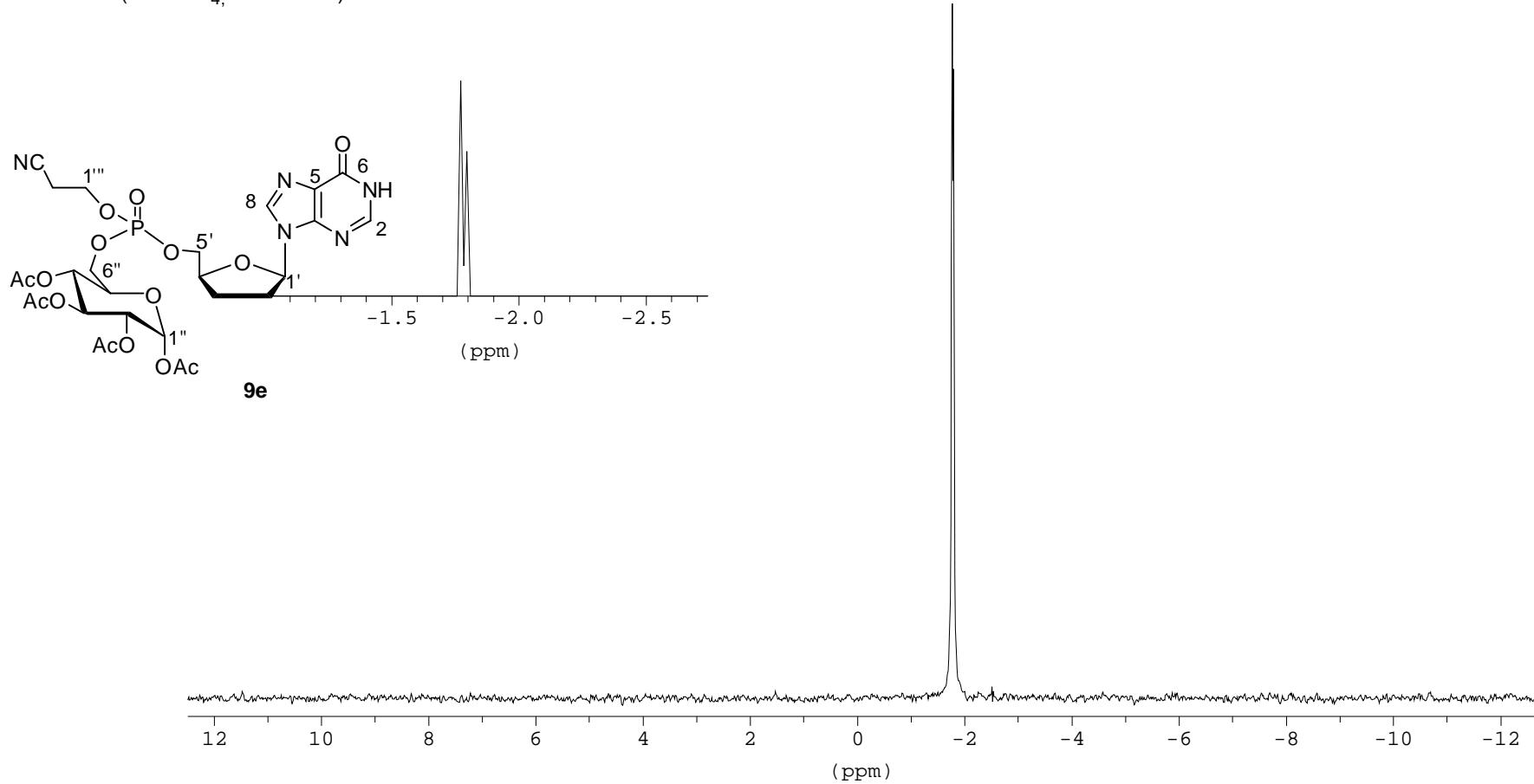
2-Cyanoethyl-(2',3'-dideoxyinosin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9e)

DEPT (MeOH-*d*₄, 150.5 MHz)

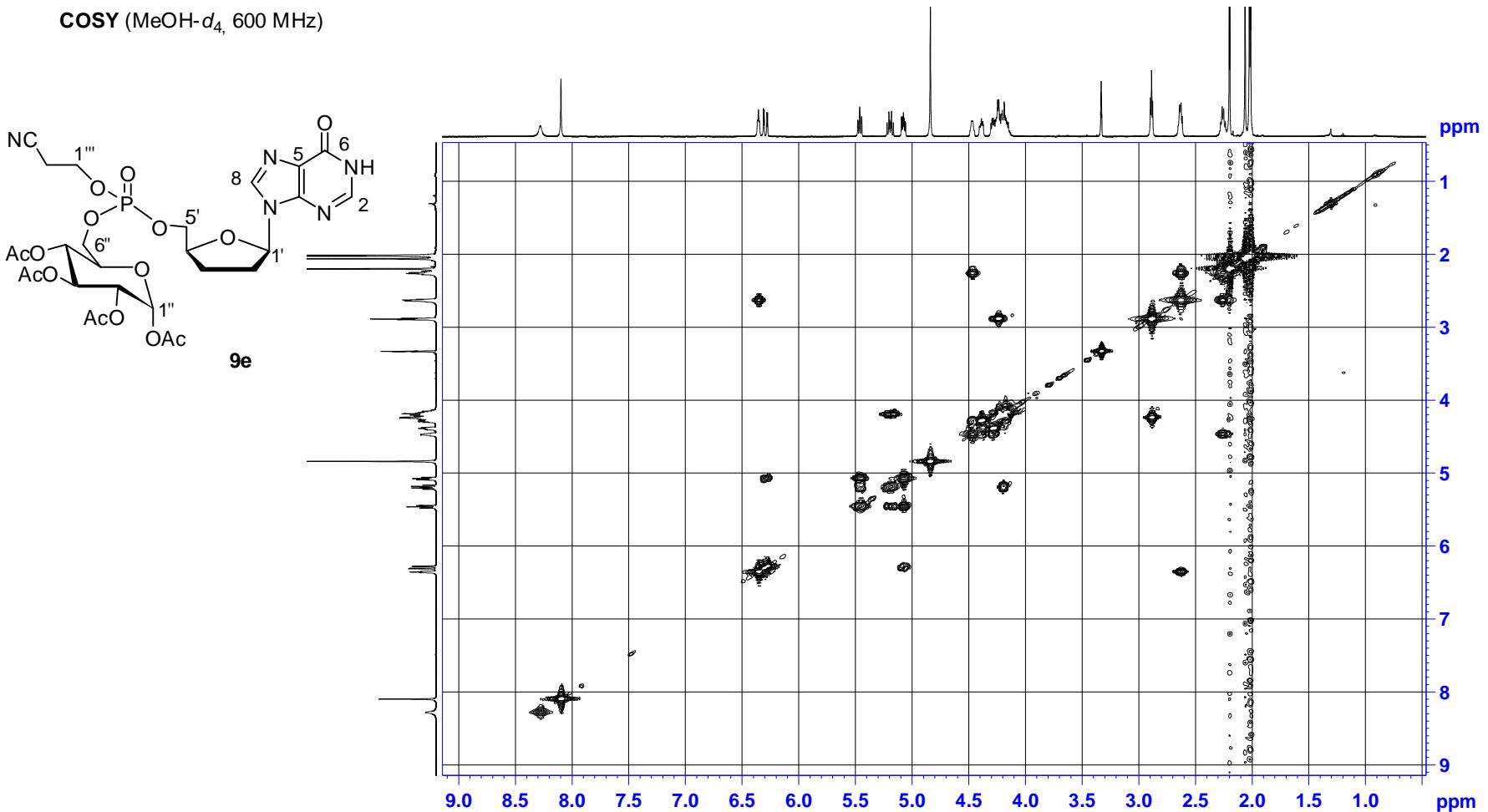


2-Cyanoethyl-(2',3'-dideoxyinosin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9e)

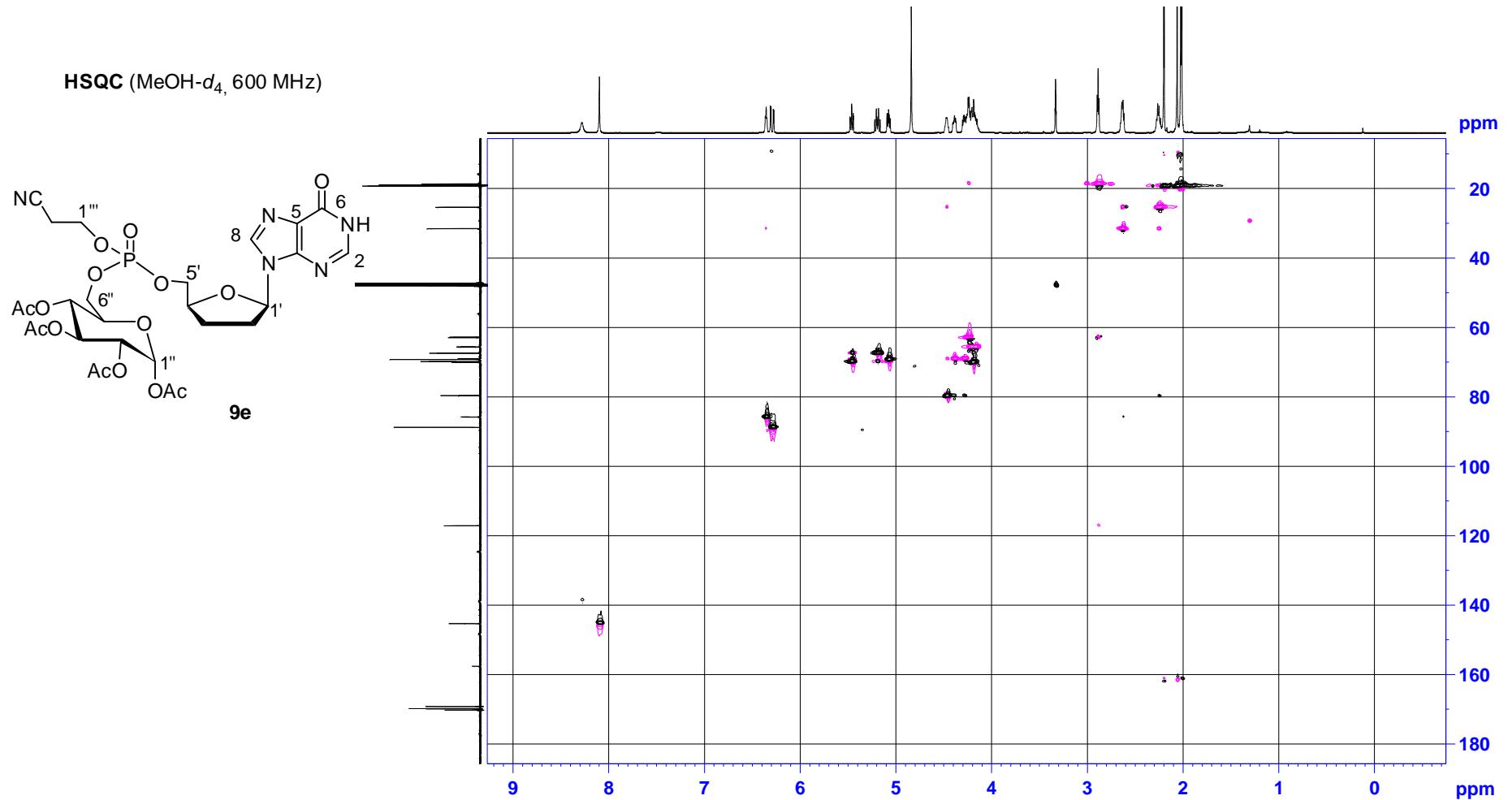
³¹P-RMN (MeOH-*d*₄, 242.9 MHz)



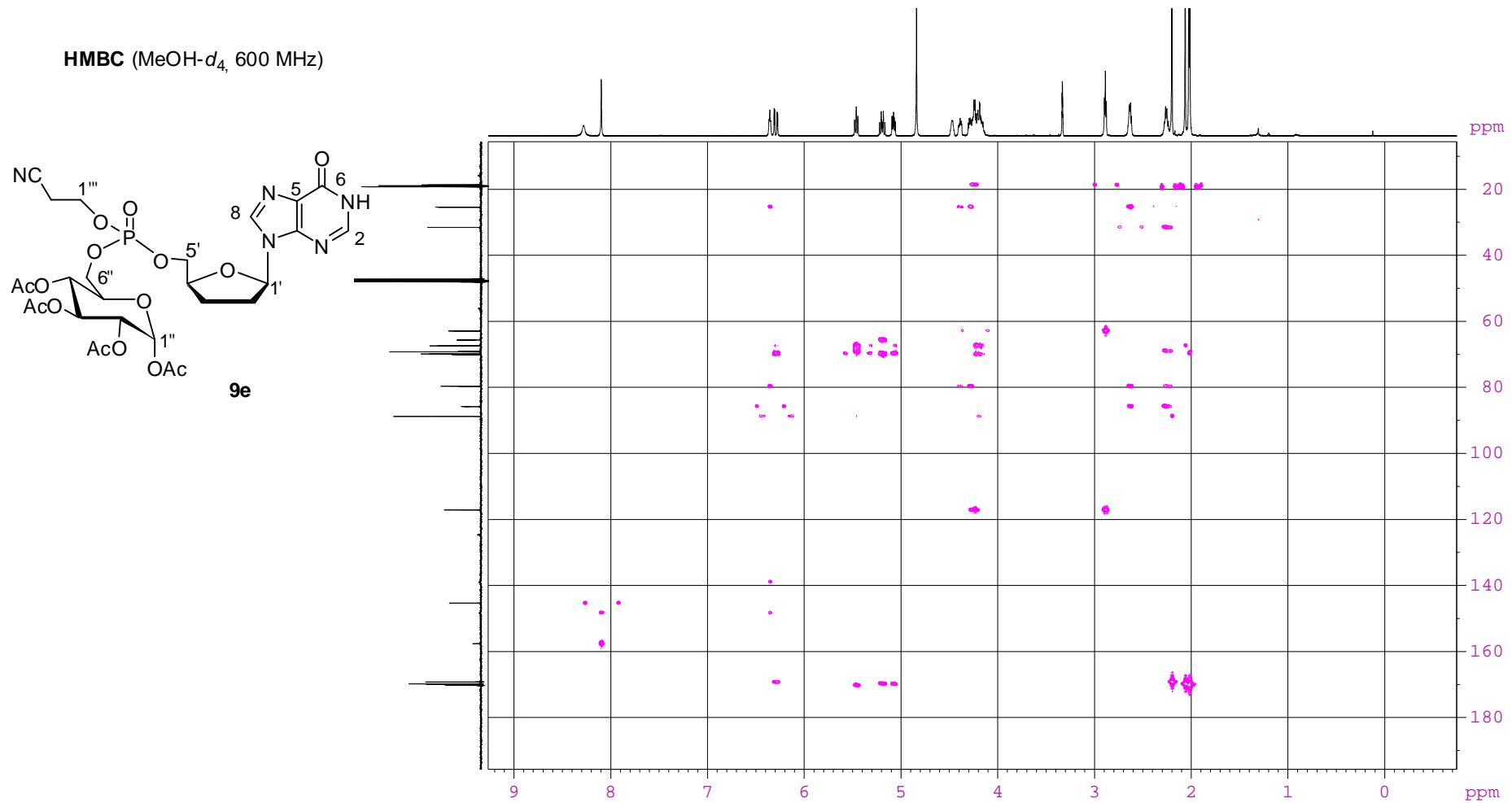
2-Cyanoethyl-(2',3'-dideoxyinosin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9e)



2-Cyanoethyl-(2',3'-dideoxyinosin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9e)

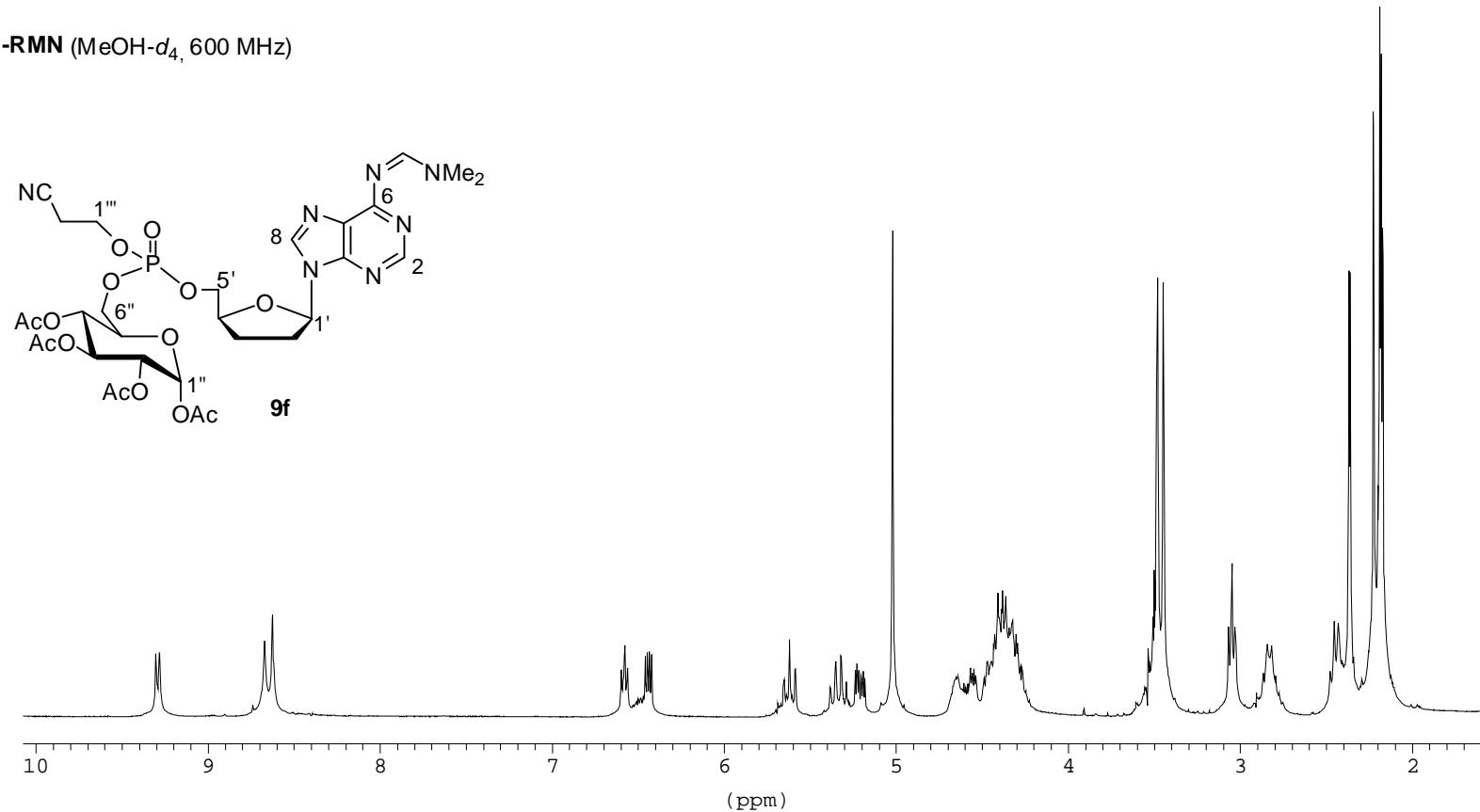


2-Cyanoethyl-(2',3'-dideoxyinosin-5'-yl)-1,2,3,4-tetra-O-acetyl- α -D-glucopyranos-6-yl phosphate (9e)



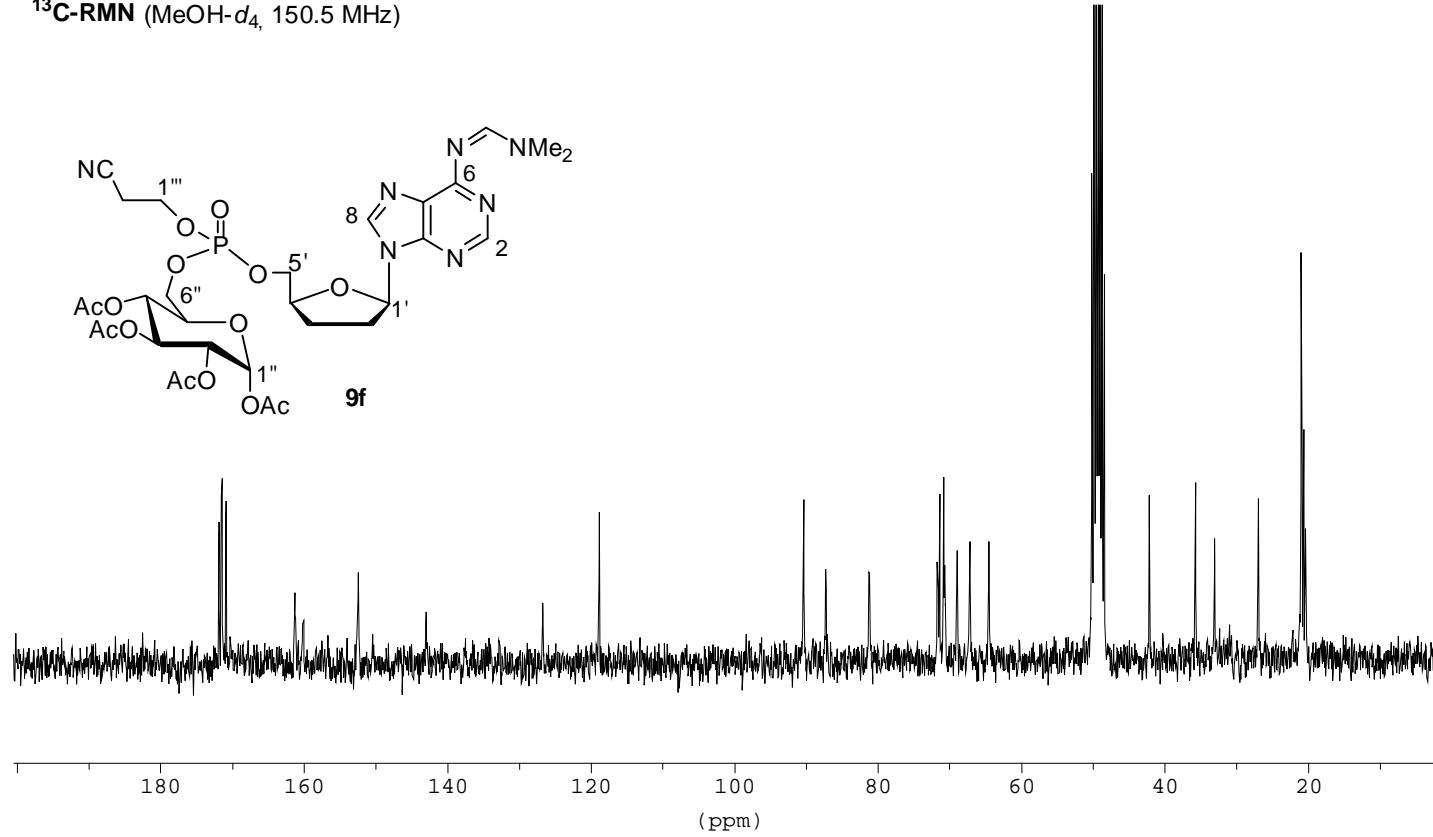
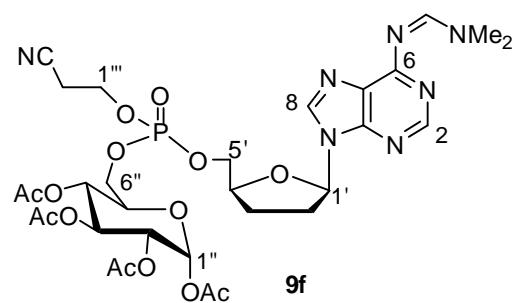
**2-Cyanoethyl-[*N*⁶-(dimethylamino)methylen-2',3'-dideoxyadenosin-5'-yl]-1,2,3,4-tetra-*O*-acetyl-
α-D-glucopyranos-6-yl phosphate (9f)**

¹H-RMN (MeOH-*d*₄, 600 MHz)



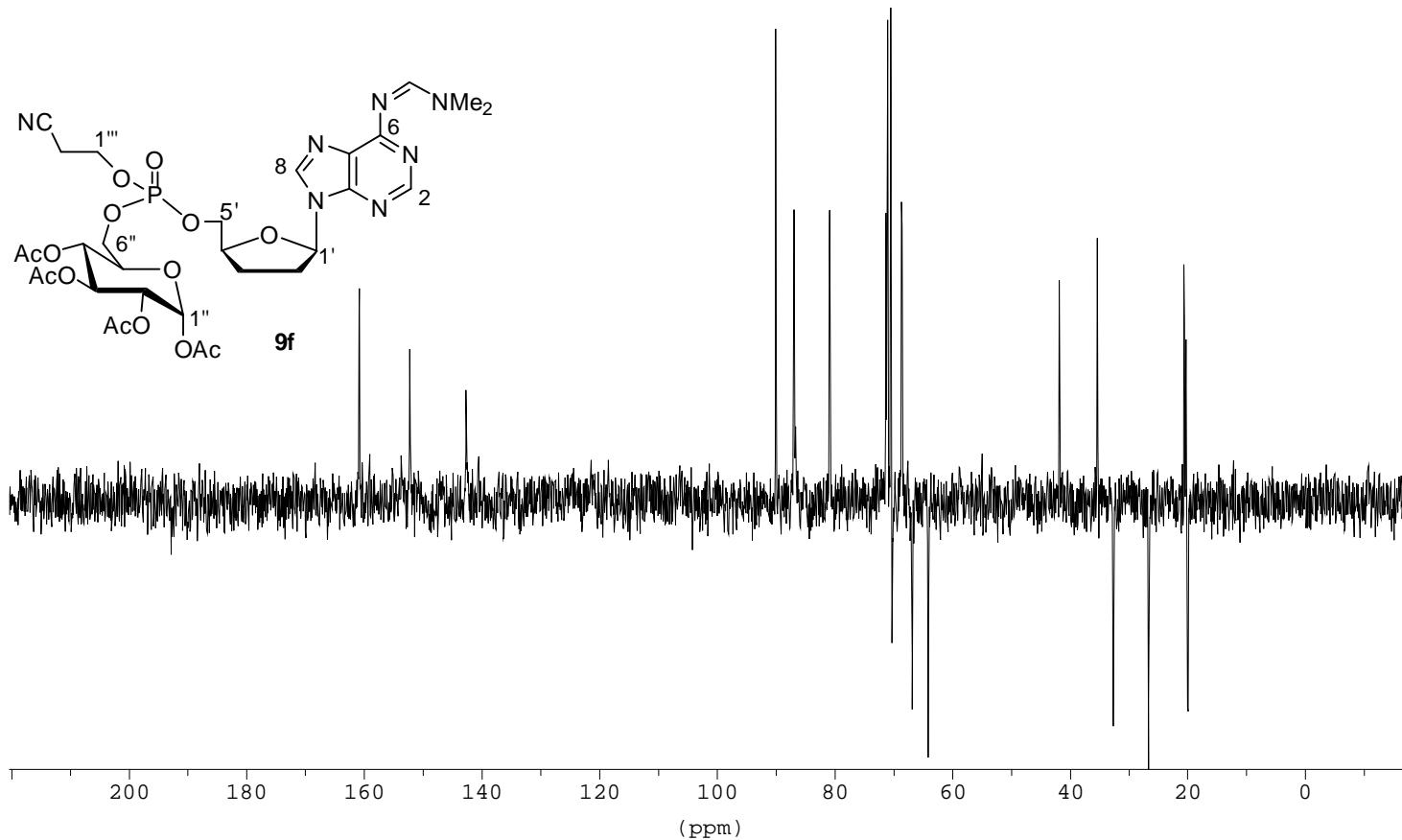
**2-Cyanoethyl-[*N*⁶-(dimethylamino)methylen-2',3'-dideoxyadenosin-5'-yl]-1,2,3,4-tetra-*O*-acetyl-
α-D-glucopyranos-6-yl phosphate (9f)**

¹³C-RMN (MeOH-*d*₄, 150.5 MHz)



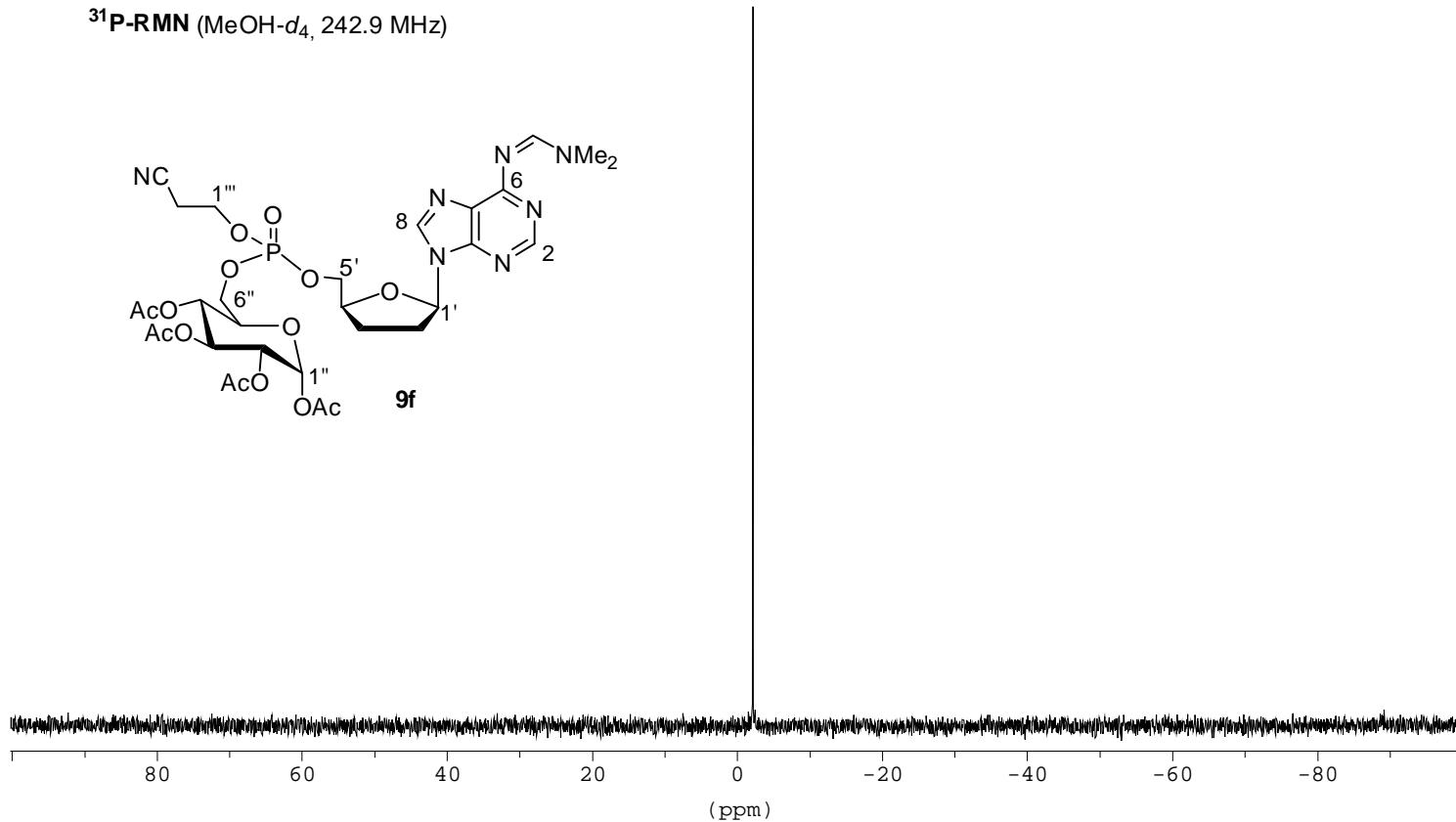
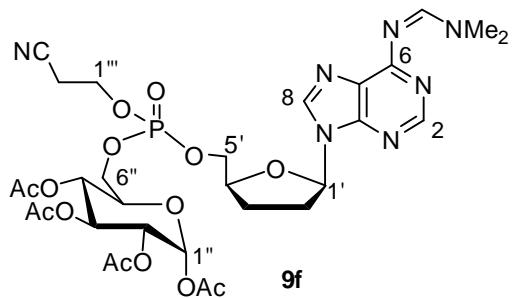
**2-Cyanoethyl-[*N*⁶-(dimethylamino)methylen-2',3'-dideoxyadenosin-5'-yl]-1,2,3,4-tetra-*O*-acetyl-
α-D-glucopyranos-6-yl phosphate (9f)**

DEPT (MeOH-*d*₄, 150.5 MHz)



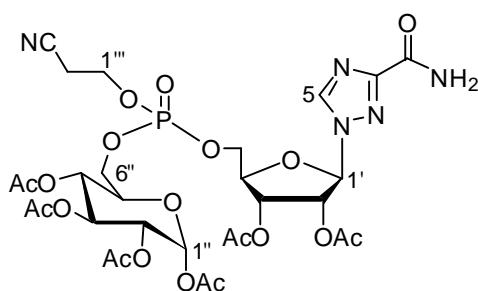
**2-Cyanoethyl-[*N*⁶-(dimethylamino)methylen-2',3'-dideoxyadenosin-5'-yl]-1,2,3,4-tetra-*O*-acetyl-
α-D-glucopyranos-6-yl phosphate (9f)**

³¹P-RMN (MeOH-*d*₄, 242.9 MHz)

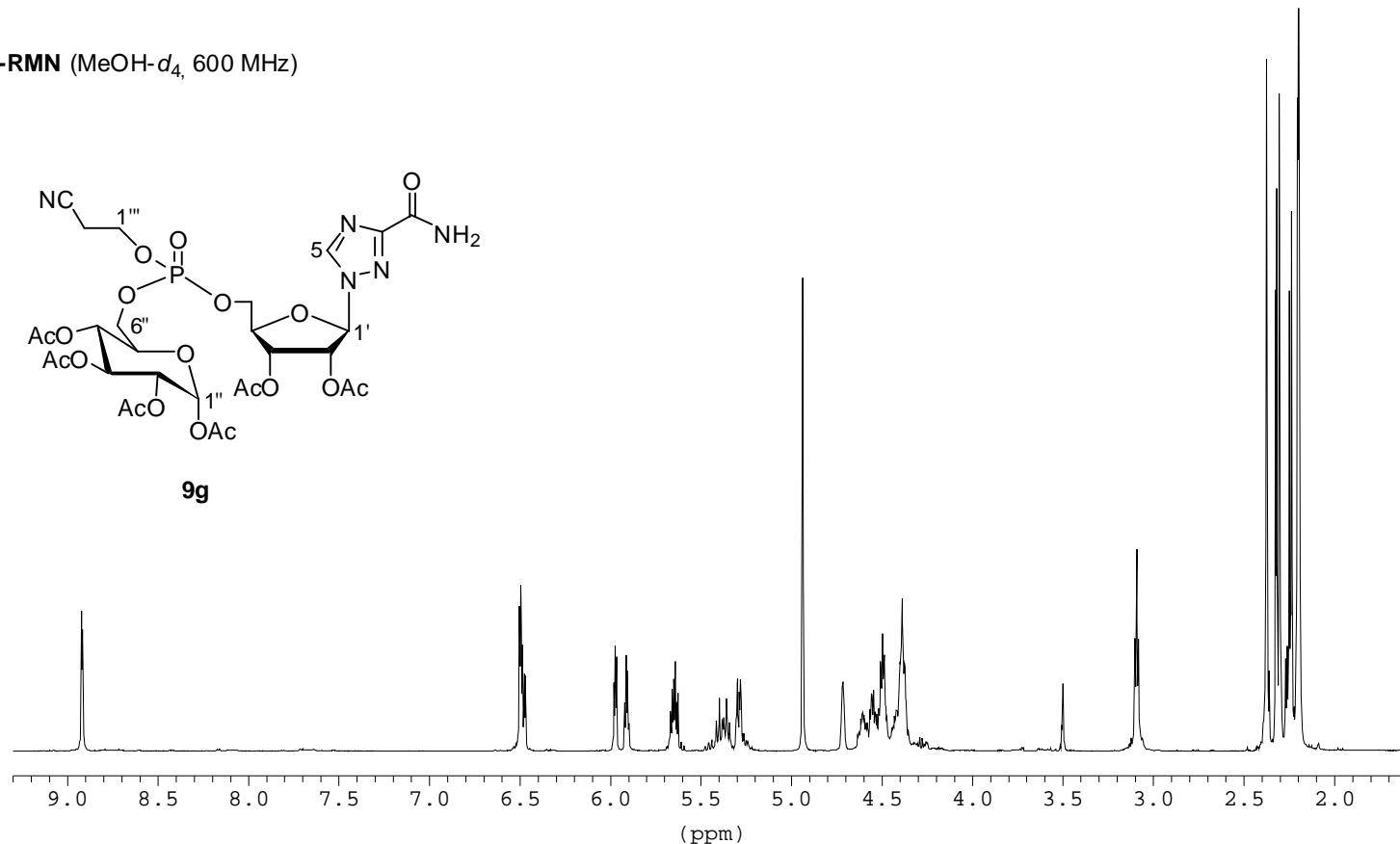


2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(3-carbamoyl-1,2,4-triazoyl)- β -D-ribofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9g)

¹H-RMN (MeOH-*d*₄, 600 MHz)

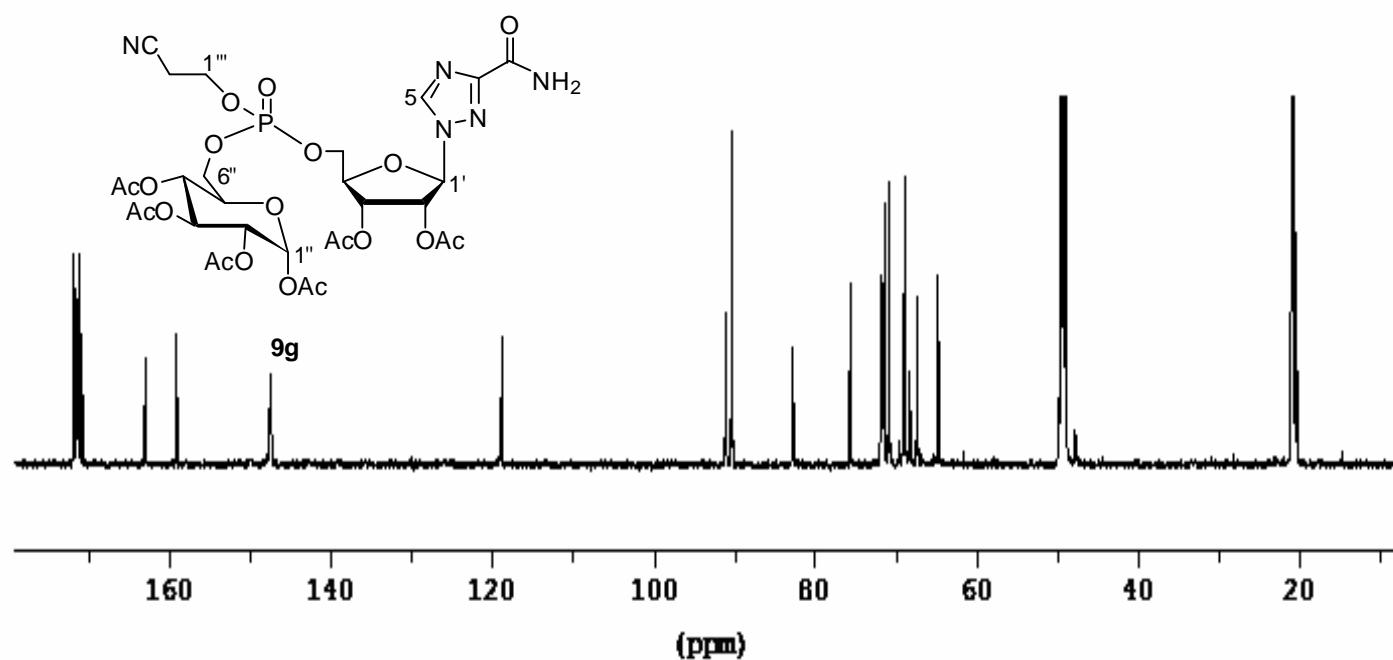


9g



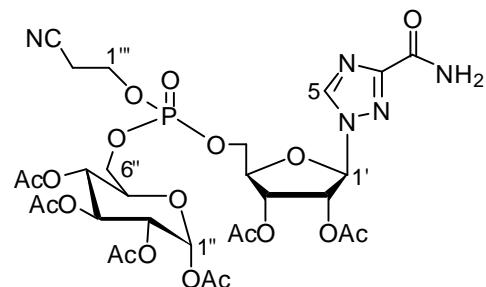
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(3-carbamoyl-1,2,4-triazoyl)- β -D-ribofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9g)

^{13}C -RMN (MeOH- d_4 , 150.5 MHz)

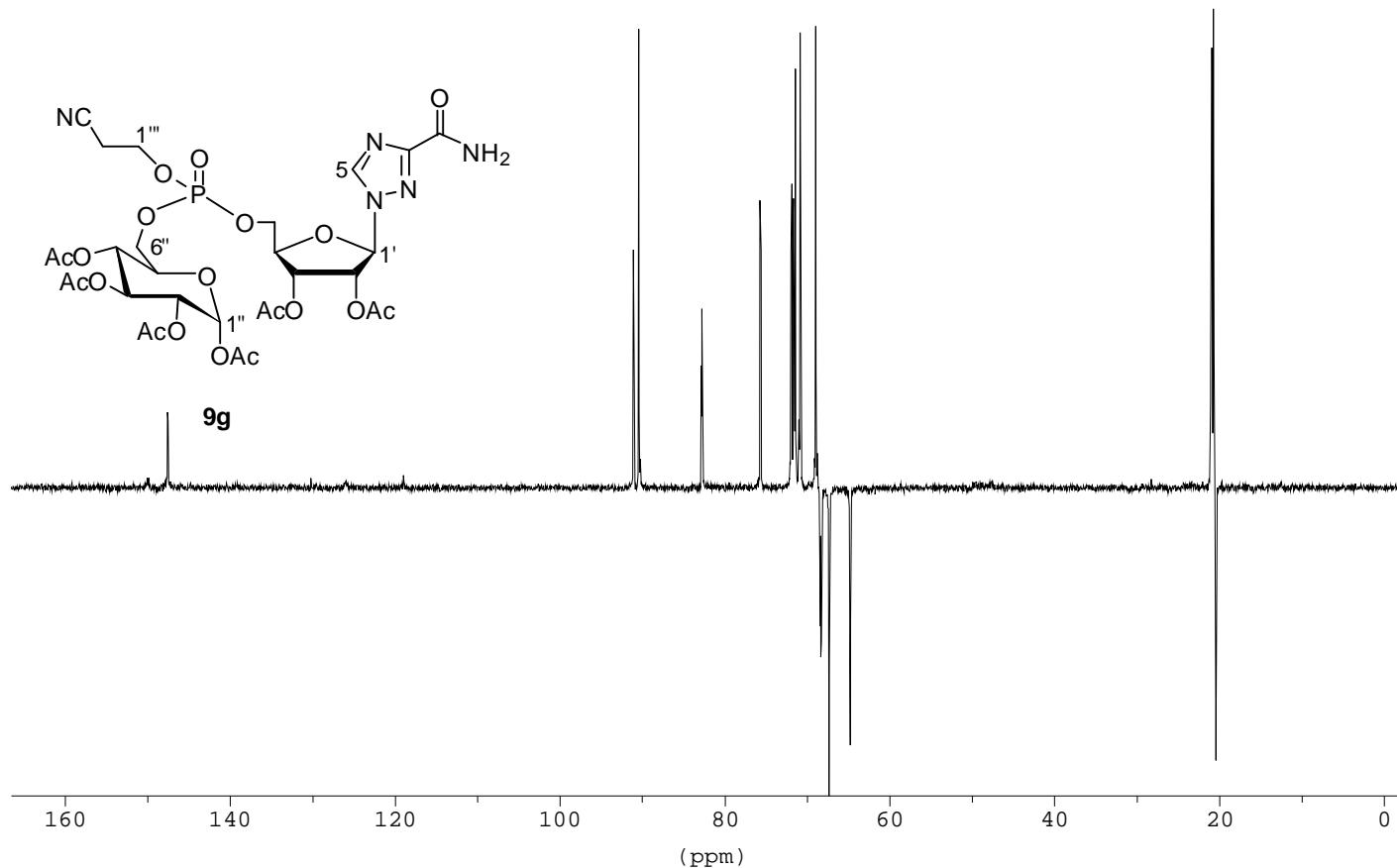


2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(3-carbamoyl-1,2,4-triazoyl)- β -D-ribofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9g)

DEPT (MeOH-*d*₄, 150.5 MHz)

