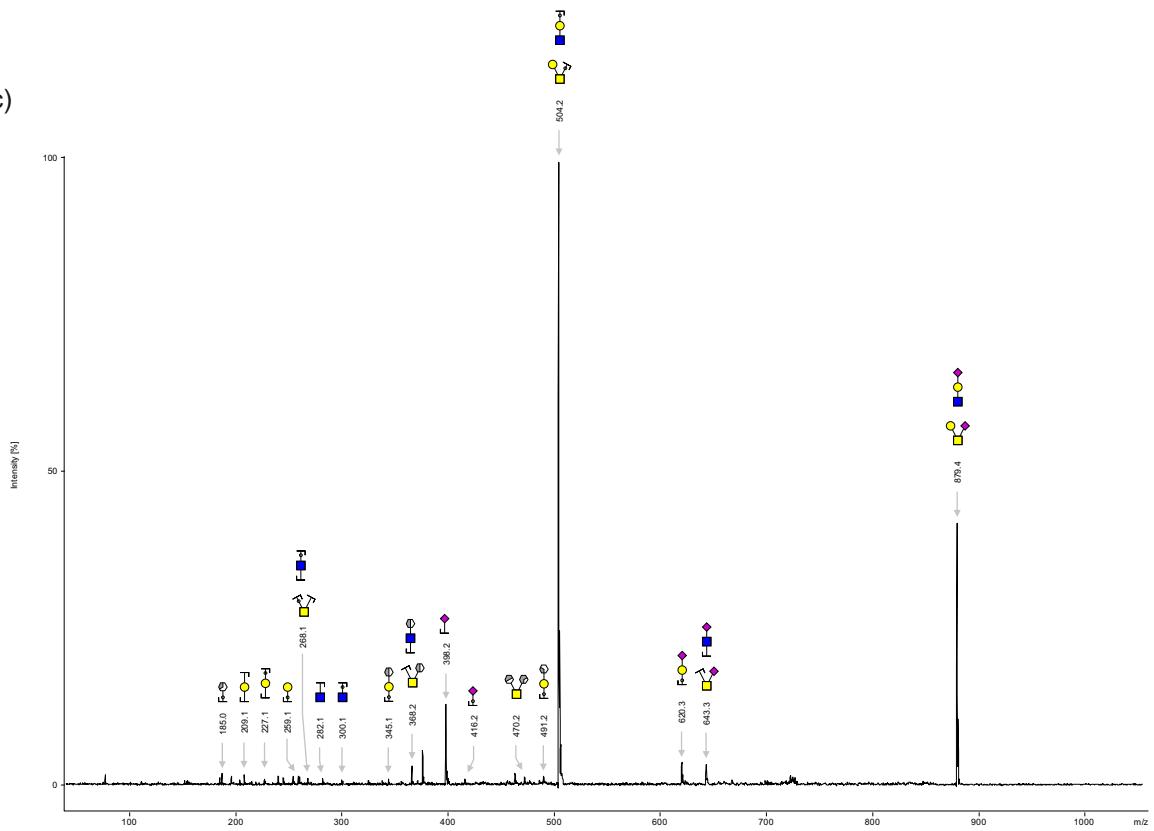
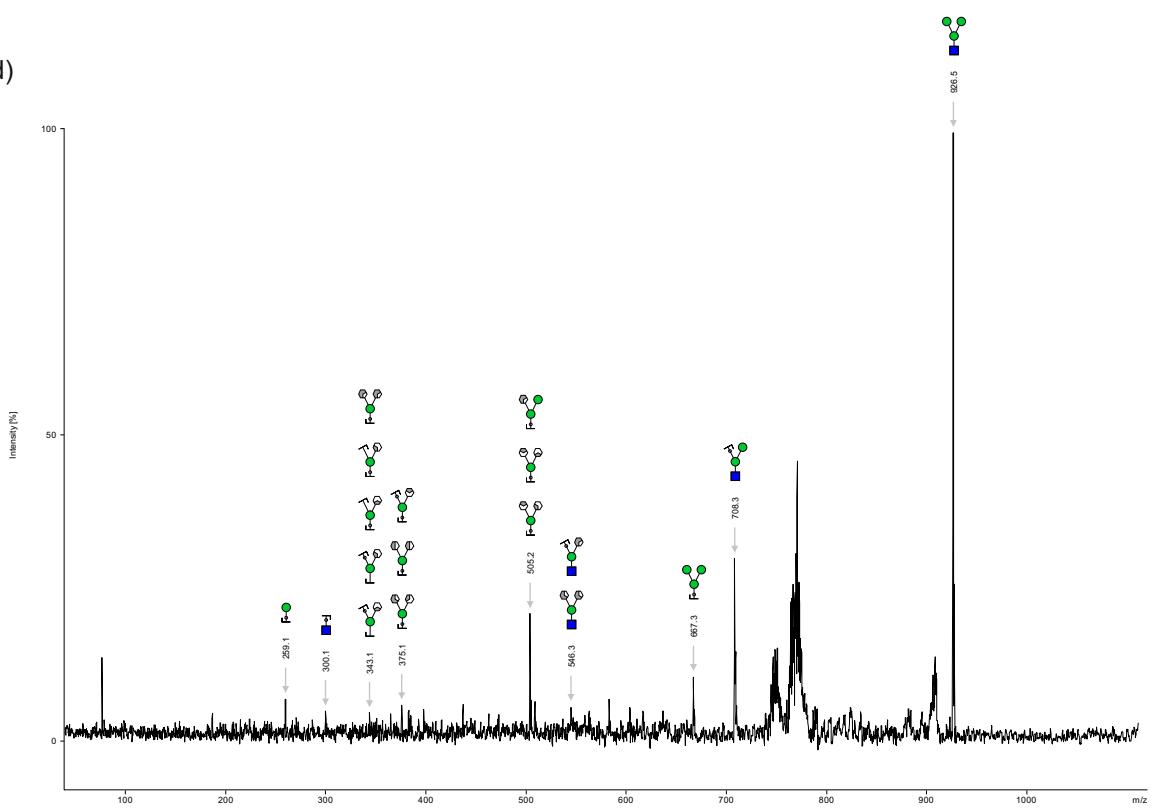
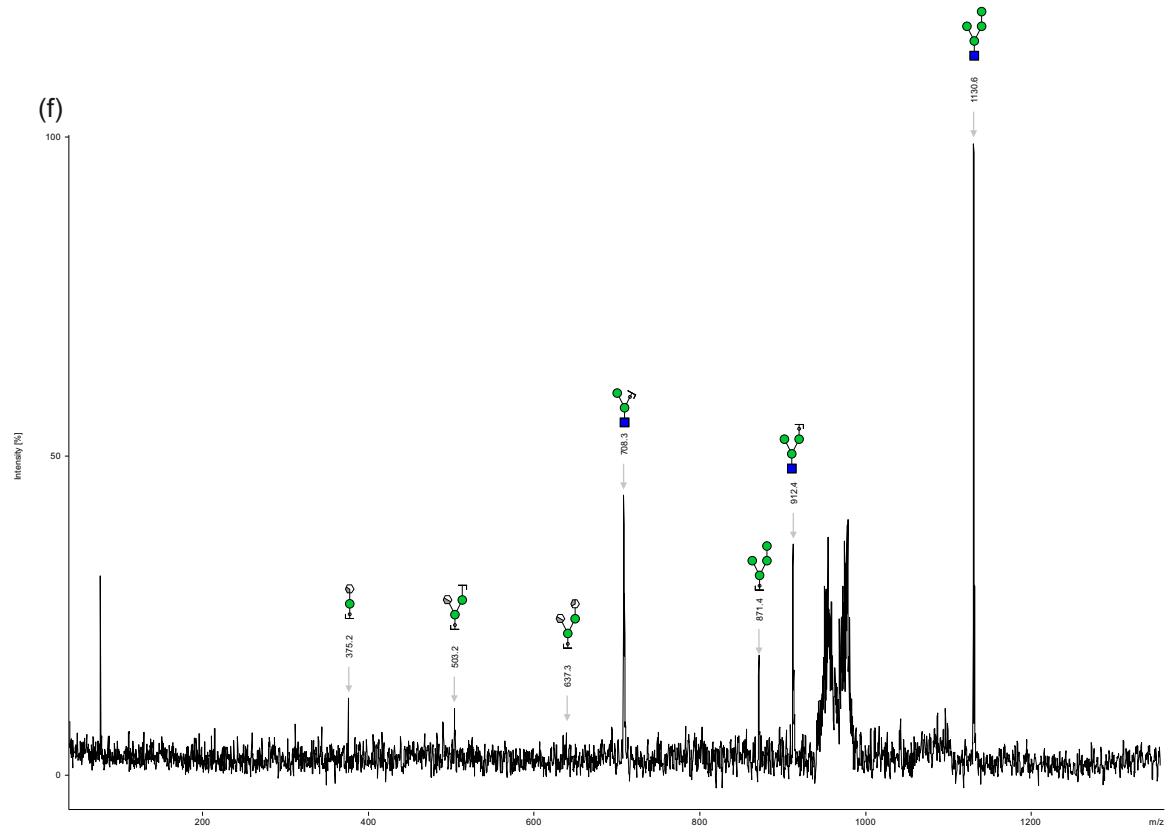
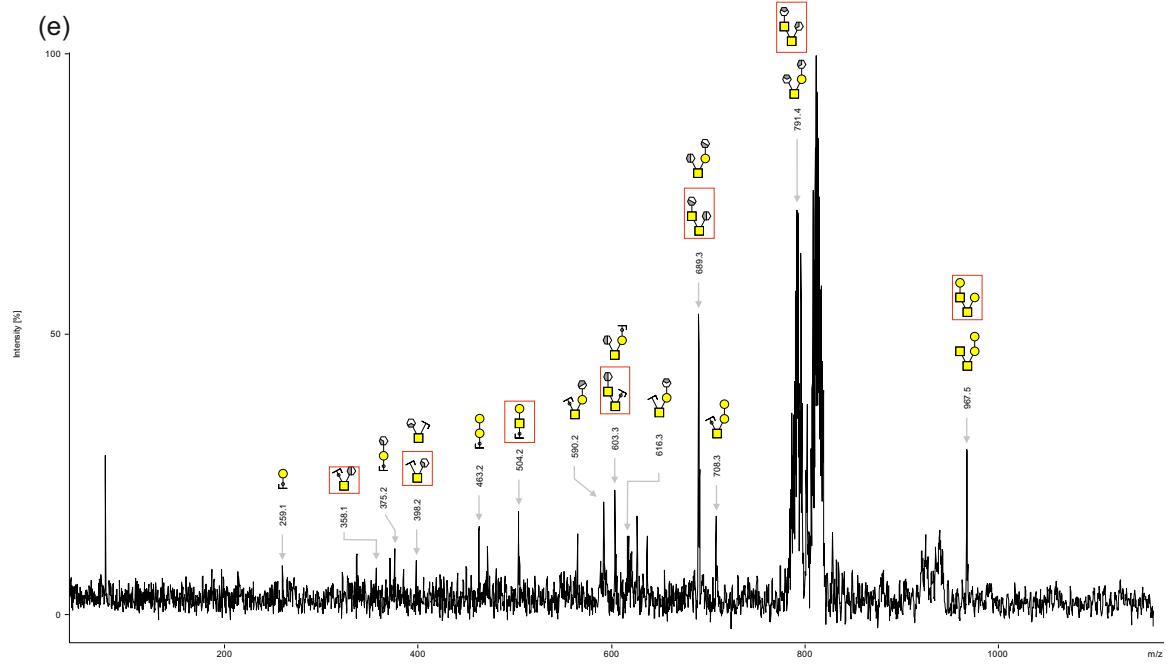


