

Transfer of "goods" from plants to humans: Fundamental and applied biochemical investigations on retaining glycosidases Kytidou, K.

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Transfer of *"goods"* from plants to humans

Fundamental and applied biochemical investigations on retaining glycosidases

Kassiani Kytidou

Transfer of "goods" from plants to humans; Fundamental and applied biochemical investigations on retaining glycosidases *Kassiani Kytidou* Doctoral thesis, Leiden University 2020

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Transfer of "goods" from plants to humans: Fundamental and applied biochemical investigations on retaining glycosidases

Proefschrift

ter verkrijging van de graad van Doctor aan de Universiteit Leiden, op gezag van Rector Magnificus prof.mr. C.J.J.M. Stolker, volgens besluit van het College voor Promoties te verdedigen op donderdag 25 Juni 2020 klokke 11:15 uur

door

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To my unofficial lab partners, Dimitris and Foivos

To my family,

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List of Abbreviations and acronyms

FD = Fabry disease ERT = Enzyme replacement therapy α -GAL A or GLA = human alpha galactosidase $(\alpha$ -)NAGA(L) = α -N-acetyl-galactosaminidase $(\alpha$ -)NAGAEL = modified α -N-acetyl-galactosaminidase α -GAL = any alpha galactosidase, including plant enzymes 4MU = 4-Methylumbelliferyl ABP = Activity-based probe CBB = Coomassie brilliant blue ConA = Concanavalin AGb3 = globotriaosylceramide GlcCer = glucosylceramide Cer = ceramide lvsoGb3 = globotriaosylsphingosine LacCer = lactosylceramide lysoLacCer = lactosylsphingosine mAb = monoclonal antibodies Gal-DNJ = deoxygalactonojirimycin GBA = human lysosomal glucocerebrosidase GAA = human lysosomal alpha glucosidase βGLUR = human lysosomal beta glucuronidase αMAN = human alpha mannosidase α FUC = human alpha fucosidase β GAL = human beta galactosidase HPSE = Heparanase MPR = Mannose-6-phosphate receptor Man-6-P = Mannose-6-phosphate moieties MR = Molecular Replacement PD = Parkinson disease GD = Gaucher disease SDS-PAGE = sodium dodecyl sulfate polyacrylamide gel electrophoresis HPLC = High Performance Liquid Chromatography LC-MS/MS = Liquid chromatography-mass spectrometry with two mass spectrometers in tandem HRP = horseradish peroxidase NBD = nitrobenzoxadiazole NBD-Gb3 = NBD-C12-globotriaosylceramide PBS = Phosphate buffered saline DAPI = 4',6-diamidino-2-phenylindole BY2 cells= Bright Yellow 2 Nicotiana tabacum cells BSA = Bovine serum albumin

RT = real time A1.1 = *Nicotiana benthamiana* alpha galactosidase, identified in current thesis B56 = *Nicotiana tabacum* beta glucosidase, identified in current thesis

List of activity-based probes coding

TB474 = Cy5 labelled alpha galactosidase activity-based probe ME741 = Biotinylated alpha galactosidase activity-based probe ME569 = Cy5 labelled epoxide beta glucosidase activity-based probe ME869 = TAMRA/Biotinylated labelled epoxide beta glucosidase activity-based probe JJB111 = Biotinylated labelled aziridine beta glucosidase activity-based probe JJB367 = Cy5 labelled aziridine beta glucosidase activity-based probe JJB75 = Bodipy-red labelled aziridine beta glucosidase activity-based probe TB652 = Cy5 labelled aziridine beta glucosidase activity-based probe JJB381 = Cy5 labelled aziridine alpha fucosidase activity-based probe JJB383 = Cy5 labelled aziridine alpha glucosidase activity-based probe JJB392 = Cy5 labelled aziridine beta glucosidase activity-based probe JJB383 = Cy5 labelled aziridine beta glucosidase activity-based probe JJB384 = Cy5 labelled aziridine beta glucosidase activity-based probe JJB392 = Cy5 labelled aziridine beta glucosidase activity-based probe TB482 = Cy5 labelled aziridine beta glucosidase activity-based probe TB482 = Cy5 labelled aziridine alpha mannosidase activity-based probe