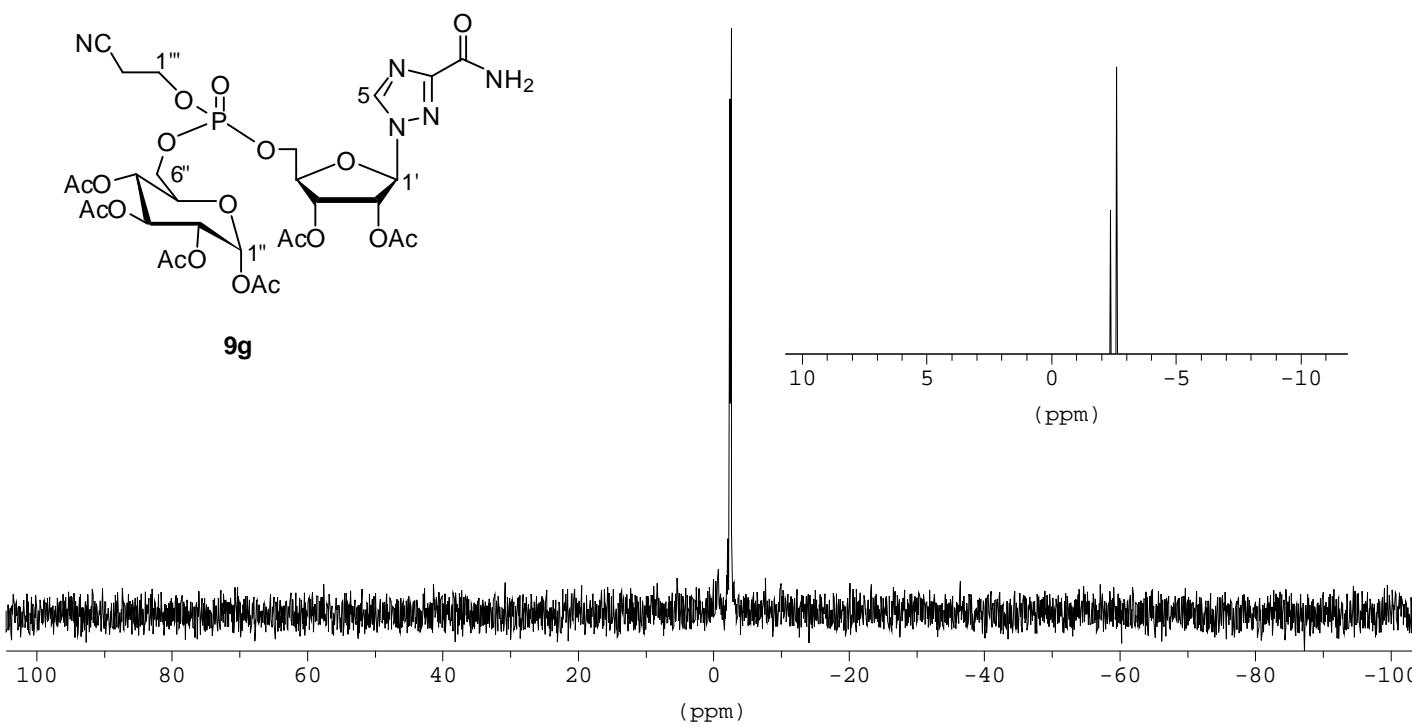


9g



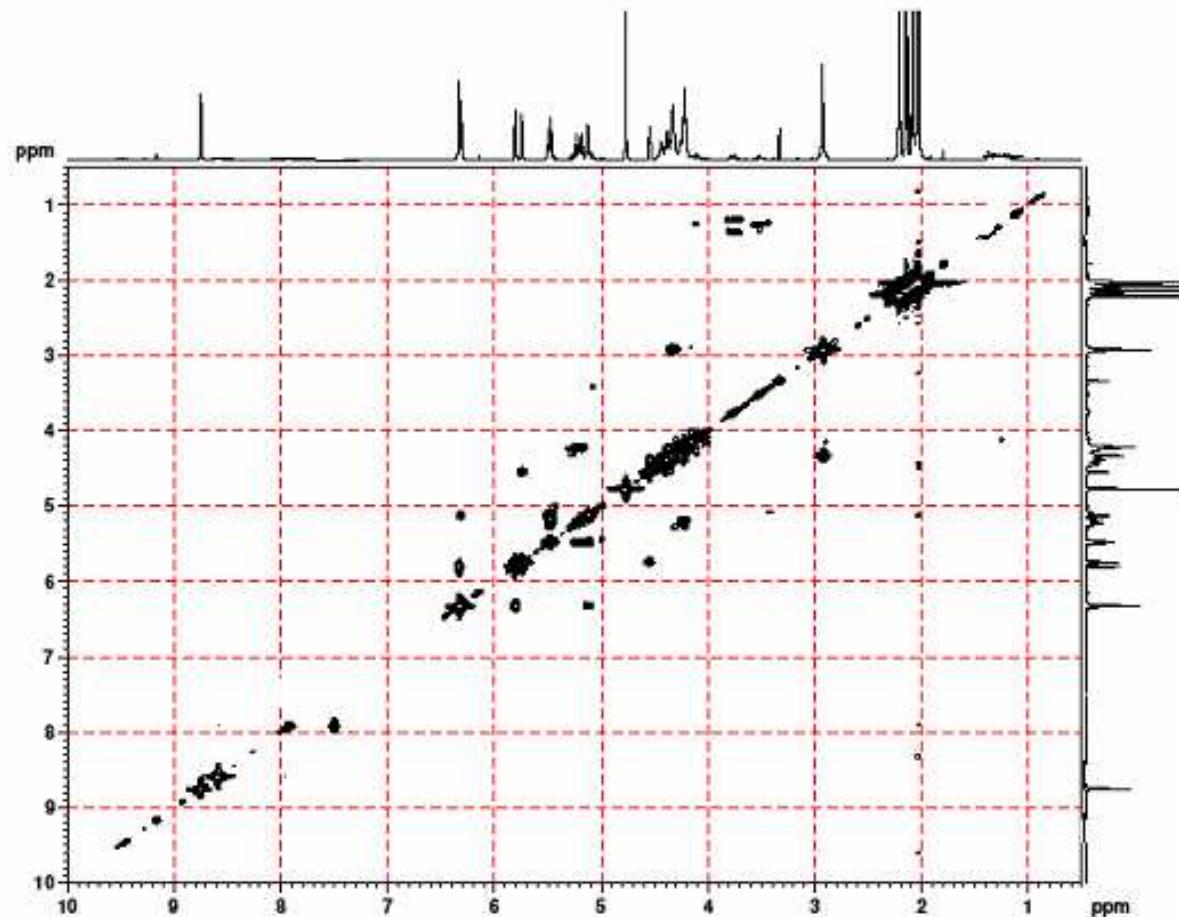
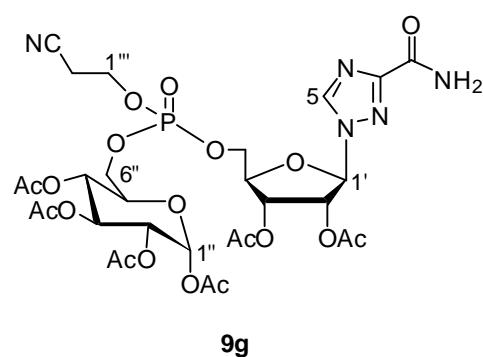
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(3-carbamoyl-1,2,4-triazoyl)- β -D-ribofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9g)

^{31}P -RMN (MeOH- d_4 , 242.9 MHz)



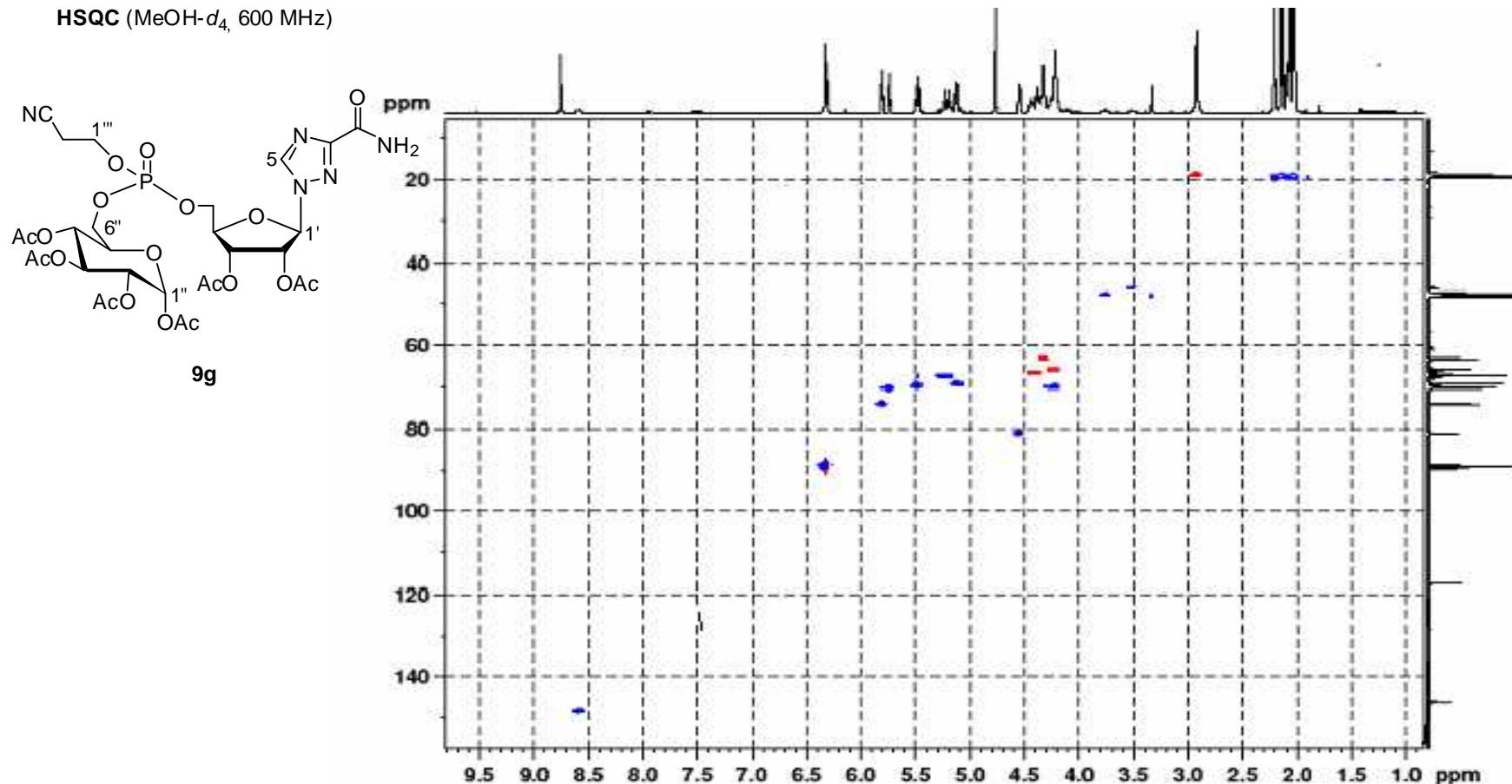
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(3-carbamoyl-1,2,4-triazoyl)- β -D-ribofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9g)

COSY (MeOH-*d*₄, 600 MHz)

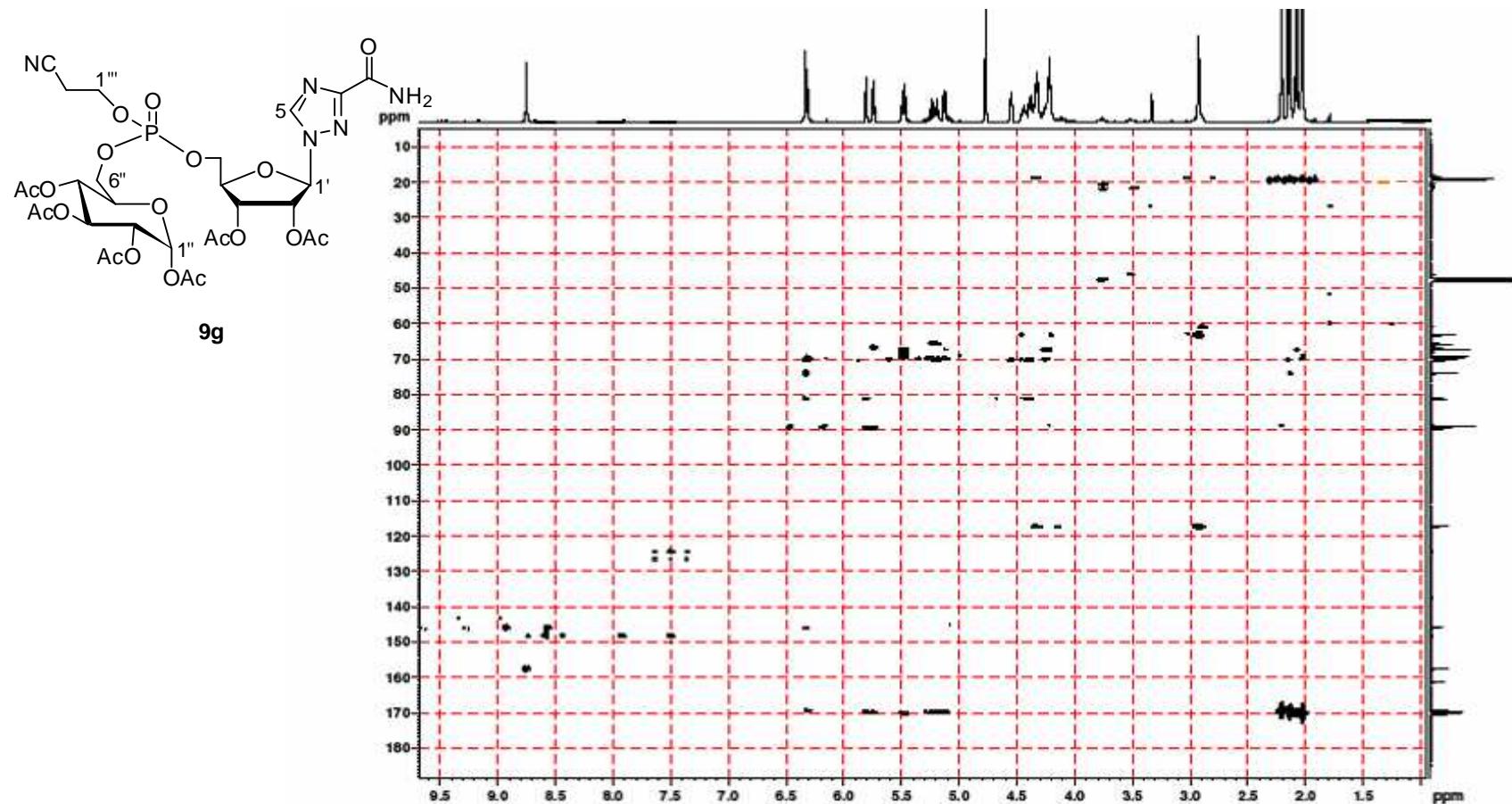


2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(3-carbamoyl-1,2,4-triazoyl)- β -D-ribofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9g)

HSQC ($\text{MeOH}-d_4$, 600 MHz)

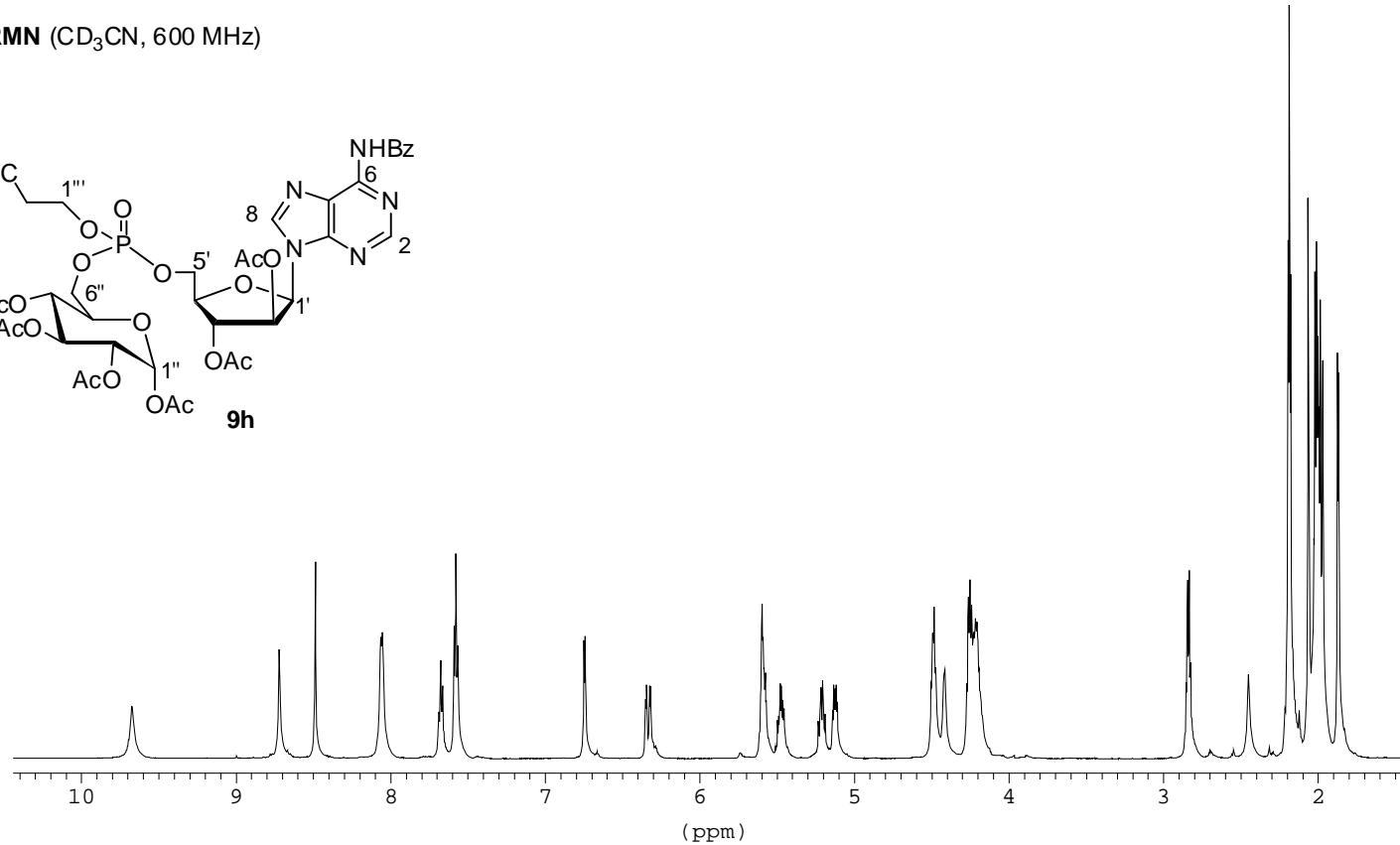
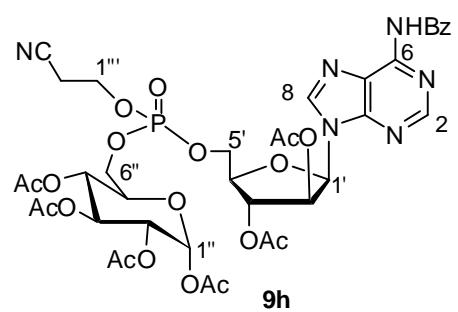


2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(3-carbamoyl-1,2,4-triazoyl)- β -D-ribofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9g)



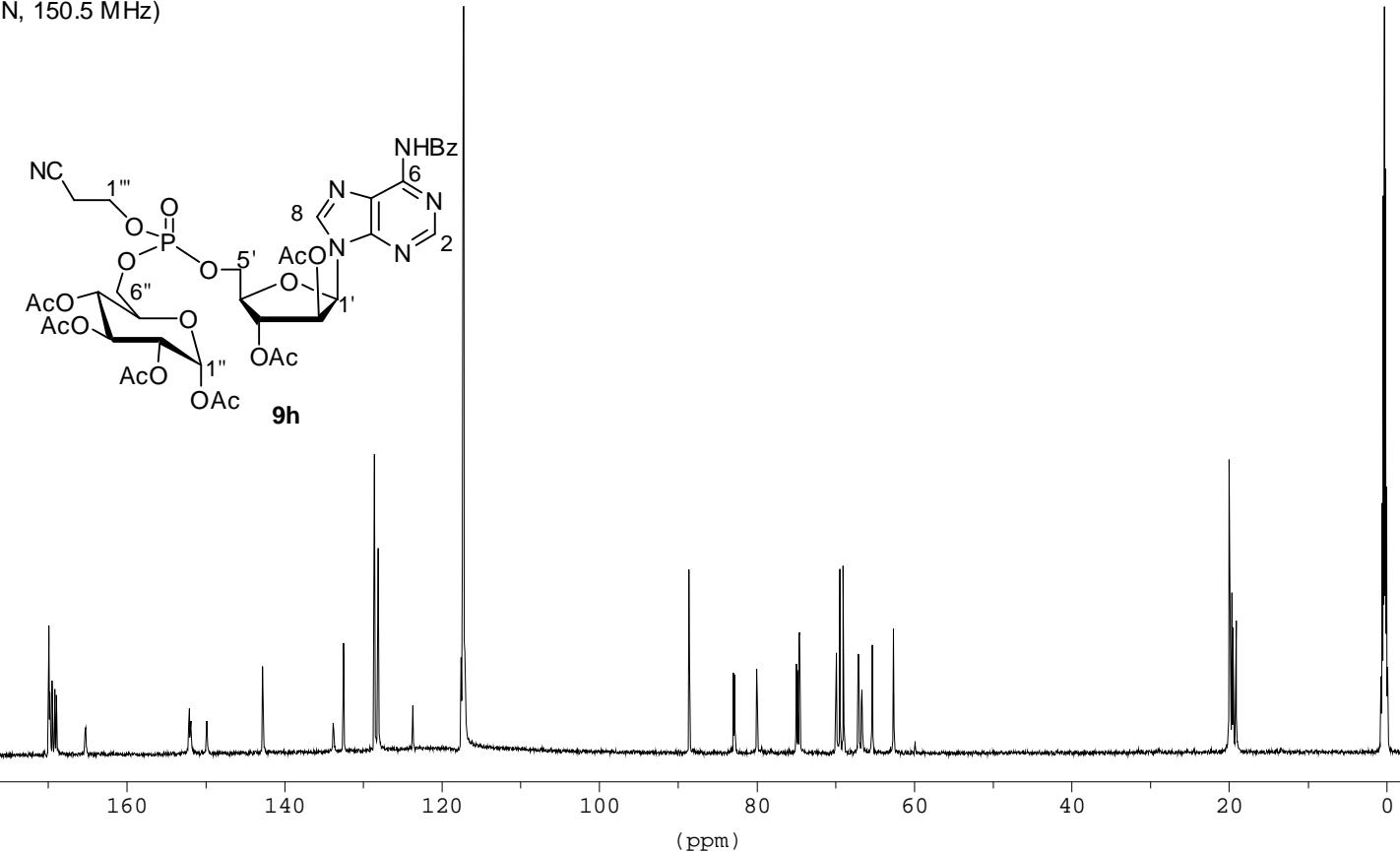
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁶-benzoyladeninyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9h)

¹H-RMN (CD₃CN, 600 MHz)

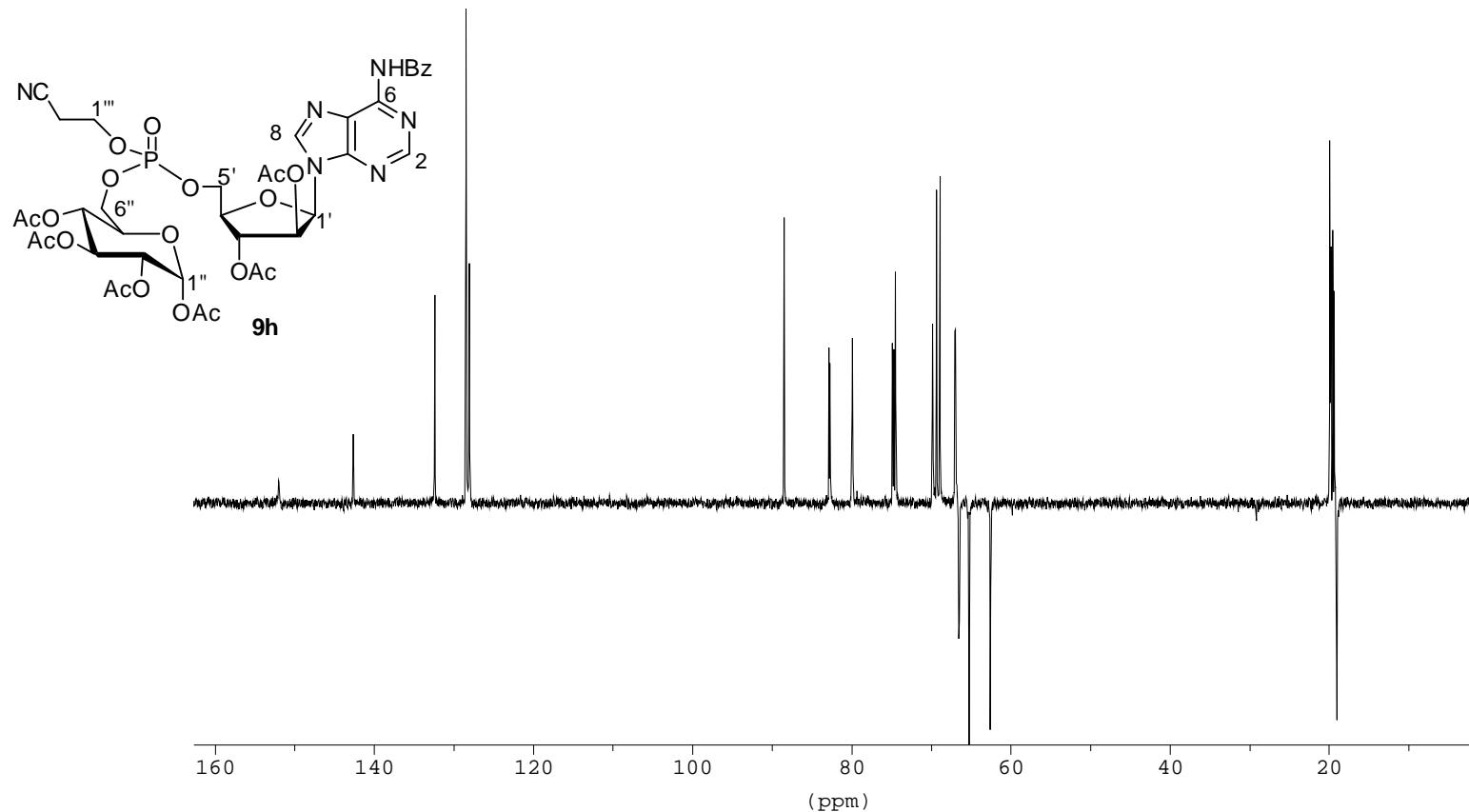


2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁶-benzoyladeninyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9h)

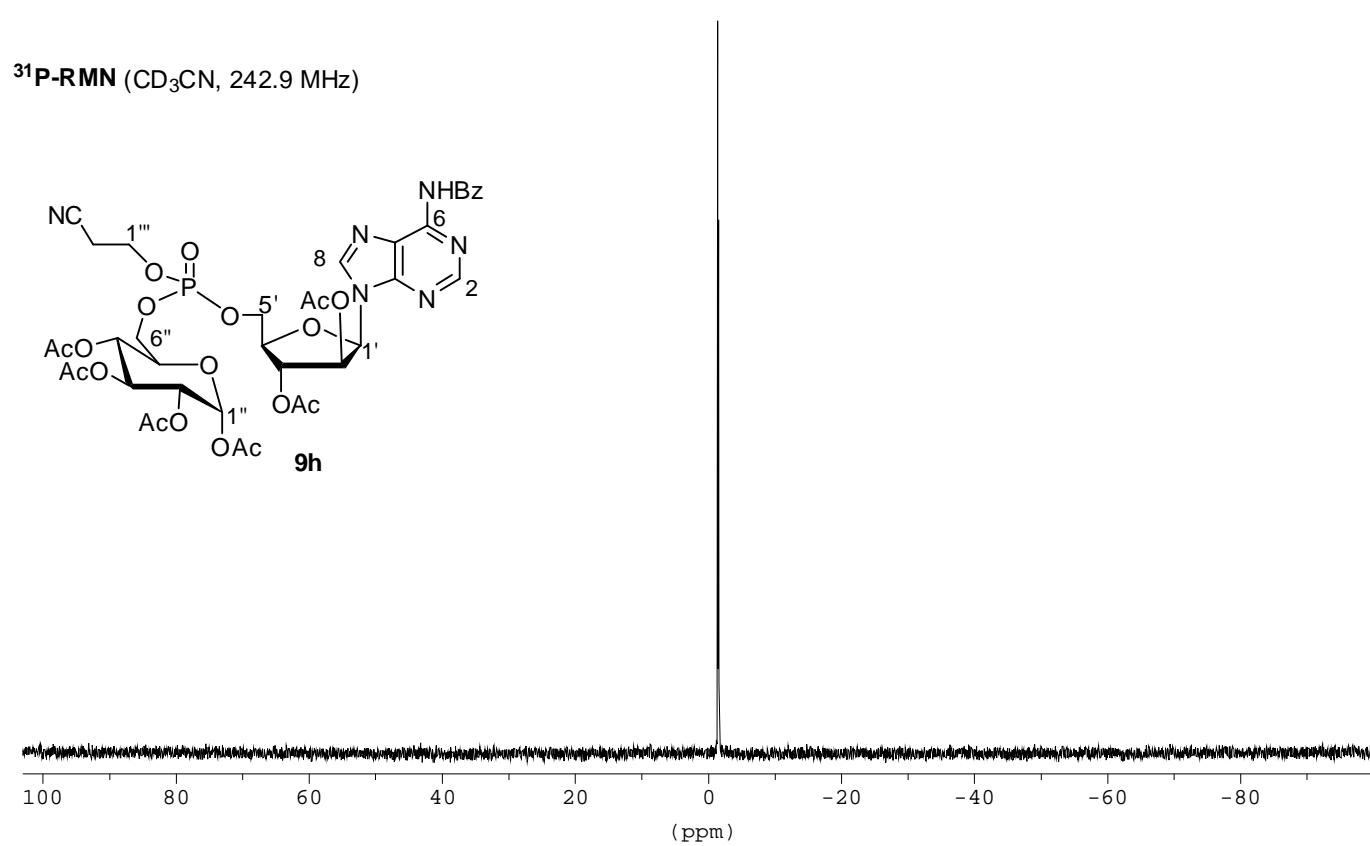
¹³C-RMN (CD₃CN, 150.5 MHz)



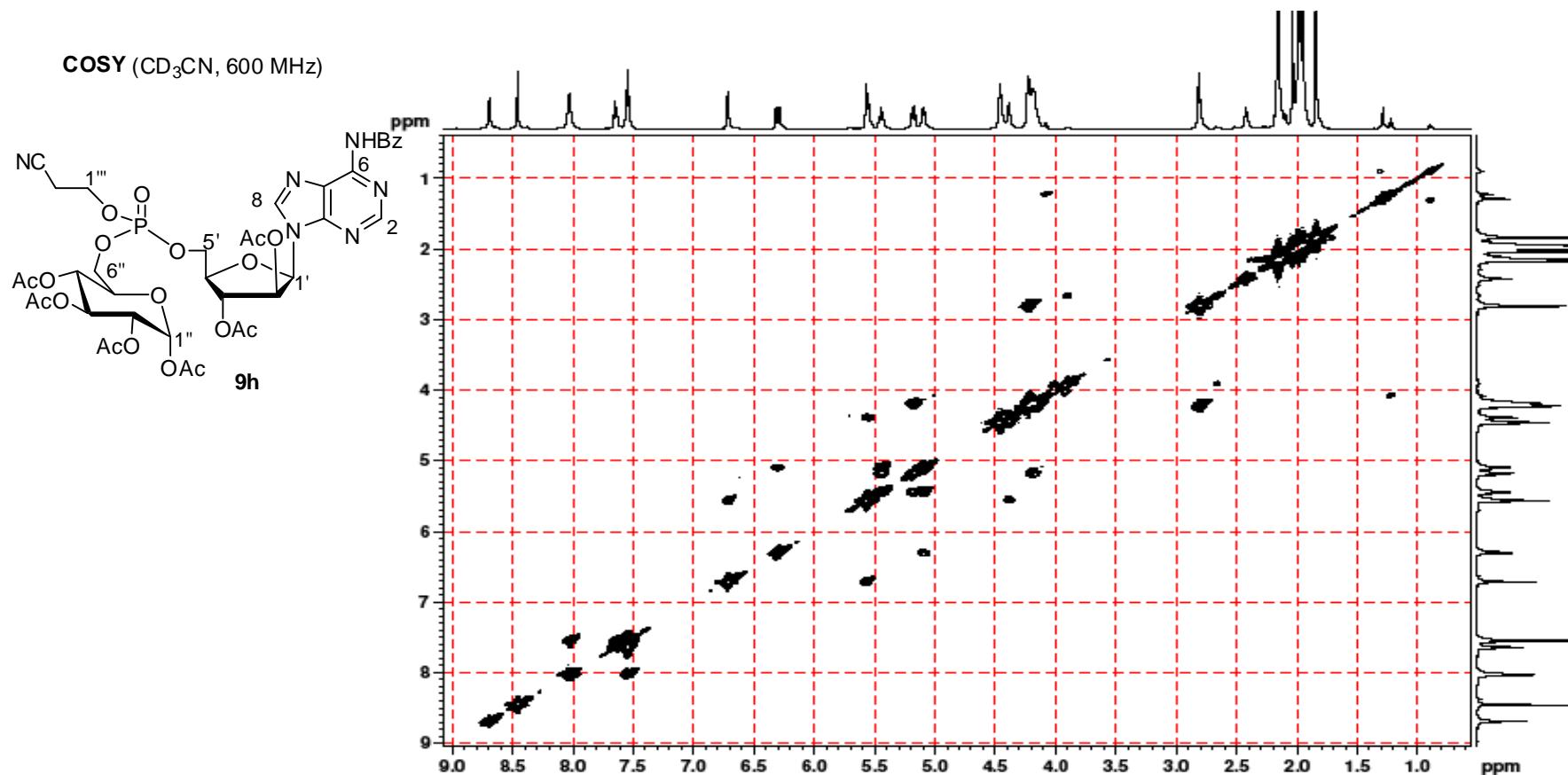
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁶-benzoyladeninyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9h)



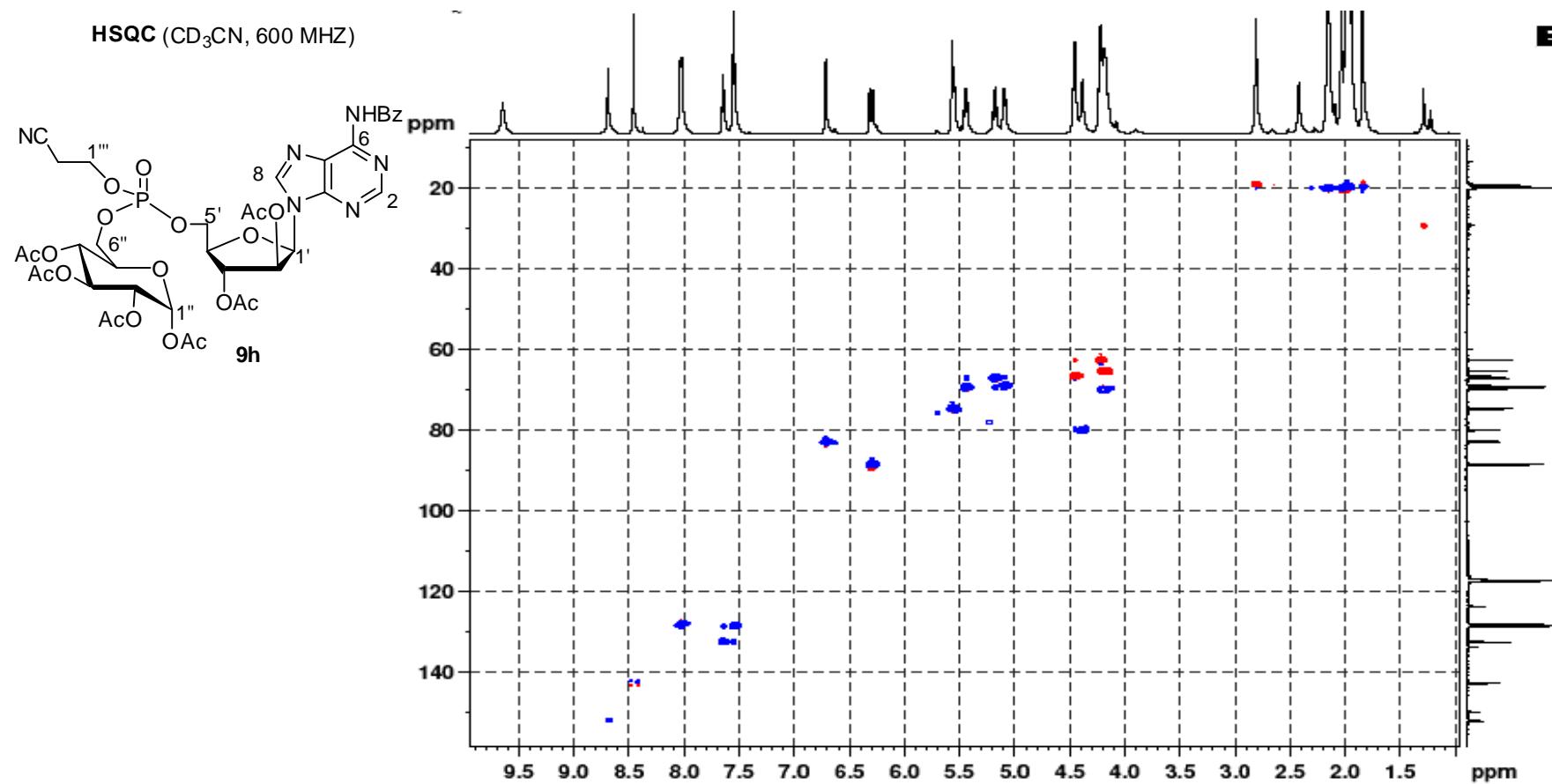
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁶-benzoyladeninyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9h)



2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁶-benzoyladeninyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9h)

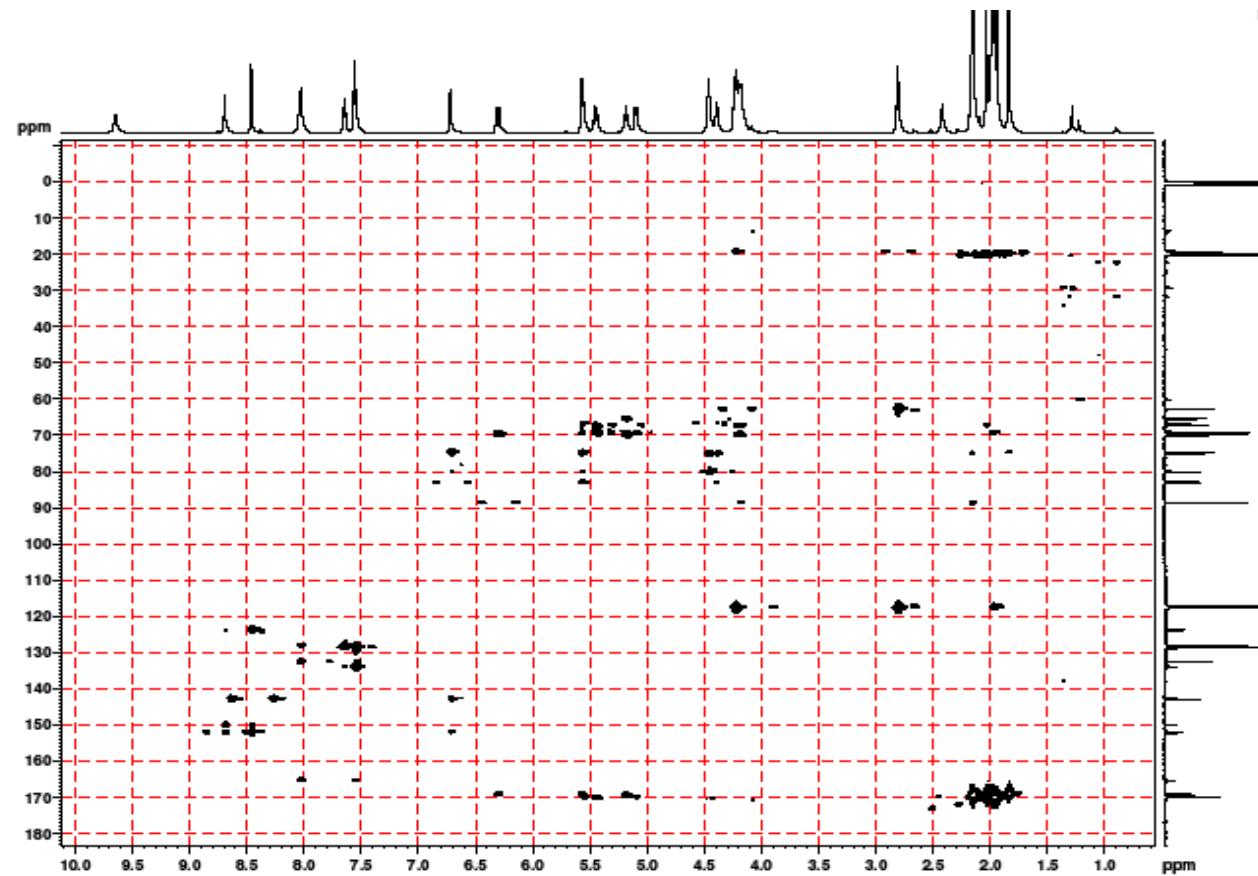
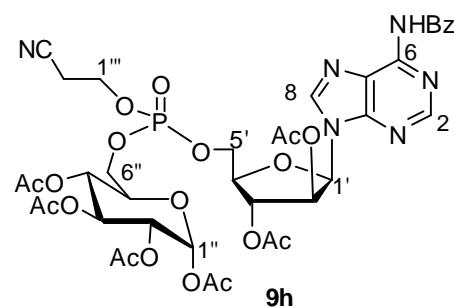


2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁶-benzoyladeninyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9h)



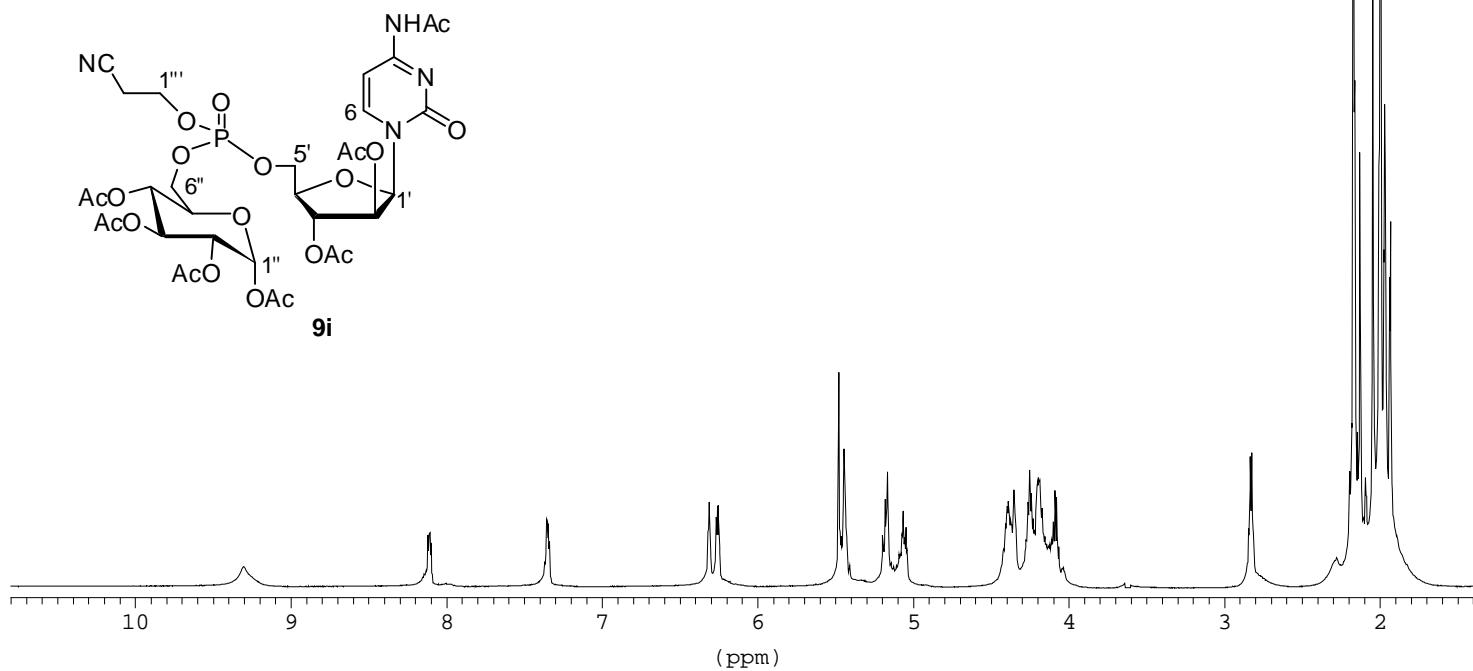
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁶-benzoyladeninyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9h)

HMBC (CD_3CN , 600 MHz)