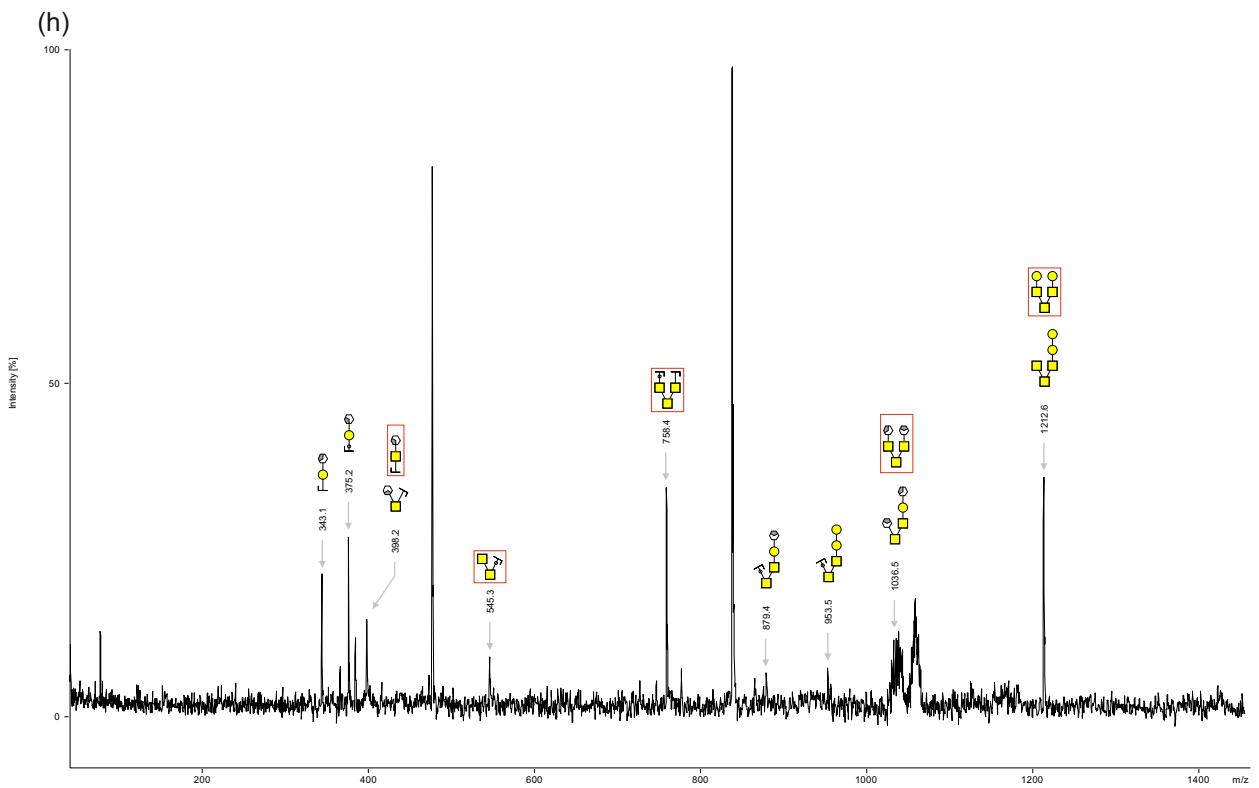
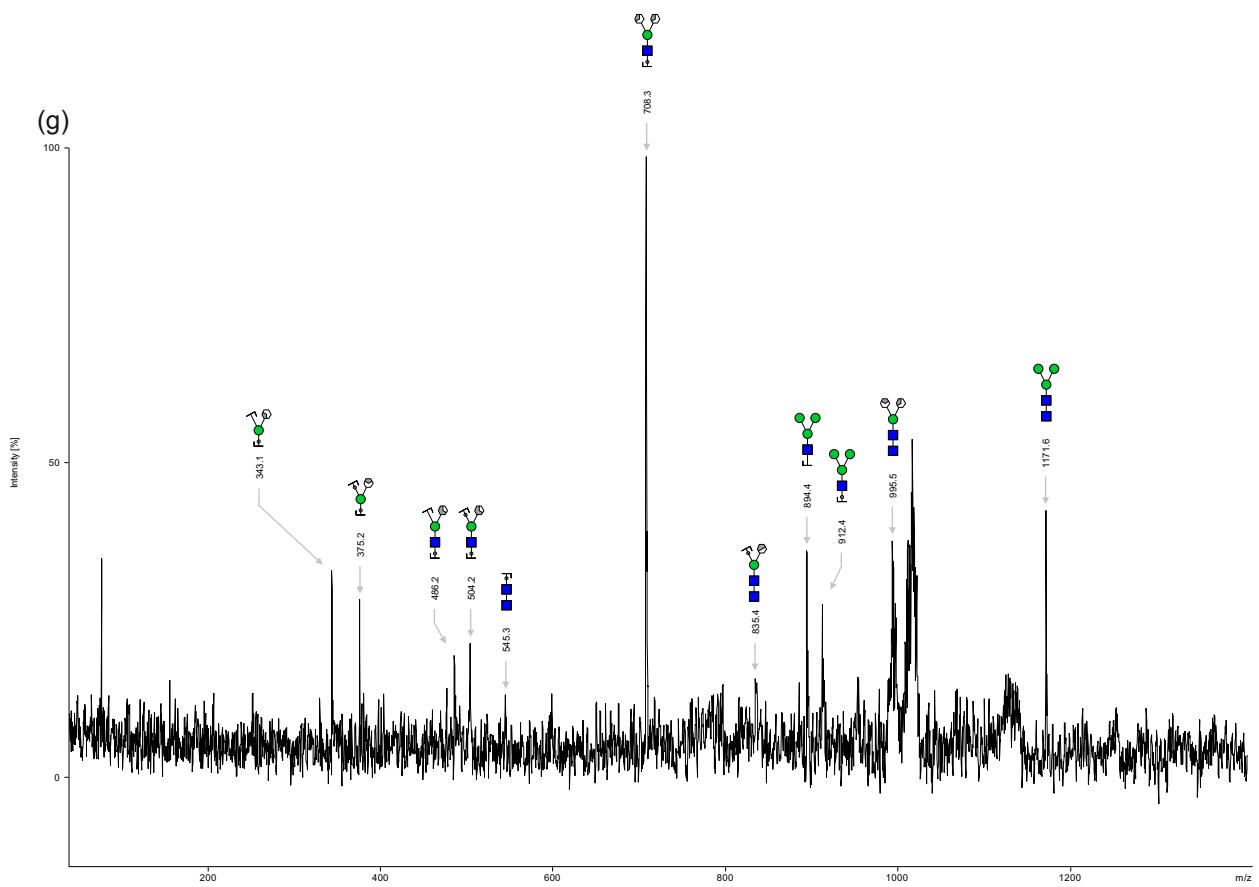
(c)



