

Supplementary materials

GlycoMine^{struct}: a new bioinformatics tool for highly accurate mapping of the human N-linked and O-linked glycoproteomes by incorporating structural features

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Supplemental methods

In this study, we used six measures, including Sensitivity (or true positive rate (TPR)), Specificity, Precision (PRE), Accuracy (ACC), Mathew's Correlation Coefficient (MCC), and the area under the curve (AUC) to evaluate the prediction performance of GlycoMine^{struct}. The first five measures are defined as follows:

Sensitivity:

$$\text{Sensitivity} = \text{TP}/(\text{TP} + \text{FN}),$$

Specificity:

$$\text{Specificity} = \frac{\text{TN}}{\text{TN} + \text{FP}},$$

Precision:

$$\text{Precision} = \frac{\text{TP}}{\text{TP} + \text{FP}},$$

Accuracy:

$$\text{Accuracy} = \frac{\text{TP} + \text{TN}}{\text{TP} + \text{TN} + \text{FP} + \text{FN}}$$

Mathew's Correlation Coefficient (MCC):

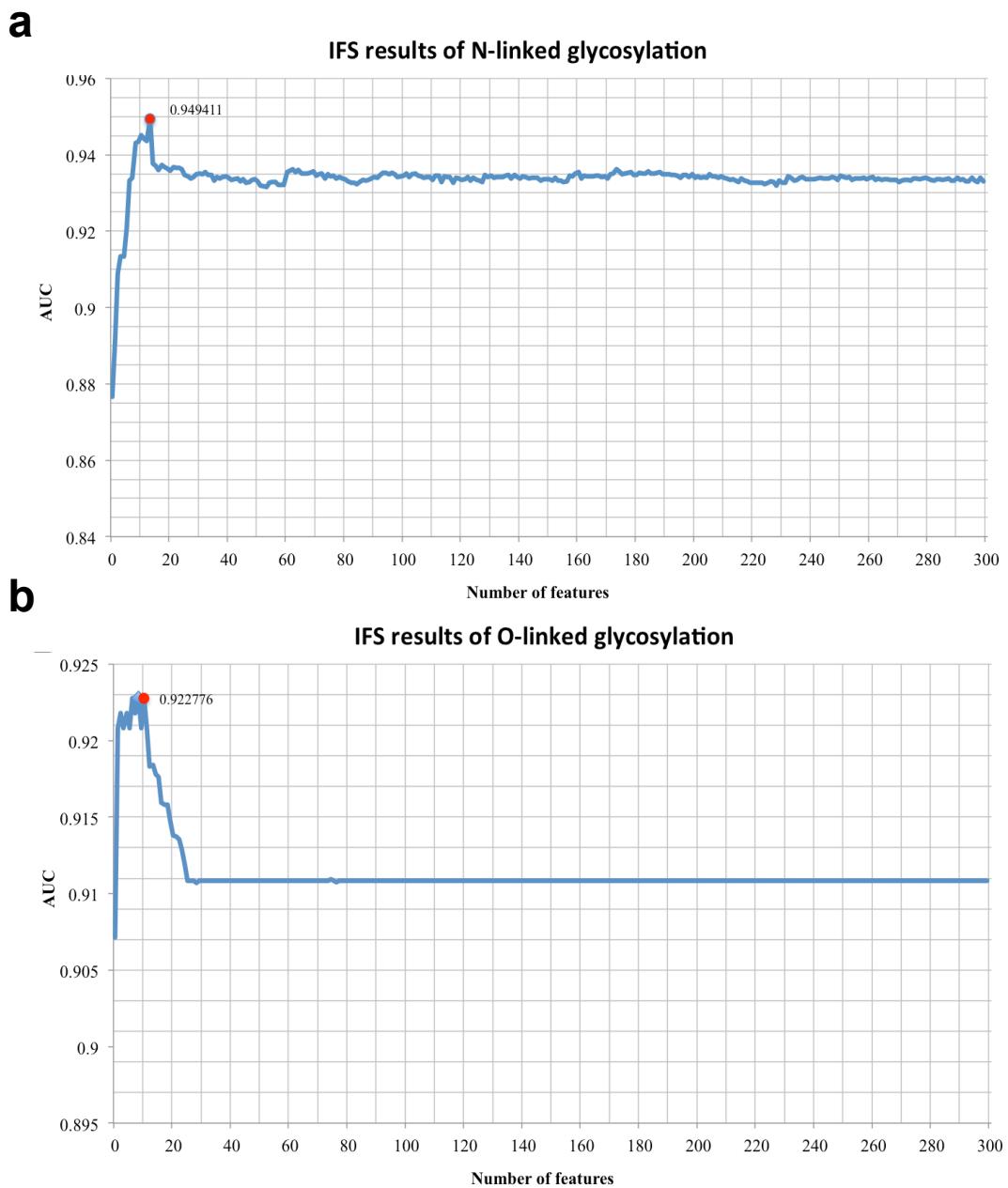
$$\text{MCC} = \frac{\text{TP} \times \text{TN} - \text{FP} \times \text{FN}}{\sqrt{(\text{TP} + \text{FP})(\text{TP} + \text{FN})(\text{TN} + \text{FP})(\text{TN} + \text{FN})}}$$

where TP , TN , FP and FN represents the numbers of true positives, true negatives, false positives and false negatives, respectively.