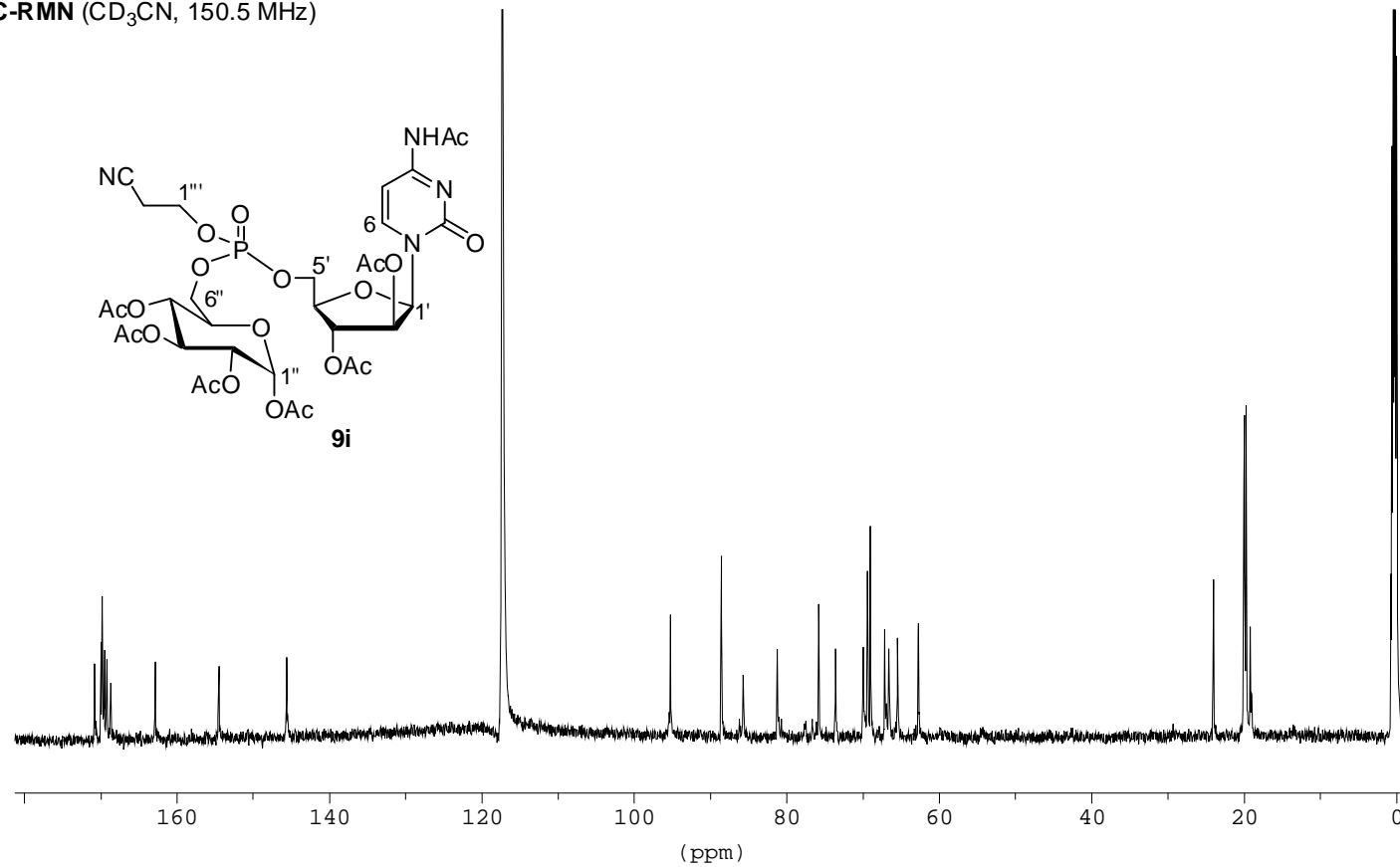
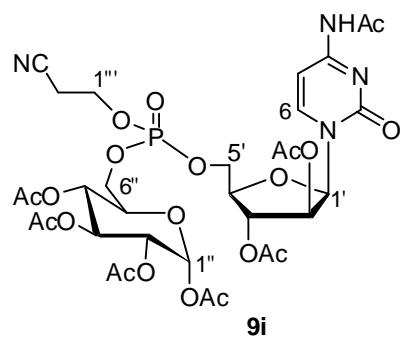
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁴-benzoylcytosinyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9i)

¹H-RMN (CD₃CN, 600 MHz)



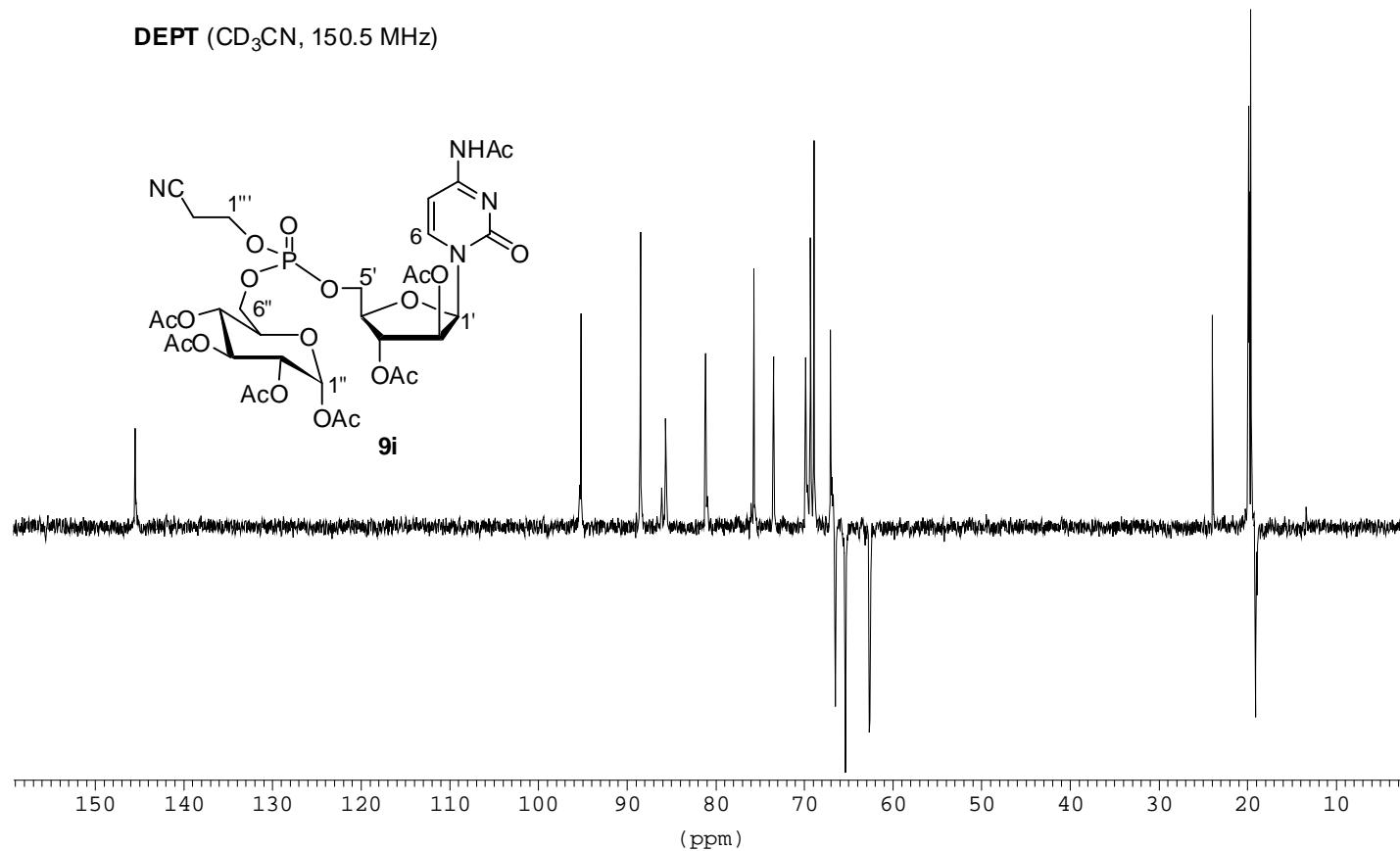
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁴-benzoylcytosinyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9i)

¹³C-RMN (CD₃CN, 150.5 MHz)



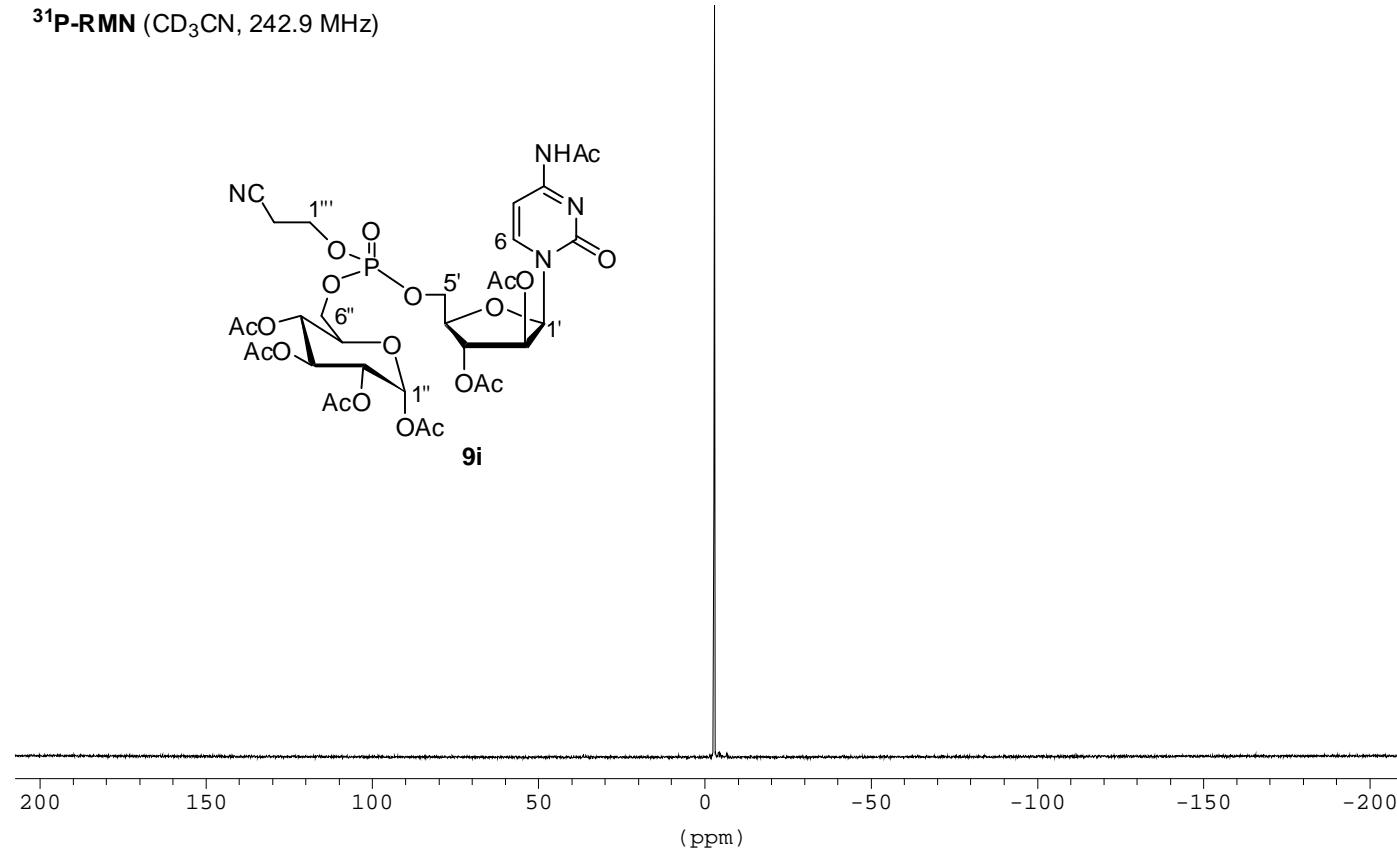
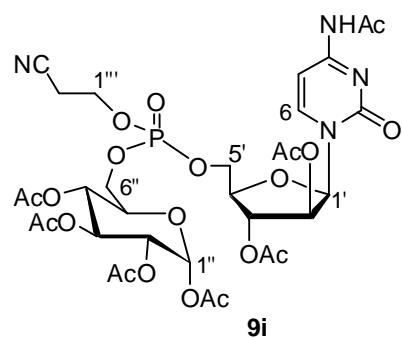
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁴-benzoylcytosinyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9i)

DEPT (CD_3CN , 150.5 MHz)



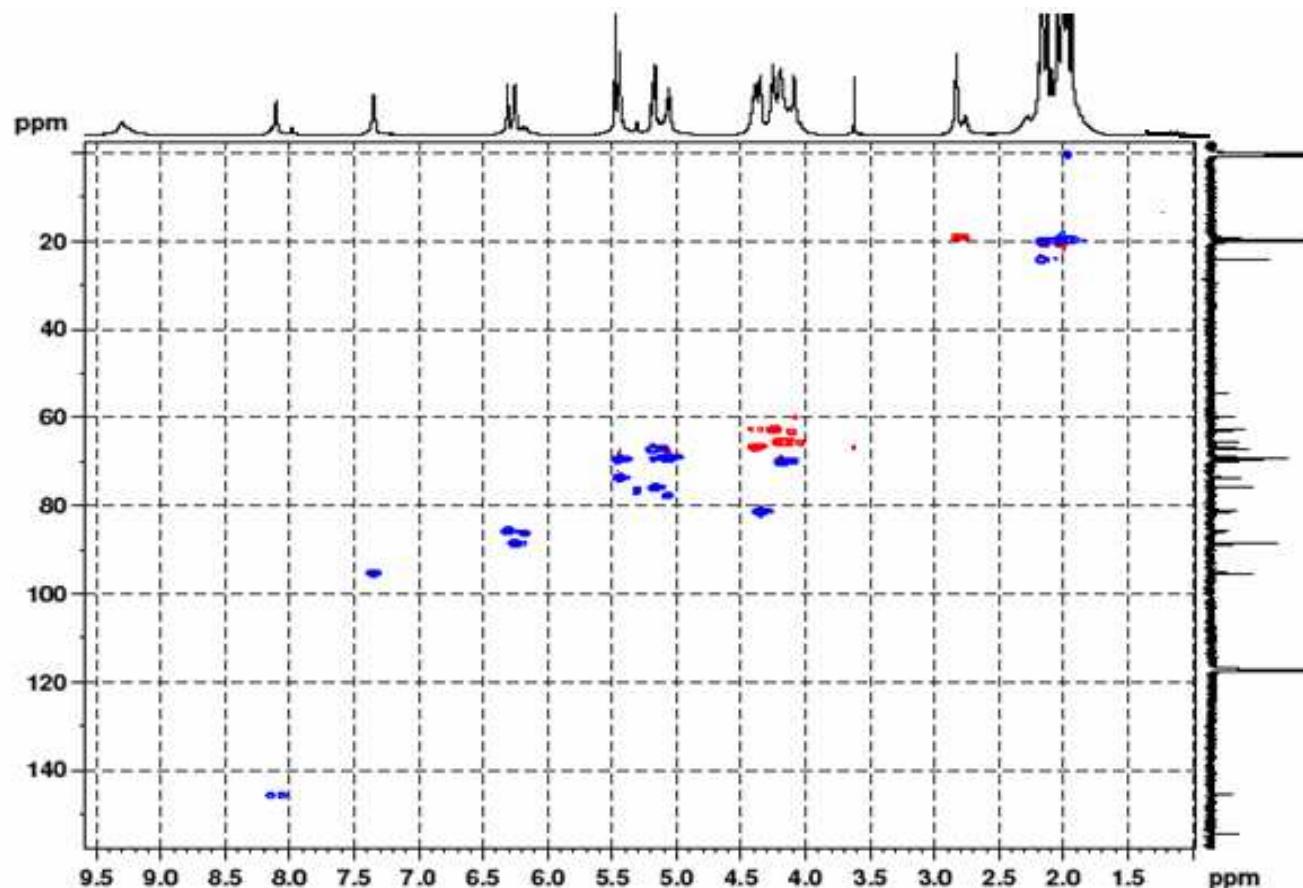
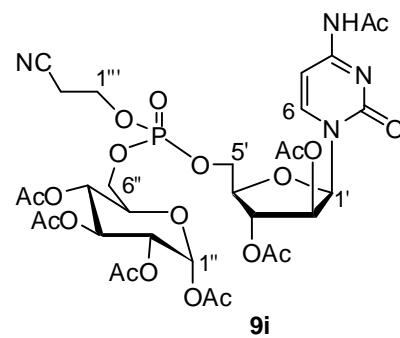
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁴-benzoylcytosinyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9i)

³¹P-RMN (CD₃CN, 242.9 MHz)



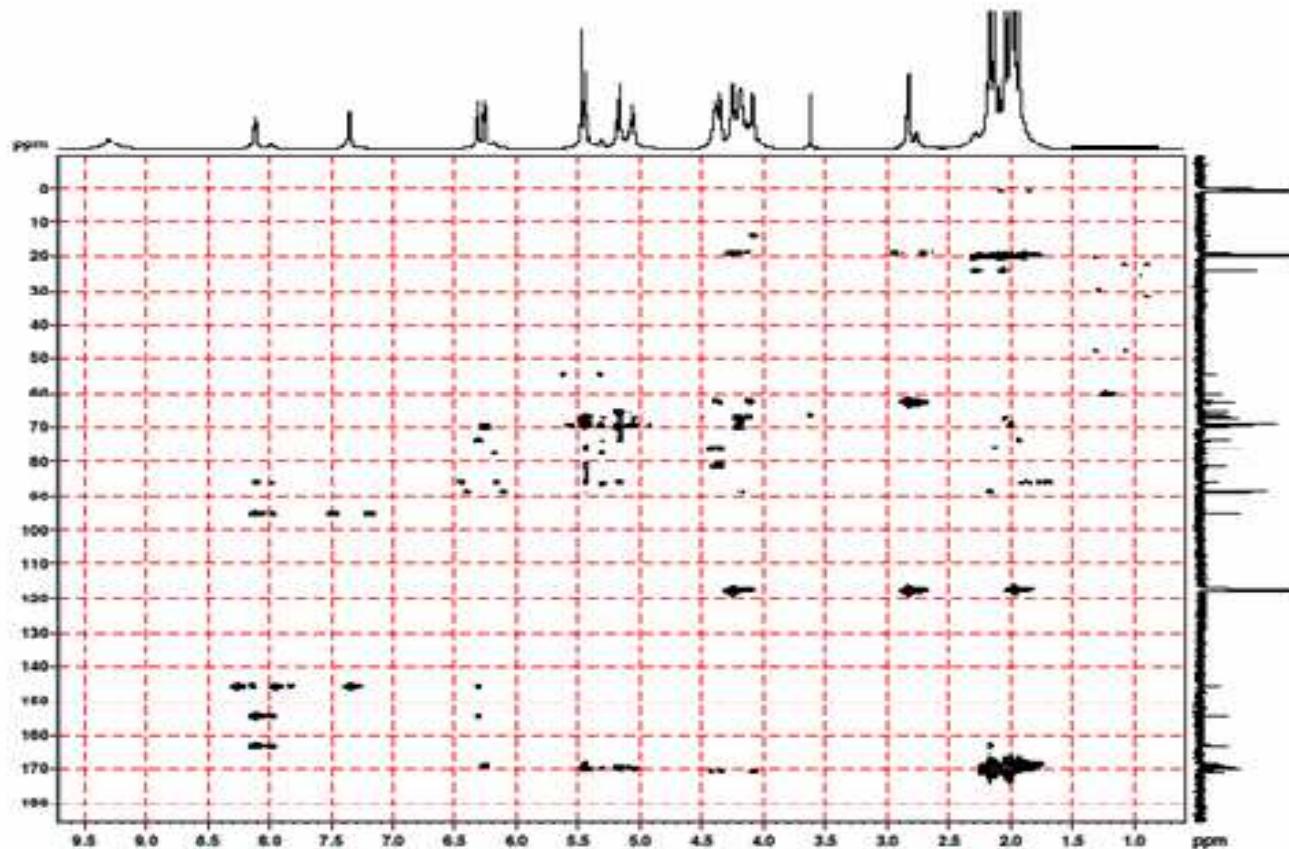
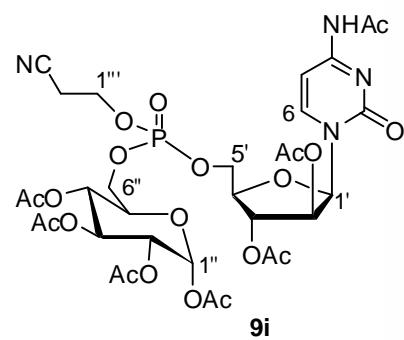
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁴-benzoylcytosinyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9i)

HSQC (CD₃CN, 600 MHz)



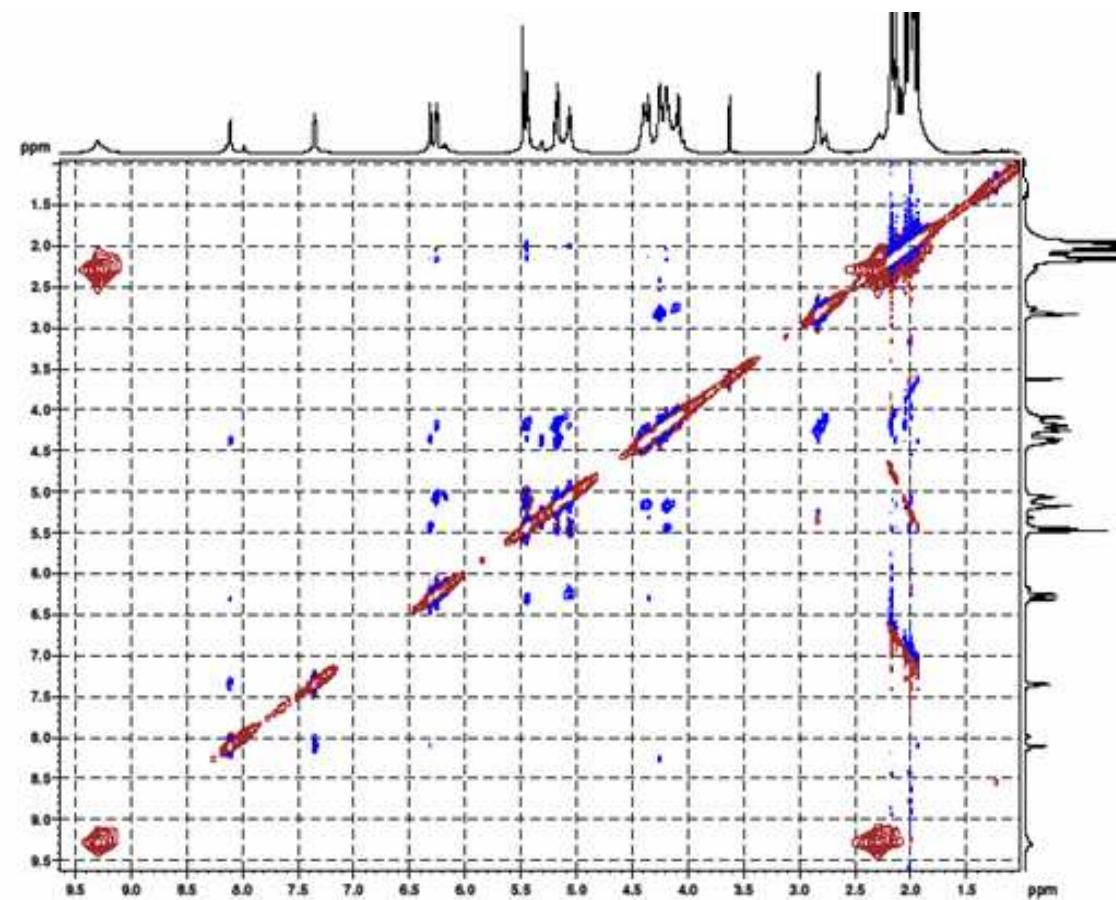
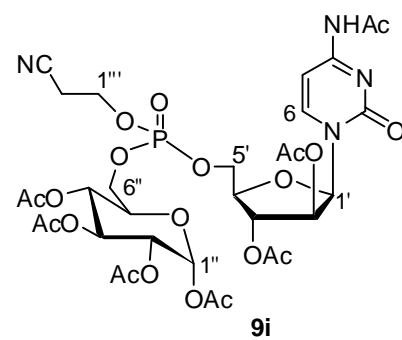
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁴-benzoylcytosinyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9i)

HMBC (CD₃CN, 600 MHz)



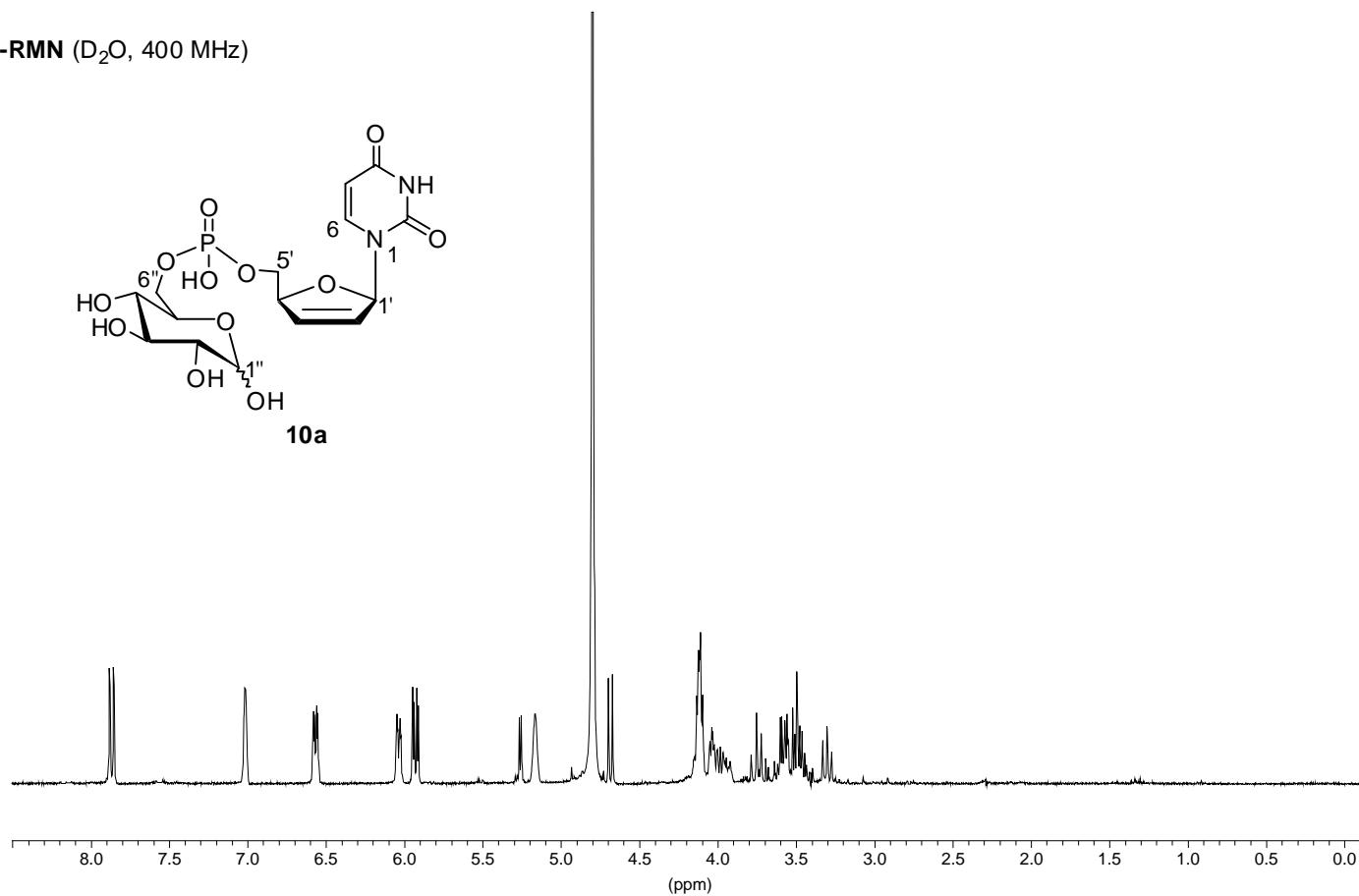
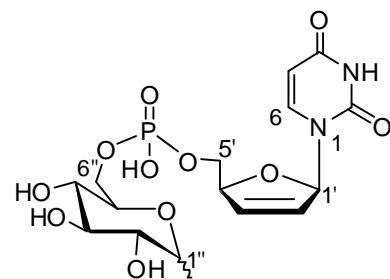
2-Cyanoethyl-[2',3'-di-*O*-acetyl-1'-(*N*⁴-benzoylcytosinyl)- β -D-arabinofuranos-5'-yl]-1,2,3,4-tetra-*O*-acetyl- α -D-glucopyranos-6-yl phosphate (9i)

NOESY (CD₃CN, 600 MHz)



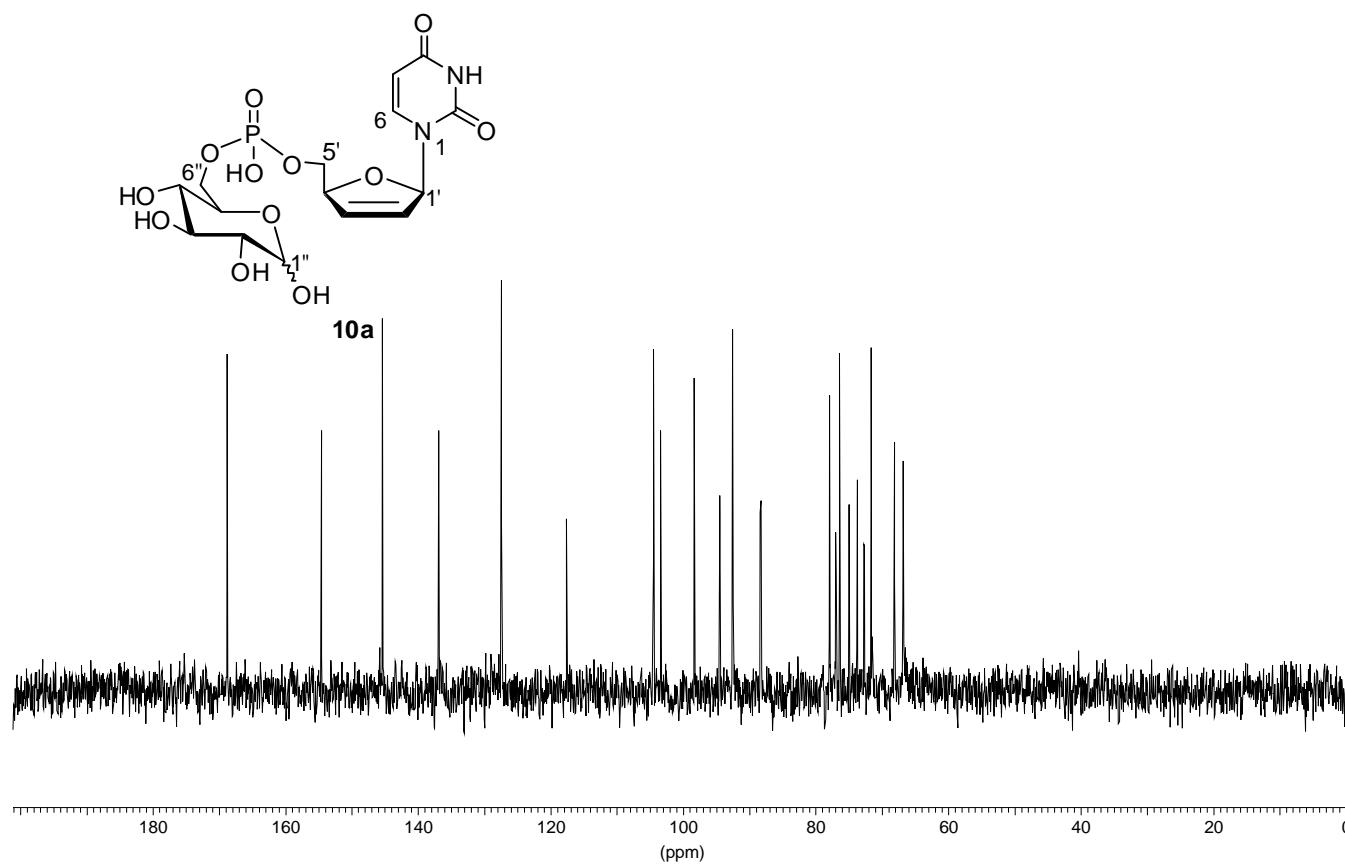
2',3'-Didehydro-2',3'-dideoxyuridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10a)

¹H-RMN (D_2O , 400 MHz)



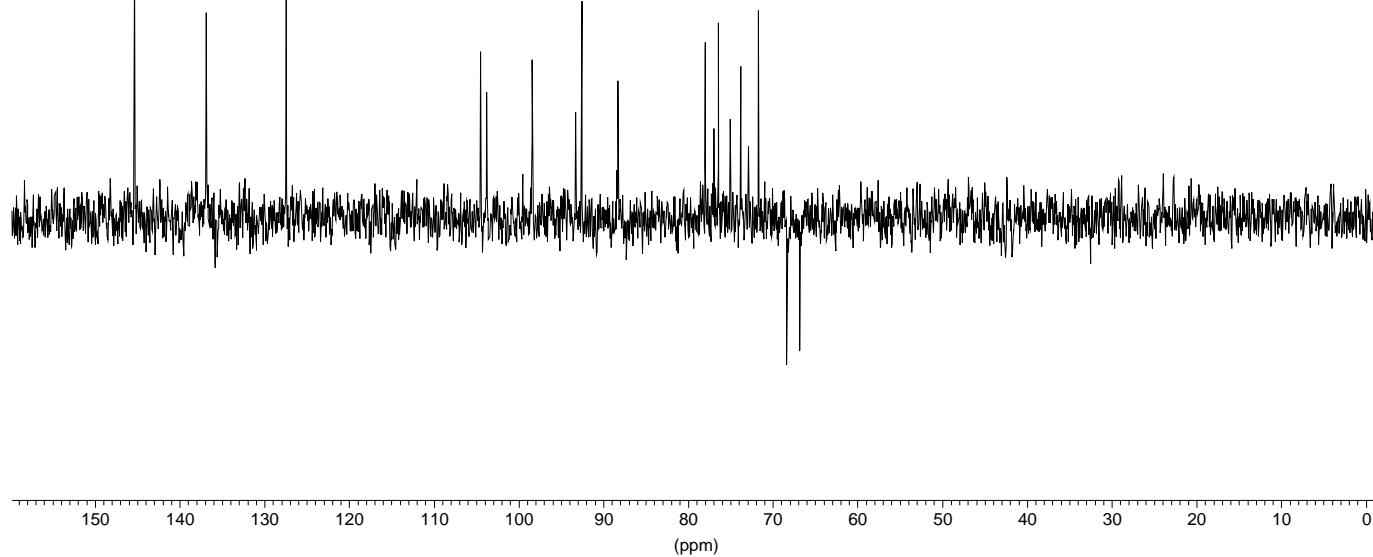
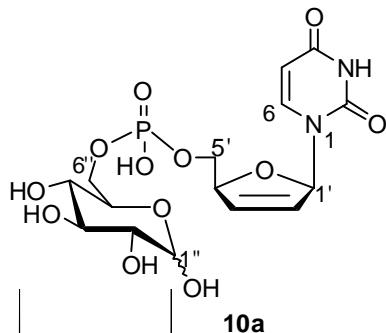
2',3'-Didehydro-2',3'-dideoxyuridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10a)

^{13}C -RMN (D_2O , 100.6 MHz)



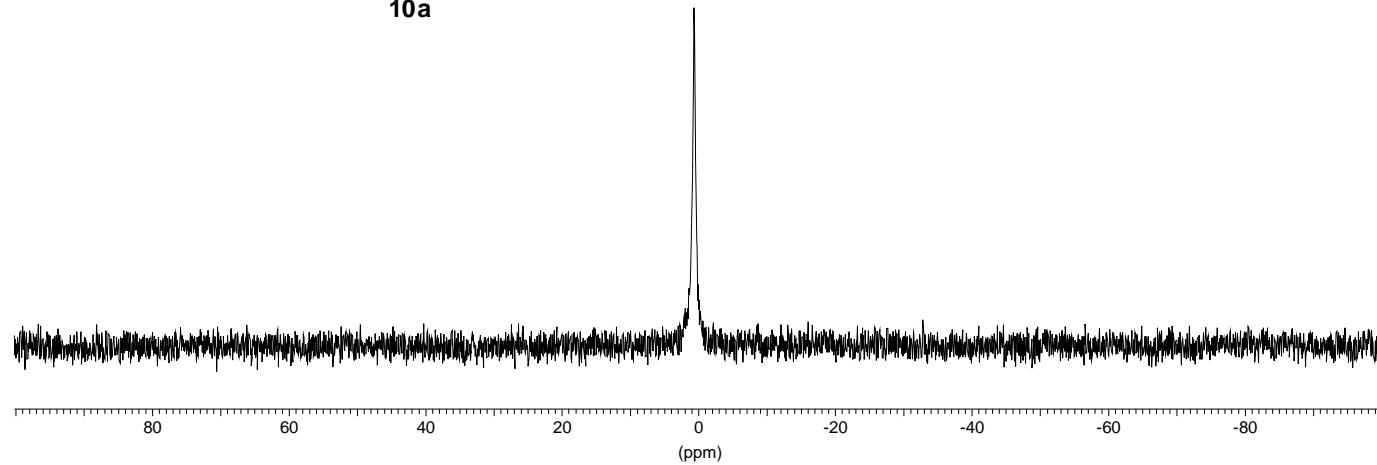
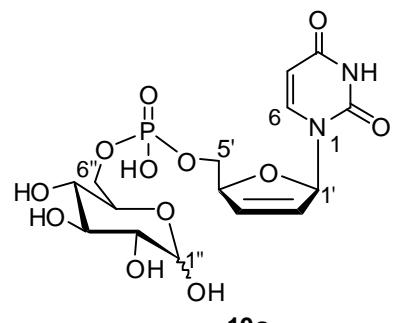
2',3'-Didehydro-2',3'-dideoxyuridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10a)

DEPT (D_2O , 100.6 MHz)

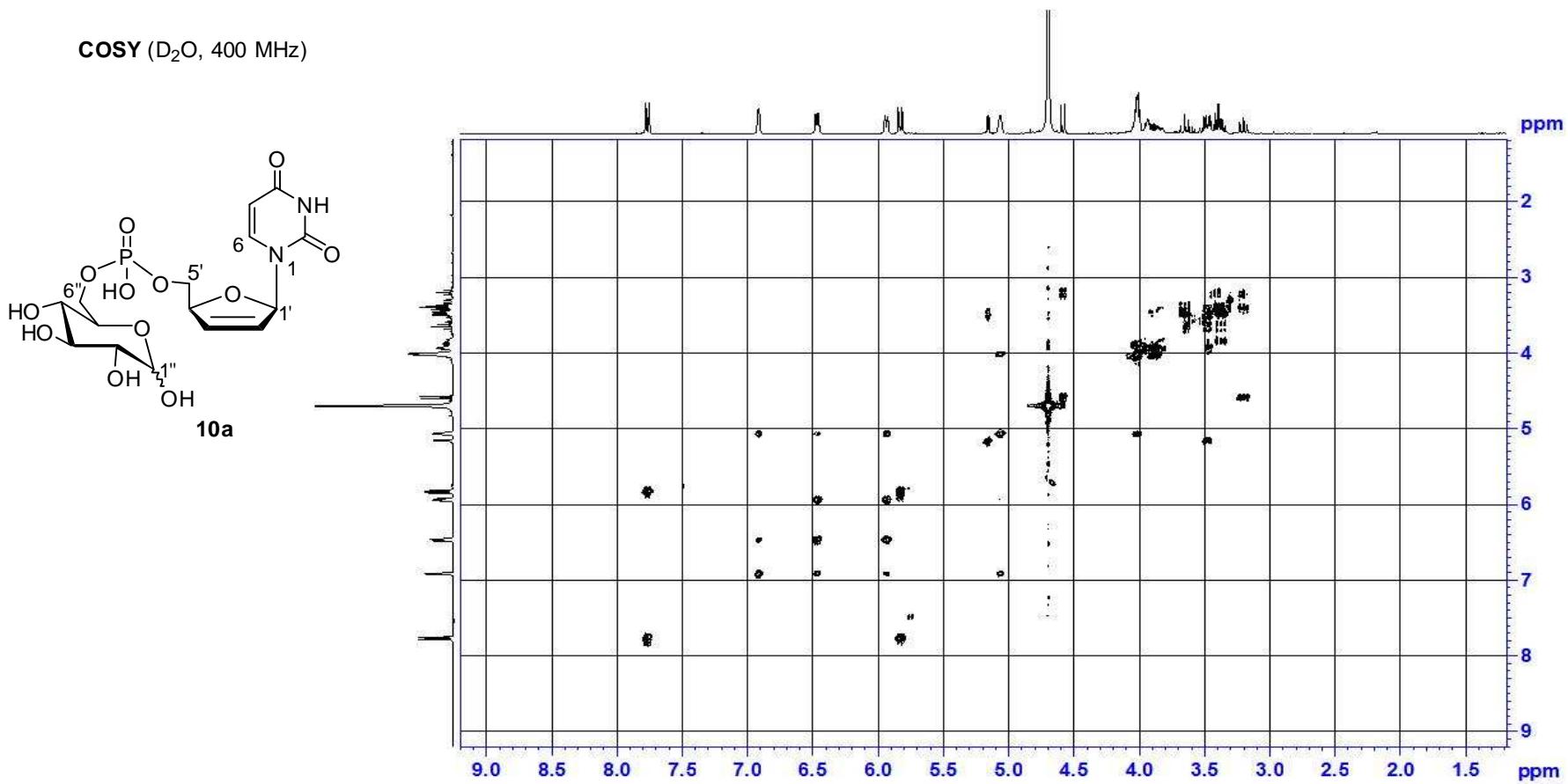


2',3'-Didehydro-2',3'-dideoxyuridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10a)

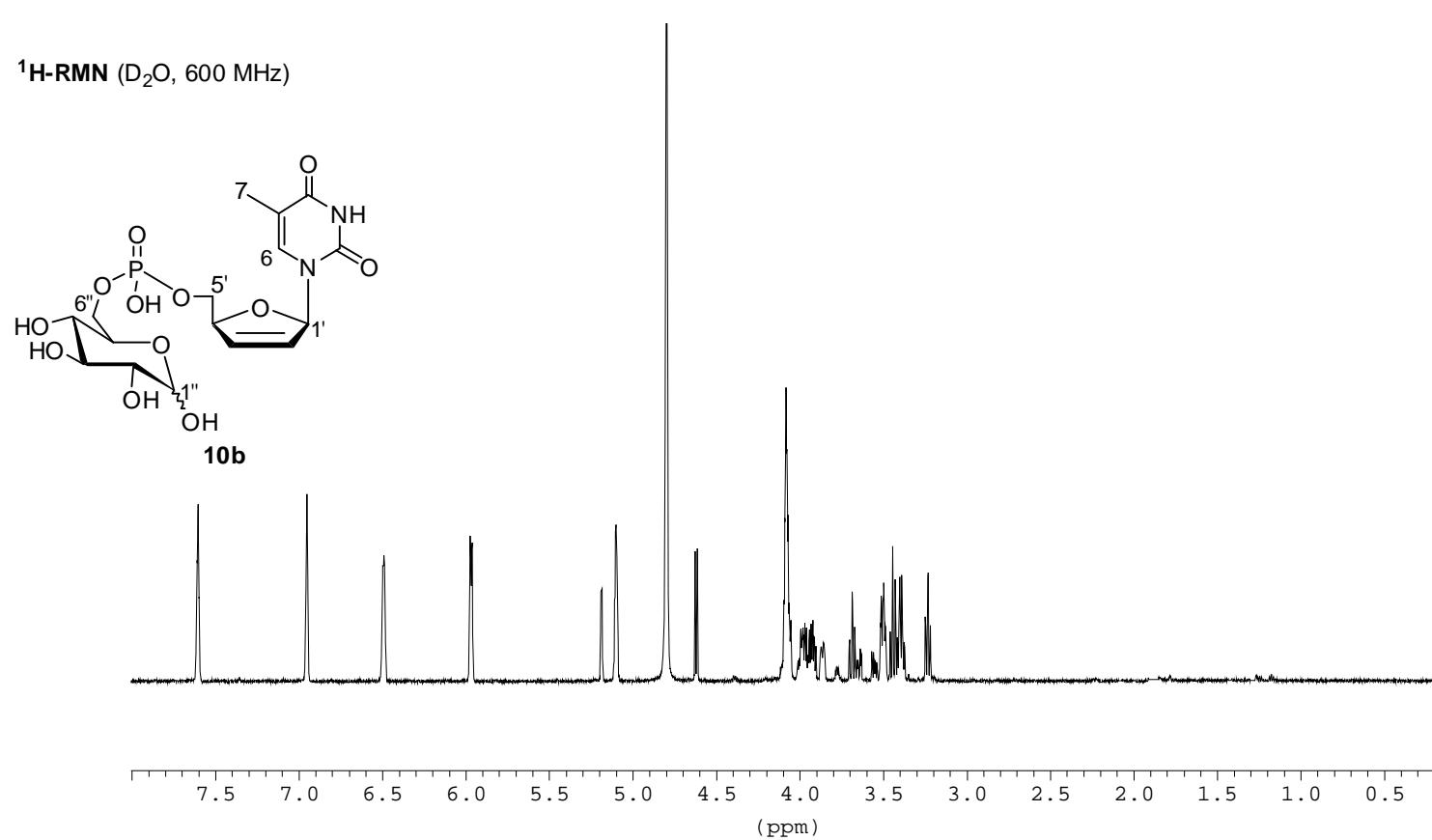
^{31}P -RMN (D_2O , 161.9 MHz)