(d)







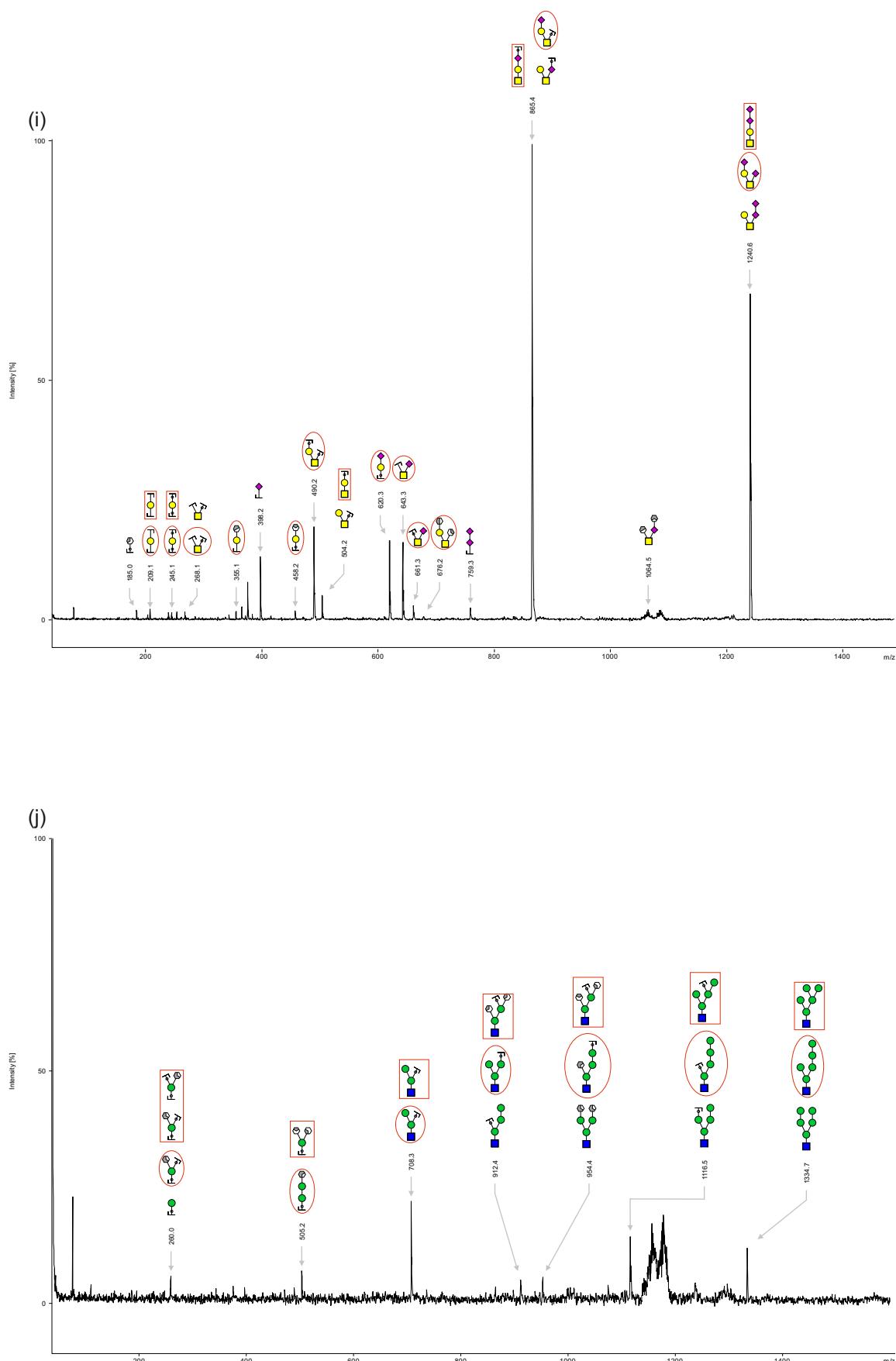
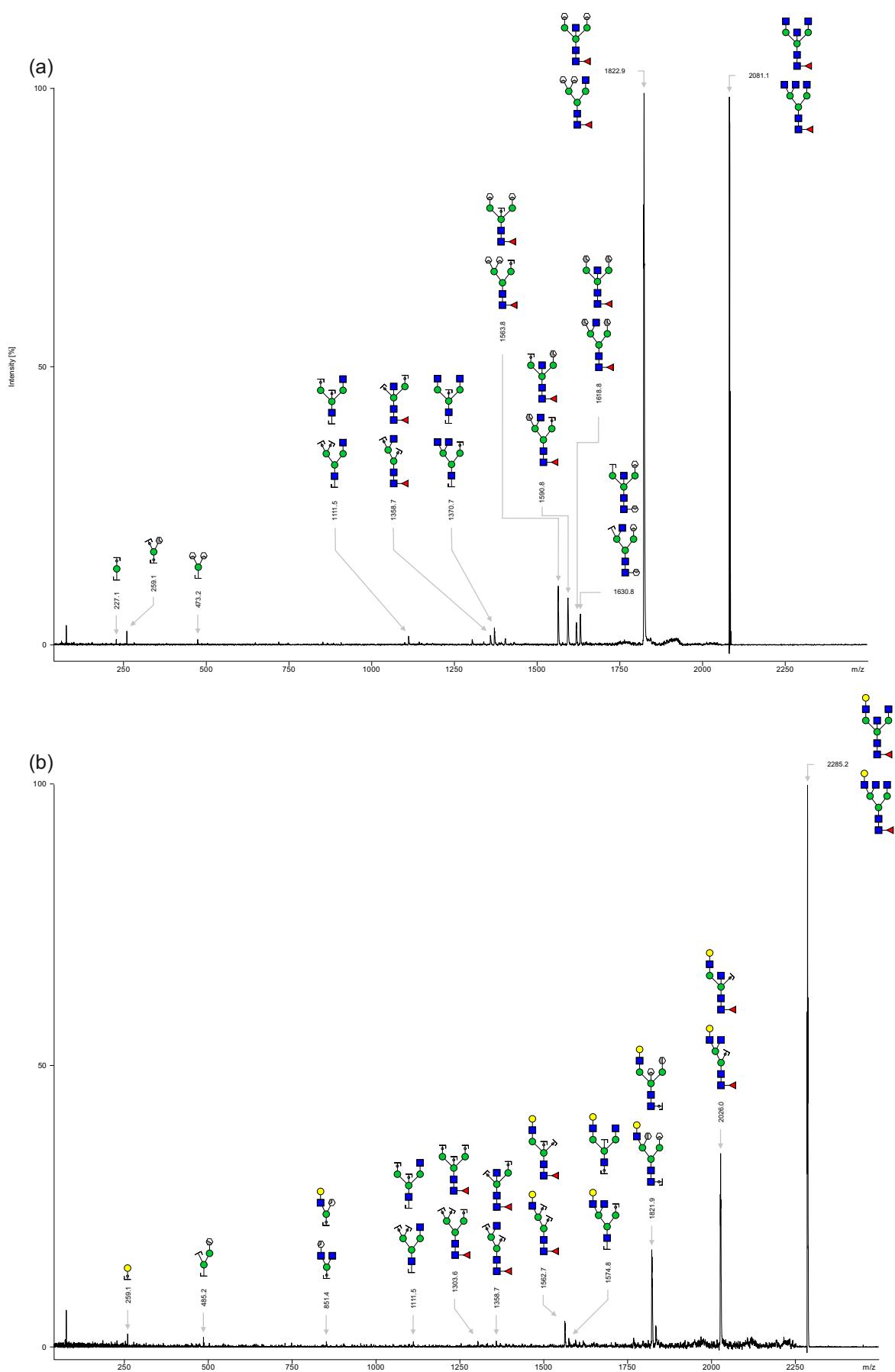
