

Chemical synthesis of fragments of galactosaminogalactan and pel polysaccharides Zhang, Y.

Citation

Zhang, Y. (2021, November 9). Chemical synthesis of fragments of galactosaminogalactan and pel polysaccharides. Retrieved from https://hdl.handle.net/1887/3239151

Version: Publisher's Version

Licence agreement concerning inclusion of doctoral thesis License:

in the Institutional Repository of the University of Leiden

Downloaded from: https://hdl.handle.net/1887/3239151

Note: To cite this publication please use the final published version (if applicable).

Chemical Synthesis of Fragments of Galactosaminogalactan and Pel Polysaccharides

Proefschrift

ter verkrijging van

de graad van doctor aan de Universiteit Leiden, op gezag van rector magnificus prof.dr.ir. H. Bijl, volgens besluit van het college voor promoties te verdedigen op dinsdag 9 november 2021

klokke 11:15 uur

door

Yongzhen Zhang

张永振

Geboren te Feixian, Shandong, China in 1989

Promotiecommissie

Promotoren:

Prof. dr. J. D. C. Codée

Prof. dr. G. A. van der Marel

Overige leden:

Prof. dr. H. S. Overkleeft, Leiden University

Prof. dr. G. J. Boons, Utrecht University

Prof. dr. D. Bonnaffé, University Paris sud

Dr. M. Artola, Leiden University

Dr. D. V. Filippov, Leiden University

To my family and my friends.

路漫漫其修远兮, 吾将上下而求索。

Long, long had been my road and far, far was the journey;

I would go up and down to seek my heart's desire.

Table of contents

List of abbreviations ·····i
Chapter 11
General Introduction
Chapter 2
Synthesis of α -galactose, α -galactosamine and α -N-acetyl galactosamine galactosaminogalactan homo-oligomers from Aspergillus fumigatus
Chapter 3
Synthesis of GAG hetero-oligomers featuring α -galactose, α -galactosamine and α - N -acety galactosamines linkages
Chapter 4 137
Synthesis of an azido-GAG heptasaccharide featuring α-GalN ₃ and α-GalNAc linkages
Chapter 5 163
Assembly of a library of Pel oligosaccharides featuring α -glucosamine and α -galactosamine linkages
Chapter 6
Summary and Future Prospects
Chinese Summary 222
Curriculum Vitae······225
List of Publications ······ 220
Acknowledgements · · · · · · 22