2',3'-Didehydro-2',3'-dideoxyuridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10a)

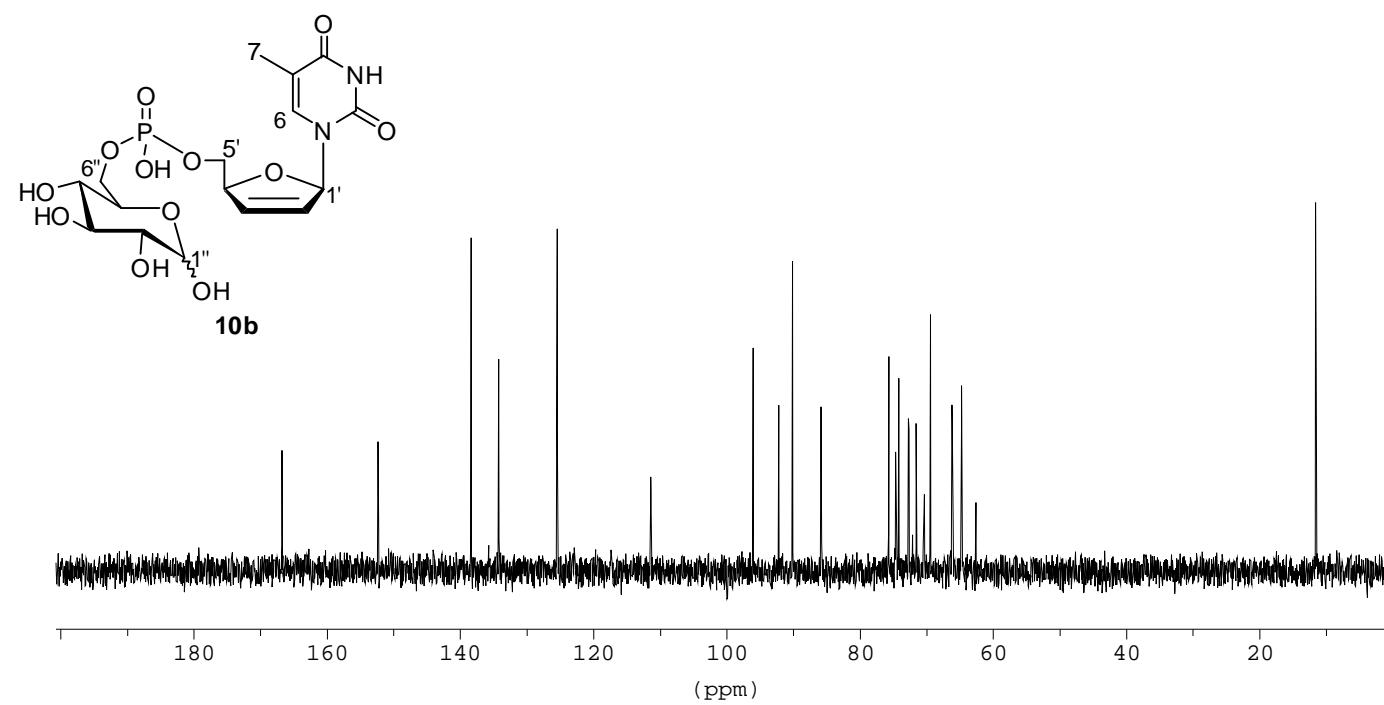


2',3'-Didehydro-2',3'-dideoxythymidine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10b)



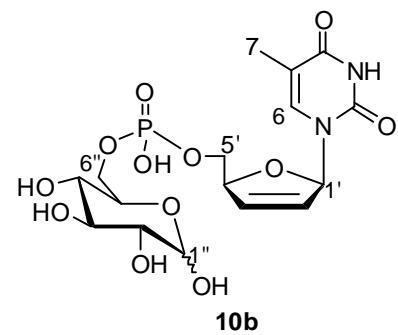
2',3'-Didehydro-2',3'-dideoxythymidine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10b)

¹³C-RMN (D_2O , 150.5 MHz)

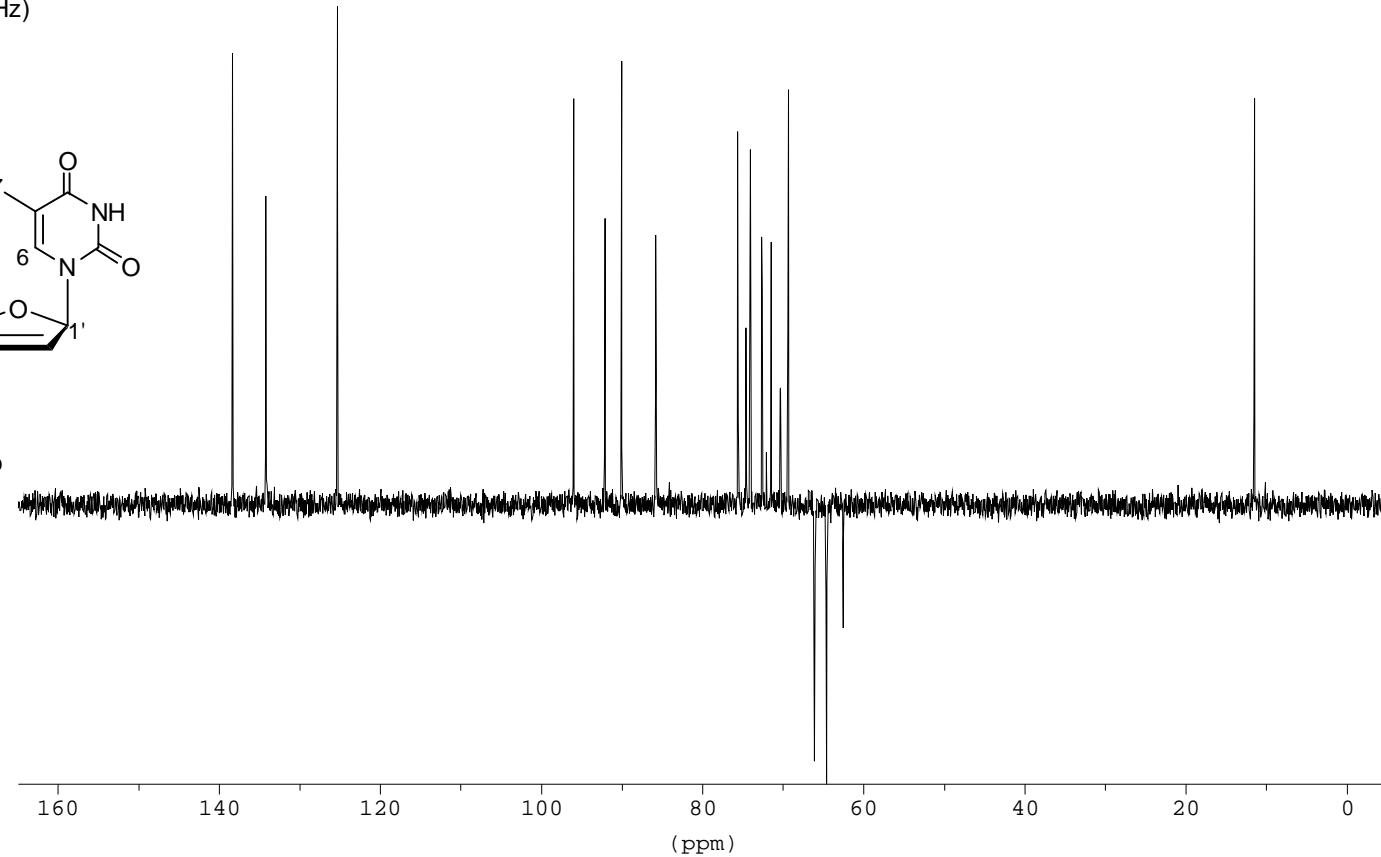


2',3'-Didehydro-2',3'-dideoxythymidine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10b)

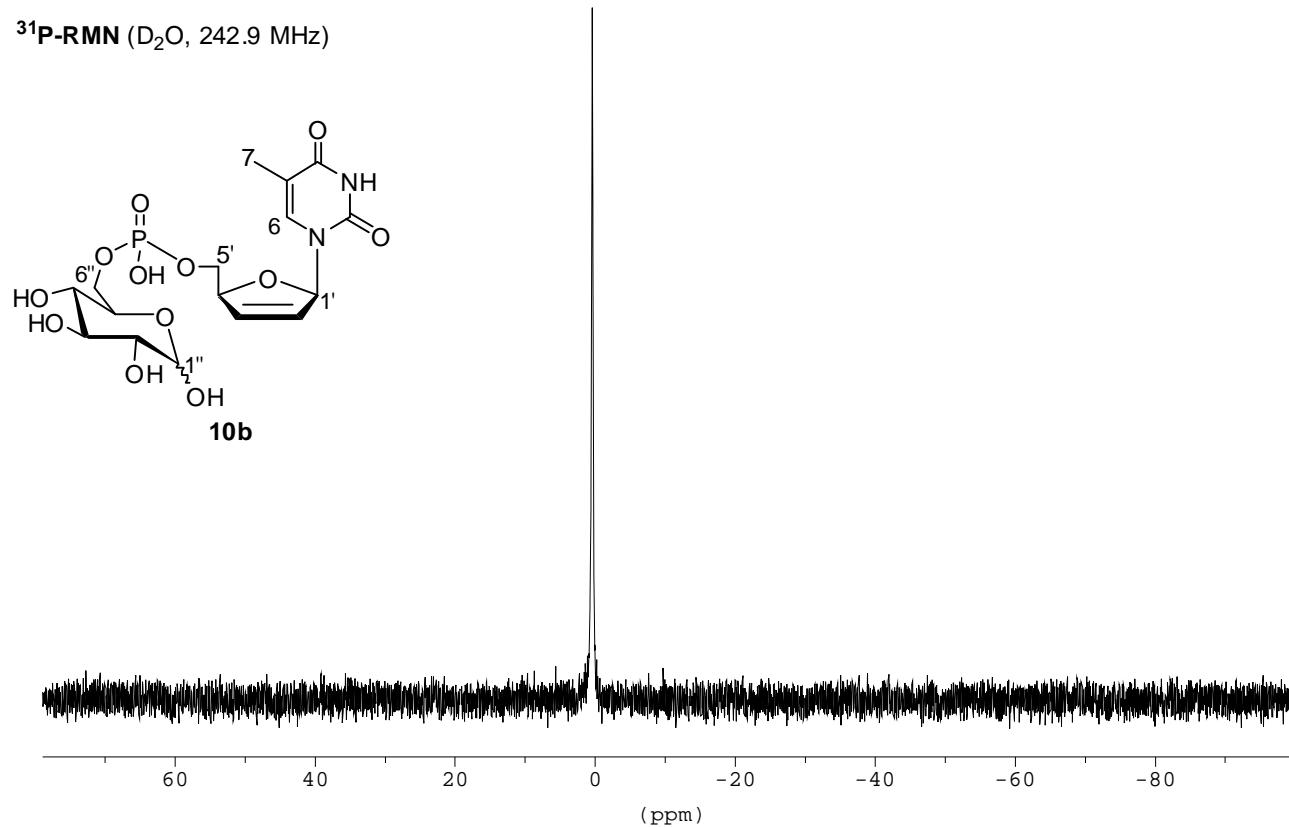
DEPT(D_2O , 150.5 MHz)



10b

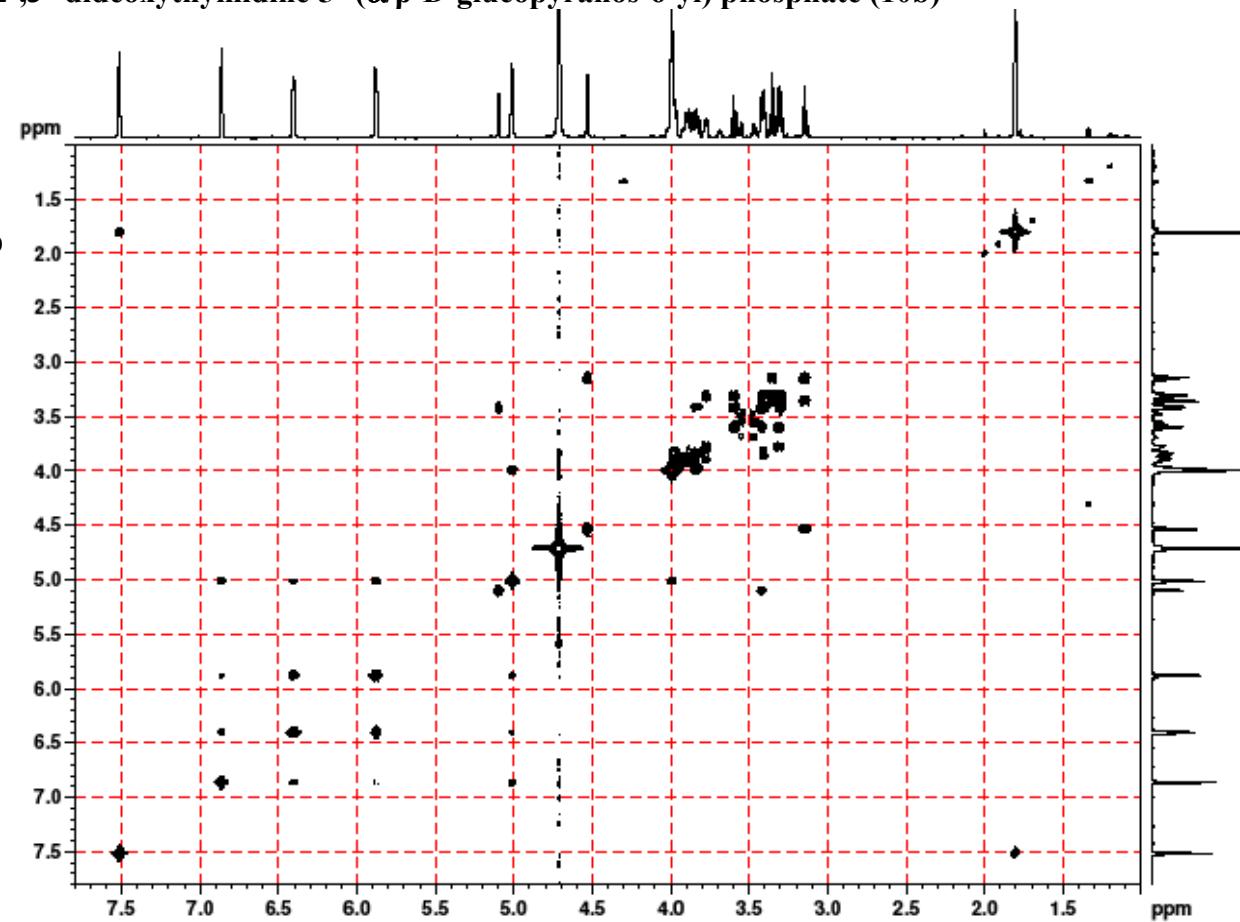
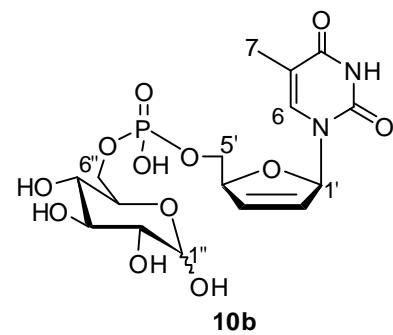


2',3'-Didehydro-2',3'-dideoxythymidine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10b)



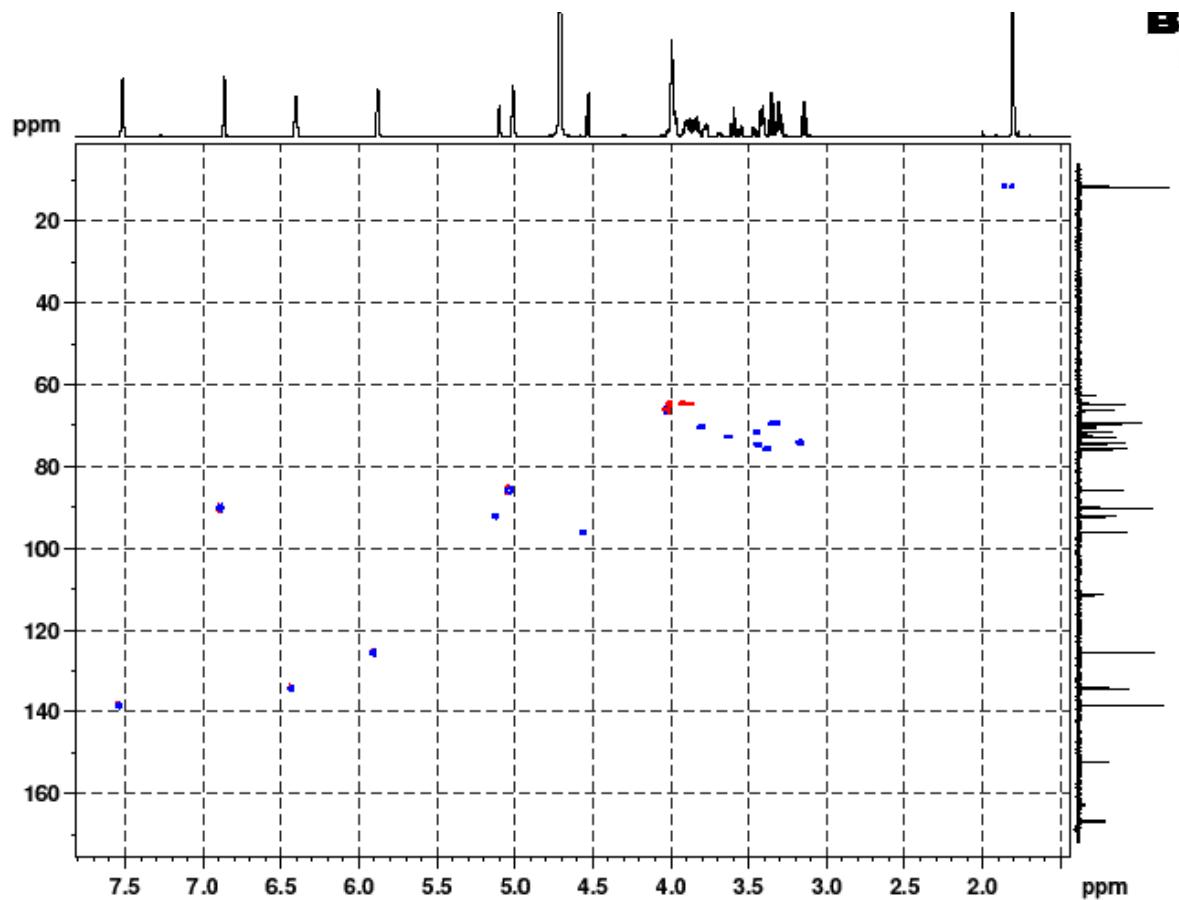
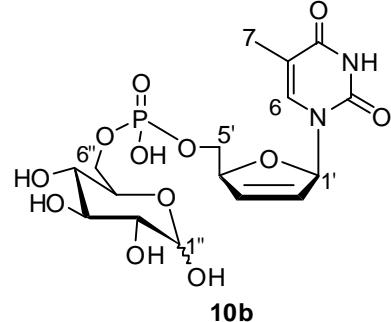
2',3'-Didehydro-2',3'-dideoxythymidine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10b)

COSY (D_2O , 600 MHz)



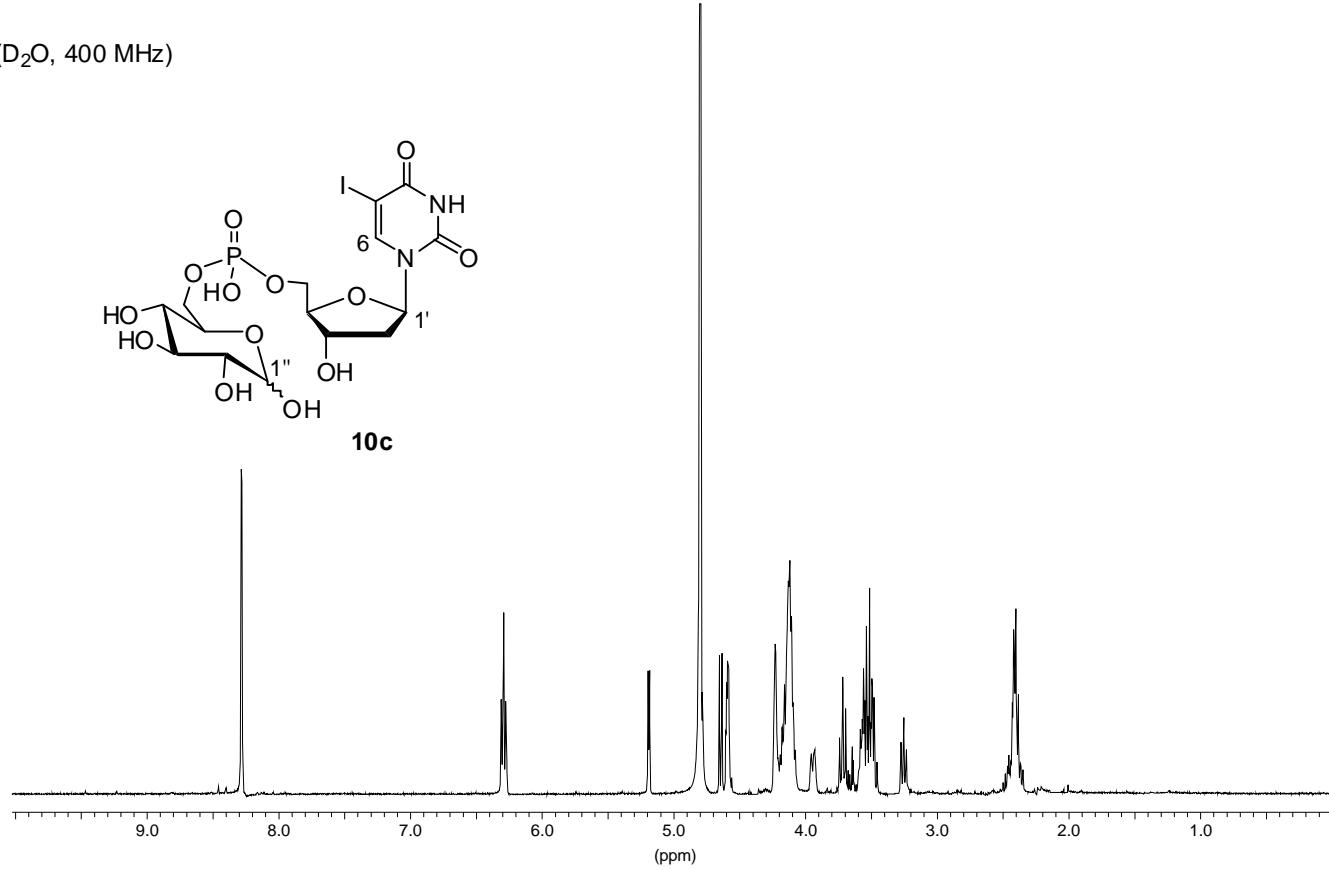
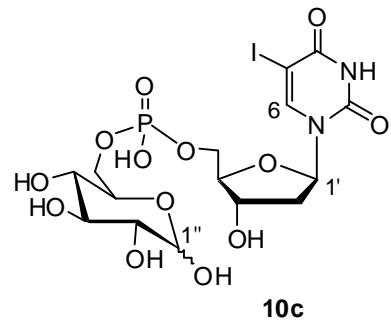
2',3'-Didehydro-2',3'-dideoxythymidine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10b)

HSQC (D_2O , 600 MHz)



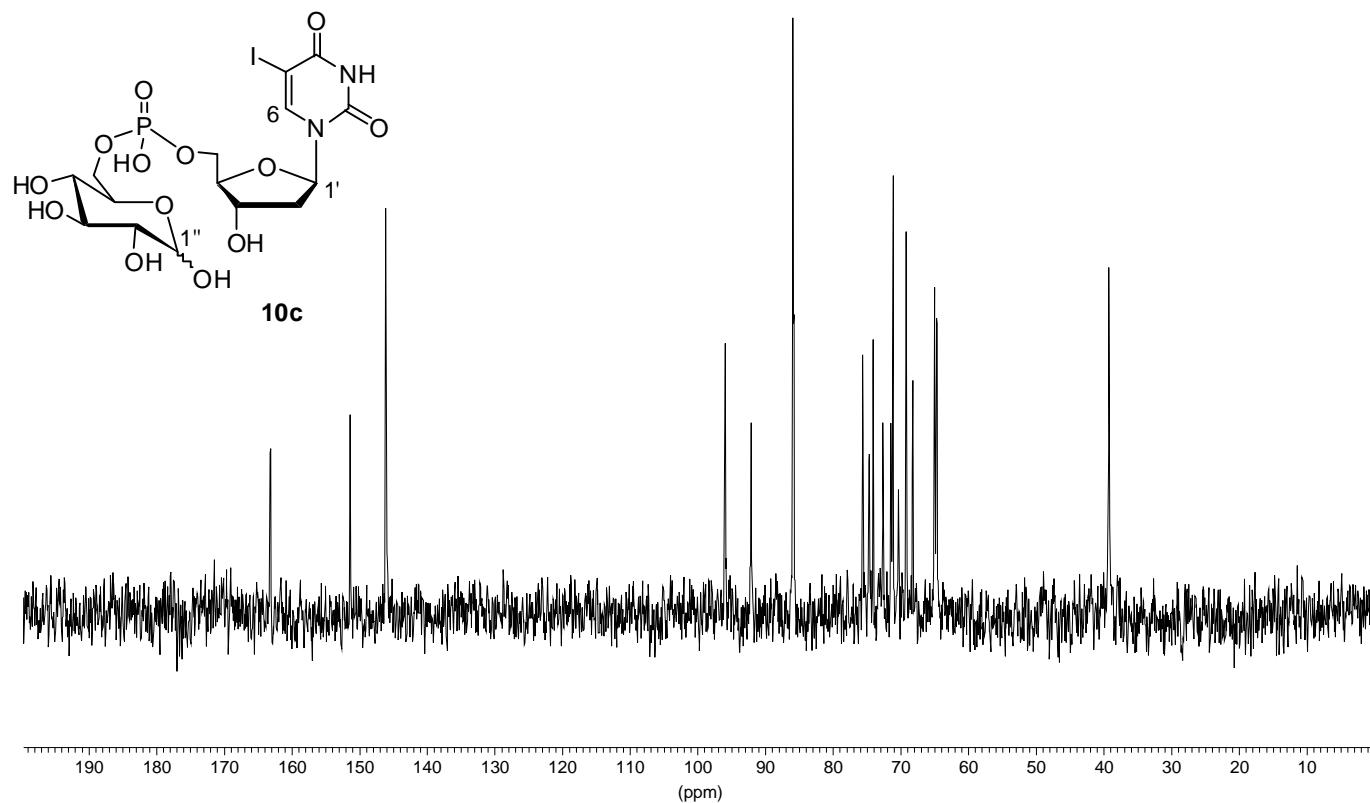
2'-Deoxy-5-iodouridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10c)

$^1\text{H-RMN}$ (D_2O , 400 MHz)

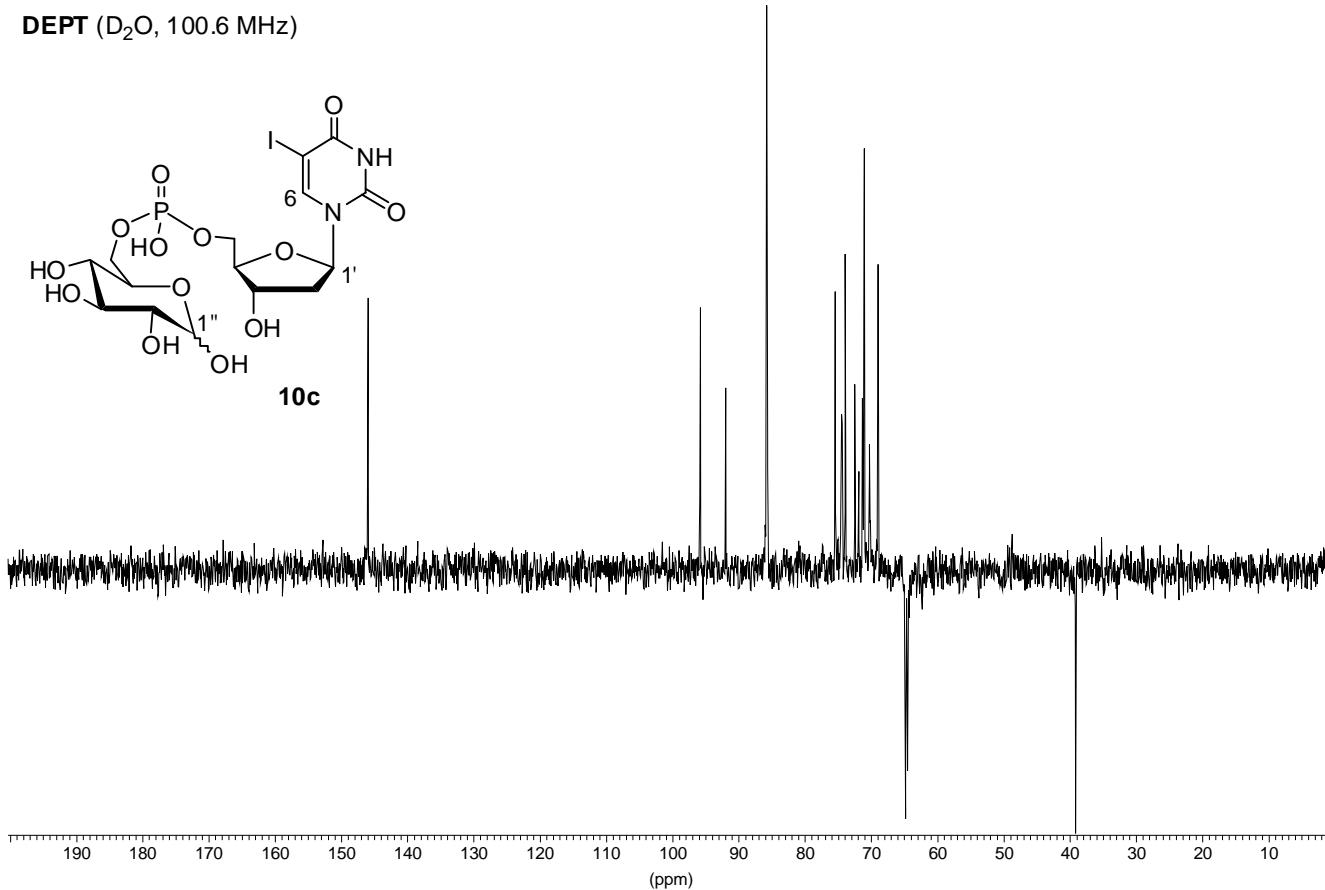


2'-Deoxy-5-iodouridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10c)

^{13}C -RMN (D_2O , 100.6 MHz)

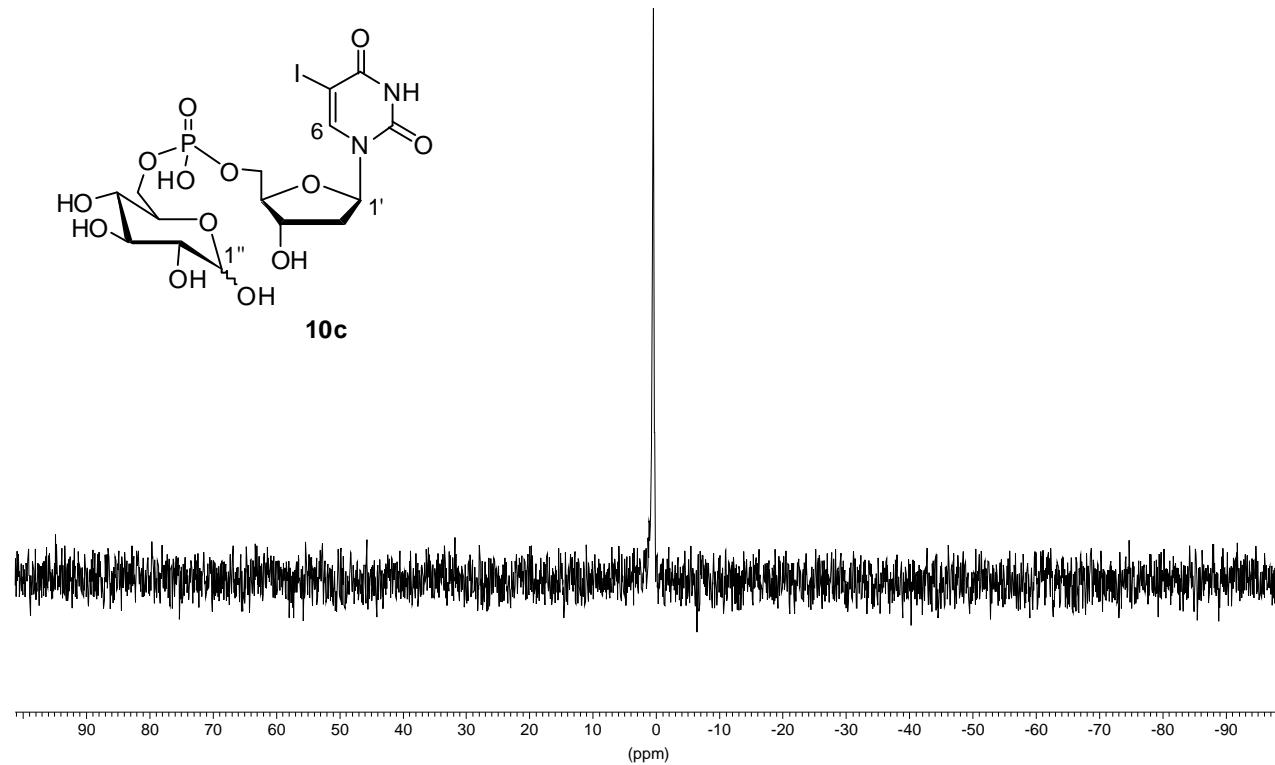
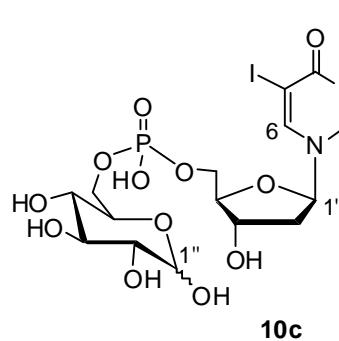


2'-Deoxy-5-iodouridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10c)

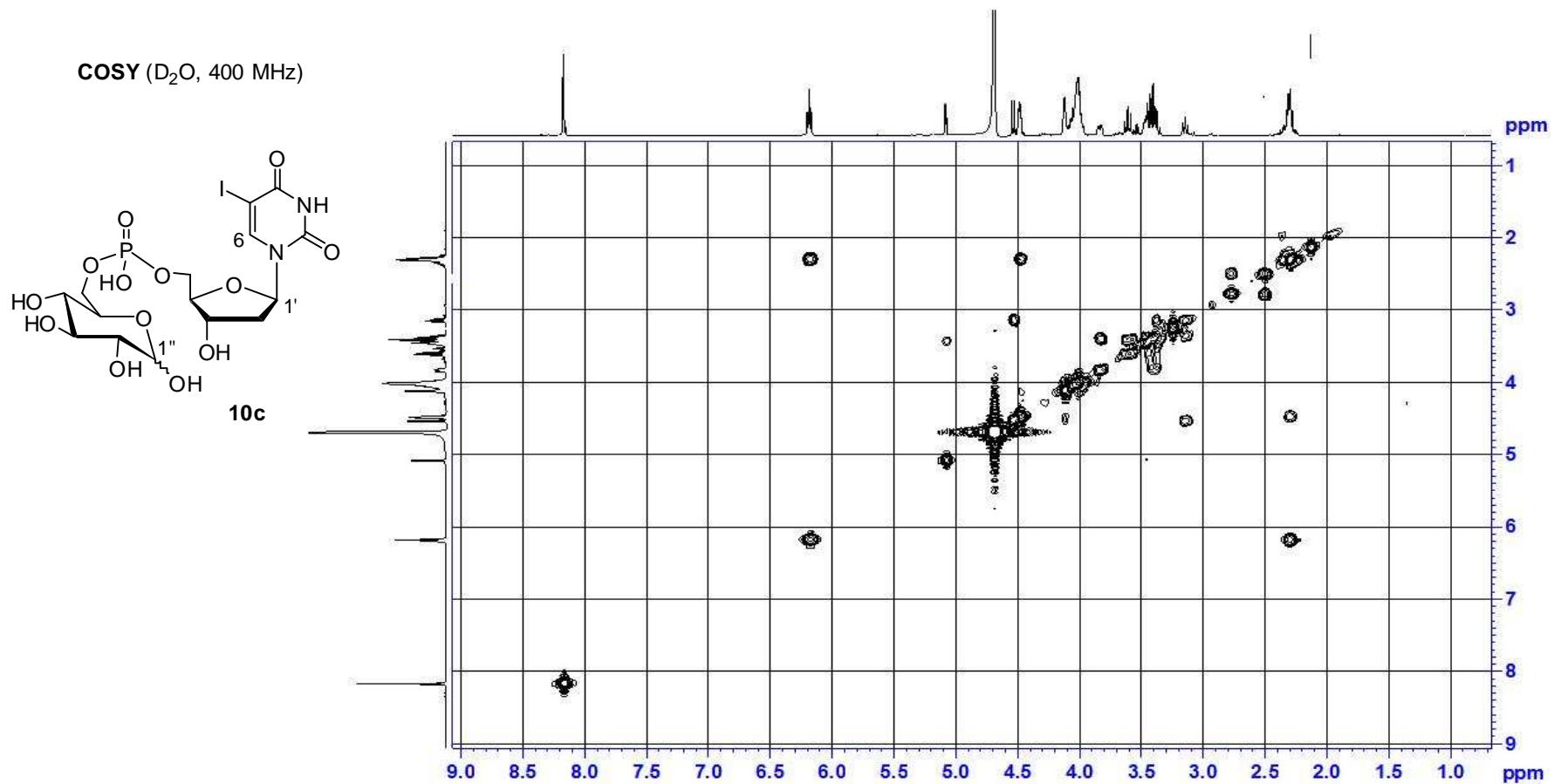


2'-Deoxy-5-iodouridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10c)

$^{31}\text{P-RMN}$ (D_2O , 161.9 MHz)

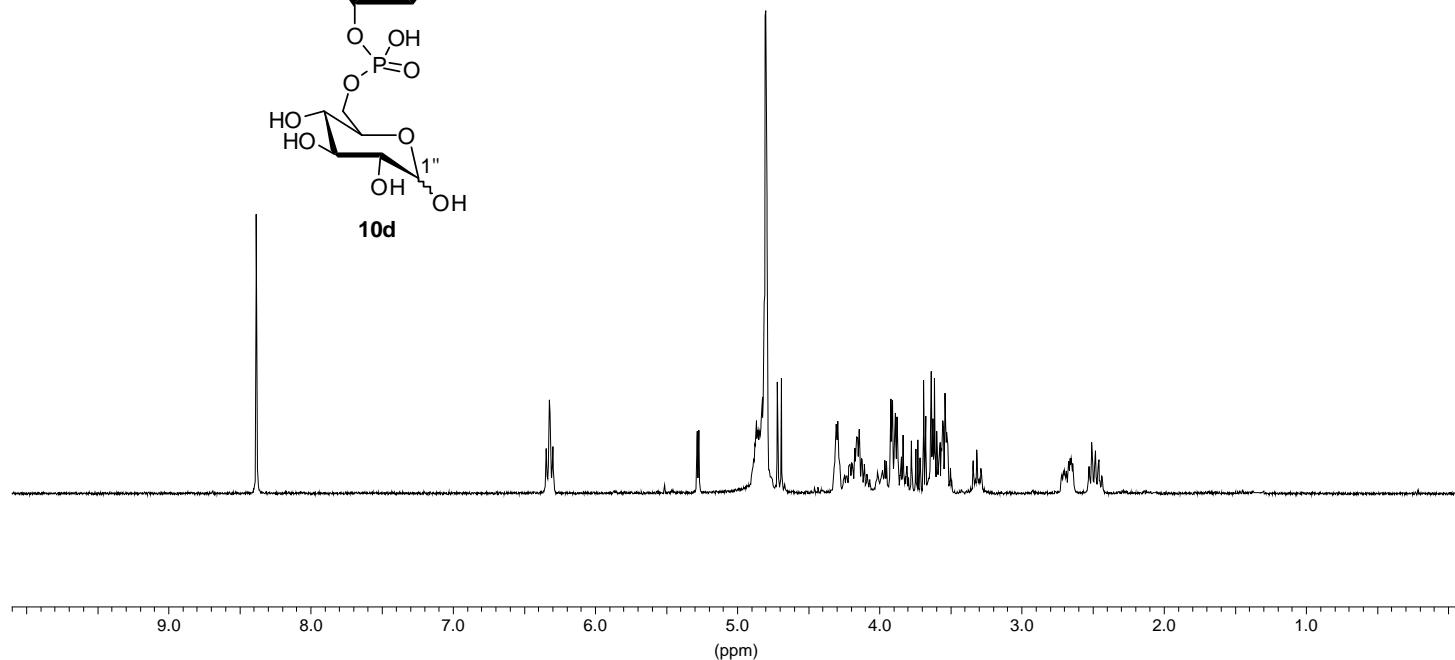
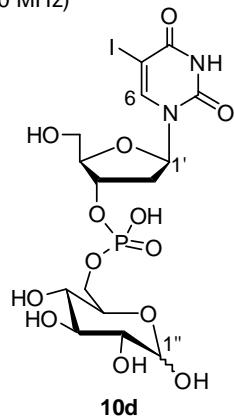


2'-Deoxy-5-iodouridine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10c)

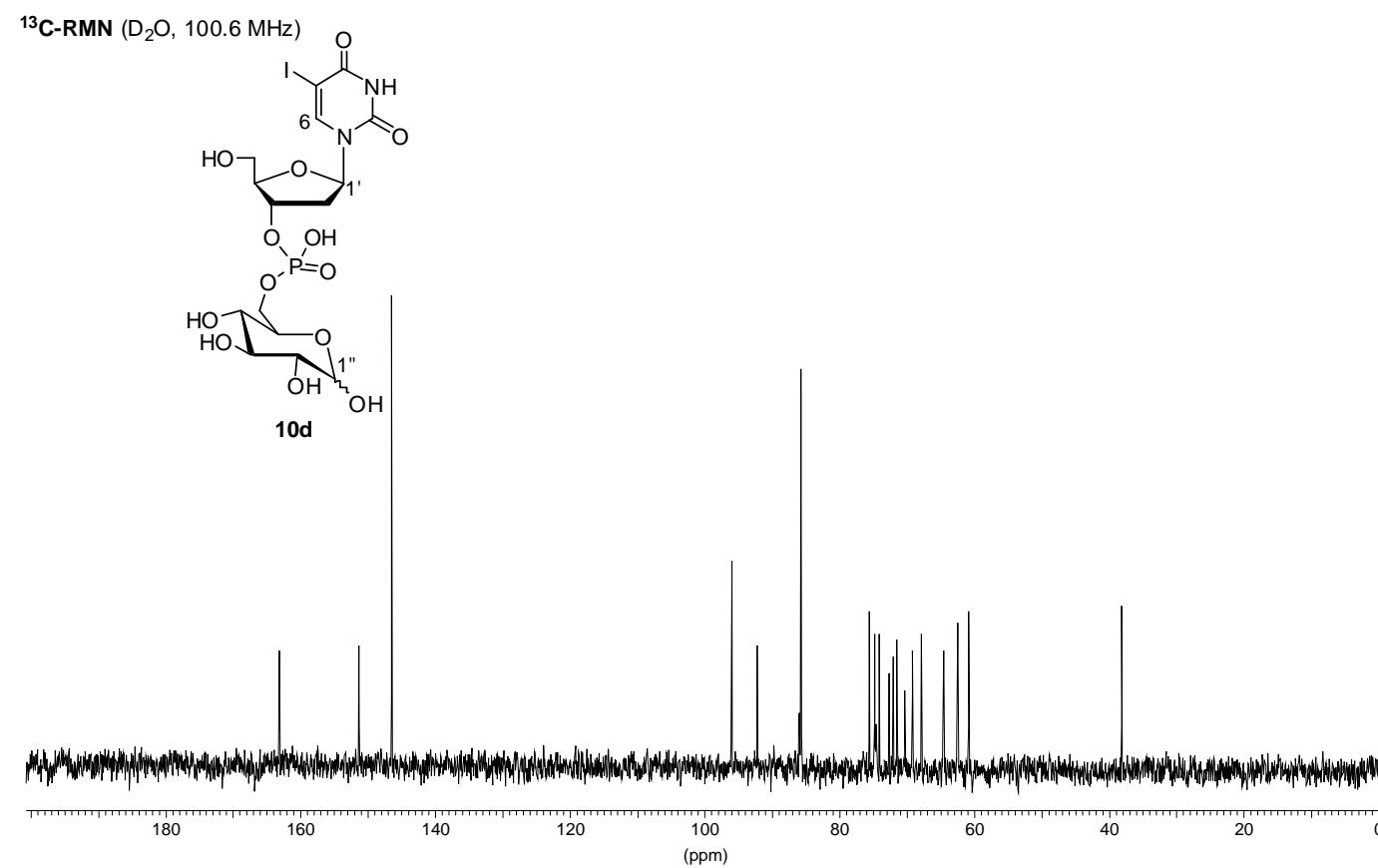


2'-Deoxy-5-iodouridine 3'-(α / β -D-glucopyranos-6-yl) phosphate (10d)