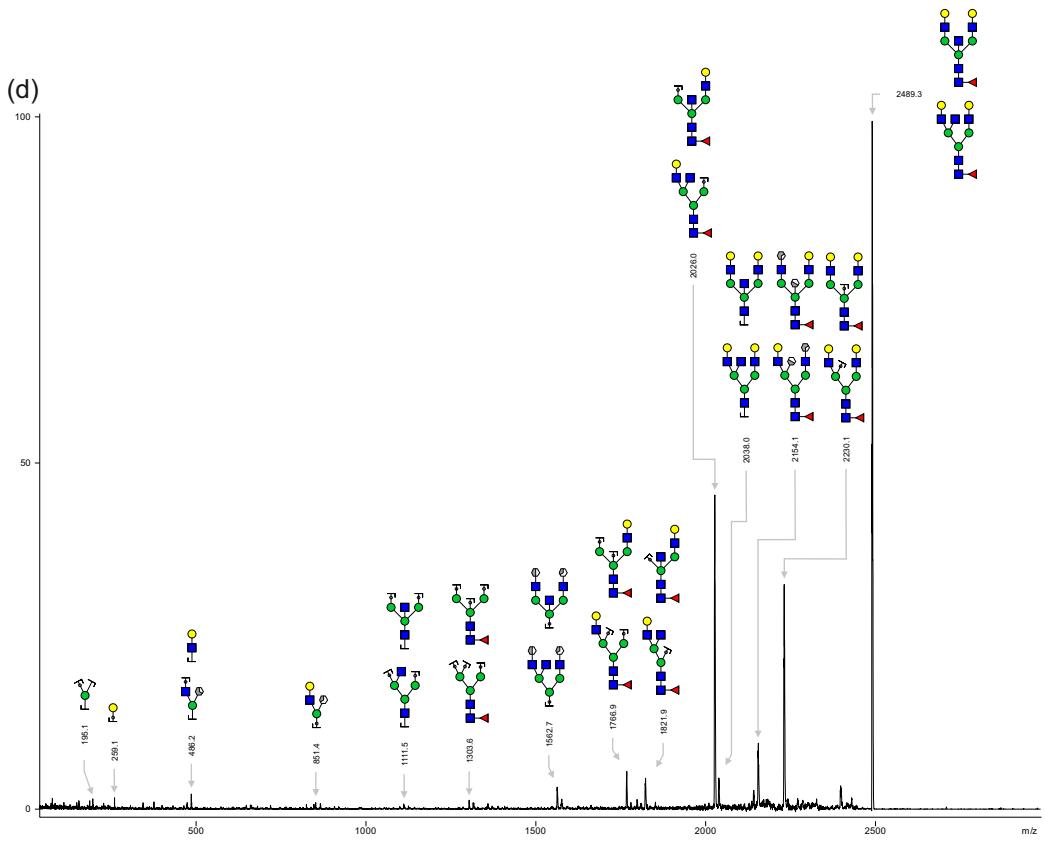
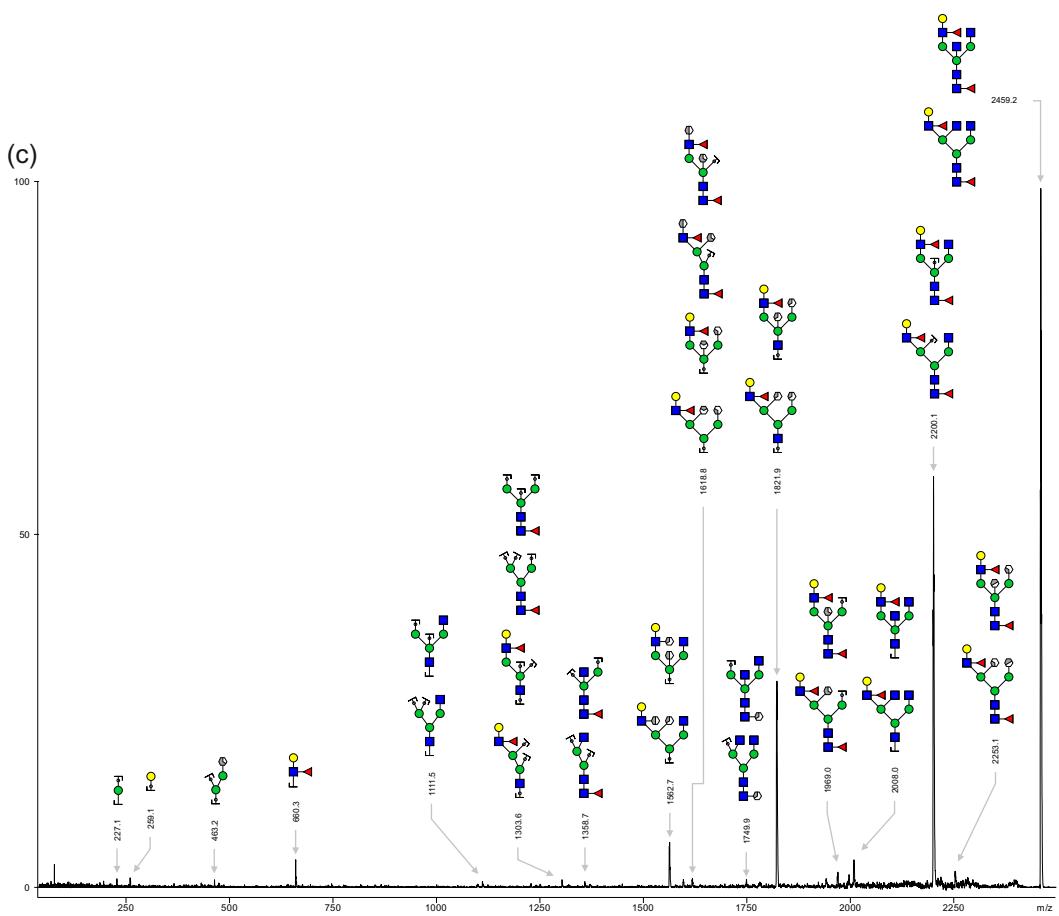
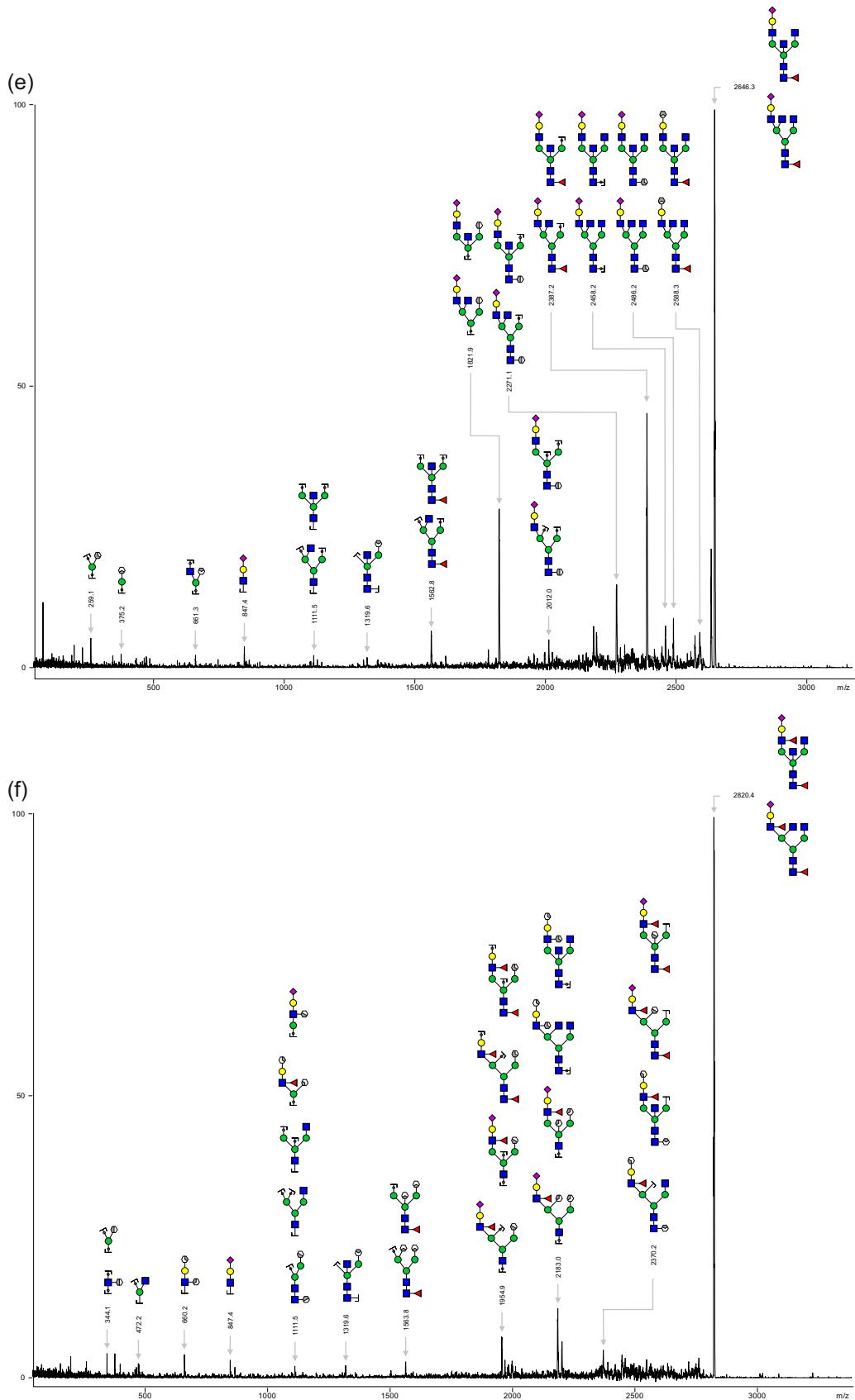