List of abbreviations

Ac	acetyl	Glc	glucose
ACN	acetonitrile	GlcN	glucosamine
aq.	aqueous	GlcN ₃	2-azido-2-deoxy glucose
atm	1 atmosphere = 10 ⁵ Pa	GlcNAc	N-acetyl glucosamine
Bn	benzyl	h	hour
Boc	t-butyloxy carbonyl	HB	hydrogen bond
bs	broad singlet	HFIP	1,1,1,3,3,3-hexafluoro-2-
BSP	1-benzenesulfinyl piperidine		propanol
Bt	benzotriazole	HMBC	heteronuclear multiple-bond
Bu	butyl		correlation
t-Bu	tert-butyl	HPLC	high performance liquid chromatography
Bz	benzoyl		
COSY	correlation spectroscopy	HRMS	high-resolution mass spectroscopy
δ	chemical shift		
d	doublet	HSQC	heteronuclear single quantum coherence
DCM	dichloromethane		
dd	doublet of doublets	Hz	Hertz
DMAP	N,N-4-	IR	infrared
	dimethylaminopyridine	J	coupling constant
DME	dimethylformomide		
DMF	dimethylformamide	LC-MS	liquid chromatography-mass
DMF DMSO	dimethyl sulfoxide	LC-MS	liquid chromatography-mass spectrometry
	dimethyl sulfoxide 2,6- <i>di-tert</i> -butyl-4-	LC-MS	
DMSO DTBMP	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine		spectrometry
DMSO DTBMP DTBS	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene	Lev	spectrometry levulinoyl
DMSO DTBMP	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine	Lev m	spectrometry levulinoyl multiplet molar matrix-associated laser
DMSO DTBMP DTBS EA/	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene	Lev m M	spectrometry levulinoyl multiplet molar
DMSO DTBMP DTBS EA/ EtOAc	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene ethyl acetate equivalents	Lev m M	spectrometry levulinoyl multiplet molar matrix-associated laser
DMSO DTBMP DTBS EA/ EtOAc eq ESI	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene ethyl acetate equivalents electrospray ionization	Lev m M MALDI	spectrometry levulinoyl multiplet molar matrix-associated laser desorption ionization
DMSO DTBMP DTBS EA/ EtOAc eq ESI Et	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene ethyl acetate equivalents electrospray ionization ethyl	Lev m M MALDI	spectrometry levulinoyl multiplet molar matrix-associated laser desorption ionization mannose
DMSO DTBMP DTBS EA/ EtOAc eq ESI Et GAG	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene ethyl acetate equivalents electrospray ionization ethyl galactosaminogalactan	Lev m M MALDI Man Me	spectrometry levulinoyl multiplet molar matrix-associated laser desorption ionization mannose methyl
DMSO DTBMP DTBS EA/ EtOAc eq ESI Et GAG Gal	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene ethyl acetate equivalents electrospray ionization ethyl galactosaminogalactan galactose	Lev m M MALDI Man Me min	spectrometry levulinoyl multiplet molar matrix-associated laser desorption ionization mannose methyl minute mehyl(phenyl)formamide mass spectrometry
DMSO DTBMP DTBS EA/ EtOAc eq ESI Et GAG Gal GalN	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene ethyl acetate equivalents electrospray ionization ethyl galactosaminogalactan galactose galactosamine	Lev m M MALDI Man Me min MPF	spectrometry levulinoyl multiplet molar matrix-associated laser desorption ionization mannose methyl minute mehyl(phenyl)formamide
DMSO DTBMP DTBS EA/ EtOAc eq ESI Et GAG Gal	dimethyl sulfoxide 2,6-di-tert-butyl-4- methylpyridine di-tert-butylsilylidene ethyl acetate equivalents electrospray ionization ethyl galactosaminogalactan galactose	Lev m M MALDI Man Me min MPF MS	spectrometry levulinoyl multiplet molar matrix-associated laser desorption ionization mannose methyl minute mehyl(phenyl)formamide mass spectrometry

NIS	N-iodosuccinimide	TBAI	tetra- <i>n</i> -butylammonium
NMR	nuclear magnetic resonance		iodide
NOESY	nuclear Overhauser effect	TBDPS	tert-butyldiphenylsilyl
	spectroscopy	TBS	tert-butyldimethylsilyl
Nu	nucleophile	TCA	trichloroacetyl
NR	no reaction	TEA	triethylamine
PE	petroleum ether	TEMPO	2,2,6,6-tetramethyl-1-
Pel	pellicle polysaccharide		piperidinyloxy
Ph	phenyl	Tf	trifluoromethanesulfonyl
Ph ₂ SO	diphenyl sulfoxide	TfOH	triflic acid
Phth	phthaloyl	Tf_2O	trifluoromethanesulfonic
PMB	para-methoxybenzyl		anhydride
ppm	parts per million	TFA	trifluoroacetic acid
Ру	pyridine	THF	tetrahydrofuran
q	quartet	TIPS	triisopropylsilyl
ч RT	room temperature	TMS	trimethylsilyl
Rf	retention factor	TLC	thin layer chromatography
		Tol	<i>p</i> -tolyl
s sat.	singlet saturated	Troc	2,2,2-
SDS-			trichloroethoxycarbonyl
SDS- PAGE	sodium dodecyl sulfate polyacrylamide gel	Ts	<i>p</i> -toluenesulfonyl
THOL	electrophoresis	TTBP	2,4,6-tri- <i>tert</i> -
t	triplet	p-TsOH	butylpyrimidine <i>p</i> -toluenesulfonic acid
TBAB	tetra-n-butylammonium	<i>p</i> -1son UDP	•
	bromide		uridine 5'-diphosphate
TBAF	tetra- <i>n</i> -butylammonium fluoride	UV	unraviolet