¹H-RMN (D_2O , 400 MHz)

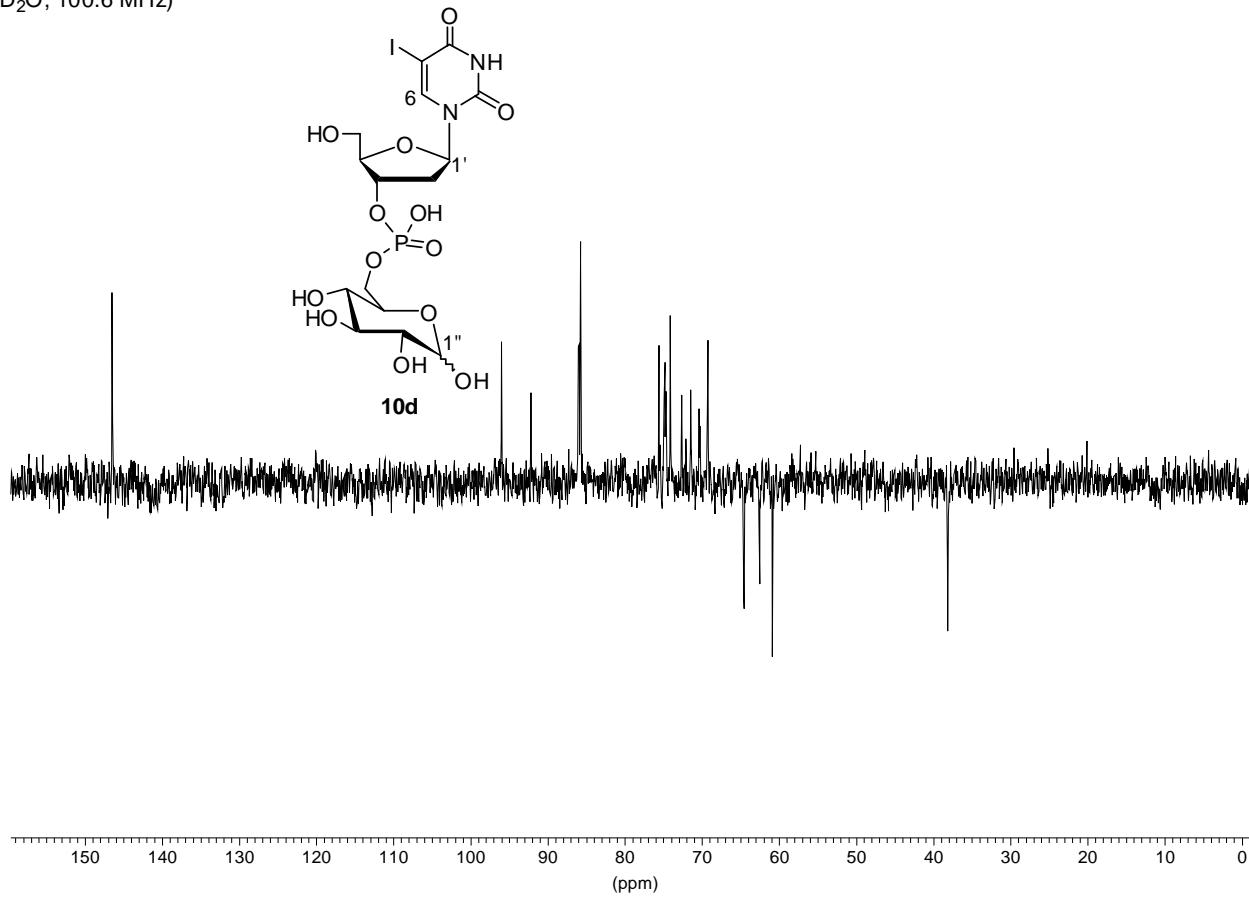


2'-Deoxy-5-iodouridine 3'-(α / β -D-glucopyranos-6-yl) phosphate (10d)

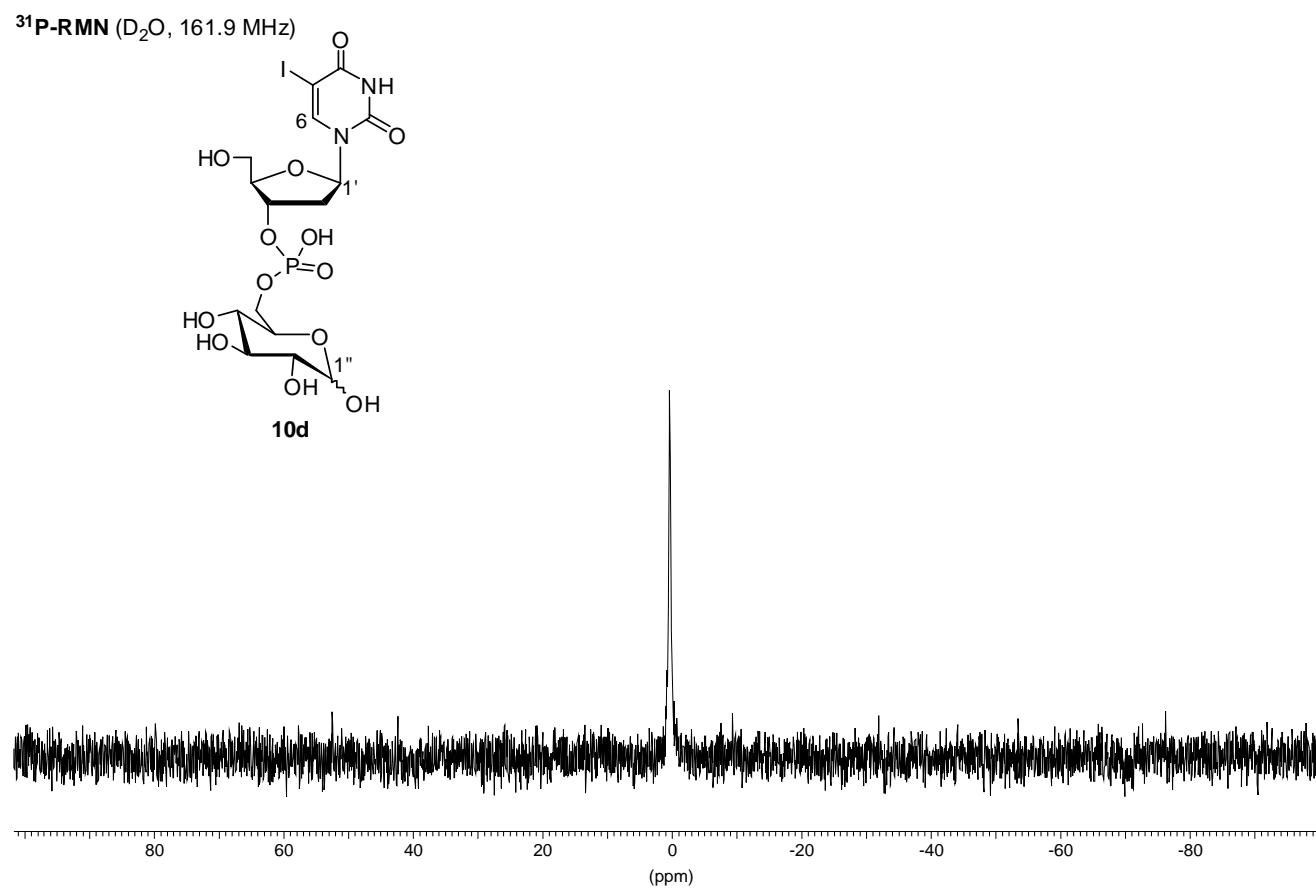


2'-Deoxy-5-iodouridine 3'-(α / β -D-glucopyranos-6-yl) phosphate (10d)

DEPT (D_2O , 100.6 MHz)

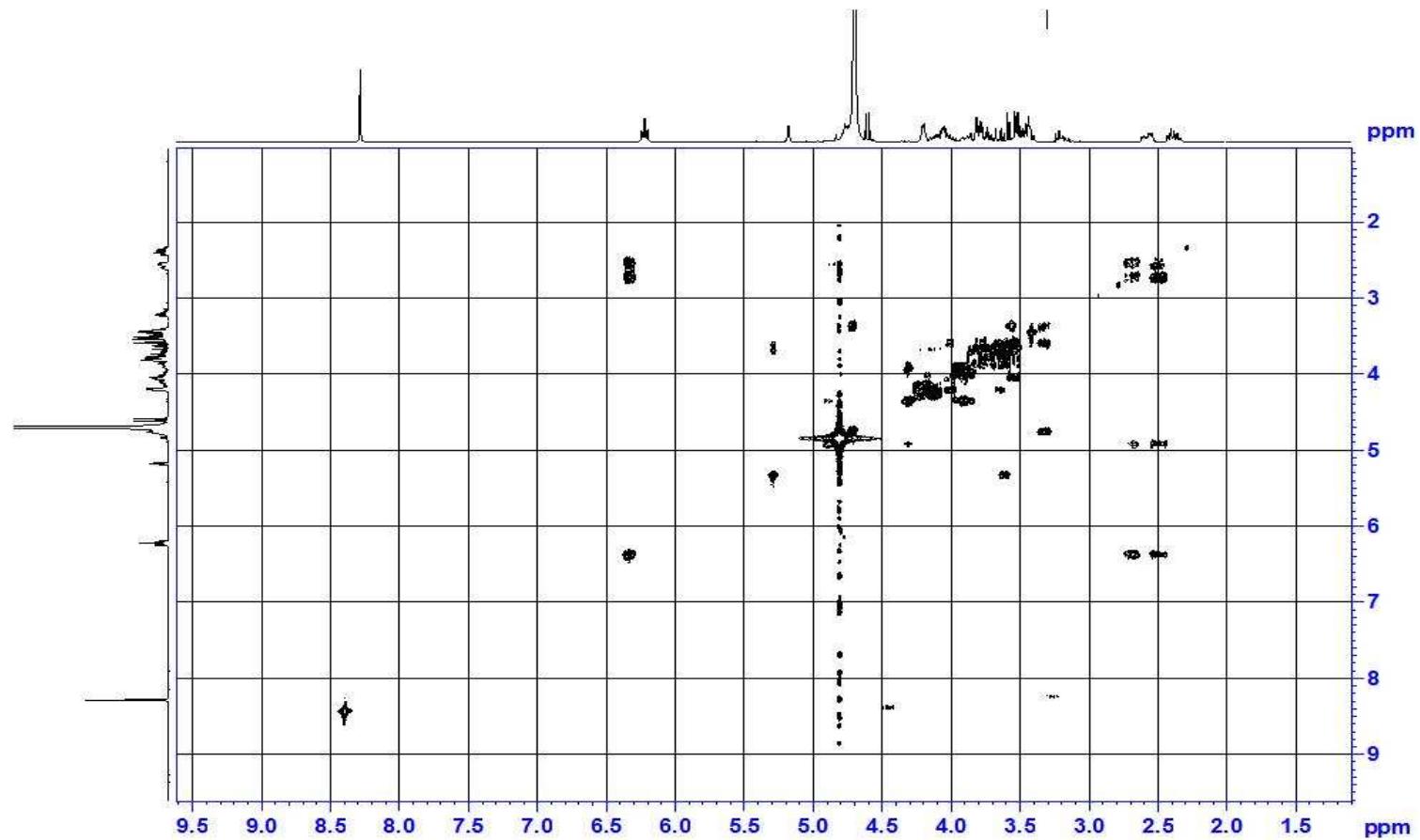
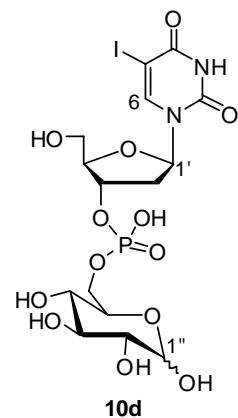


2'-Deoxy-5-iodouridine 3'-(α / β -D-glucopyranos-6-yl) phosphate (10d)

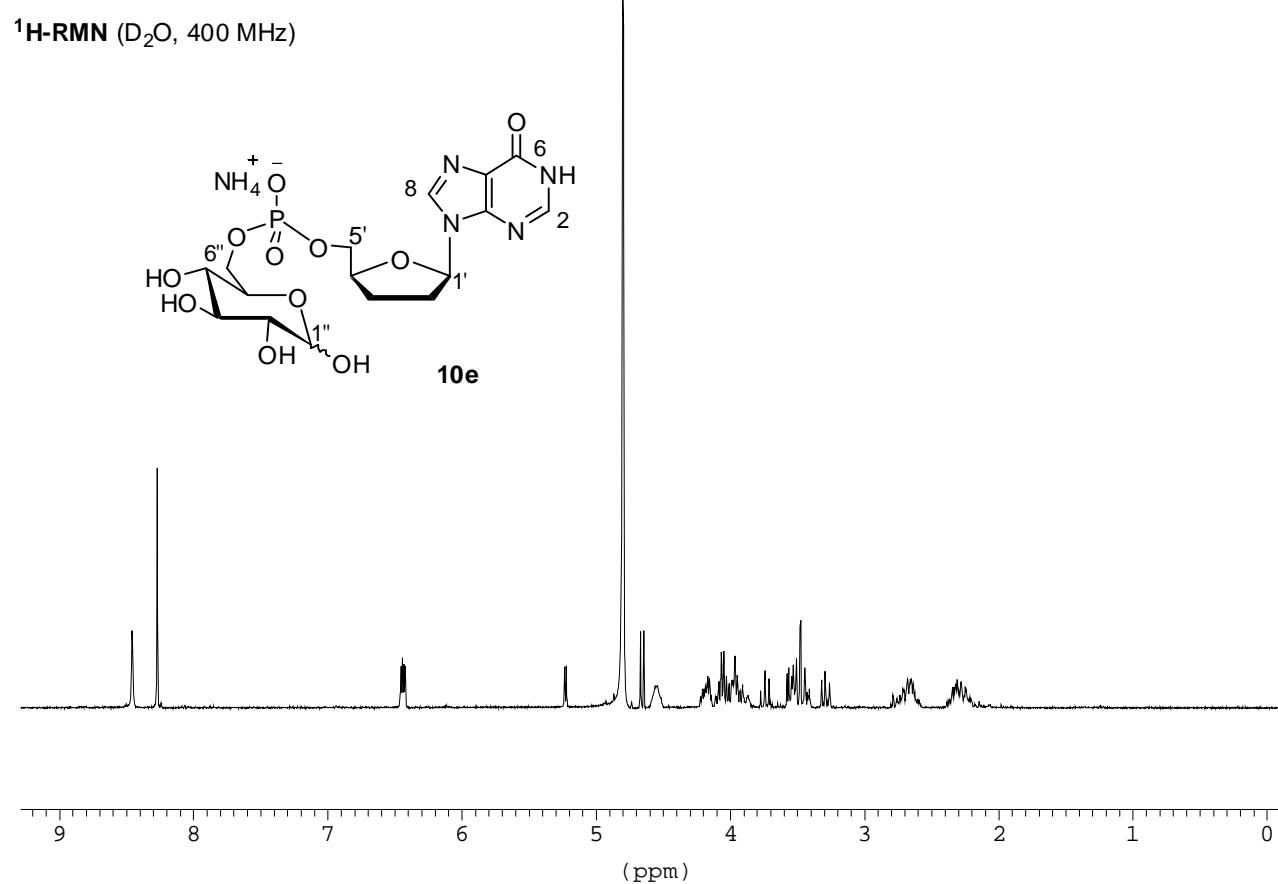


2'-Deoxy-5-iodouridine 3'-(α / β -D-glucopyranos-6-yl) phosphate (10d)

COSY (D_2O , 400 MHz)

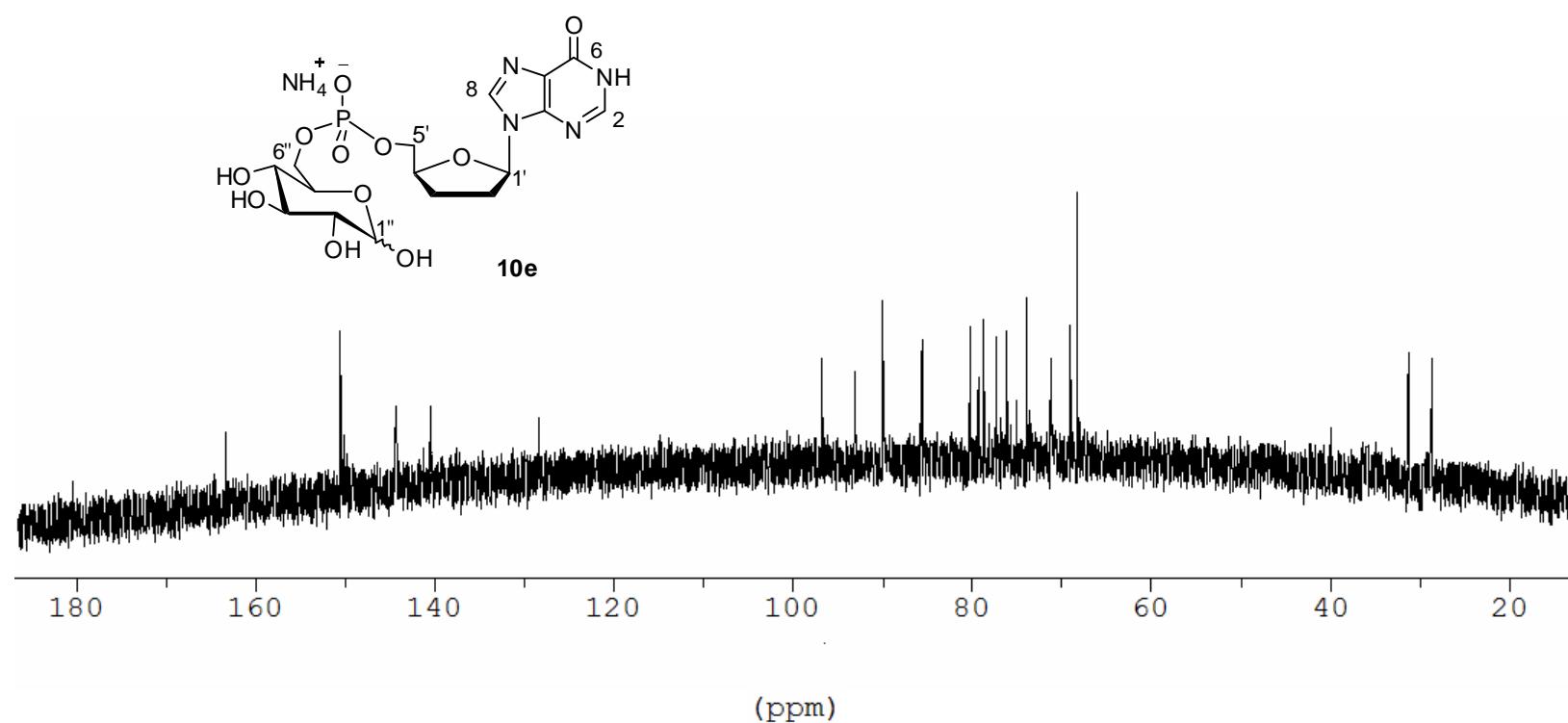


2',3'-Dideoxyinosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10e)



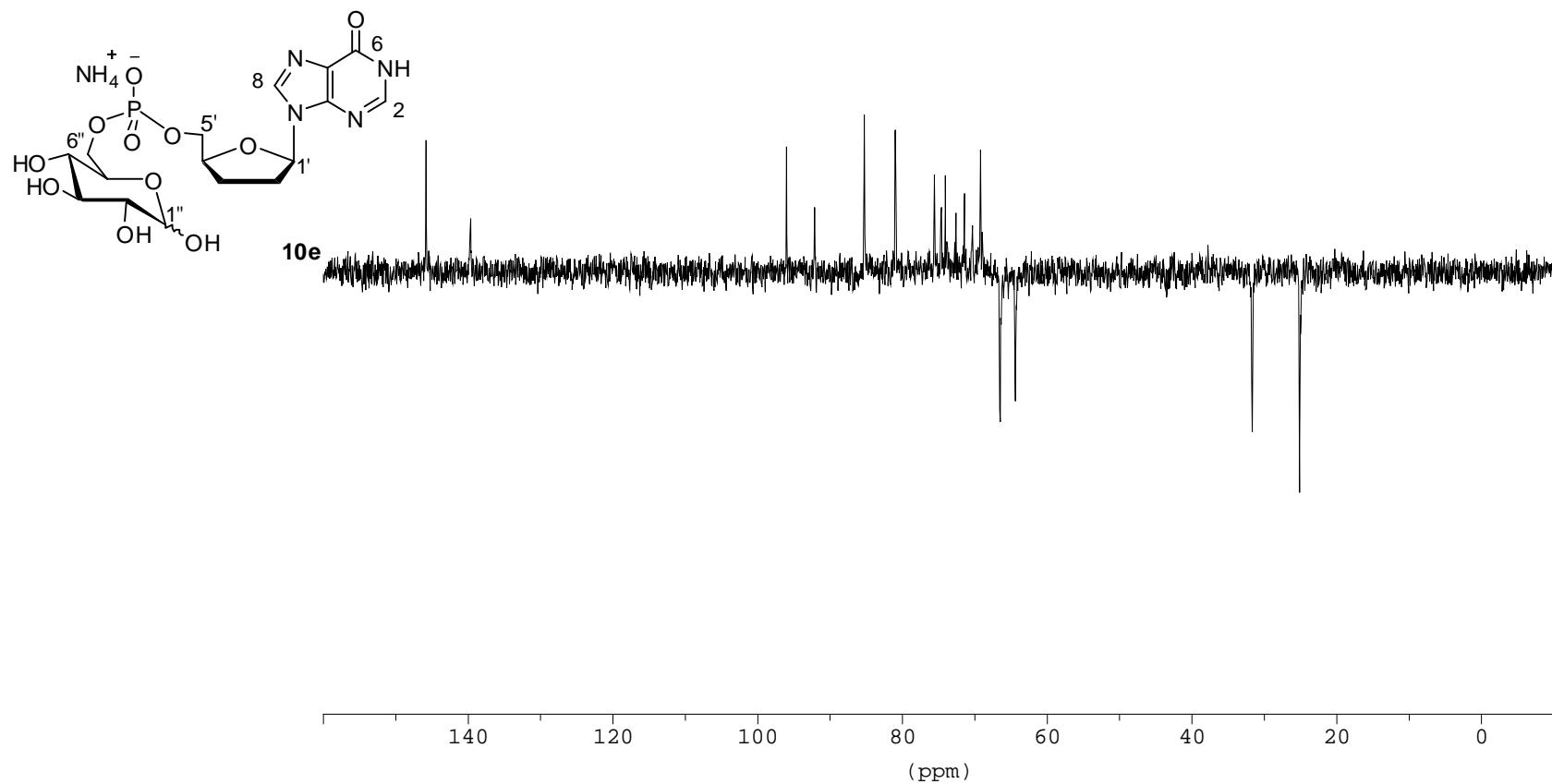
2',3'-Dideoxyinosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10e)

¹³C-RMN (D_2O , 100.6 MHz)



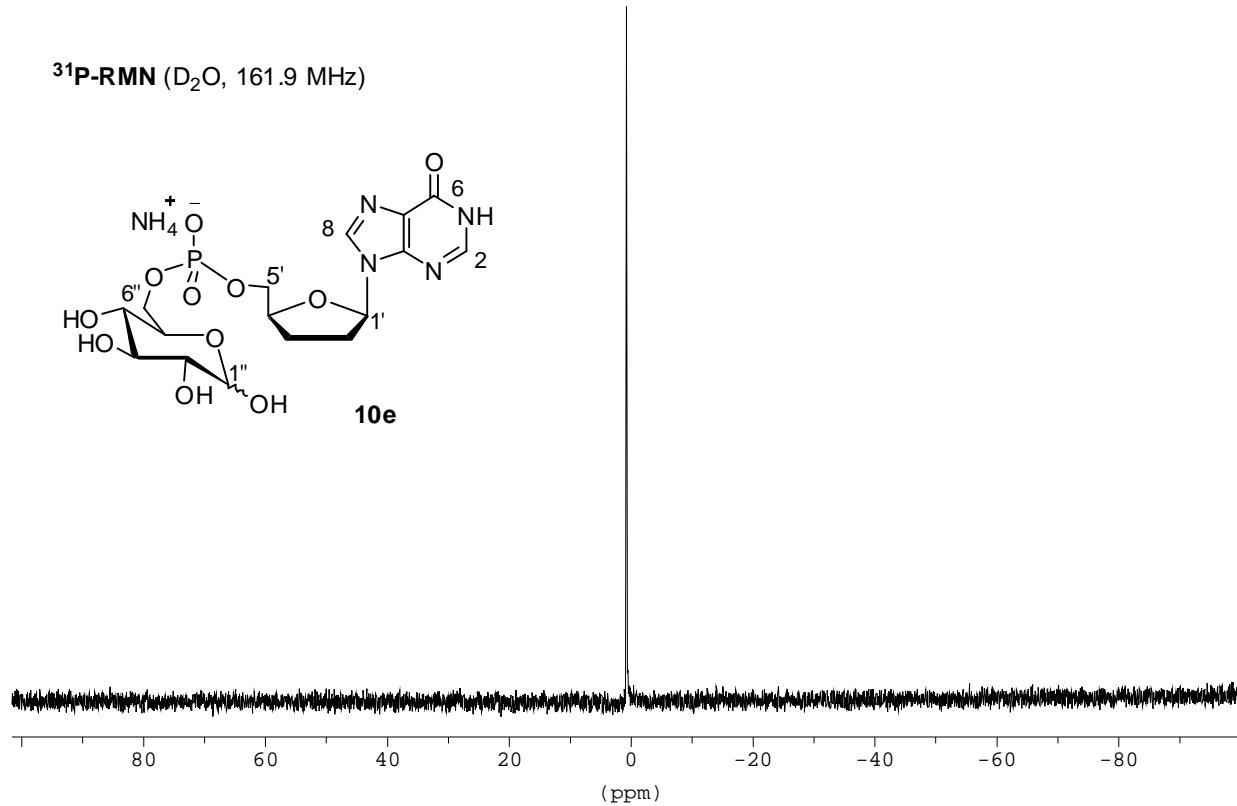
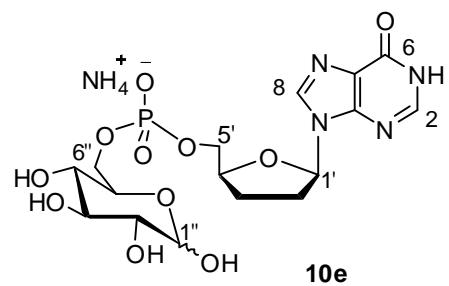
2',3'-Dideoxyinosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10e)

DEPT (D_2O , 100.6 MHz)



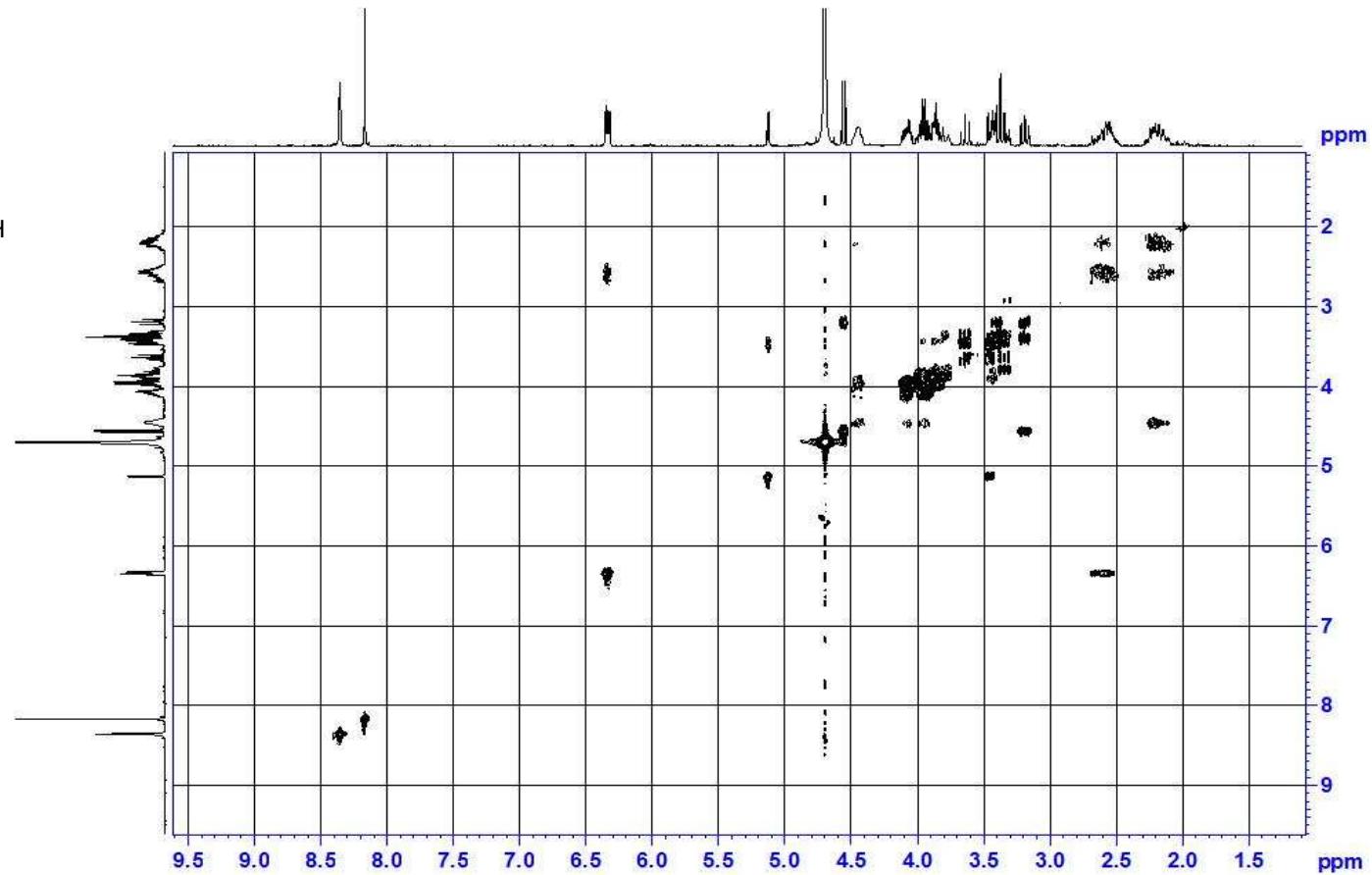
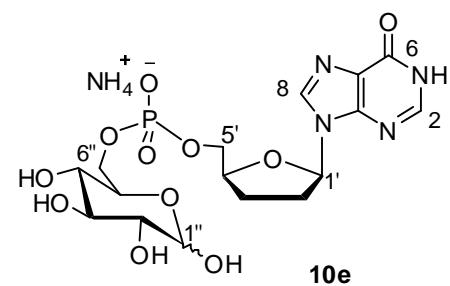
2',3'-Dideoxyinosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10e)

^{31}P -RMN (D_2O , 161.9 MHz)



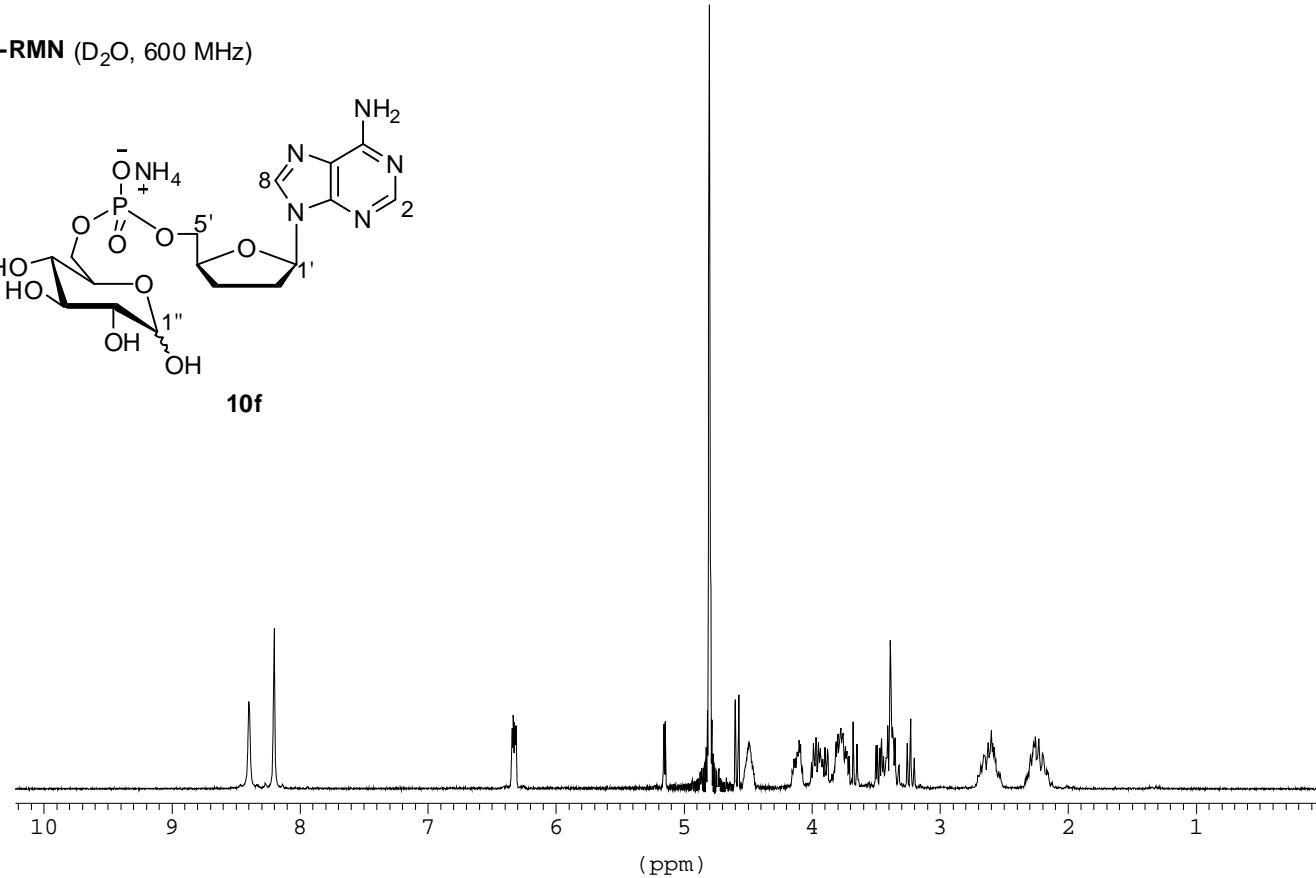
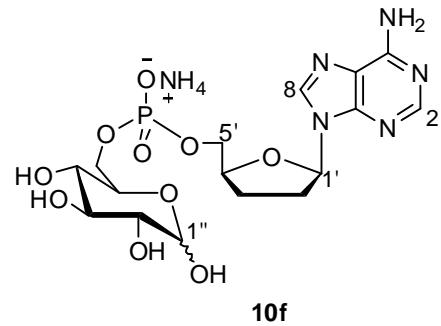
2',3'-Dideoxyinosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10e)

COSY (D_2O , 400 MHz)



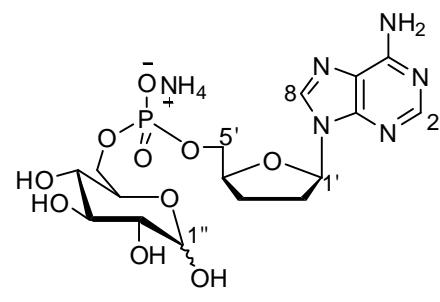
2',3'-Dideoxyadenosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10f)

¹H-RMN (D₂O, 600 MHz)

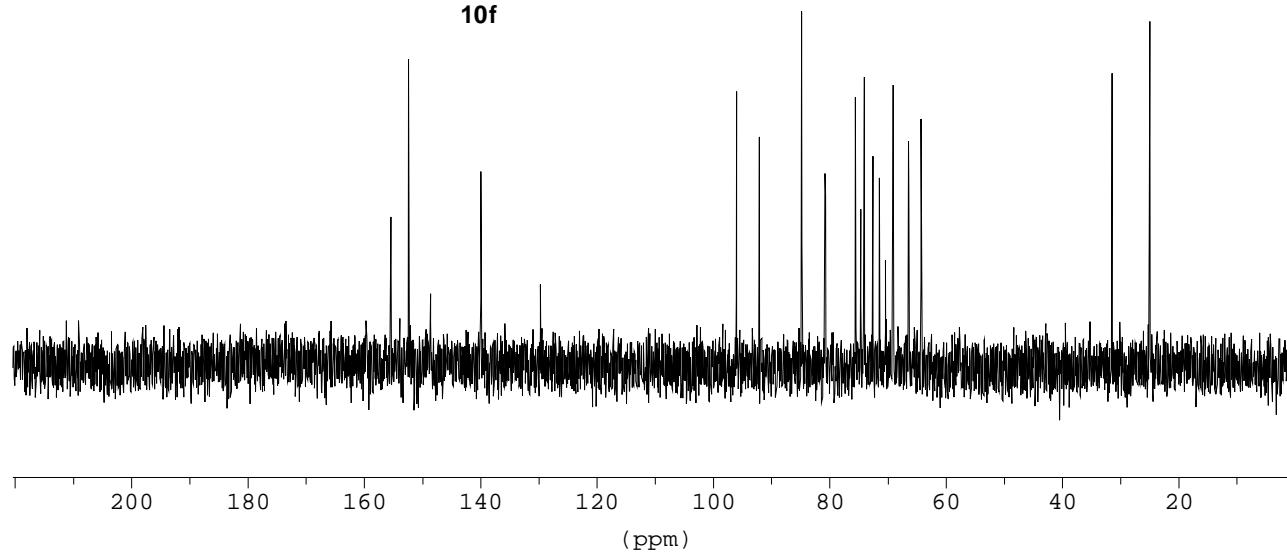


2',3'-Dideoxyadenosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10f)

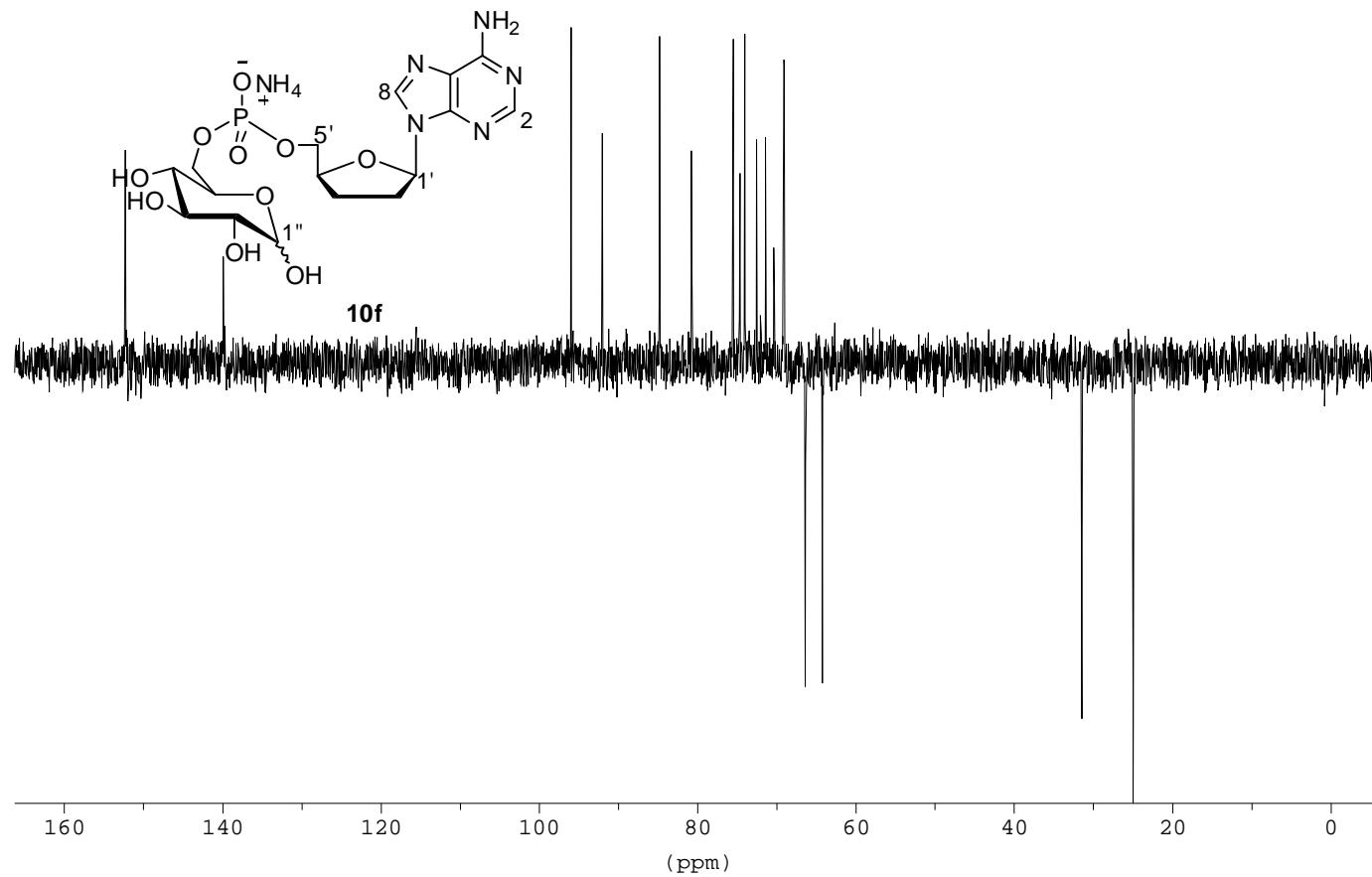
^{13}C -RMN (D_2O , 150.5 MHz)



10f

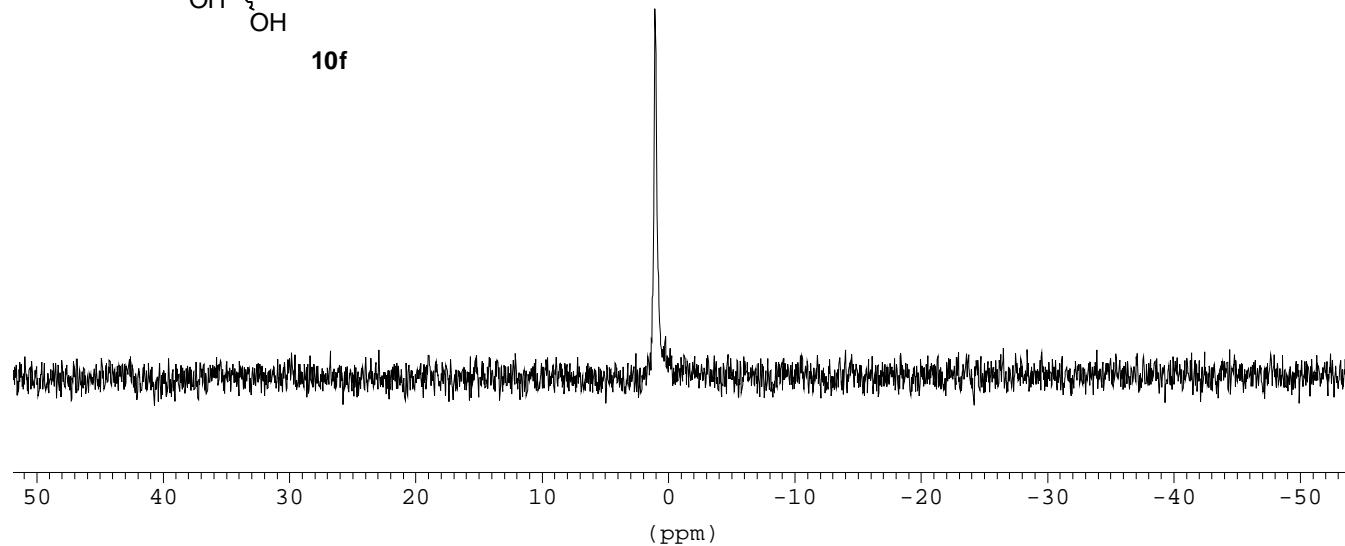
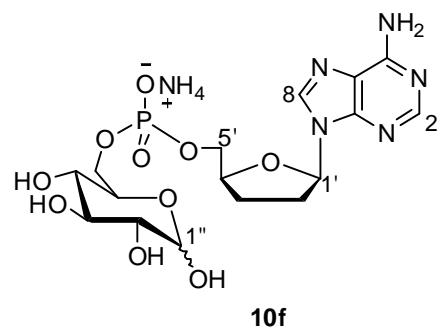


2',3'-Dideoxyadenosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10f)
DEPT (D_2O , 150.5 MHz)