Figure S1. MALDI-TOF/TOF mass spectra of the  $[M + Na]^+$  molecular ions at  $m/z$  (a) 722.4, (b) 825.4, (c) 879.4, (d) 926.5, (e) 967.5, (f) 1130.6, (g) 1171.6, (h) 1212.6, (i) 1240.6, (j) 1334.7 derived from free glycans isolated from CSF. Registered fragments may be formed by different fragmentation pathways, only one of which is depicted in the figure. Green circle, Man; yellow circle, Gal; yellow square, GalNAc; blue square, GlcNAc; red triangle, Fuc; Neu5Ac: pink diamond.







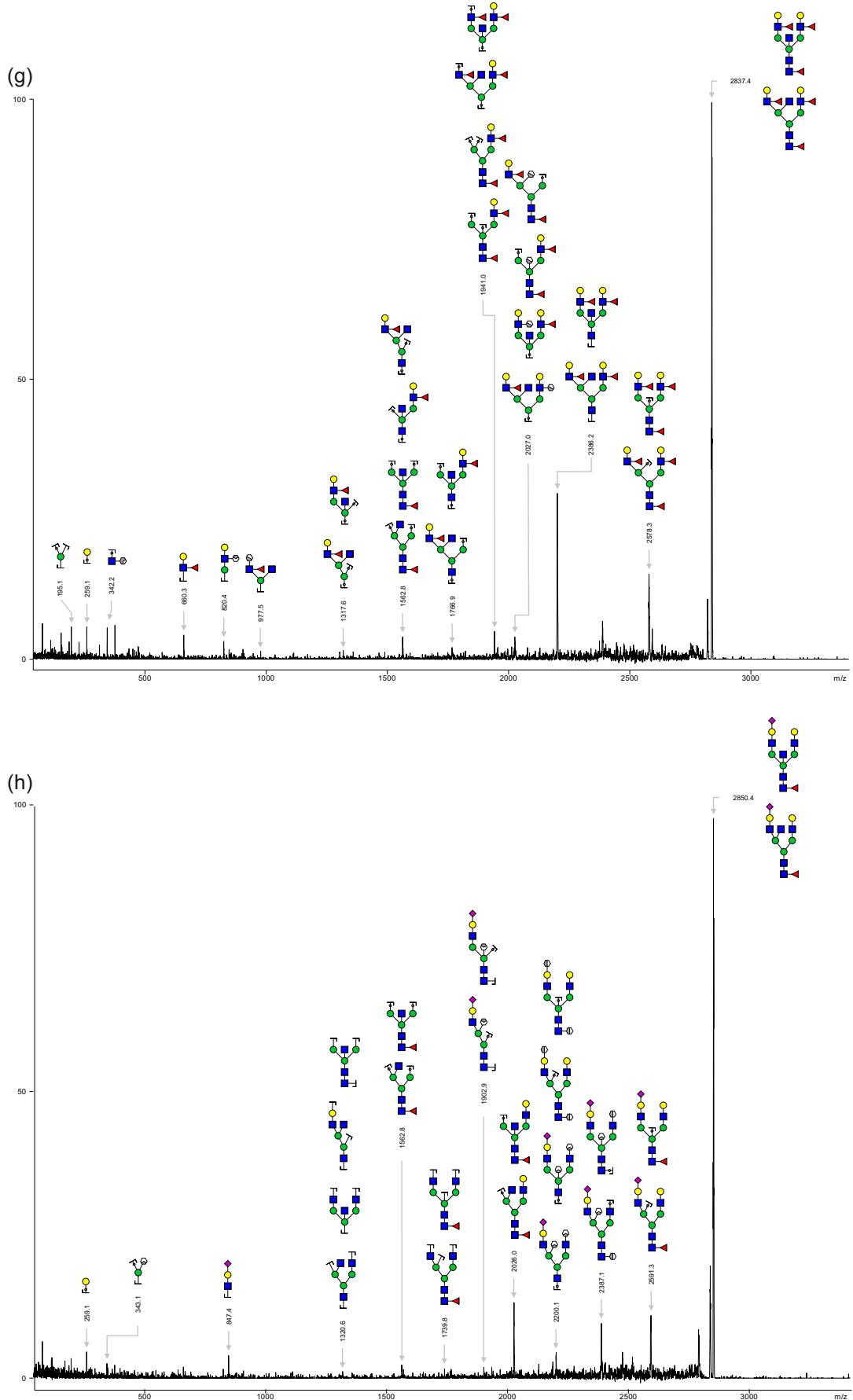


Figure S2. MALDI-TOF/TOF mass spectra of the  $[M + Na]^+$  molecular ions at  $m/z$  (a) 2081.1, (b) 2285.2 (c) 2459.2 (d) 2489.3 (e) 2646.3, (f) 2820.4 (g) 2837.4, (h) 2850.4, derived from N-glycans isolated from CSF. Registered fragments may be formed by different fragmentation pathway which is depicted in the figure. Green circle, Man; yellow circle, Gal; blue square, GlcNAc; red triangle, Fuc; Neu5Ac: pink diamond.

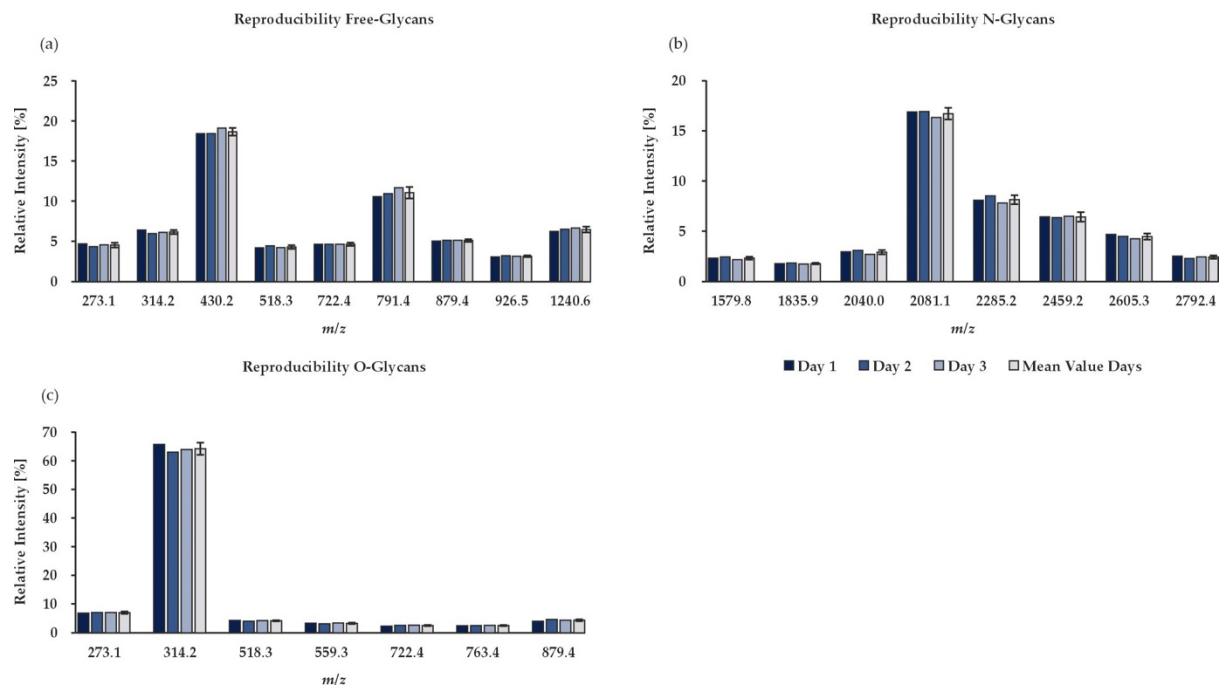


Figure S3: Inter-day reproducibility of the analytical workflow used to prepare (a) free glycans, (b) N-glycans and (c) O-glycans from CSF. Average is shown as mean  $\pm$  SD.

Supplementary Material: Tables

*Table S1: Two-way analysis of variance (ANOVA) to assess any statistical significance of gender in the dataset of free-glycans. P-values ≤ 0.05 were considered as statistically significant and p-values ≤ 0.01 as highly significant.*

<b><i>m/z</i></b>	<b>Composition</b>	<b>Significance Levene-Test</b>	<b>Significance - Group</b>	<b>Significance – Sex</b>	<b>Significance – Group * Sex</b>
<b>273.1</b>	Hex <sub>1</sub>	0.002	0.000	0.799	0.658
<b>314.2</b>	HexNAc <sub>1</sub>	0.098	0.003	0.217	0.457
<b>430.2</b>	Neu5Ac <sub>1</sub>	0.034	0.682	0.462	0.708
<b>518.3</b>	Hex <sub>1</sub> HexNAc <sub>1</sub>	0.000	0.000	0.546	0.262
<b>559.3</b>	HexNAc <sub>2</sub>	0.001	0.000	0.165	0.276
<b>675.3</b>	Neu5Ac <sub>1</sub> HexNAc <sub>1</sub>	0.003	0.000	0.083	0.214
<b>722.4</b>	Hex <sub>2</sub> HexNAc <sub>1</sub>	0.004	0.000	0.500	0.770
<b>733.4</b>	Fuc <sub>1</sub> HexNAc <sub>2</sub>	0.009	0.000	0.482	0.289
<b>791.4</b>	Neu5Ac <sub>2</sub>	0.001	0.025	0.584	0.584
<b>825.4</b>	Fuc <sub>2</sub> Hex <sub>2</sub>	0.031	0.033	0.901	0.974
<b>879.4</b>	NeuAc <sub>1</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.209	0.000	0.733	0.364
<b>926.5</b>	Hex <sub>3</sub> HexNAc <sub>1</sub>	0.003	0.000	0.592	0.802
<b>967.5</b>	Hex <sub>2</sub> HexNAc <sub>2</sub>	0.000	0.000	0.215	0.675
<b>1130.6</b>	Hex <sub>4</sub> HexNAc <sub>1</sub>	0.000	0.000	0.919	0.954
<b>1171.6</b>	Hex <sub>3</sub> HexNAc <sub>2</sub>	0.000	0.002	0.200	0.818
<b>1212.6</b>	Hex <sub>2</sub> HexNAc <sub>3</sub>	0.006	0.143	0.238	0.857
<b>1240.6</b>	Neu5Ac <sub>2</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.927	0.032	0.789	0.494
<b>1334.7</b>	Hex <sub>5</sub> HexNAc <sub>1</sub>	0.002	0.000	0.974	0.971
<b>1601.8</b>	Neu5Ac <sub>3</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.930	0.000	0.451	0.516

*Table S2: Two-way analysis of variance (ANOVA) to evaluate the statistical significance of gender in the dataset of N-glycans. P-values ≤ 0.05 were considered as statistically significant and p-values ≤ 0.01 as highly significant.*

<i>m/z</i>	Composition	Significance Levene-Test	Significance Patient Groups	Significance – Sex	Significance – Group * Sex
<b>1579.8</b>	Hex:HexNAc <sub>2</sub>	0.001	0.002	0.001	0.282
<b>1661.8</b>	Hex:HexNAc <sub>4</sub>	0.007	0.000	0.357	0.030
<b>1783.9</b>	Hex:HexNAc <sub>2</sub>	0.001	0.025	0.438	0.566
<b>1835.9</b>	Fuc:Hex <sub>3</sub> HexNAc <sub>4</sub>	0.000	0.012	0.422	0.013
<b>2040.0</b>	Fuc:Hex <sub>4</sub> HexNAc <sub>4</sub>	0.005	0.188	0.041	0.129
<b>2070.0</b>	Hex:HexNAc <sub>4</sub>	0.002	0.506	0.780	0.937
<b>2081.1</b>	Fuc:Hex <sub>3</sub> HexNAc <sub>5</sub>	0.000	0.006	0.023	0.109
<b>2111.1</b>	Hex <sub>4</sub> HexNAc <sub>5</sub>	0.118	0.004	0.770	0.540
<b>2214.1</b>	Fuc:Hex <sub>4</sub> HexNAc <sub>4</sub>	0.354	0.006	0.214	0.010
<b>2244.1</b>	Fuc:Hex <sub>5</sub> HexNAc <sub>4</sub>	0.008	0.003	0.006	0.736
<b>2285.2</b>	Fuc:Hex <sub>4</sub> HexNAc <sub>5</sub>	0.000	0.137	0.574	0.359
<b>2431.2</b>	Neu5Ac <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	0.025	0.254	0.723	0.312
<b>2459.2</b>	Fuc:Hex <sub>4</sub> HexNAc <sub>5</sub>	0.016	0.000	0.000	0.972
<b>2489.3</b>	Fuc:Hex <sub>5</sub> HexNAc <sub>5</sub>	0.009	0.009	0.045	0.766
<b>2592.3</b>	Fuc:Hex <sub>5</sub> HexNAc <sub>4</sub>	0.051	0.023	0.939	0.731
<b>2605.3</b>	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	0.002	0.000	0.014	0.971
<b>2646.3</b>	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	0.008	0.000	0.189	0.966
<b>2792.4</b>	Neu5Ac <sub>2</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	0.105	0.076	0.576	0.724
<b>2820.4</b>	Neu5Ac <sub>1</sub> Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	0.273	0.048	0.835	0.950
<b>2837.4</b>	Fuc:Hex <sub>5</sub> HexNAc <sub>5</sub>	0.026	0.001	0.188	0.448
<b>2850.4</b>	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	0.003	0.000	0.977	0.657
<b>2966.5</b>	Neu5Ac <sub>2</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	0.000	0.000	0.060	0.866

*Table S3: Two-way analysis of variance (ANOVA) to evaluate the statistical significance of gender in the dataset of O-glycans. P-values ≤ 0.05 were considered as statistically significant and p-values ≤ 0.01 as highly significant.*

<b>m/z</b>	<b>Composition</b>	<b>Significance</b>	<b>Significance –</b>	<b>Significance –</b>	<b>Significance –</b>
		<b>Levene-Test</b>	<b>Patient Groups</b>	<b>Sex</b>	<b>Group * Sex</b>
<b>273.1</b>	Hex <sub>1</sub>	0.000	0.000	0.036	0.006
<b>314.2</b>	HexNAc <sub>1</sub>	0.145	0.002	0.104	0.197
<b>518.3</b>	Hex <sub>1</sub> HexNAc <sub>1</sub>	0.002	0.000	0.416	0.633
<b>559.3</b>	HexNAc <sub>2</sub>	0.309	0.000	0.856	0.823
<b>675.3</b>	NeuAc <sub>1</sub> HexNAc <sub>1</sub>	0.871	0.001	0.941	0.563
<b>722.4</b>	Hex <sub>2</sub> HexNAc <sub>1</sub>	0.962	0.000	0.747	0.720
<b>763.4</b>	Hex <sub>1</sub> HexNAc <sub>2</sub>	0.383	0.000	0.987	0.777
<b>804.4</b>	HexNAc <sub>3</sub>	0.132	0.000	0.668	0.436
<b>825.4</b>	Fuc <sub>2</sub> Hex <sub>2</sub>	0.102	0.000	0.923	0.270
<b>879.4</b>	NeuAc <sub>1</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.000	0.000	0.551	0.423
<b>967.5</b>	Hex <sub>2</sub> HexNAc <sub>2</sub>	0.057	0.001	0.907	0.540
<b>1124.6</b>	Neu5Ac <sub>1</sub> Hex <sub>1</sub> HexNAc <sub>2</sub>	0.066	0.003	0.029	0.918
<b>1240.6</b>	NeuAc <sub>2</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.000	0.000	0.091	0.575

Table S4: N-Glycans detected in this study by MALDI-TOF-MS.