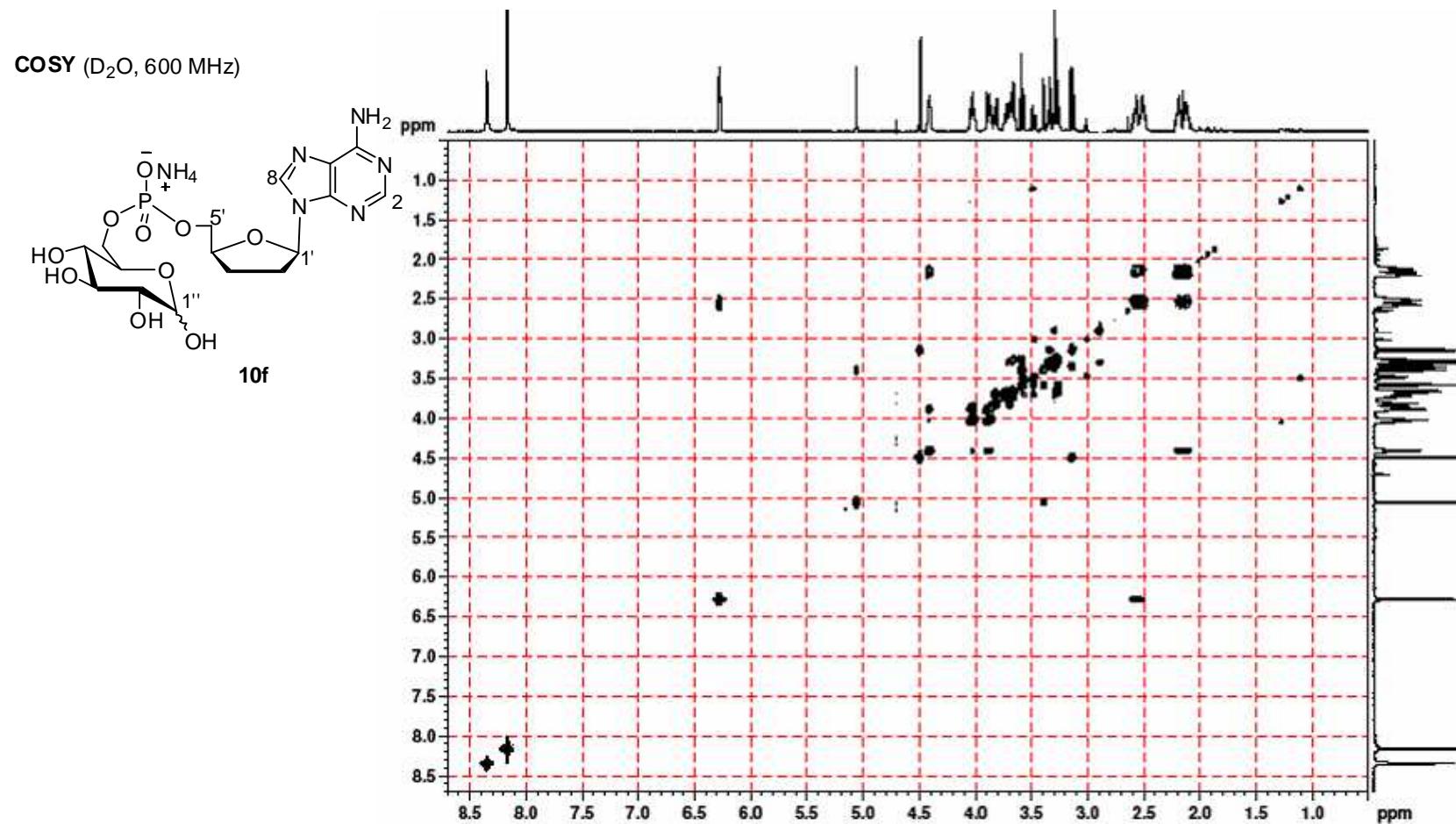


2',3'-Dideoxyadenosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10f)

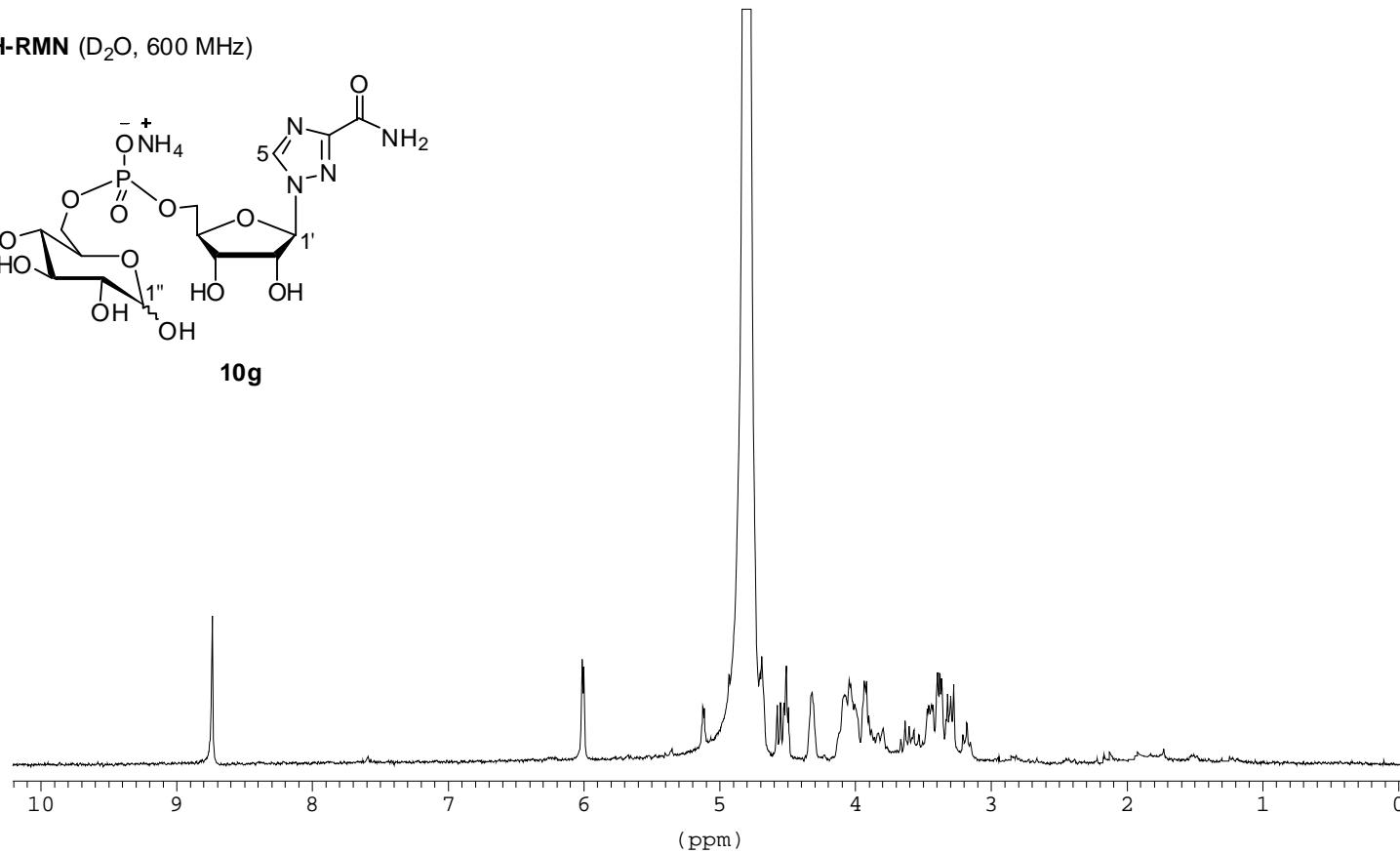
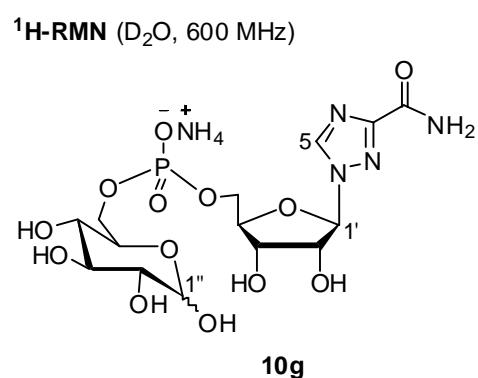
$^{31}\text{P-RMN}$ (D_2O , 242.9 MHz)



2',3'-Dideoxyadenosine 5'-(α / β -D-glucopyranos-6-yl) phosphate (10f)

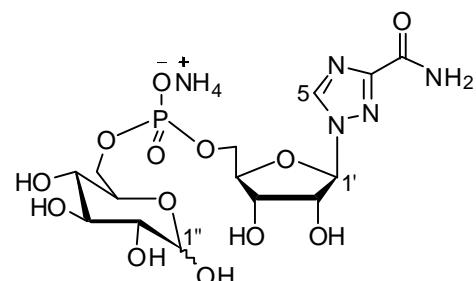


1'-(3-Carbamoyl-1,2,4-triazoyl)- β -D-ribofuranose 5'-(α/β -D-glucopyranos-6-yl) phosphate (10g)

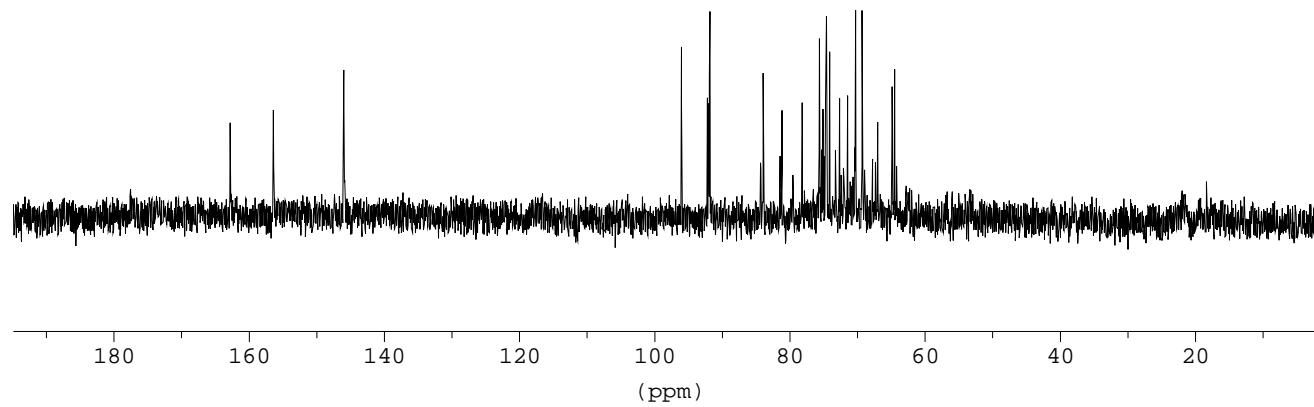


1'-(3-Carbamoyl-1,2,4-triazoyl)- β -D-ribofuranose 5'-(α/β -D-glucopyranos-6-yl) phosphate (10g)

¹³C-RMN (D_2O , 150.5 MHz)

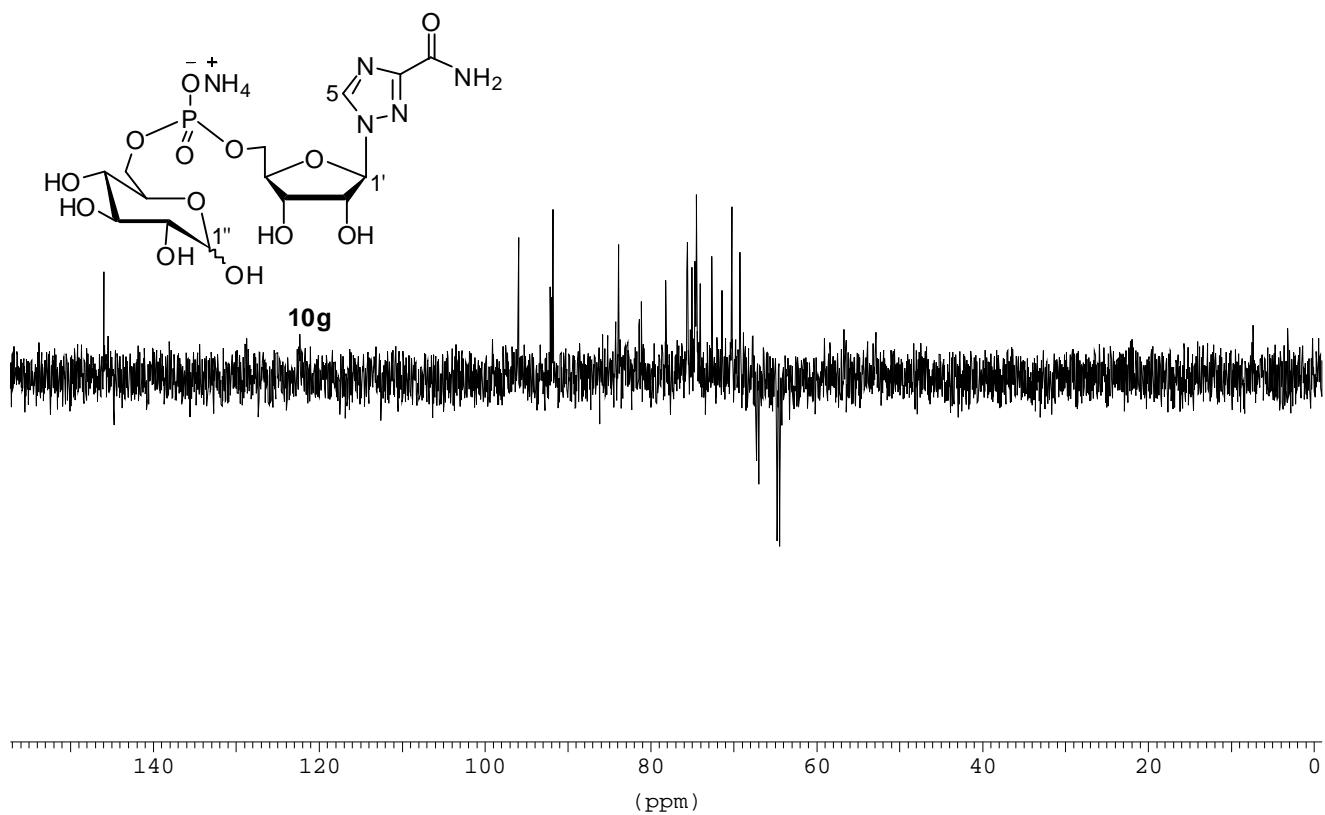


10g

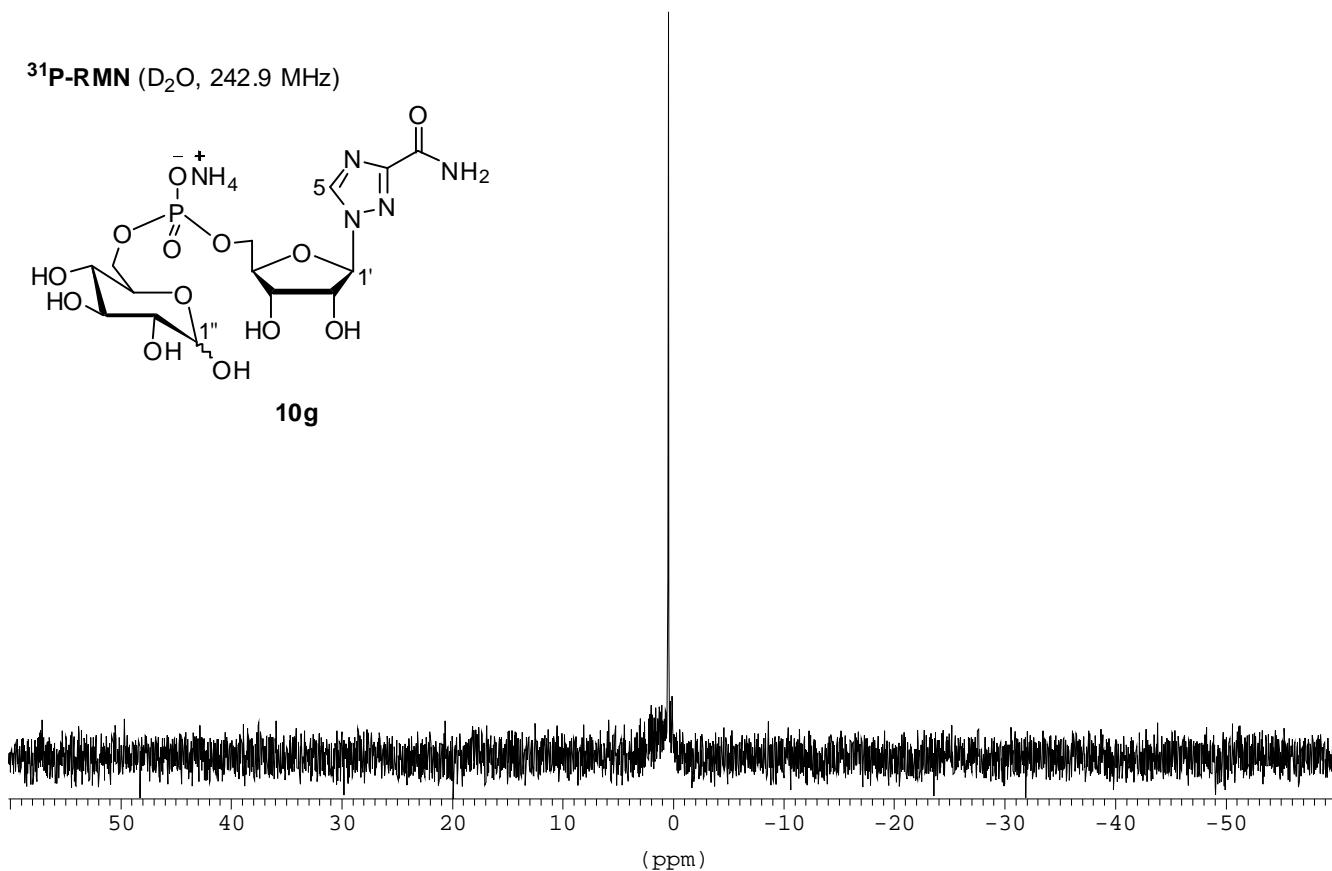


1'-(3-Carbamoyl-1,2,4-triazoyl)- β -D-ribofuranose 5'-(α/β -D-glucopyranos-6-yl) phosphate (10g)

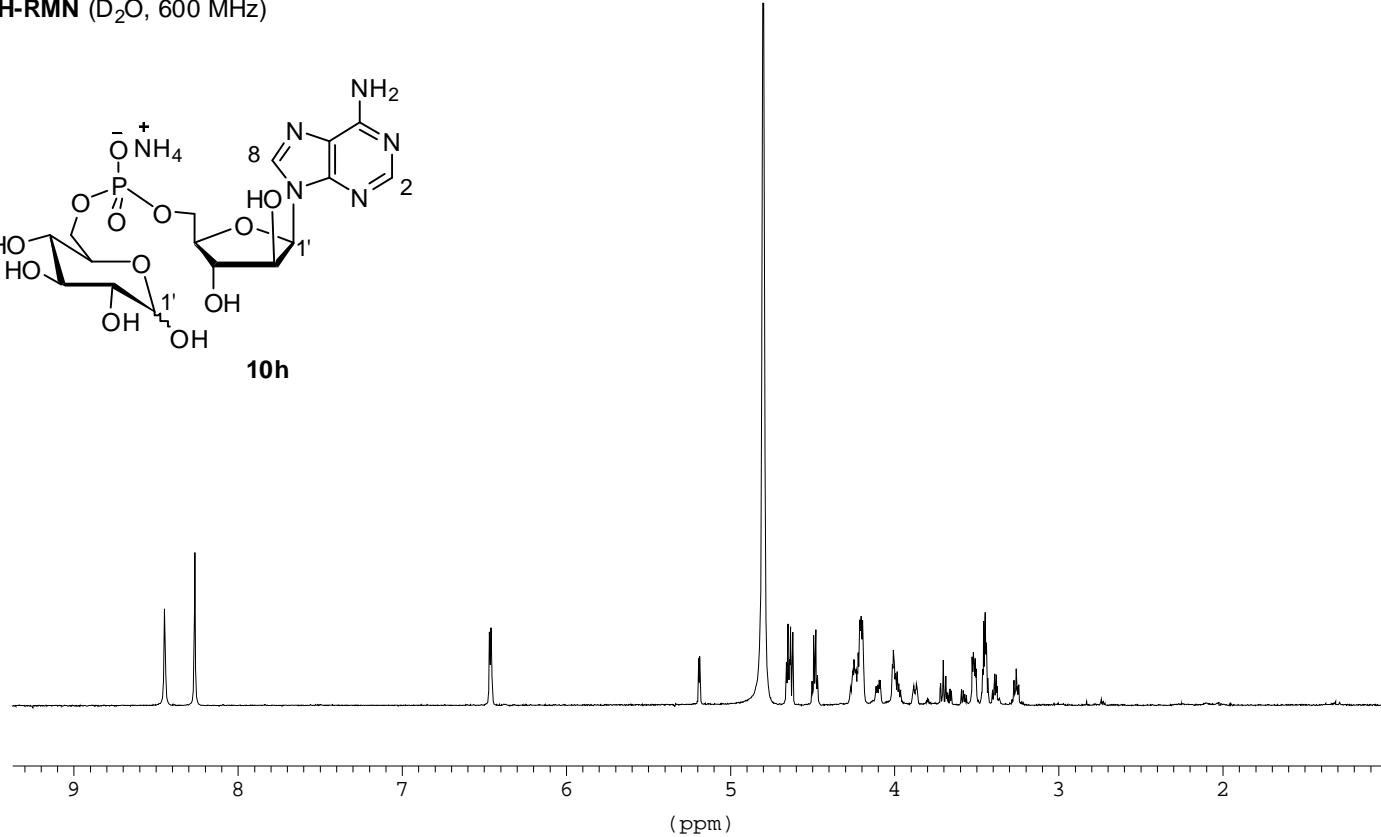
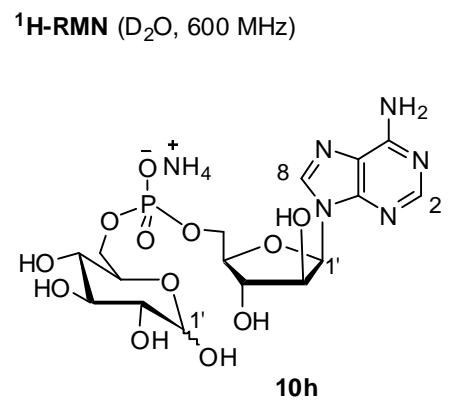
DEPT (D_2O , 150.5 MHz)



1'-(3-Carbamoyl-1,2,4-triazoyl)- β -D-ribofuranose 5'-(α/β -D-glucopyranos-6-yl) phosphate (10g)

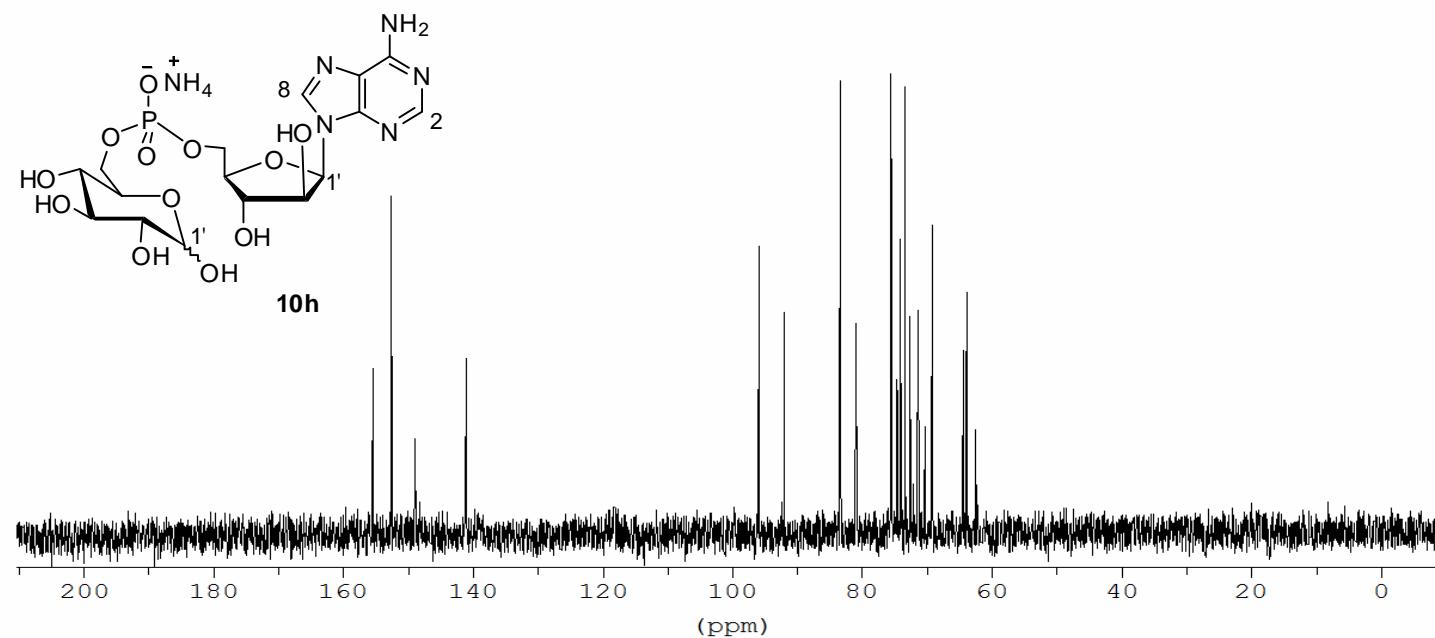


1'-Adeninyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10h)

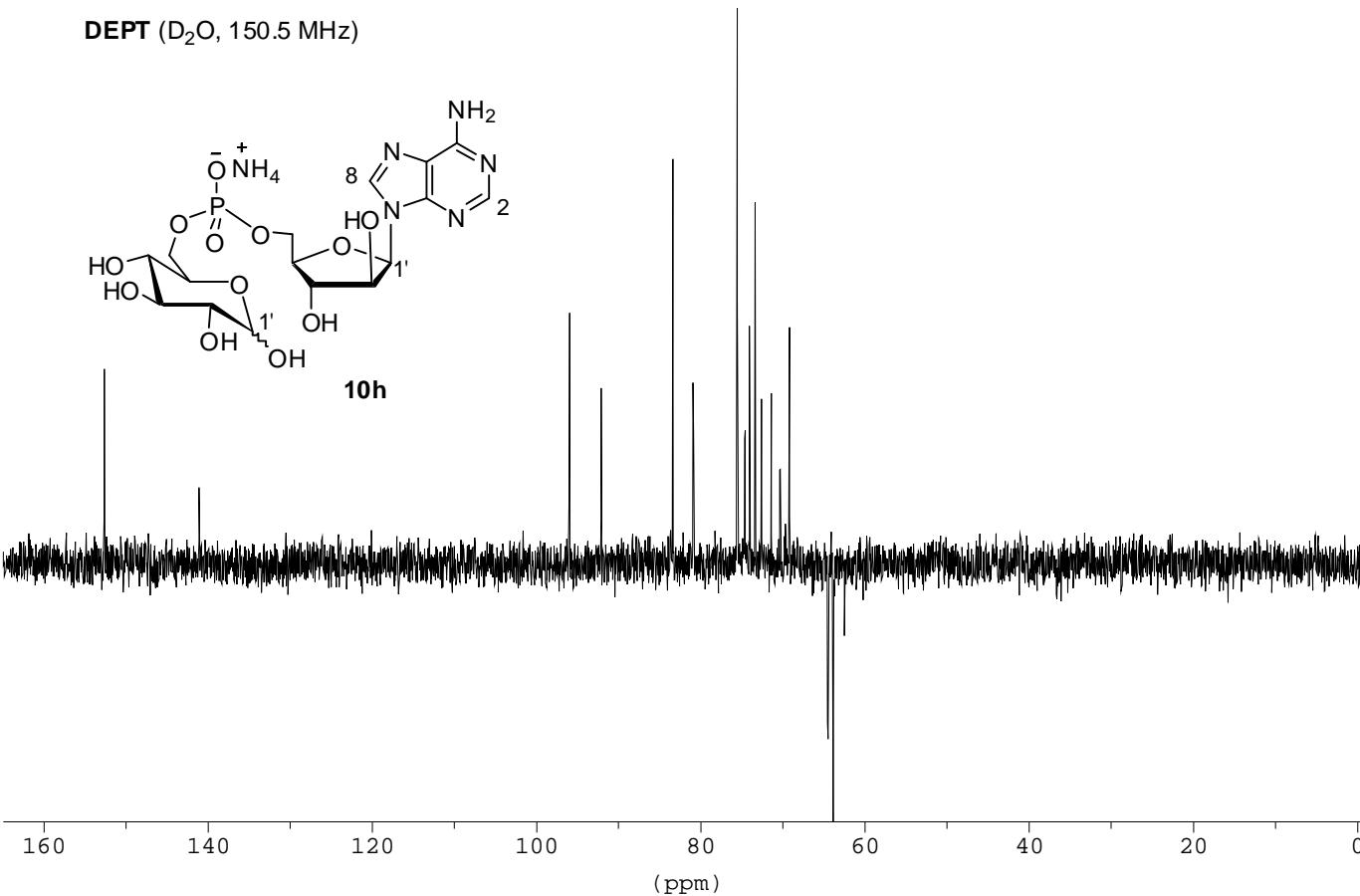


1'-Adeninyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10h)

^{13}C -RMN (D_2O , 150.5 MHz)

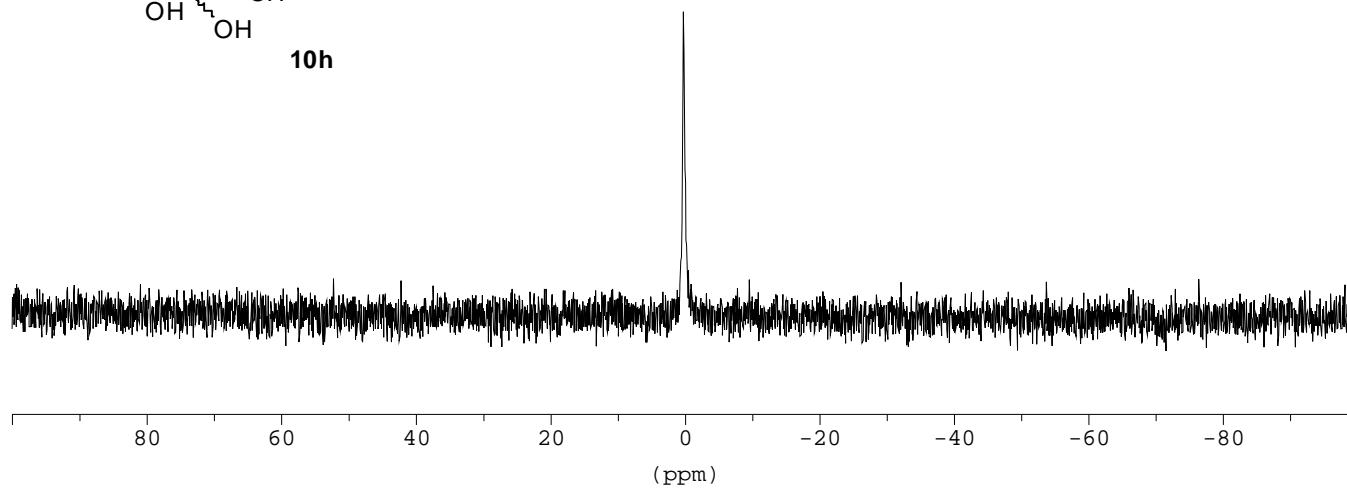
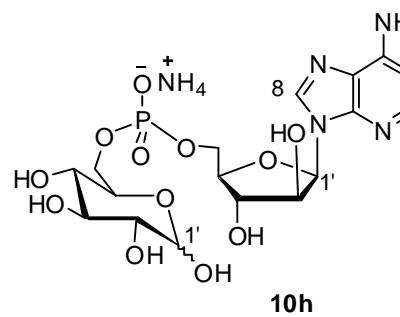


1'-Adeninyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10h)

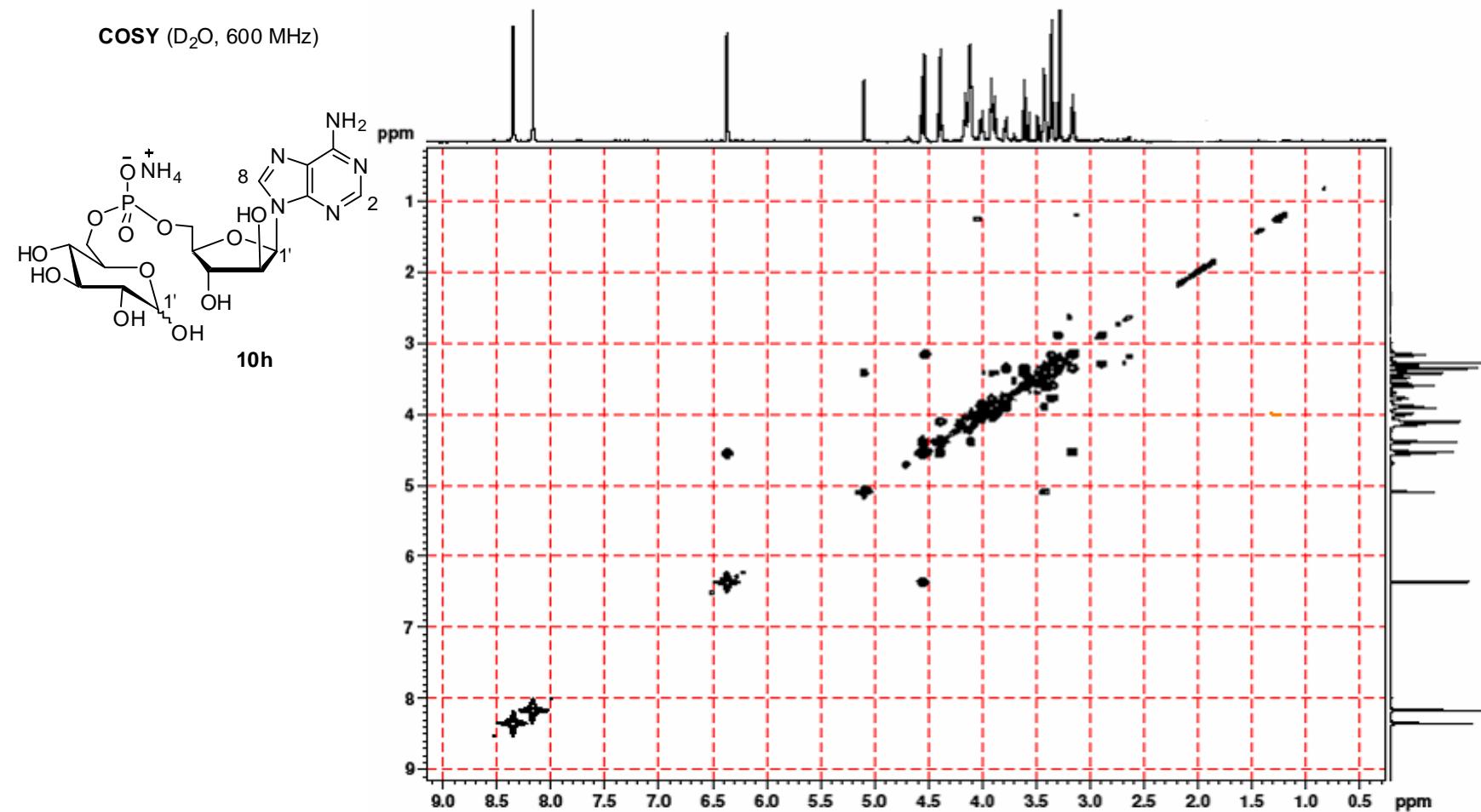


1'-Adeninyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10h)

$^{31}\text{P-RMN}$ (D_2O , 242.9 MHz)

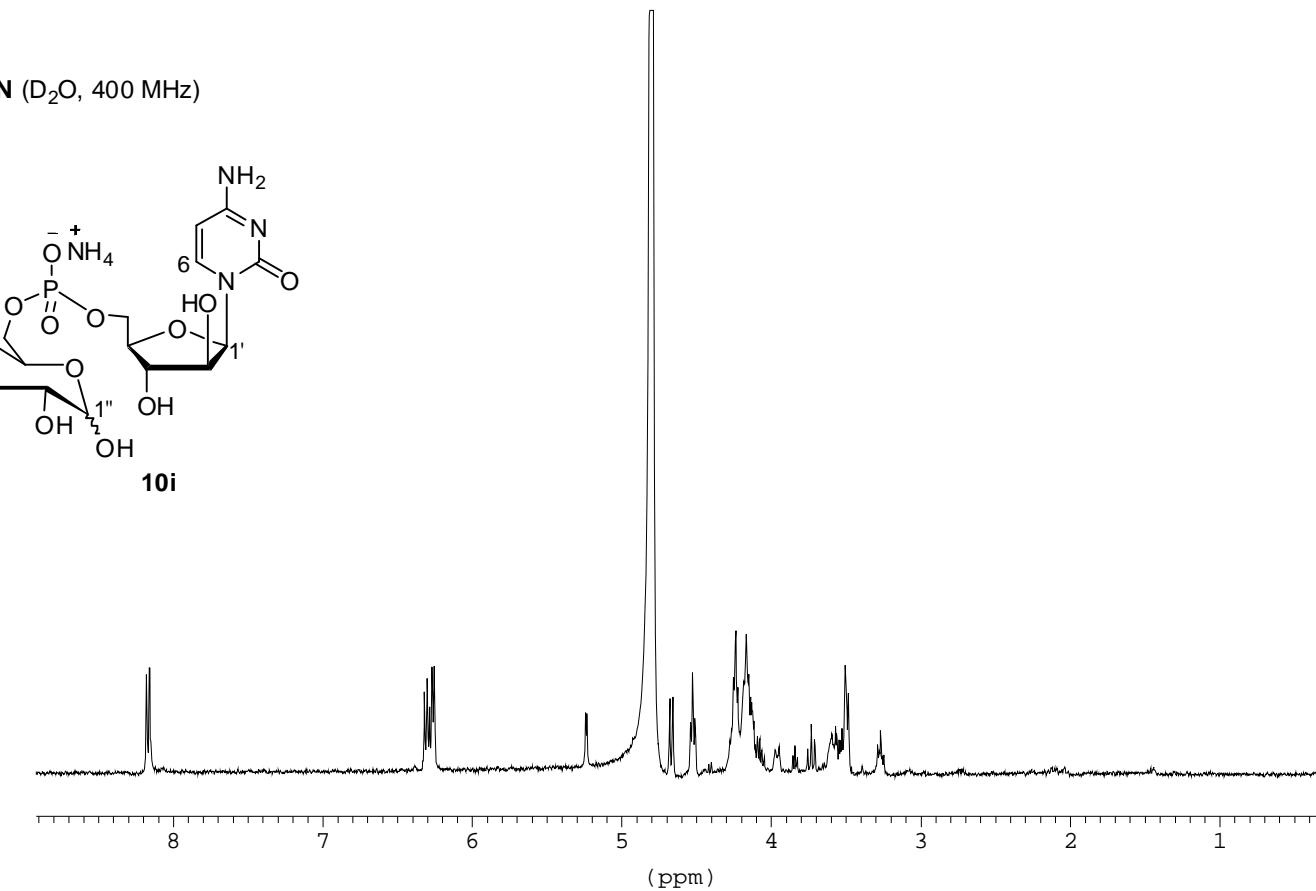
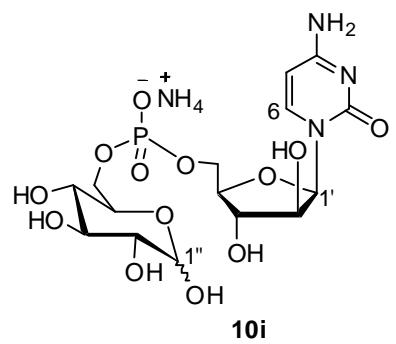


1'-Adeninyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10h)



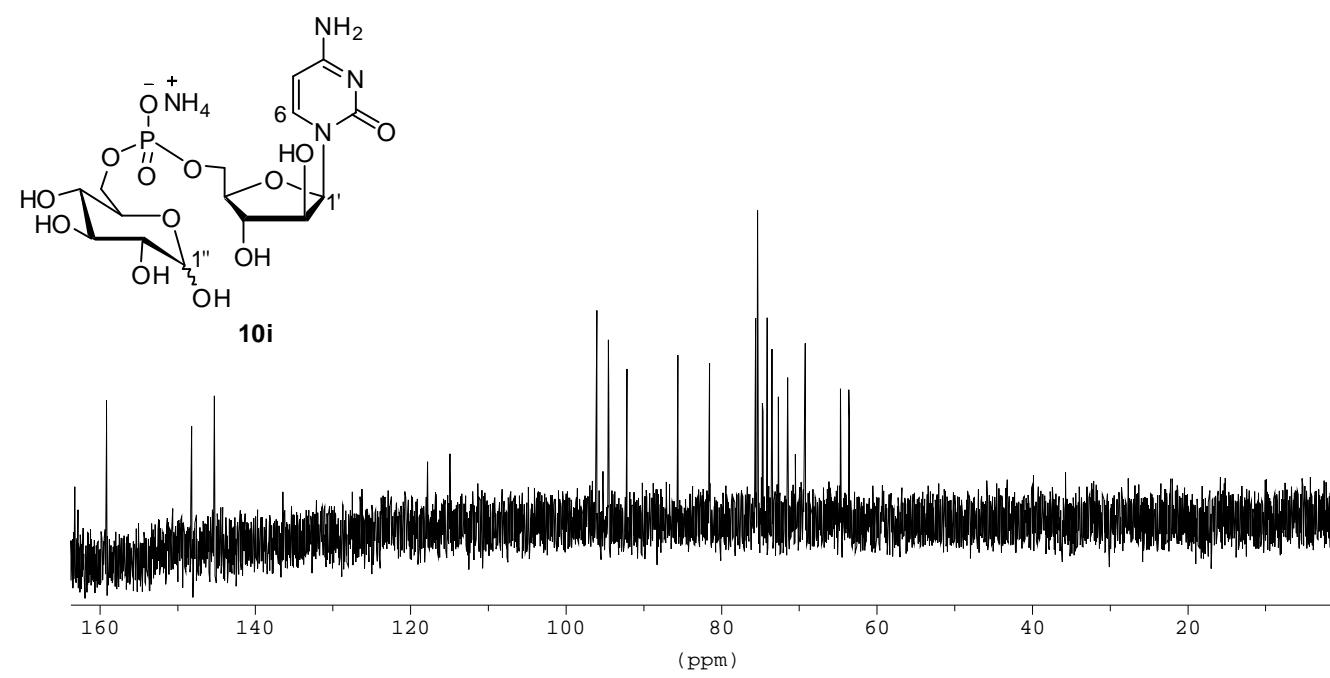
1'-Cytosinyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10i)

$^1\text{H-RMN}$ (D_2O , 400 MHz)

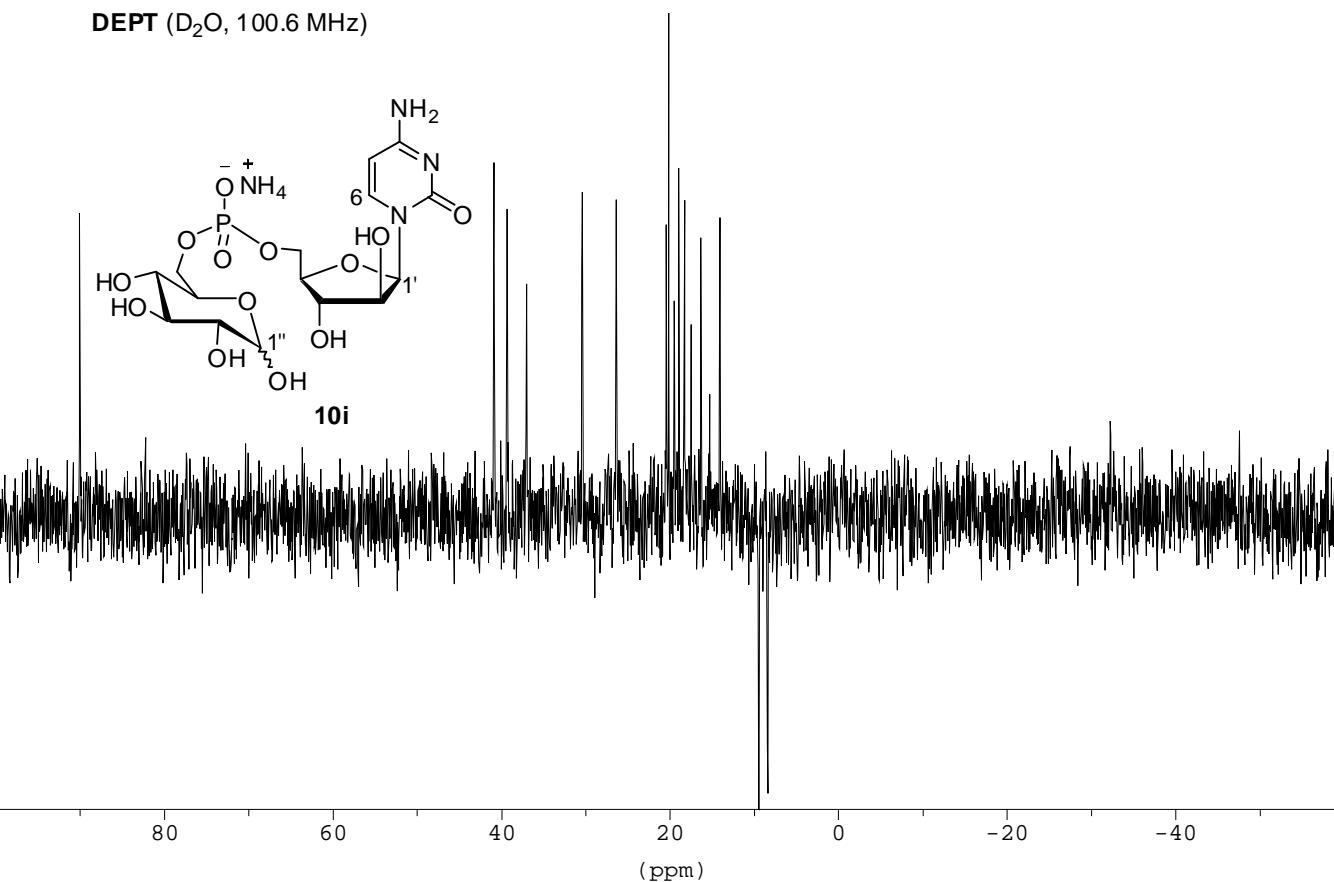


1'-Cytosinyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10i)