<i>m/z</i>	Composition	
1171.6	Hex <sub>3</sub> HexNAc <sub>2</sub>	
1345.7	Fuc <sub>1</sub> Hex <sub>3</sub> HexNAc <sub>2</sub>	
1375.7	Hex <sub>4</sub> HexNAc <sub>2</sub>	
1579.8	Hex <sub>5</sub> HexNAc <sub>2</sub>	
1590.8	Fuc <sub>1</sub> Hex <sub>3</sub> HexNAc <sub>3</sub>	
1620.8	Hex <sub>4</sub> HexNAc <sub>3</sub>	
1661.8	Hex <sub>3</sub> HexNAc <sub>4</sub>	
1753.9	Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>2</sub>	
1783.9	Hex <sub>6</sub> HexNAc <sub>2</sub>	
1794.9	Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>3</sub>	
1835.9	Fuc <sub>1</sub> Hex <sub>3</sub> HexNAc <sub>4</sub>	
1865.9	Hex <sub>4</sub> HexNAc <sub>4</sub>	
1907.0	Hex <sub>3</sub> HexNAc <sub>5</sub>	
1982.0	Neu5Ac <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>3</sub>	
1988.0	Hex <sub>7</sub> HexNAc <sub>2</sub>	
1999.0	Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>3</sub>	
2010.0	Fuc <sub>2</sub> Hex <sub>3</sub> HexNAc <sub>5</sub>	
2029.0	Hex <sub>6</sub> HexNAc <sub>3</sub>	
2040.0	Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>4</sub>	
2070.0	Hex <sub>5</sub> HexNAc <sub>4</sub>	
2081.1	Fuc <sub>1</sub> Hex <sub>3</sub> HexNAc <sub>5</sub>	
2111.1	Hex <sub>4</sub> HexNAc <sub>5</sub>	
2152.1	Hex <sub>3</sub> HexNAc <sub>6</sub>	
2156.1	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>3</sub>	
2192.1	Hex <sub>8</sub> HexNAc <sub>2</sub>	
2214.1	Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>4</sub>	
2227.1	Neu5Ac <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>4</sub>	
2244.1	Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	
2285.2	Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	
2315.2	Hex <sub>5</sub> HexNAc <sub>5</sub>	
2326.2	Fuc <sub>1</sub> Hex <sub>3</sub> HexNAc <sub>6</sub>	
2360.2	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>3</sub>	
2390.2	Neu5Ac <sub>1</sub> Hex <sub>6</sub> HexNAc <sub>3</sub>	
2396.2	Hex <sub>9</sub> HexNAc <sub>2</sub>	
2401.2	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>4</sub>	
2418.2	Fuc <sub>2</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	
2431.2	Neu5Ac <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	
2459.2	Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	

2472.2	Neu5Ac <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	
2489.3	Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	
2519.3	Hex <sub>6</sub> HexNAc <sub>5</sub>	
2530.3	Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>6</sub>	
2560.3	Hex <sub>5</sub> HexNAc <sub>6</sub>	
2564.3	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>6</sub> HexNAc <sub>3</sub>	
2592.3	Fuc <sub>3</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	
2605.3	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	
2646.3	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	
2663.3	Fuc <sub>2</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	
2676.3	Neu5Ac <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	
2693.4	Fuc <sub>1</sub> Hex <sub>6</sub> HexNAc <sub>5</sub>	
2704.4	Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>6</sub>	
2717.4	Neu5Ac <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>6</sub>	
2734.4	Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>6</sub>	
2749.5	Neu5Ac <sub>1</sub> Fuc <sub>3</sub> Hex <sub>4</sub> HexNAc <sub>4</sub>	
2764.4	Hex <sub>6</sub> HexNAc <sub>6</sub>	
2792.4	Neu5Ac <sub>2</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	
2820.4	Neu5Ac <sub>1</sub> Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	
2837.4	Fuc <sub>3</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	
2850.4	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	
2880.4	Neu5Ac <sub>1</sub> Hex <sub>6</sub> HexNAc <sub>5</sub>	
2891.5	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>6</sub>	
2908.5	Fuc <sub>2</sub> Hex <sub>5</sub> HexNAc <sub>6</sub>	
2921.5	Neu5Ac <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>6</sub>	
2966.5	Neu5Ac <sub>2</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	

Table S5: ROC curves were generated for free glycans using 89 patients with AD, 86 disease control patients and 87 healthy control patients.

<i>m/z</i>	Composition	AD – Disease Control	AD – Healthy Control	Disease Control – Healthy Control
<b>273.1</b>	Hex <sub>1</sub>	0.85	0.76	0.45
<b>314.2</b>	HexNAc <sub>1</sub>	0.55	0.39	0.37
<b>430.2</b>	Neu5Ac <sub>1</sub>	0.51	0.55	0.53
<b>518.3</b>	Hex <sub>1</sub> HexNAc <sub>1</sub>	0.24	0.15	0.44
<b>559.3</b>	HexNAc <sub>2</sub>	0.41	0.25	0.34
<b>675.3</b>	NeuAc <sub>1</sub> HexNAc <sub>1</sub>	0.29	0.13	0.30
<b>722.4</b>	Hex <sub>2</sub> HexNAc <sub>1</sub>	0.38	0.17	0.33
<b>733.4</b>	Fuc <sub>1</sub> HexNAc <sub>2</sub>	0.27	0.26	0.50
<b>791.4</b>	Neu5Ac <sub>2</sub>	0.46	0.53	0.57
<b>825.4</b>	Fuc <sub>2</sub> Hex <sub>2</sub>	0.39	0.39	0.49
<b>879.4</b>	NeuAc <sub>1</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.21	0.30	0.59
<b>926.5</b>	Hex <sub>3</sub> HexNAc <sub>1</sub>	0.32	0.25	0.46
<b>967.5</b>	Hex <sub>2</sub> HexNAc <sub>2</sub>	0.34	0.28	0.50
<b>1130.6</b>	Hex <sub>4</sub> HexNAc <sub>1</sub>	0.33	0.33	0.53
<b>1171.6</b>	Hex <sub>3</sub> HexNAc <sub>2</sub>	0.39	0.44	0.57
<b>1212.6</b>	Hex <sub>2</sub> HexNAc <sub>3</sub>	0.50	0.44	0.44
<b>1240.6</b>	NeuAc <sub>2</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.39	0.48	0.58
<b>1334.7</b>	Hex <sub>5</sub> HexNAc <sub>1</sub>	0.35	0.42	0.58
<b>1601.8</b>	NeuAc <sub>3</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.46	0.71	0.72

Table S6: Correlations were calculated between the free glycans Hex<sub>1</sub> (*m/z* 273.1), HexNAc<sub>1</sub>Hex<sub>1</sub>Neu5Ac<sub>1</sub> (*m/z* 879.4) and CSF parameters using the using the Spearman's rank correlation coefficient, *p*<0.05 was considered as statistically significant.

Hex <sub>1</sub>							
	MMSE	Age of patients	Aβ40	Aβ42	Aβ-Ratio	pTau	tTau
<b>AD</b>	0,232	0,550	0,001	0,147	0,238	0,031	0,024
<b>HC</b>	0,792	0,943	0,002	0,031	0,598	0,000	0,015
<b>DC</b>	0,001	0,645	0,000	0,000	0,239	0,264	0,888
HexNAc <sub>1</sub> Hex <sub>1</sub> Neu5Ac <sub>1</sub>							
<b>AD</b>	0,178	0,430	0,822	0,742	0,959	0,577	0,749
<b>HC</b>	0,508	0,321	0,000	0,068	0,585	0,004	0,060
<b>DC</b>	0,051	0,400	0,009	0,029	0,559	0,961	0,227

Table S7: Mean relative areas and standard deviations of N-glycans isolated from the three patient cohorts as judged by MALDI-TOF-MS. AD: Alzheimer's disease, DC: disease control and HC: healthy control.

m/z	Composition	AD patients [%]		Disease control patients [%]		Healthy control patients [%]	
		Males	Females	Males	Females	Males	Females
1579.8	Hex <sub>5</sub> HexNAc <sub>2</sub>	6.13 ± 1.74	6.74 ± 1.69	5.84 ± 1.94	7.36 ± 2.66	7.23 ± 2.23	7.80 ± 2.79
1661.8	Hex <sub>3</sub> HexNAc <sub>4</sub>	1.12 ± 0.36	1.00 ± 0.36	1.21 ± 0.43	1.40 ± 0.54	1.10 ± 0.34	1.17 ± 0.32
1783.9	Hex <sub>6</sub> HexNAc <sub>2</sub>	0.99 ± 0.19	1.05 ± 0.19	1.12 ± 0.36	1.10 ± 0.29	1.11 ± 0.23	1.13 ± 0.24
1835.9	Fuc <sub>1</sub> Hex <sub>3</sub> HexNAc <sub>4</sub>	4.65 ± 1.13	4.10 ± 1.16	4.74 ± 2.21	5.46 ± 2.29	4.82 ± 1.69	4.13 ± 1.11
2040.0	Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>4</sub>	3.98 ± 0.79	3.38 ± 0.93	3.89 ± 1.65	4.02 ± 1.19	4.22 ± 1.30	3.75 ± 1.07
2070.0	Hex <sub>5</sub> HexNAc <sub>4</sub>	0.99 ± 0.21	0.98 ± 0.19	0.97 ± 0.29	0.99 ± 0.29	1.01 ± 0.23	1.02 ± 0.17
2081.1	Fuc <sub>1</sub> Hex <sub>3</sub> HexNAc <sub>5</sub>	17.38 ± 3.73	17.68 ± 2.52	15.20 ± 5.71	17.96 ± 3.55	18.41 ± 3.74	18.91 ± 4.20
2111.1	Hex <sub>4</sub> HexNAc <sub>5</sub>	1.09 ± 0.33	1.05 ± 0.36	1.22 ± 0.43	1.30 ± 0.35	1.14 ± 0.38	1.13 ± 0.26
2214.1	Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>4</sub>	1.40 ± 0.30	1.37 ± 0.31	1.14 ± 0.32	1.35 ± 0.32	1.30 ± 0.26	1.26 ± 0.24
2244.1	Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	1.73 ± 0.31	1.54 ± 0.29	1.64 ± 0.56	1.54 ± 0.33	1.87 ± 0.45	1.72 ± 0.37
2285.2	Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	6.64 ± 1.28	6.54 ± 0.96	5.93 ± 2.17	6.41 ± 1.44	6.31 ± 1.17	6.24 ± 1.04
2431.2	Neu5Ac <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	2.97 ± 0.85	3.24 ± 1.24	2.90 ± 1.64	2.73 ± 0.94	3.00 ± 1.12	2.74 ± 1.13
2459.2	Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	6.34 ± 1.37	7.11 ± 1.46	5.34 ± 2.22	6.19 ± 1.65	7.11 ± 1.44	8.00 ± 1.91
2489.3	Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	1.94 ± 0.40	2.11 ± 0.38	1.78 ± 0.60	1.85 ± 0.39	1.87 ± 0.34	1.97 ± 0.42
2592.3	Fuc <sub>3</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	1.09 ± 0.29	1.10 ± 0.27	0.95 ± 0.35	0.98 ± 0.36	1.07 ± 0.26	1.03 ± 0.24
2605.3	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	3.23 ± 0.60	3.04 ± 0.44	2.73 ± 0.91	2.55 ± 0.63	2.91 ± 0.57	2.68 ± 0.55
2646.3	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	2.17 ± 0.51	2.08 ± 0.49	1.82 ± 0.70	1.75 ± 0.60	1.75 ± 0.48	1.63 ± 0.42
2792.4	Neu5Ac <sub>2</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	8.58 ± 3.30	8.87 ± 3.47	8.16 ± 4.82	7.62 ± 4.13	7.66 ± 3.60	7.07 ± 4.21
2820.4	Neu5Ac <sub>1</sub> Fuc <sub>2</sub> Hex <sub>4</sub> HexNAc <sub>5</sub>	1.03 ± 0.37	1.02 ± 0.28	0.91 ± 0.37	0.89 ± 0.38	0.92 ± 0.32	0.93 ± 0.35
2837.4	Fuc <sub>3</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	1.10 ± 0.35	1.23 ± 0.34	1.03 ± 0.46	1.01 ± 0.42	1.19 ± 0.40	1.28 ± 0.34
2850.4	Neu5Ac <sub>1</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>5</sub>	2.12 ± 0.57	2.23 ± 0.59	1.90 ± 0.88	1.85 ± 0.71	1.80 ± 0.55	1.75 ± 0.60
2966.5	Neu5Ac <sub>2</sub> Fuc <sub>1</sub> Hex <sub>5</sub> HexNAc <sub>4</sub>	2.64 ± 0.85	2.35 ± 0.53	2.25 ± 1.11	2.07 ± 1.09	2.03 ± 0.71	1.87 ± 0.92
other		20.24	19.70	25.56	21.05	19.70	20.23

Table S8: Mean relative areas and standard deviations of O-glycans isolated from the three patient cohorts as judged by MALDI-TOF-MS. AD: Alzheimer's disease, DC: disease control and HC: healthy control. As m/z 273.1 and 1124.6 were gender-dependent (Supplementary Material Table S3), mean relative areas and standard deviations were calculated separately.

m/z	Composition	AD patients [%]		Disease control patients [%]		Healthy control patients [%]	
		Males	Females	Males	Females	Males	Females
<b>273.1</b>	Hex <sub>1</sub>	22.9 ± 11.1	16.3 ± 10.5	15.3 ± 5.8	15.0 ± 8.2	12.7 ± 4.9	13.2 ± 4.4
<b>1124.6</b>	Neu5Ac <sub>1</sub> Hex <sub>1</sub> HexNAc <sub>2</sub>	0.8 ± 0.6	0.7 ± 0.4	0.8 ± 0.5	0.6 ± 0.3	1.0 ± 0.7	0.9 ± 0.4
<b>314.2</b>	HexNAc <sub>1</sub>	44.2 ± 11.2		50.0 ± 10.6		45.0 ± 12.1	
<b>518.3</b>	Hex <sub>1</sub> HexNAc <sub>1</sub>	8.2 ± 2.0		10.1 ± 4.3		8.5 ± 2.5	
<b>559.3</b>	HexNAc <sub>2</sub>	5.4 ± 1.4		4.3 ± 1.3		4.9 ± 1.1	
<b>675.3</b>	Neu5Ac <sub>1</sub> HexNAc <sub>1</sub>	3.0 ± 1.3		2.8 ± 1.6		3.6 ± 1.3	
<b>722.4</b>	Hex <sub>2</sub> HexNAc <sub>1</sub>	3.6 ± 1.1		3.1 ± 1.1		4.2 ± 1.2	
<b>763.8</b>	Hex <sub>1</sub> HexNAc <sub>2</sub>	3.0 ± 1.1		2.6 ± 1.1		3.5 ± 1.2	
<b>804.4</b>	HexNAc <sub>3</sub>	2.3 ± 1.5		2.4 ± 1.5		3.3 ± 1.8	
<b>825.4</b>	Fuc <sub>2</sub> Hex <sub>2</sub>	2.3 ± 1.0		2.1 ± 1.0		2.9 ± 1.2	
<b>879.4</b>	Neu5Ac <sub>1</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	3.0 ± 1.2		2.7 ± 1.5		4.3 ± 2.1	
<b>967.5</b>	Hex <sub>2</sub> HexNAc <sub>2</sub>	2.0 ± 1.0		1.9 ± 1.1		2.5 ± 1.3	
<b>1240.6</b>	Neu5Ac <sub>2</sub> Hex <sub>1</sub> HexNAc <sub>1</sub>	0.6 ± 0.5		0.6 ± 0.6		1.2 ± 1.0	

Table S9: Glycan types and Glycosylation features used to generate the mean relative intensities presented in Figure 3. Bold: sialylated glycans.

Glycan type	Subcategory	Relative intensities of MS signals at m/z
Free	N: N-glycans	733.4, 926.5, 1130.6, 1171.6, 1334.7
	O: O-glycans	825.4, 967.5, 1212.6, <b>1240.6, 1601.8</b>
	Fragments (N-, O-glycans)	273.1, 314.2, 518.3, 559.3, <b>675.3, 722.4, 879.4</b>
	Free Sialic Acids	<b>430.2, 791.4</b>
N-glycans	high-mannose	1579.8, 1783.9
	biantennary	1661.8, 1835.9, 2040.0, 2070.0, 2214.1, 2244.1, <b>2431.2, 2592.3, 2605.3, 2792.4, 2966.5</b>
	bisected biantennary	2081.1, 2111.1, 2285.2, 2459.2, 2489.3, <b>2646.3, 2820.4, 2837.4, 2850.4</b>
O-glycans		273.1, 314.2, 518.3, 559.3, <b>675.3, 722.4, 763.4, 804.4, 825.4, 879.4, 967.5, 1124.6, 1240.6</b>

Table S10. Correlations were calculated between N-glycans groups by features and t-Tau using the Spearman's rank correlation coefficient, p<0.05 and p<0.01 were considered as statistically significant and very significant.

Group	Spearman	
	Males	Females
<b>high-mannose</b>	0.018	0.616
<b>asialylated biantennary</b>	0.183	0.106
<b>sialylated biantennary</b>	0.889	0.163
<b>asialylated bisected</b>	0.041	0.018
<b>sialylated bisected</b>	0.000	0.000