^{13}C -RMN (D_2O , 100.6 MHz)

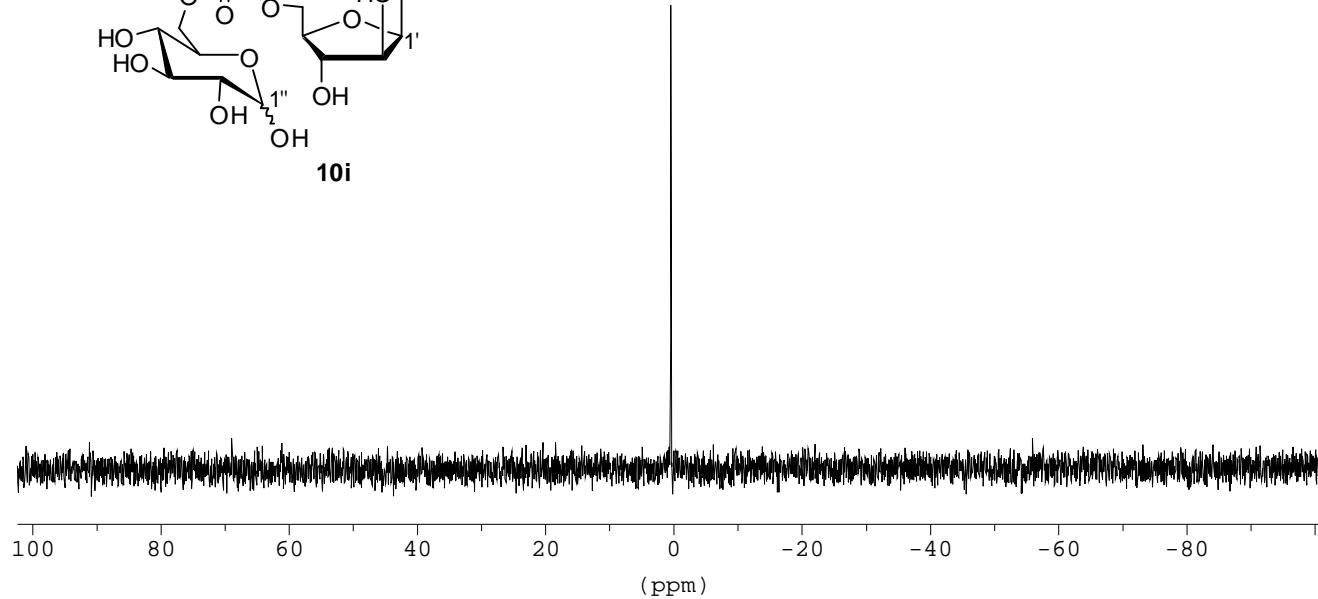
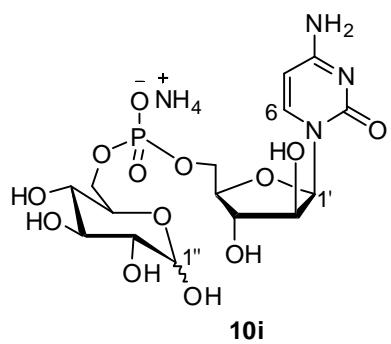


1'-Cytosinyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10i)



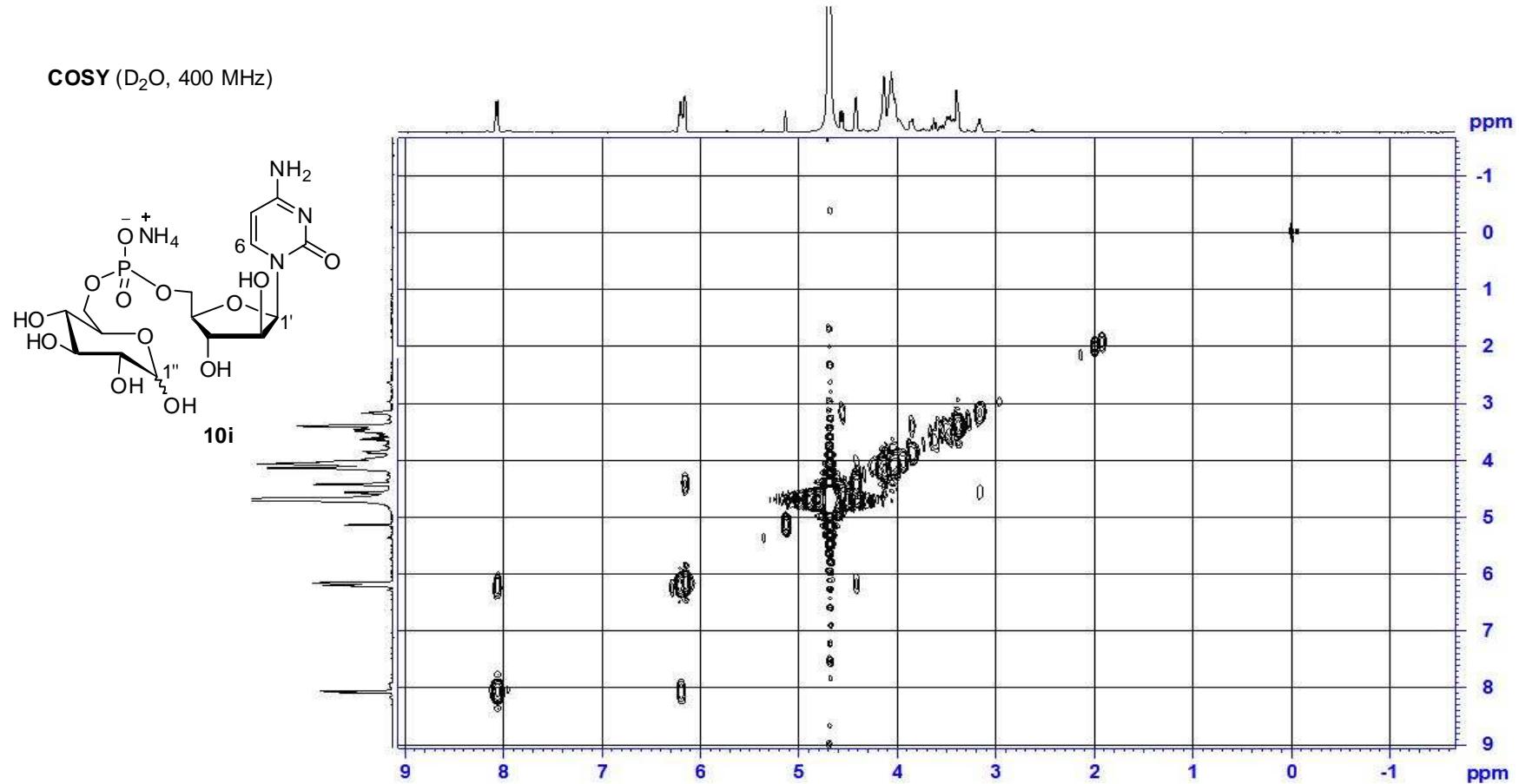
1'-Cytosinyl- β -D-arabinofuranose 5'-(α / β -D-glucopyranos-6-yl) phosphate (10i)

^{31}P -RMN (D_2O , 161.9 MHz)



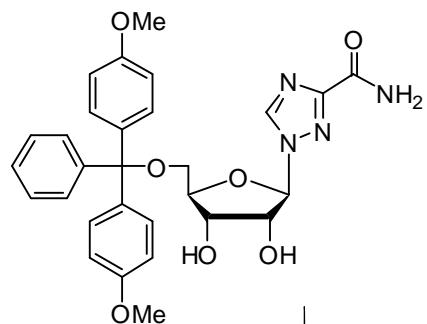
1'-Cytosinyl- β -D-arabinofuranose 5'-(α/β -D-glucopyranos-6-yl) phosphate (10i)

COSY (D_2O , 400 MHz)

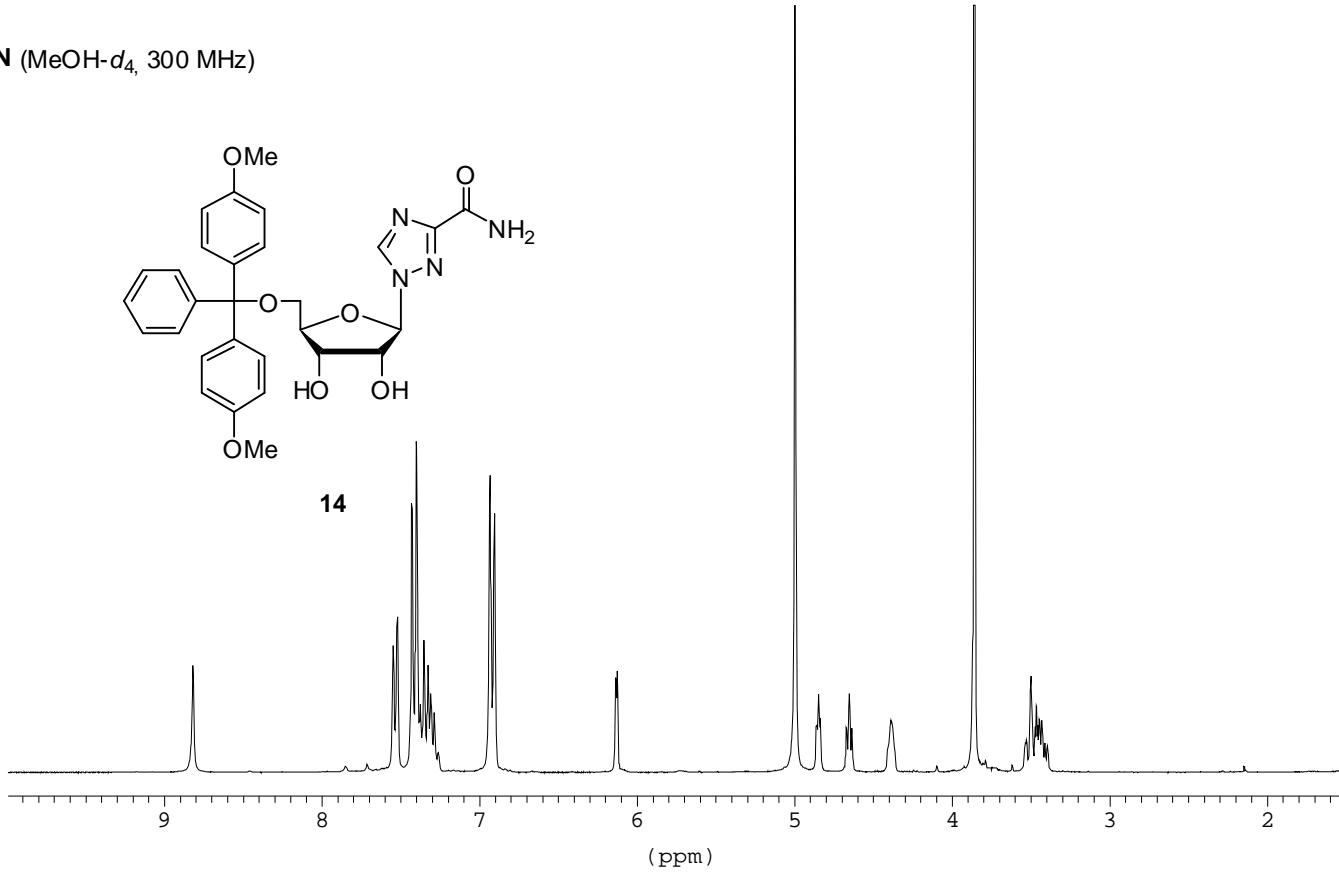


1-(5'-O-dimethoxytrityl- β -D-ribofuranosyl)-1,2,4-triazole-3-carboxamide (14)

$^1\text{H-RMN}$ ($\text{MeOH-}d_4$, 300 MHz)

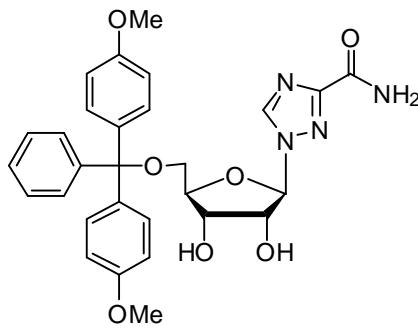


14

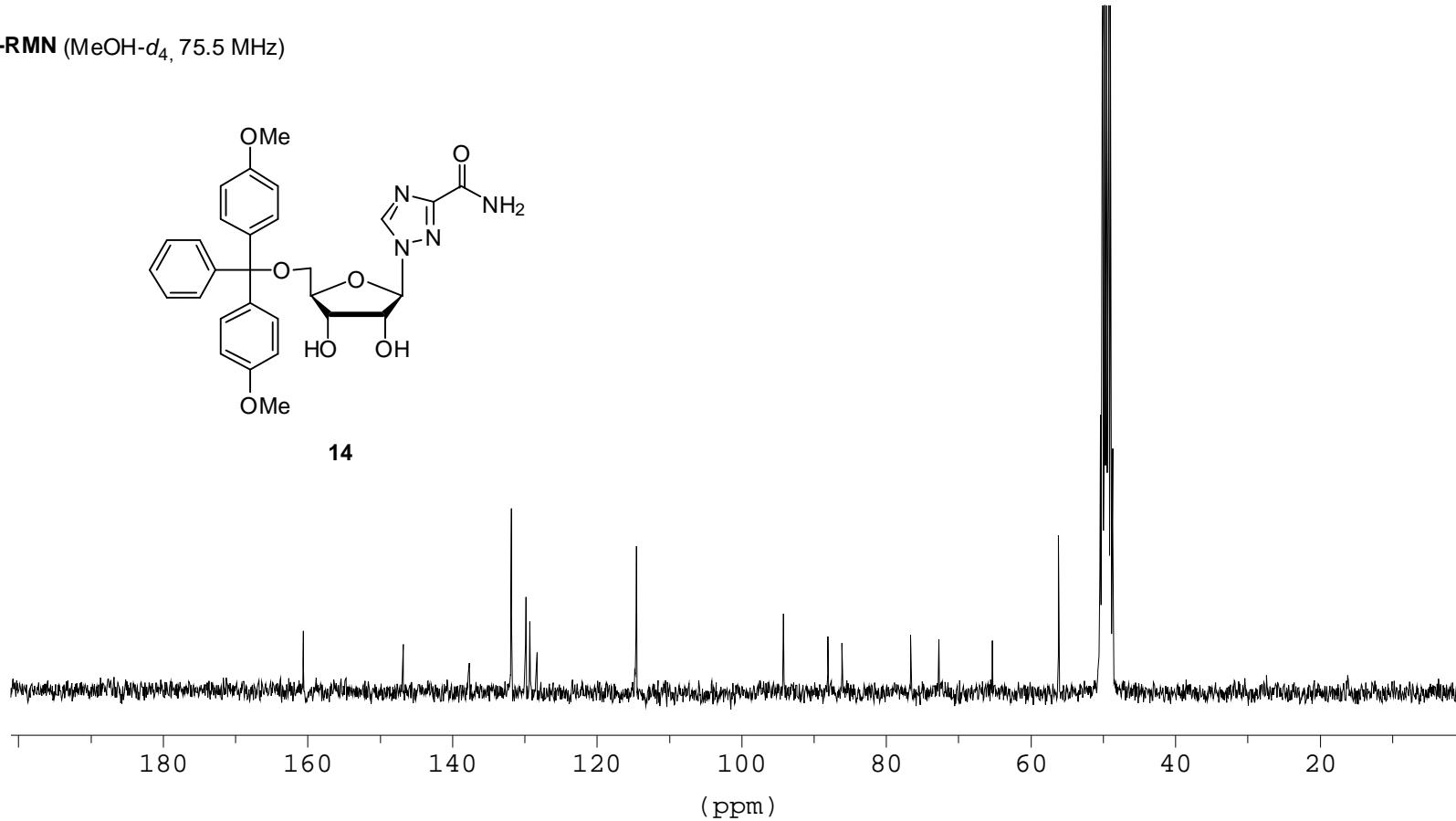


1-(5'-O-dimethoxytrityl- β -D-ribofuranosyl)-1,2,4-triazole-3-carboxamide (14)

^{13}C -RMN (MeOH- d_4 , 75.5 MHz)

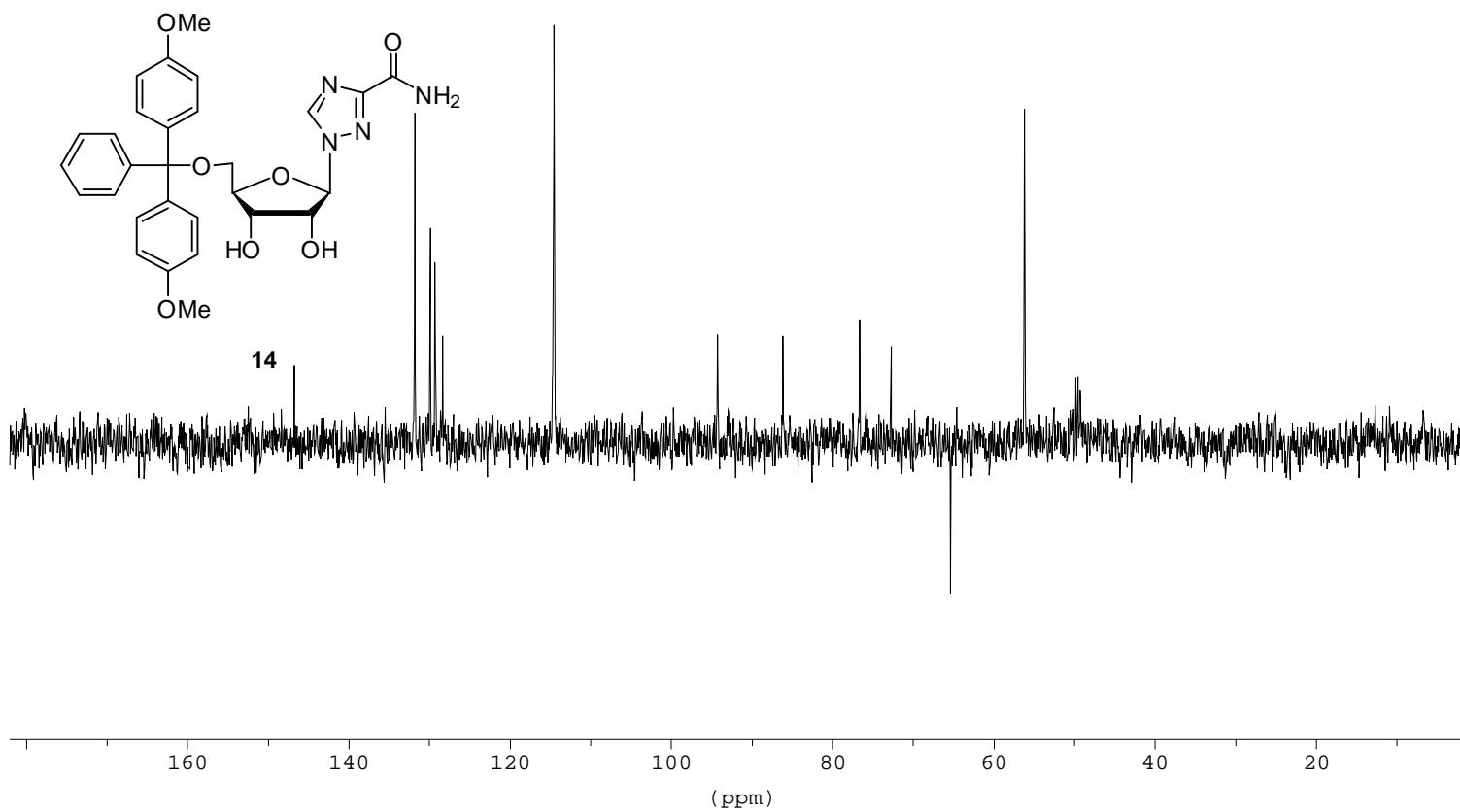


14



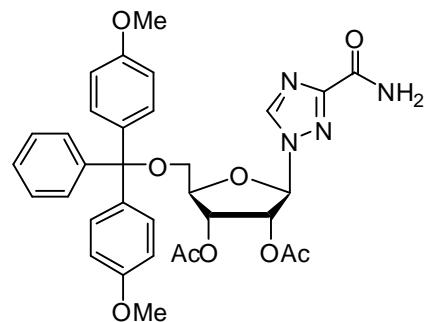
1-(5'-O-dimethoxytrityl- β -D-ribofuranosyl)-1,2,4-triazole-3-carboxamide (14)

DEPT (MeOH- d_4 , 75.5 MHz)

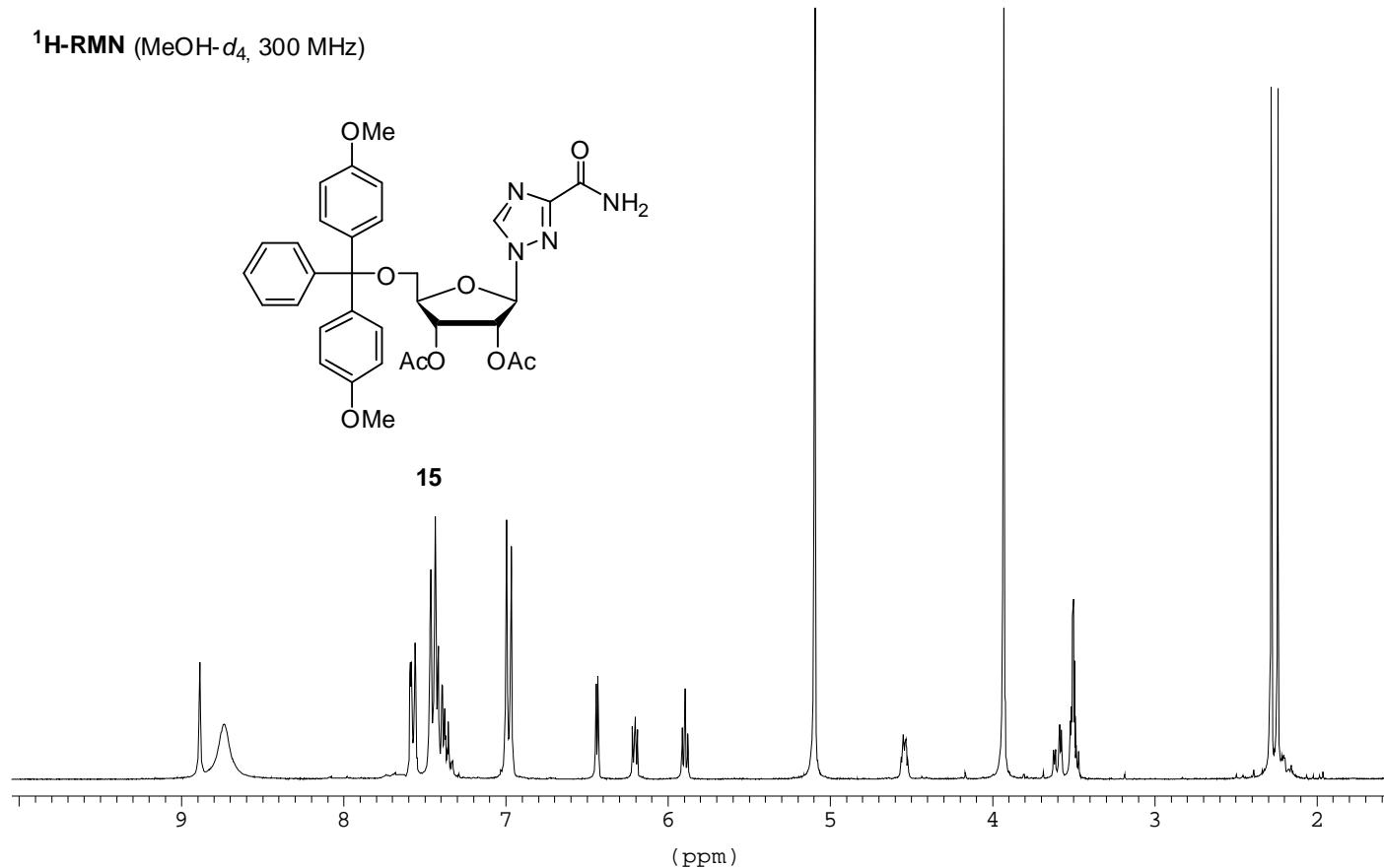


1-(2',3'-di-*O*-acetyl-5'-*O*-dimethoxytrityl- β -D-ribofuranosyl)-1,2,4-triazole-3-carboxamide (15).

¹H-RMN (MeOH-*d*₄, 300 MHz)

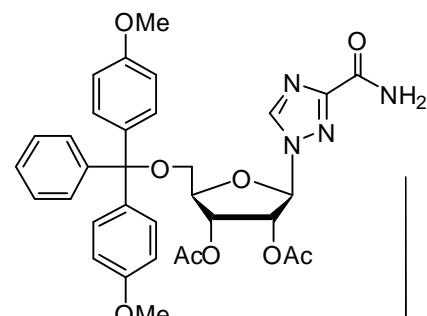


15

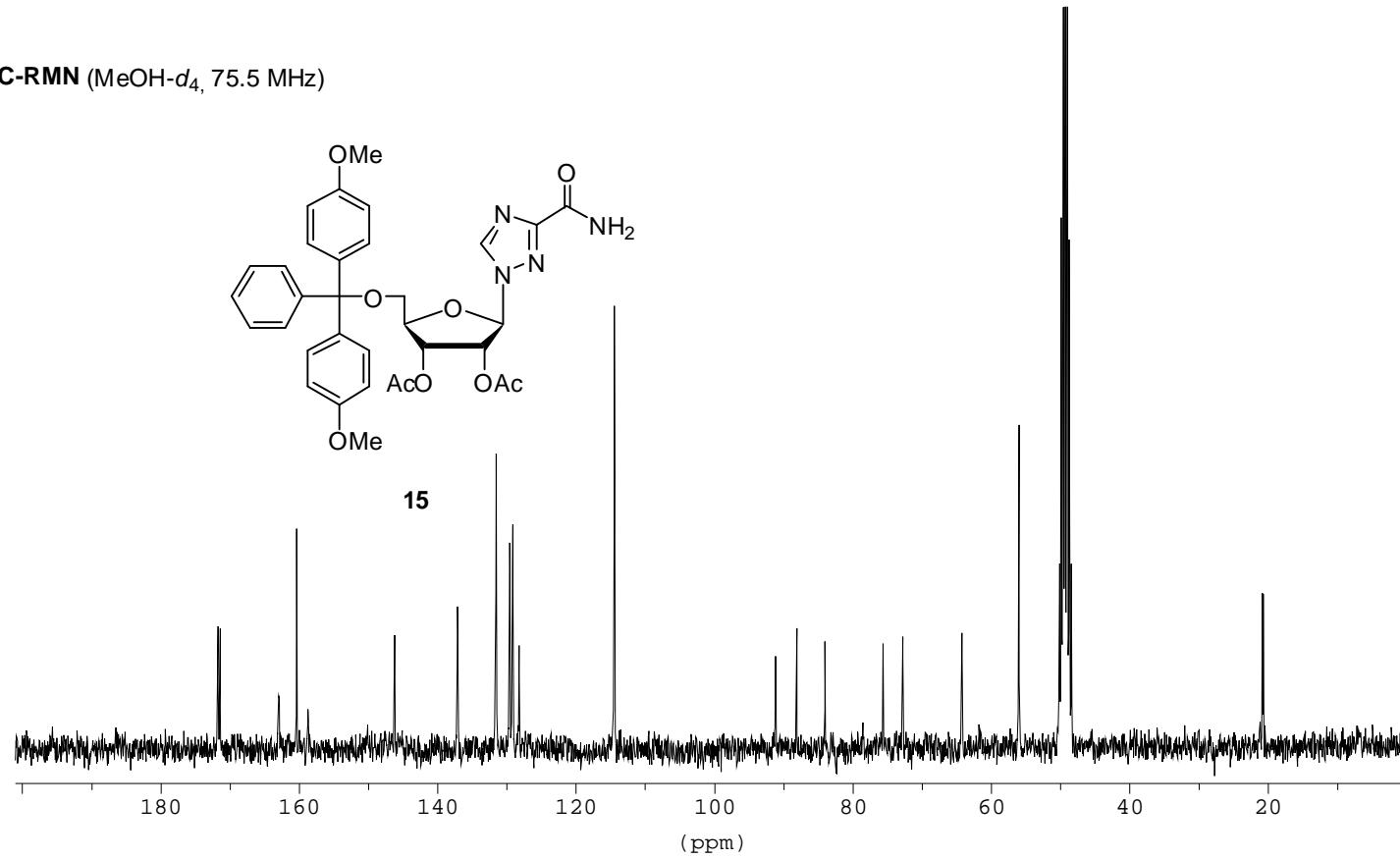


1-(2',3'-di-*O*-acetyl-5'-*O*-dimethoxytrityl- β -D-ribofuranosyl)-1,2,4-triazole-3-carboxamide (15).

¹³C-RMN (MeOH-*d*₄, 75.5 MHz)

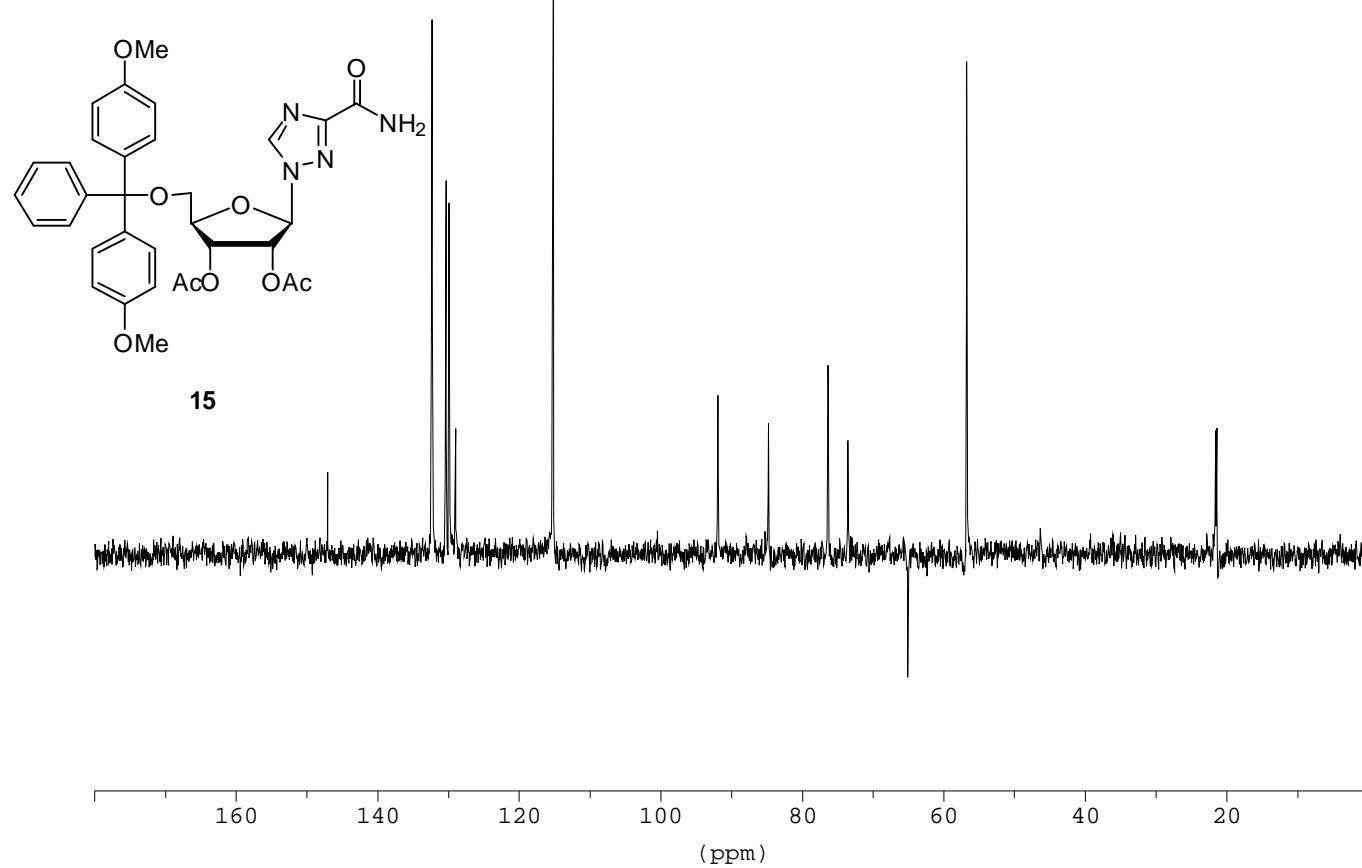


15



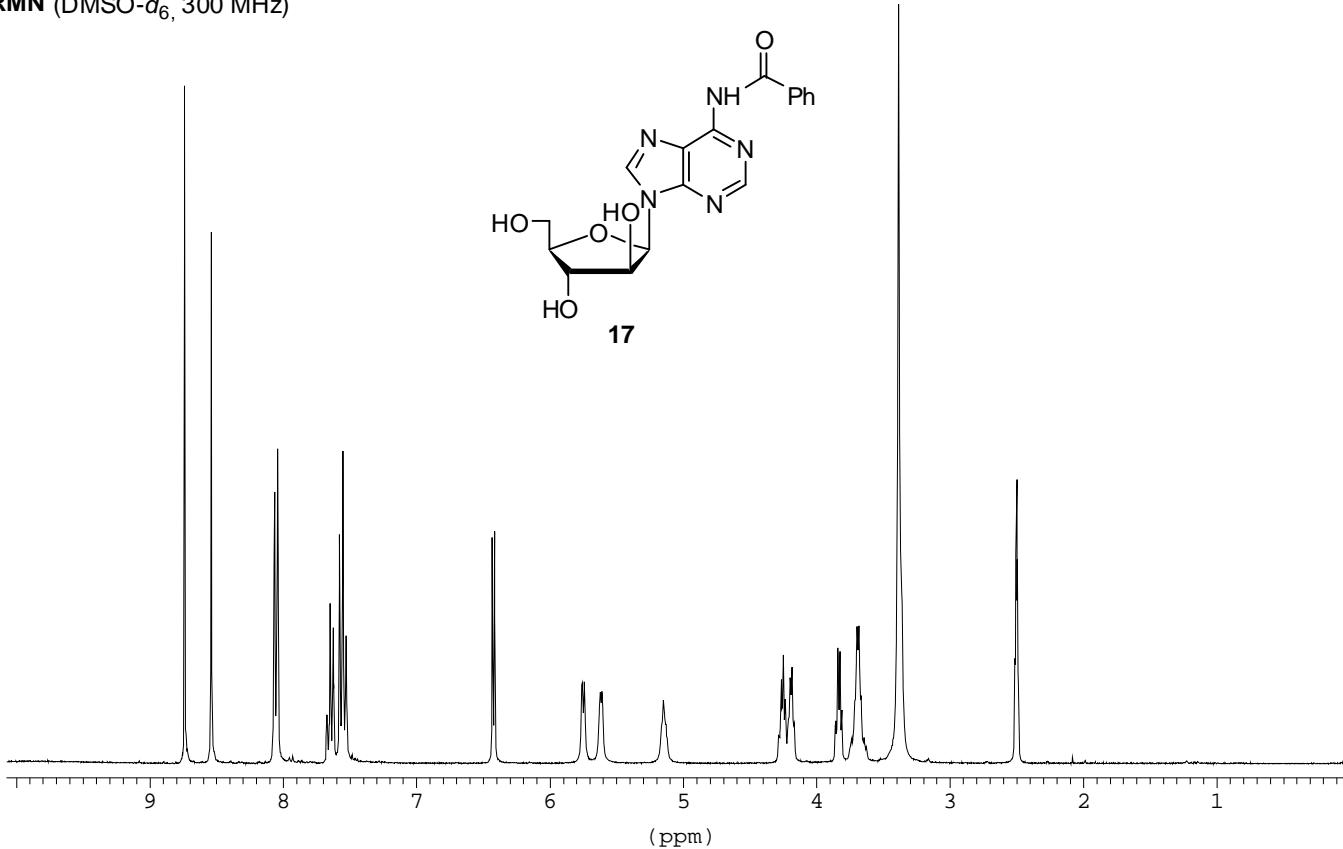
1-(2',3'-di-*O*-acetyl-5'-*O*-dimethoxytrityl- β -D-ribofuranosyl)-1,2,4-triazole-3-carboxamide (15).

DEPT (MeOH-*d*₄, 75.5 MHz)



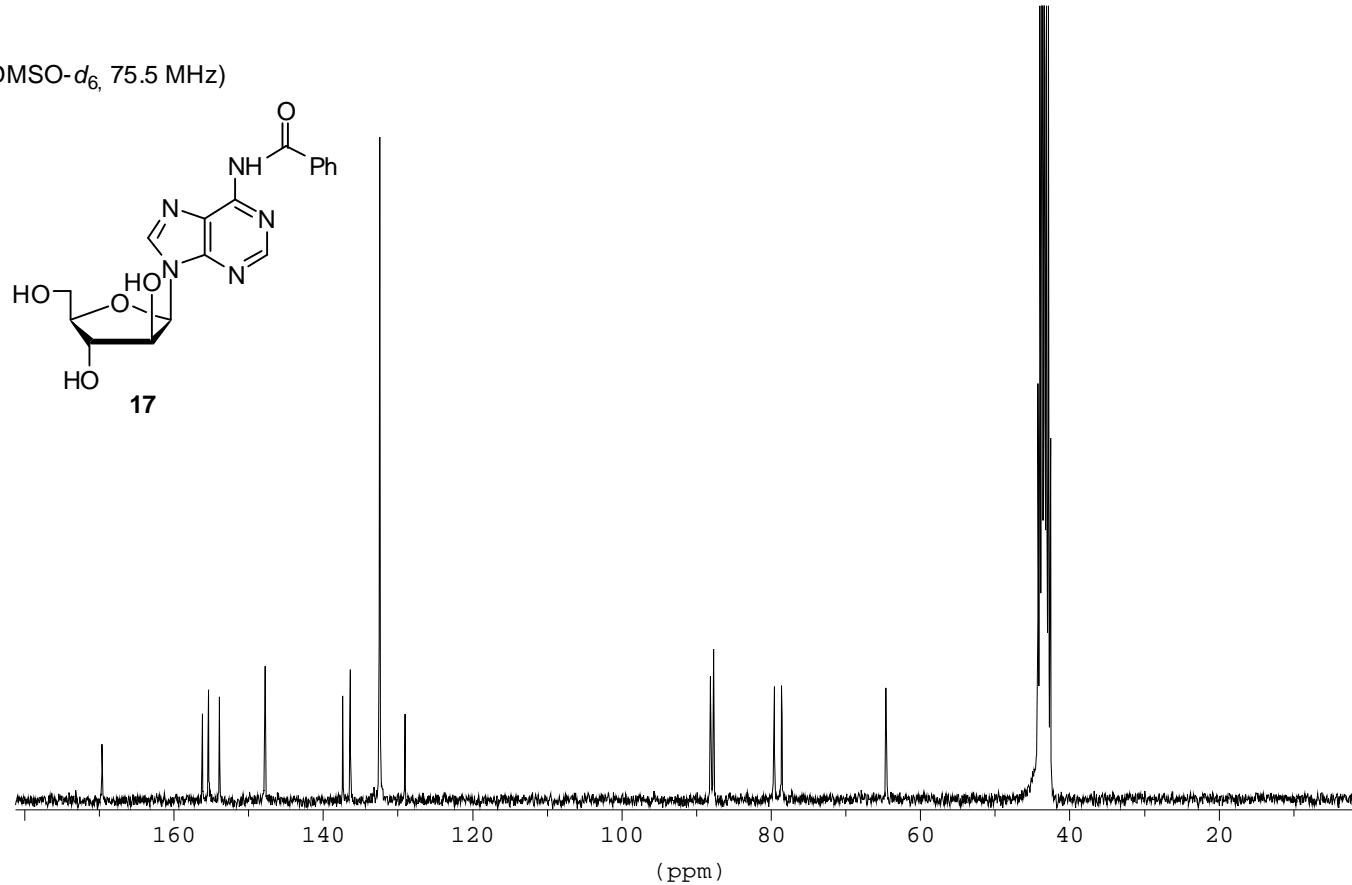
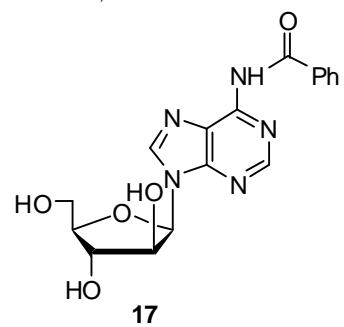
9-(β -D-arabinofuranosyl)- N^6 -benzoyladenine (17)

$^1\text{H-RMN}$ (DMSO- d_6 , 300 MHz)



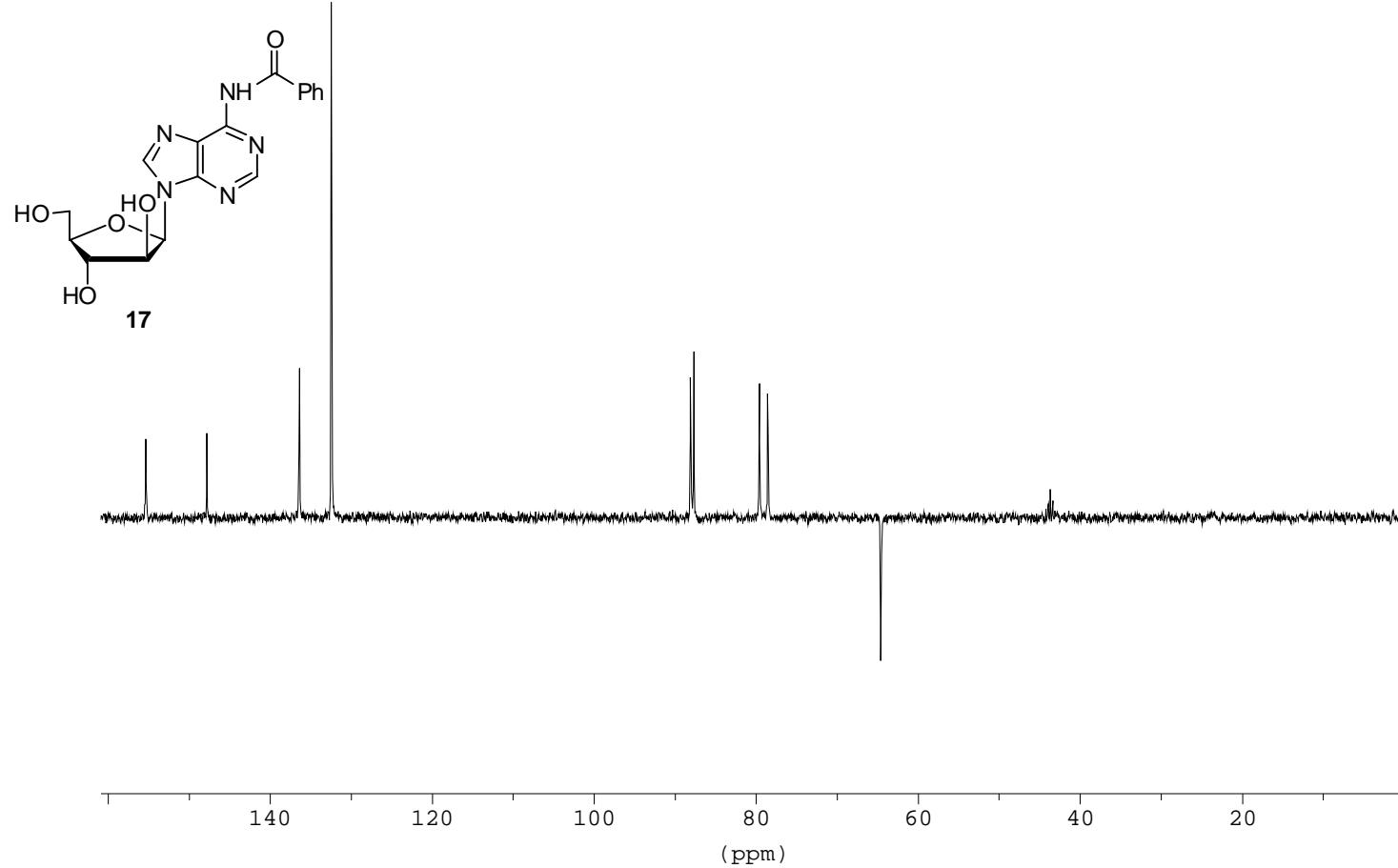
9-(β -D-arabinofuranosyl)- N^6 -benzoyladenine (17)

^{13}C -RMN (DMSO- d_6 , 75.5 MHz)



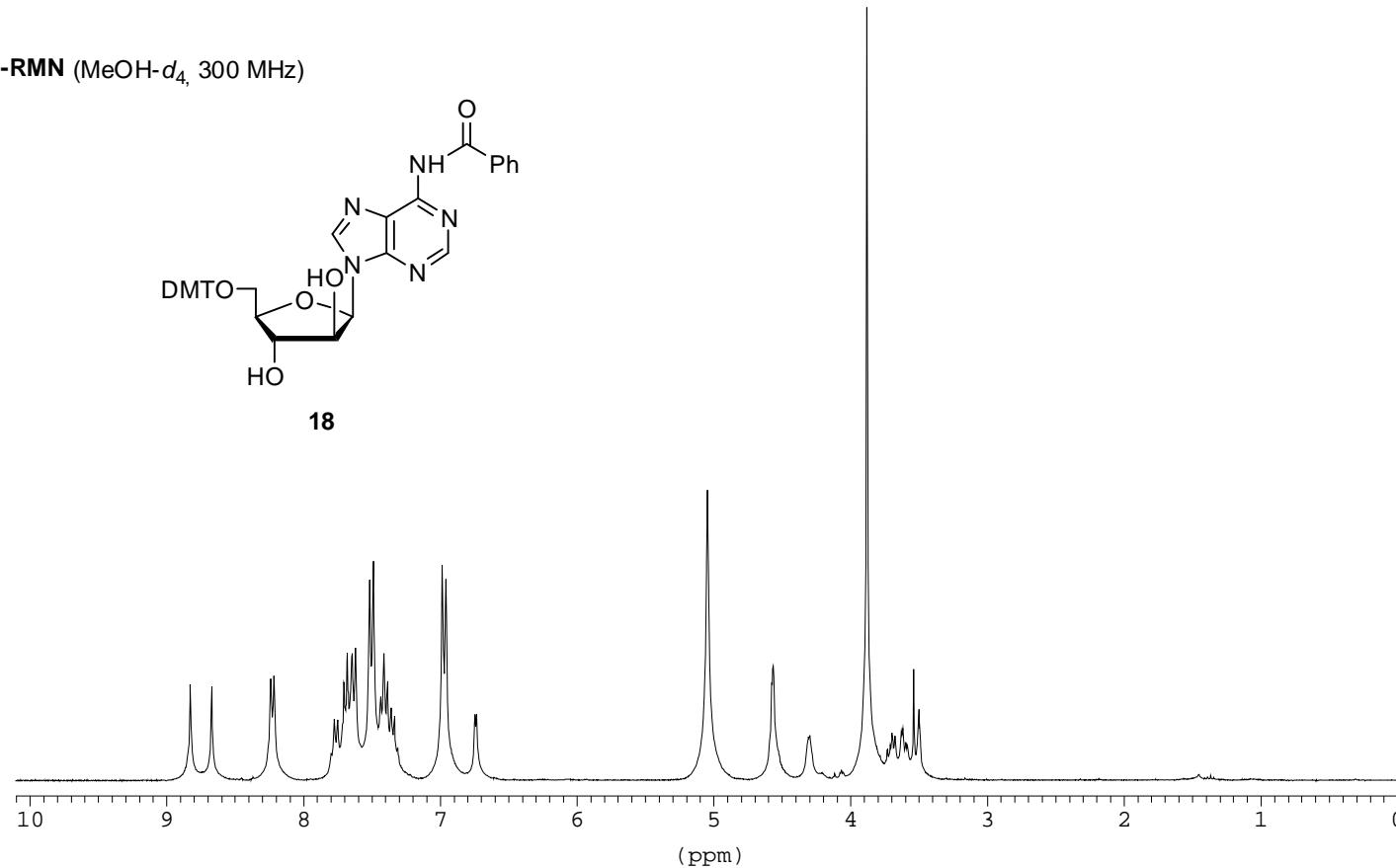
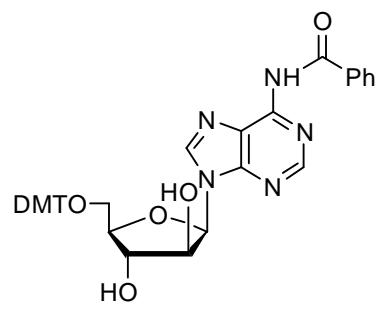
9-(β -D-arabinofuranosyl)- N^6 -benzoyladenine (17)

DEPT(DMSO- d_6 , 75.5 MHz)

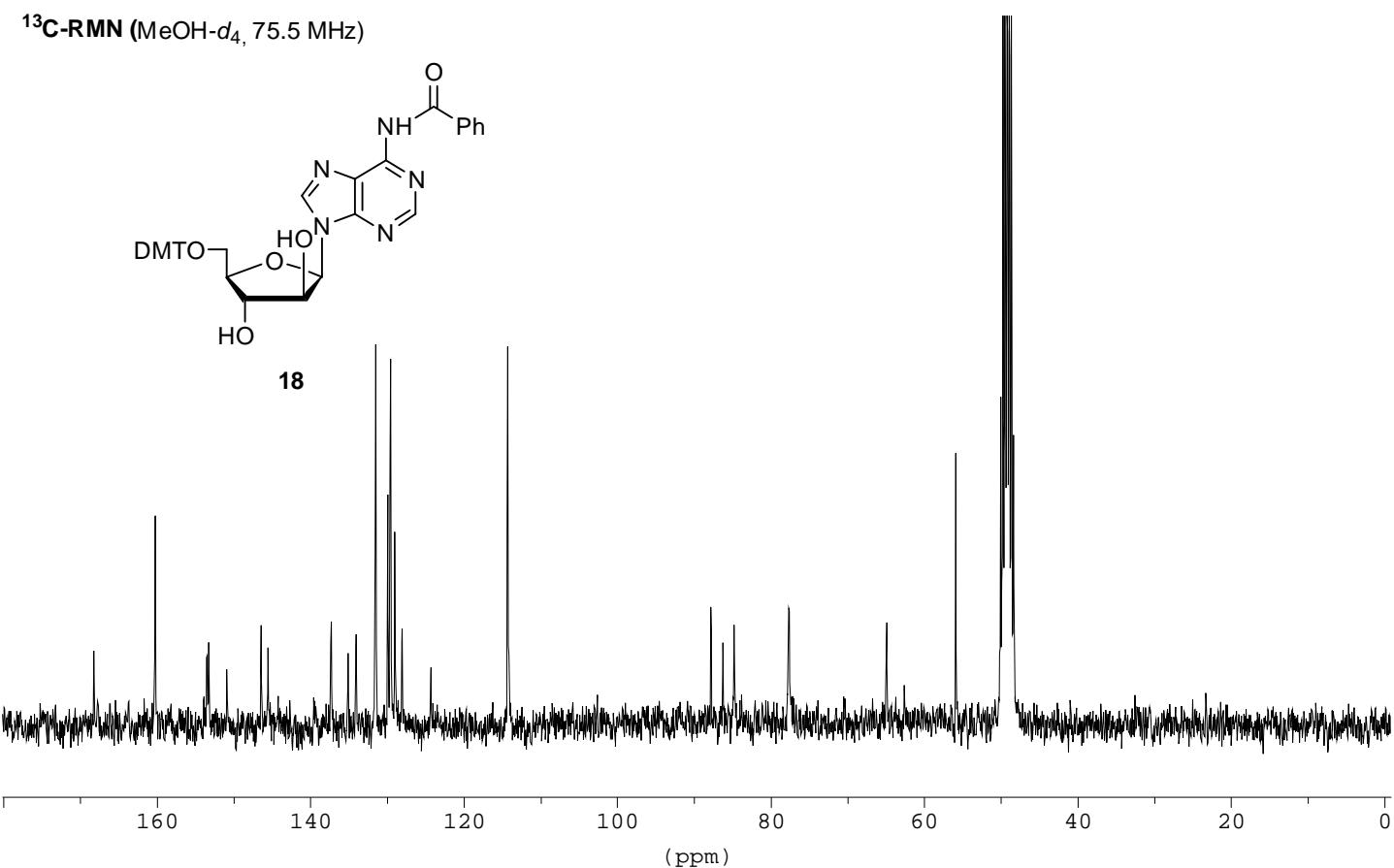


Synthesis of *N*⁶-benzoyl-9-(5'-*O*-dimethoxytrityl- β -D-arabinofuranosyl)adenine (**18**)

¹H-RMN (MeOH-*d*₄, 300 MHz)

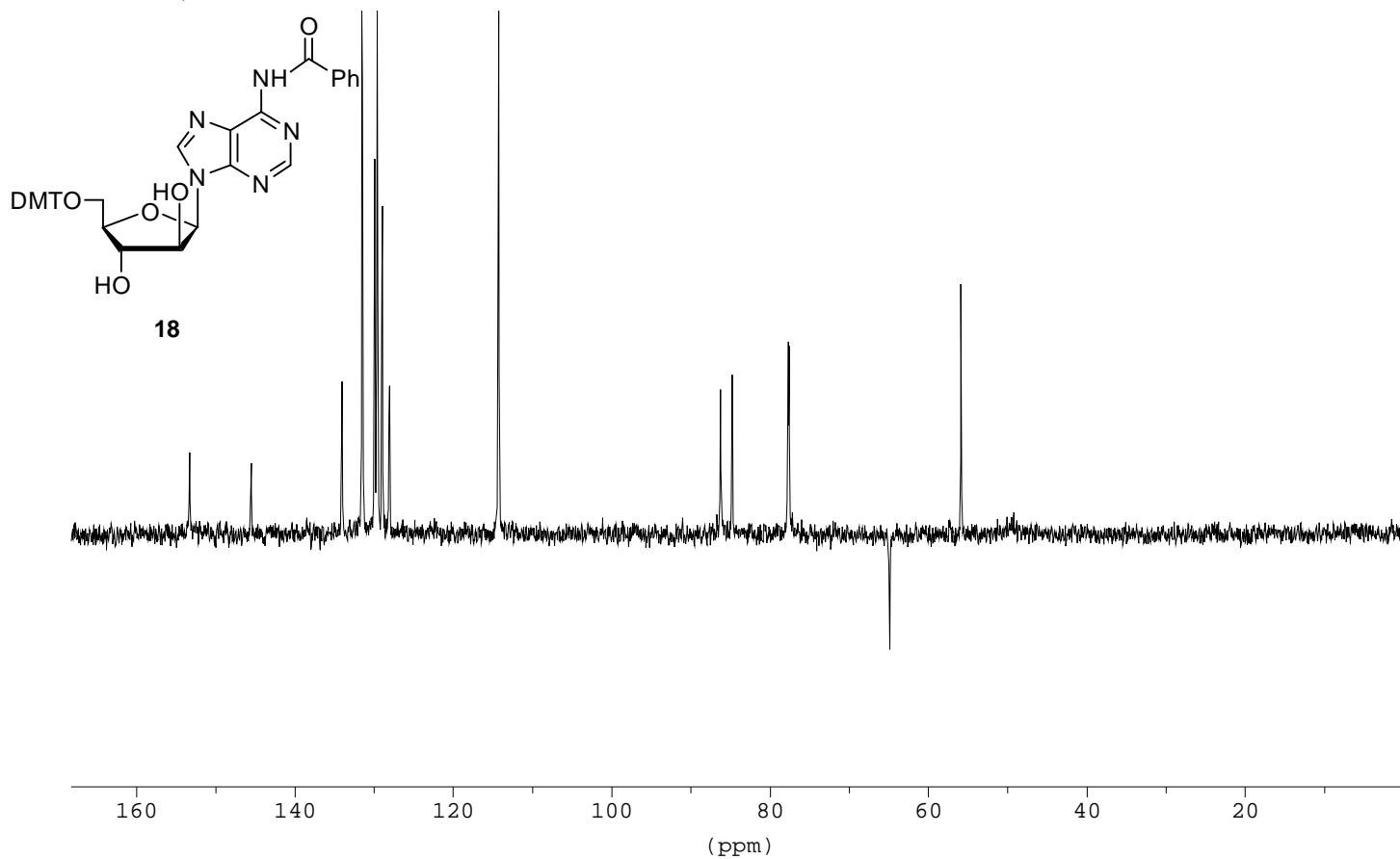


Synthesis of *N*⁶-benzoyl-9-(5'-*O*-dimethoxytrityl- β -D-arabinofuranosyl)adenine (**18**)

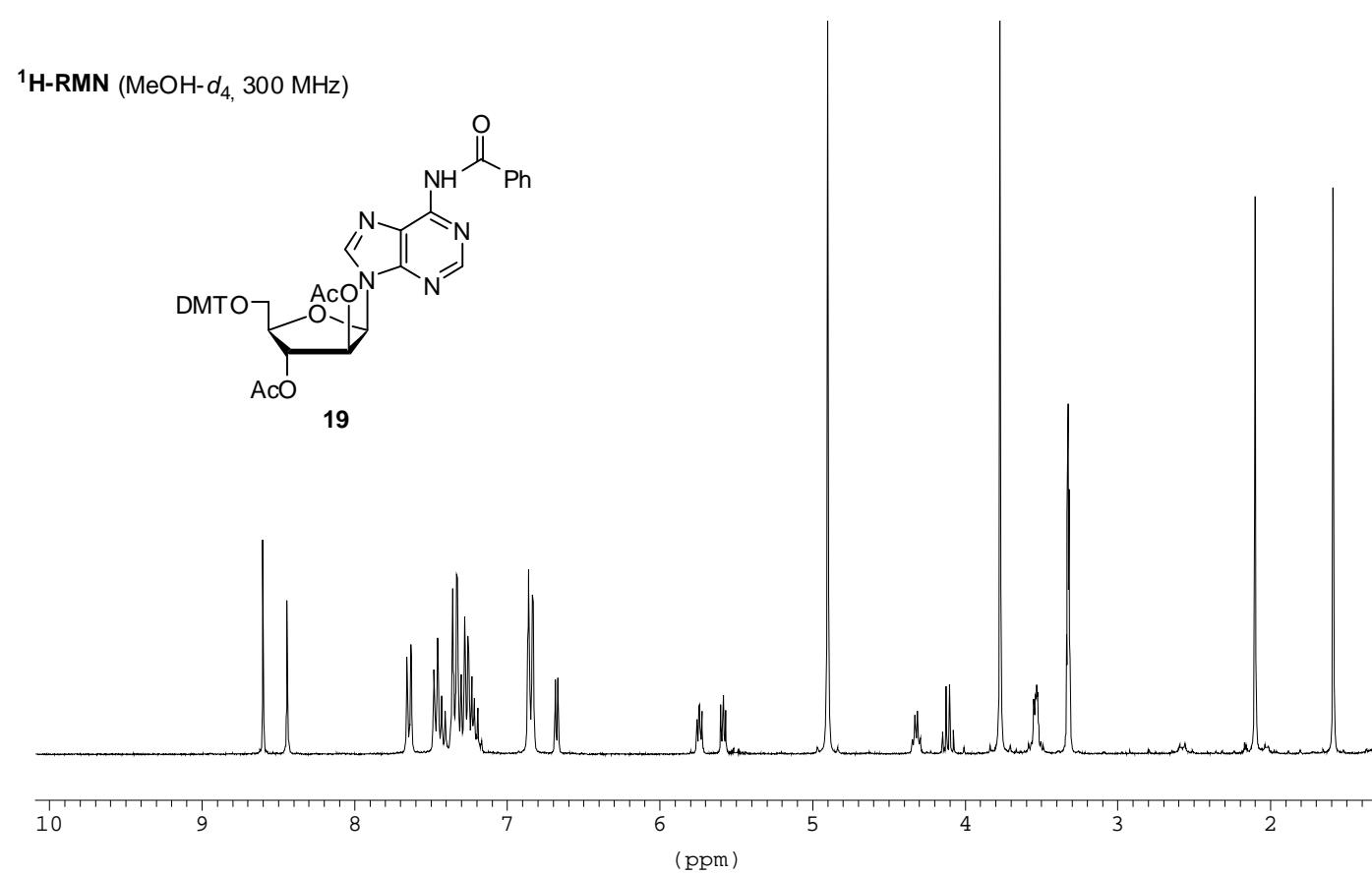


Synthesis of *N*⁶-benzoyl-9-(5'-*O*-dimethoxytrityl- β -D-arabinofuranosyl)adenine (18)

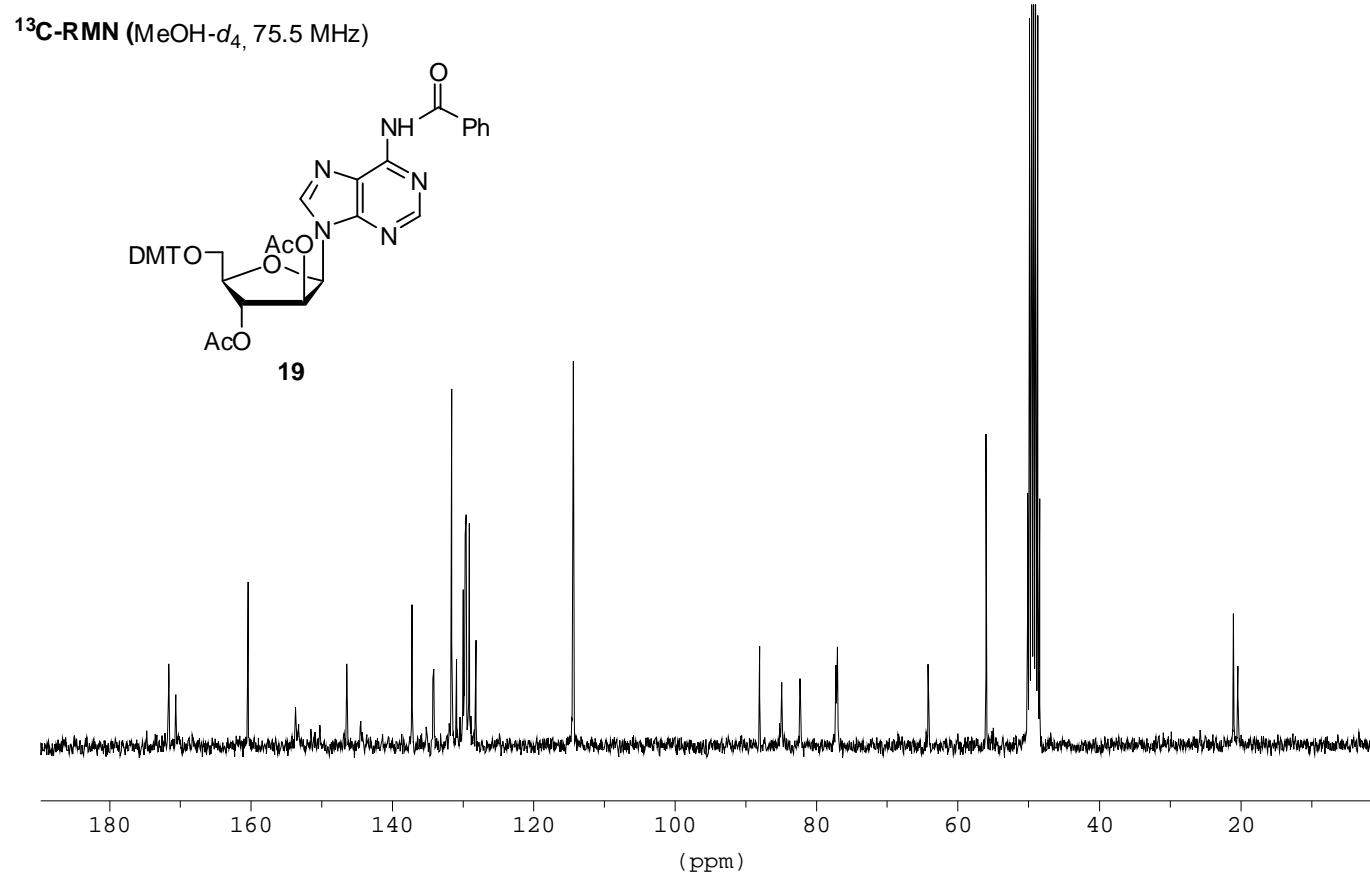
DEPT (MeOH-*d*₄, 75.5 MHz)



*N*⁶-benzoyl-9-(2',3'-di-*O*-acetyl-5'-*O*-dimethoxytrityl- β -D-arabinofuranosyl)adenine (19)

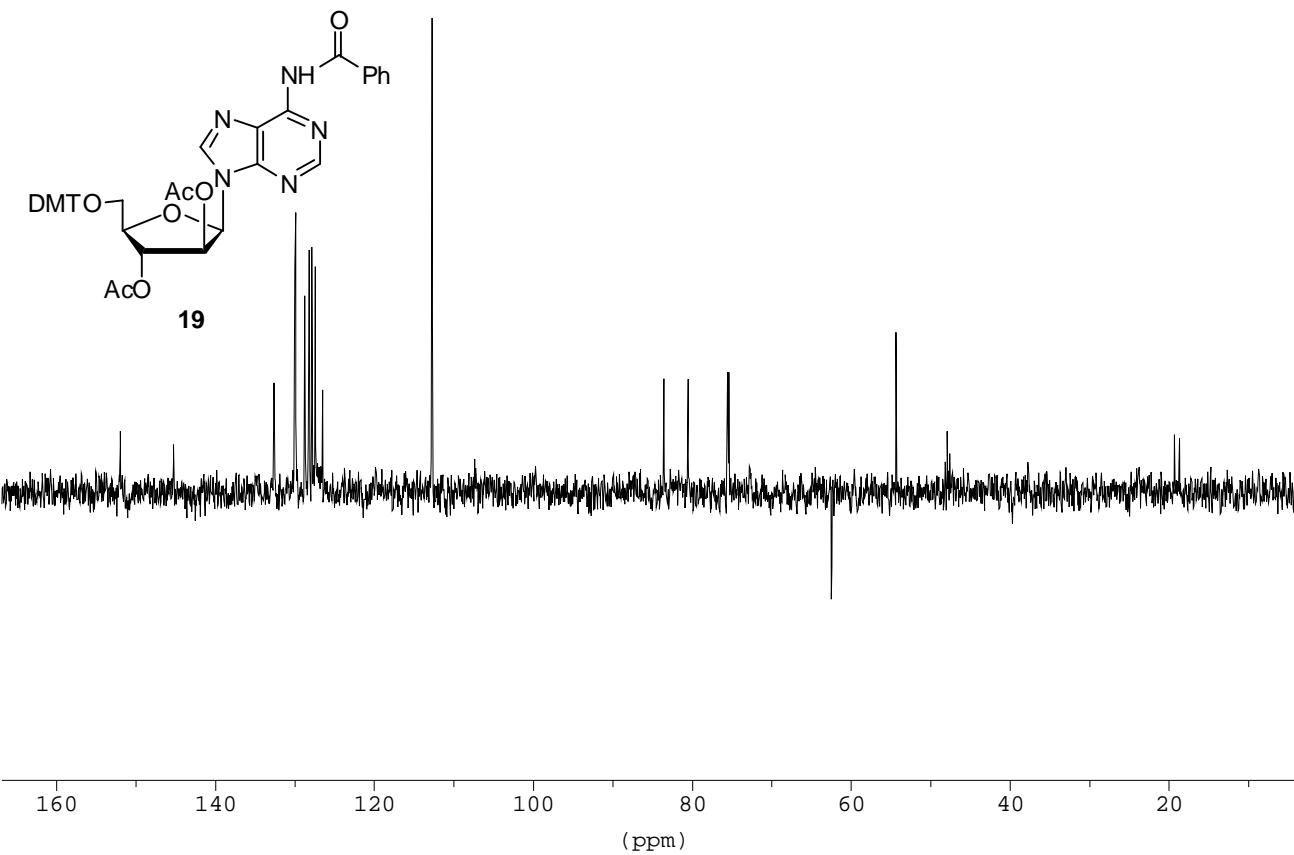


*N*⁶-benzoyl-9-(2',3'-di-*O*-acetyl-5'-*O*-dimethoxytrityl- β -D-arabinofuranosyl)adenine (19)



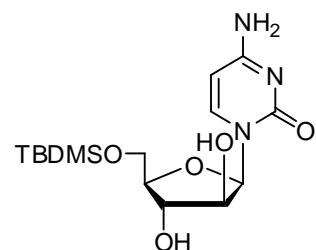
*N*⁶-benzoyl-9-(2',3'-di-*O*-acetyl-5'-*O*-dimethoxytrityl- β -D-arabinofuranosyl)adenine (19)

DEPT (MeOH-*d*₄, 75.5 MHz)

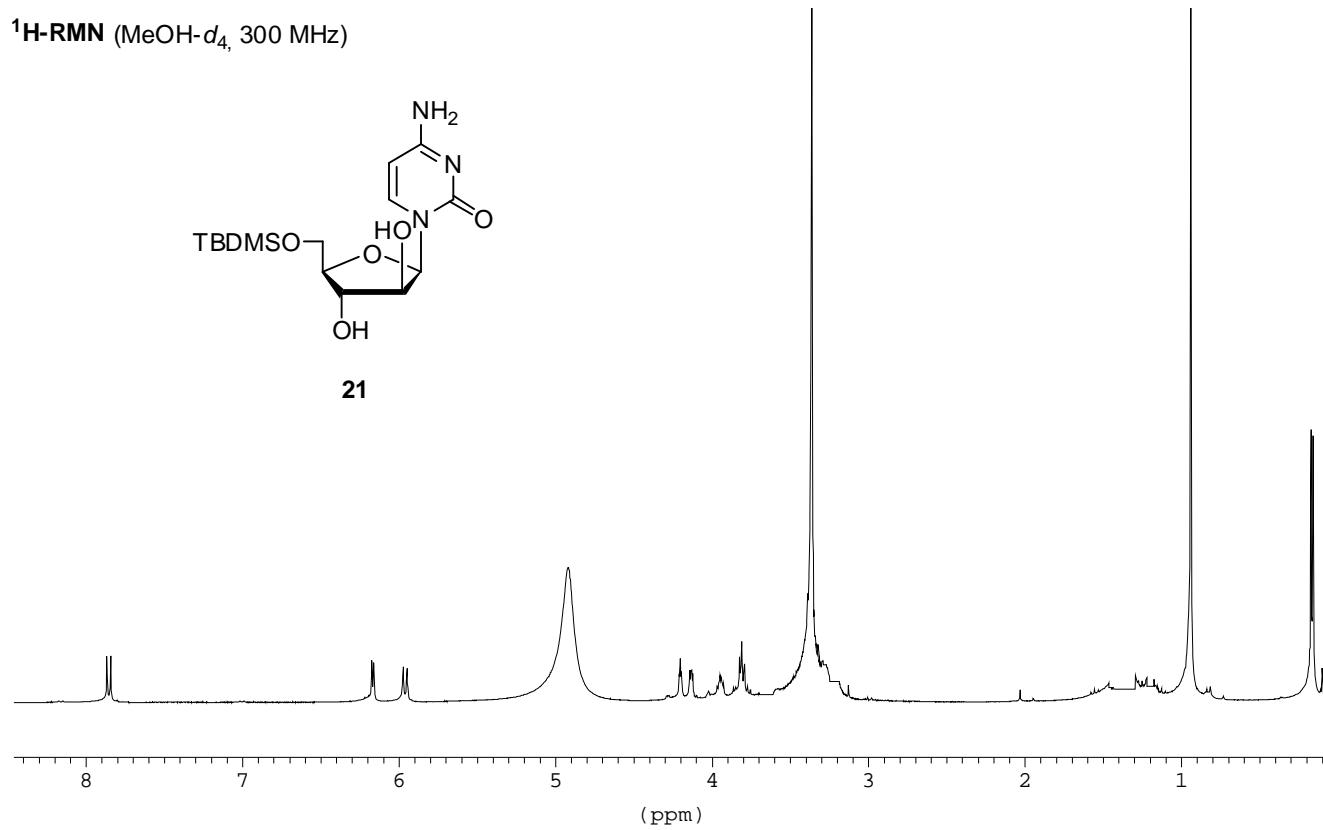


1-(5'-*O*-*tert*-butyldimethylsilyl- β -D-arabinofuranosyl)cytosine (21)

¹H-RMN (MeOH-*d*₄, 300 MHz)

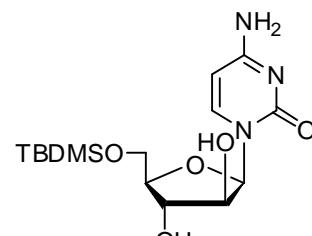


21

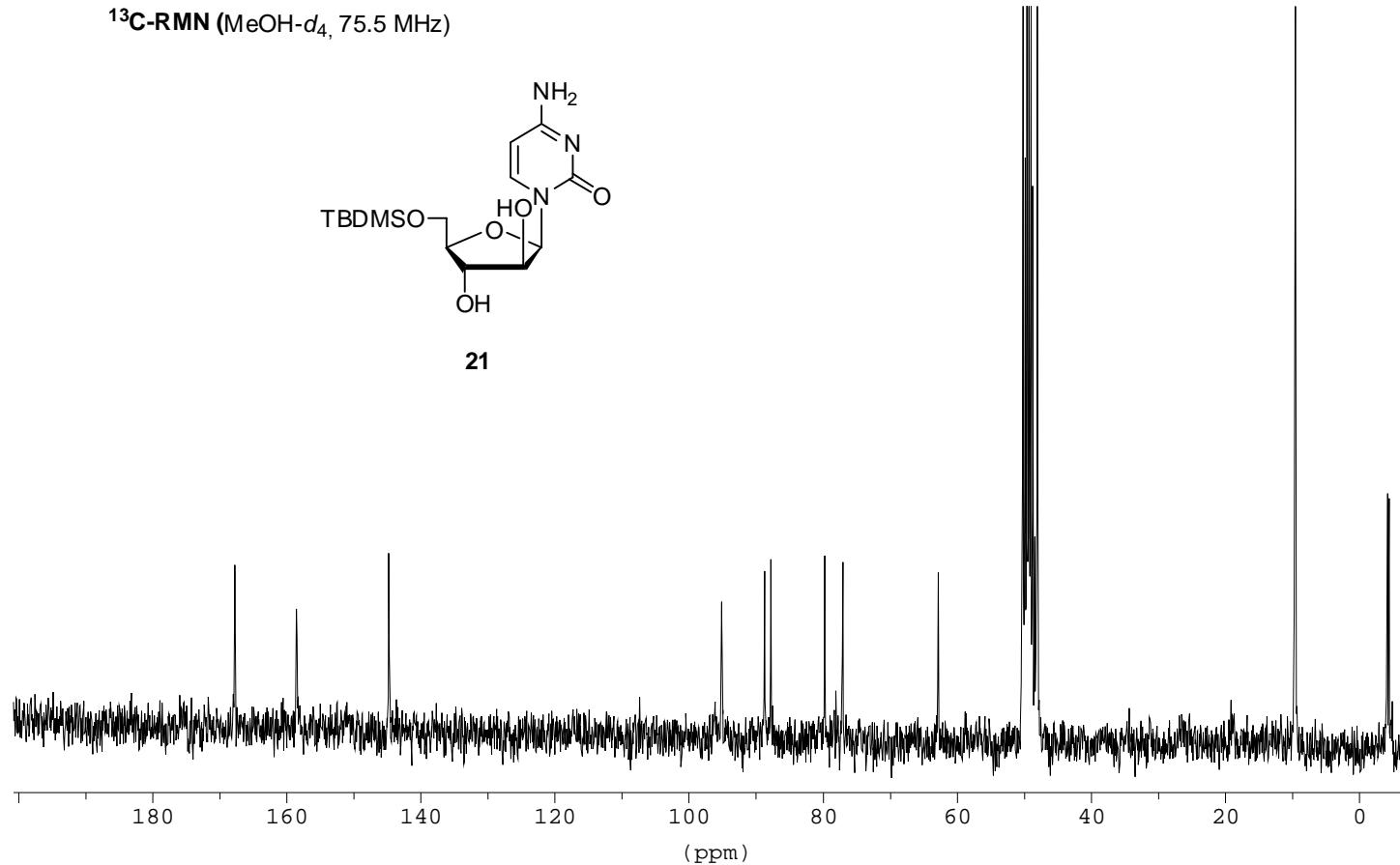


1-(5'-*O*-*tert*-butyldimethylsilyl- β -D-arabinofuranosyl)cytosine (21)

^{13}C -RMN (MeOH- d_4 , 75.5 MHz)

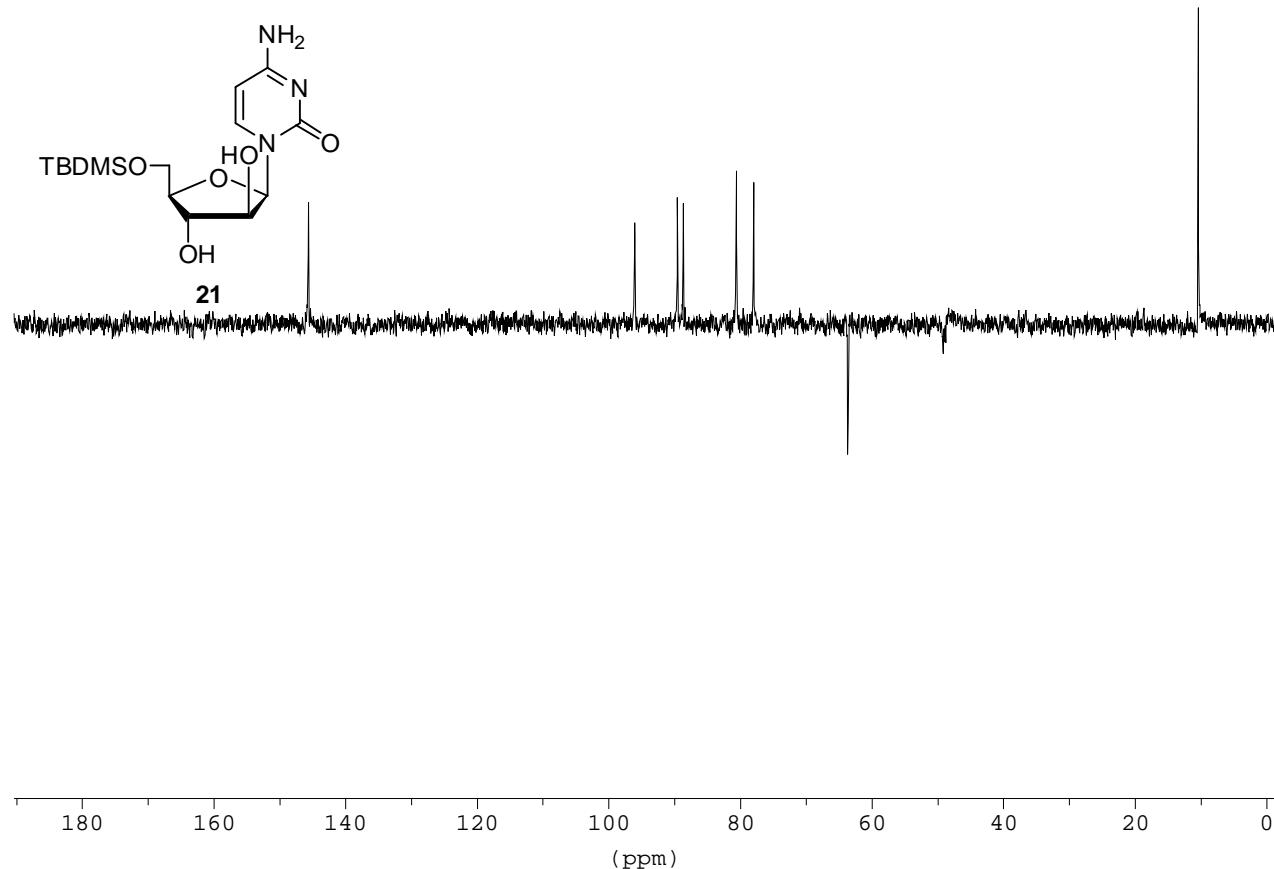


21



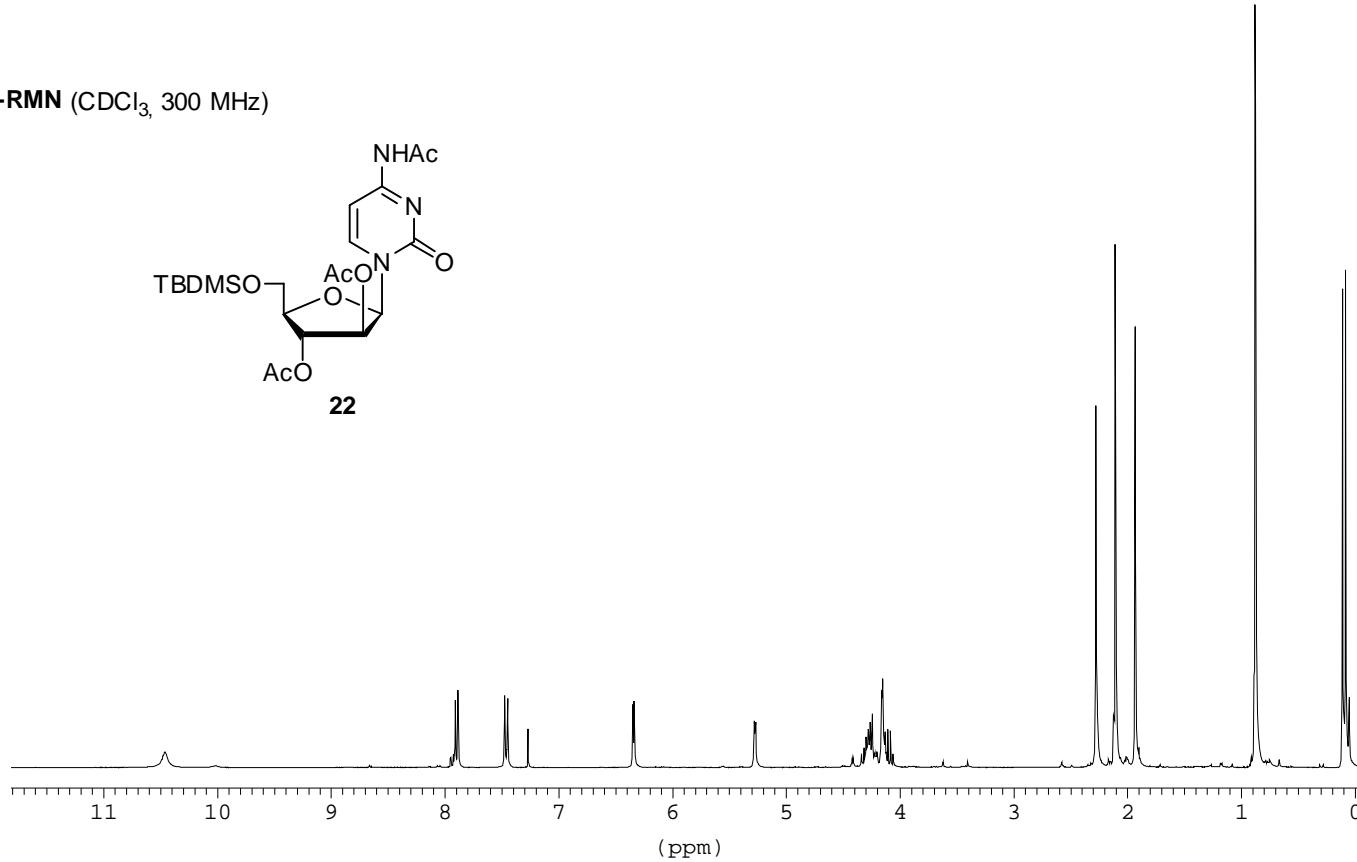
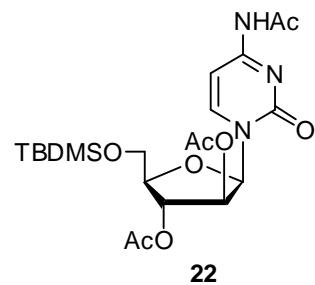
1-(5'-*O*-*tert*-butyldimethylsilyl- β -D-arabinofuranosyl)cytosine (21)

DEPT (MeOH-*d*₄, 75.5 MHz)



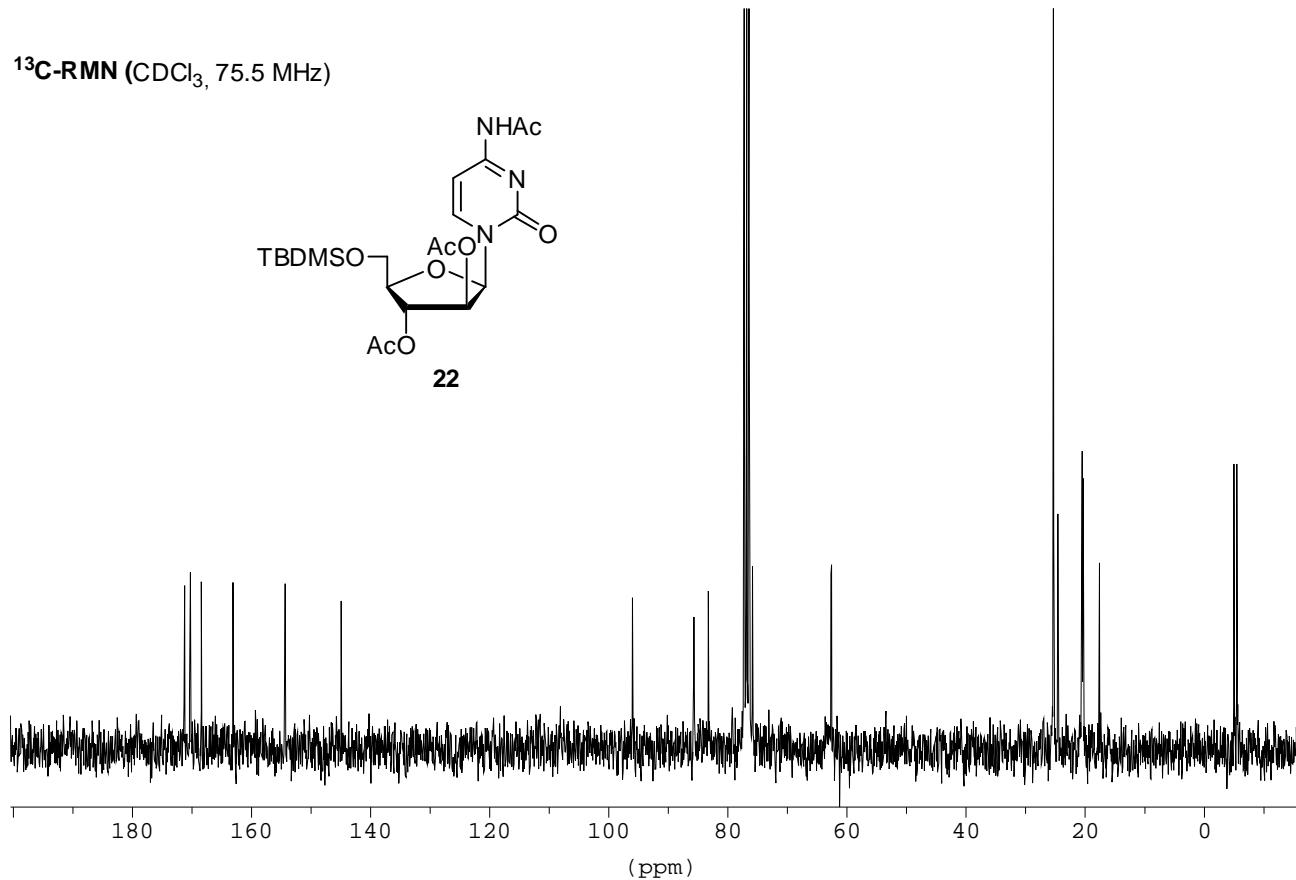
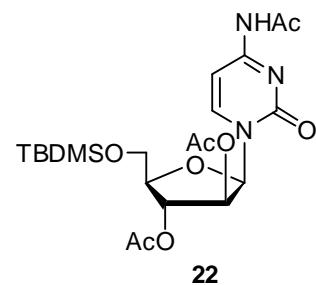
*N*⁴-acetyl-1-(2',3'-di-*O*-acetyl-5'-*O*-*tert*-butyldimethylsilyl- β -D-arabinofuranosyl)cytosine (22)

¹H-RMN (CDCl₃, 300 MHz)



*N*⁴-acetyl-1-(2',3'-di-*O*-acetyl-5'-*O*-tert-butyldimethylsilyl- β -D-arabinofuranosyl)cytosine (22)

¹³C-RMN (CDCl₃, 75.5 MHz)



*N*⁴-acetyl-1-(2',3'-di-*O*-acetyl-5'-*O*-*tert*-butyldimethylsilyl- β -D-arabinofuranosyl)cytosine (22)

DEPT (CDCl_3 , 75.5 MHz)