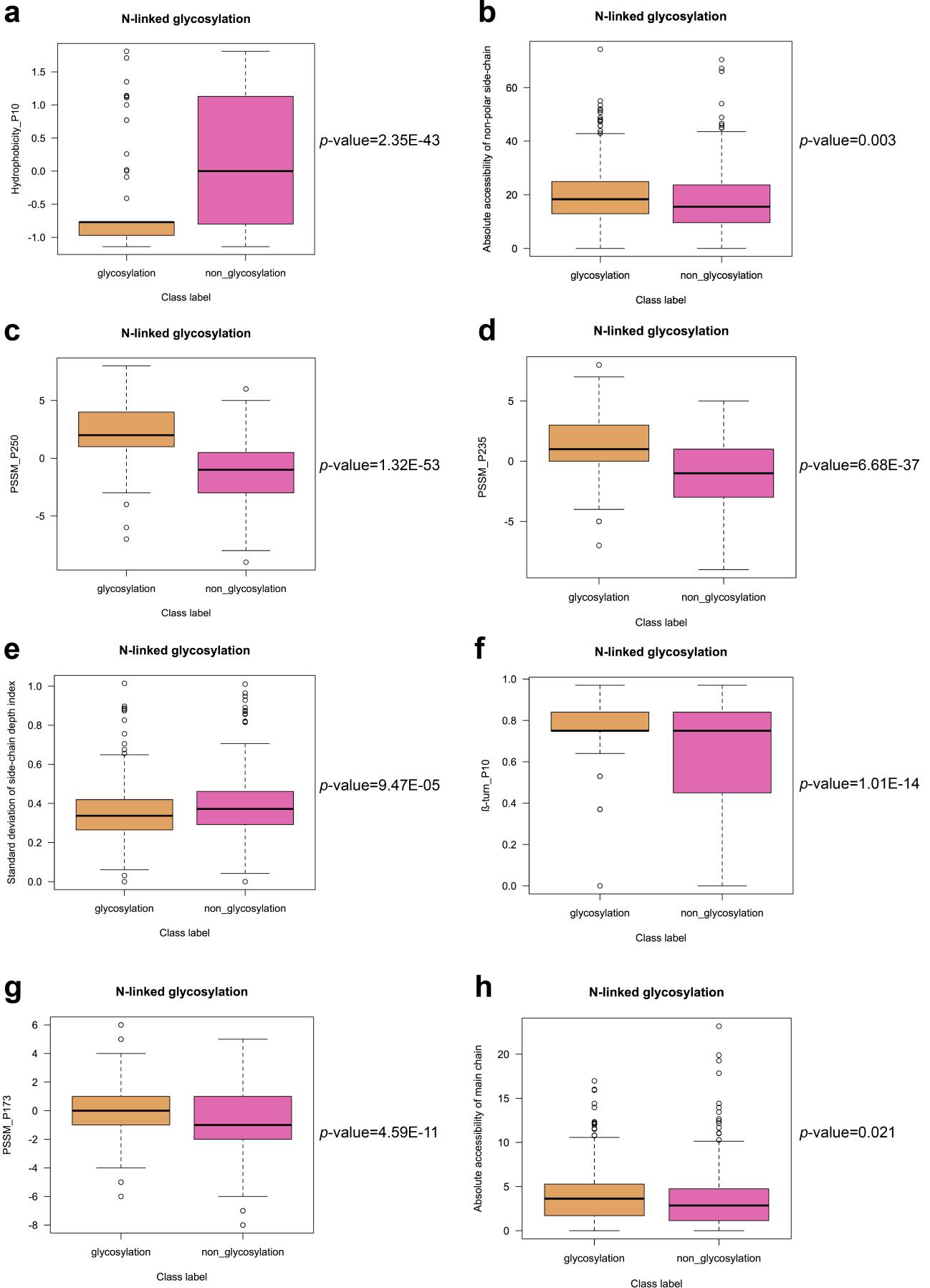
Supplemental results

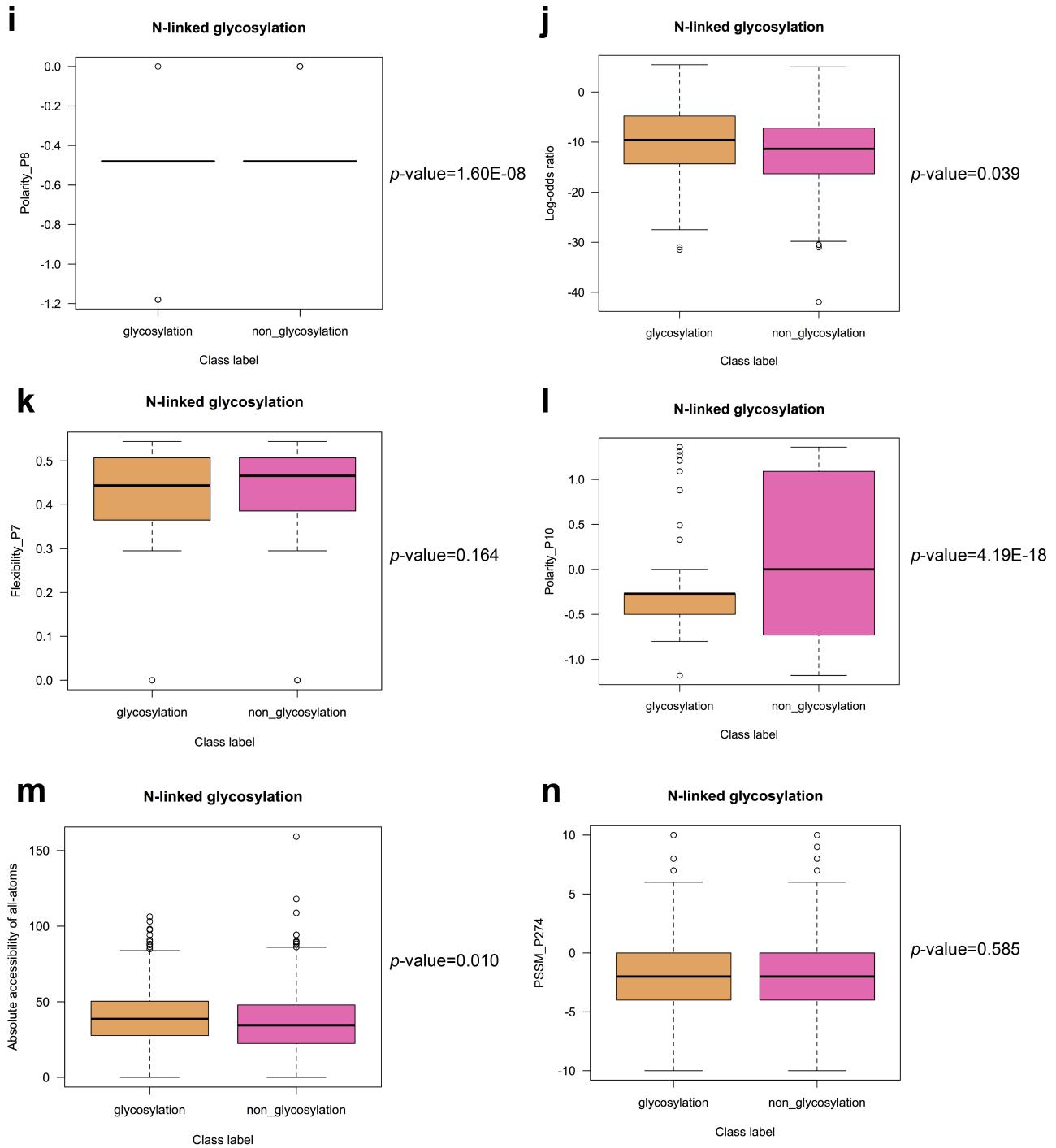
Analysis of significantly enriched Gene Ontology (GO) terms for N- and O-glycosylated proteins

We collected GO terms for all the N- and O-glycosylated proteins in our benchmark dataset and conducted analyses of significantly enriched GO terms for both types of glycosylated proteins. The GO term annotations were extracted from the UniProt database¹ using the identifier for each protein. For statistical analysis, for each type of glycosylation, we prepared two datasets, i.e. the positive dataset (with glycosylated proteins) and the background set (from human proteins). We performed the two-side hypergeometric test to find significantly enriched GO terms from the positive set compared with the background set. The results for N- and O-glycosylation are respectively shown in **Supplementary Tables S4 and S5**. The GO term annotations involve three subcategories, namely cellular component (CC), molecular function (MF) and biological process (BP). For CC, it is clear from **Supplementary Tables S4 and S5** that O-glycosylated proteins possess more GO terms (110 GO terms) compared with N-glycosylated proteins (17 GO terms). 17 significantly enrich GO terms for N-glycosylated proteins include “Endoplasmic reticulum”, “Cytosol”, “Mitochondrion”, “Mitochondrial matrix”, “Centrosome”, “Spindle pole”, “Golgi apparatus”, “Membrane”, “Cell surface”, “Extracellular region”, “Integral component of membrane”, “Lamellipodium”, “Extracellular space”, “Endoplasmic reticulum membrane”, “Endolysosome membrane”, “Beta-catenin destruction complex”, and “Endolysosome lumen”. While for O-linked glycosylation, the top ten significantly enriched GO terms are “ATP binding”, “Proteolysis”, “Negative regulation of apoptotic process”, “Glycolytic process”, and “Cellular response to lipopolysaccharide”.

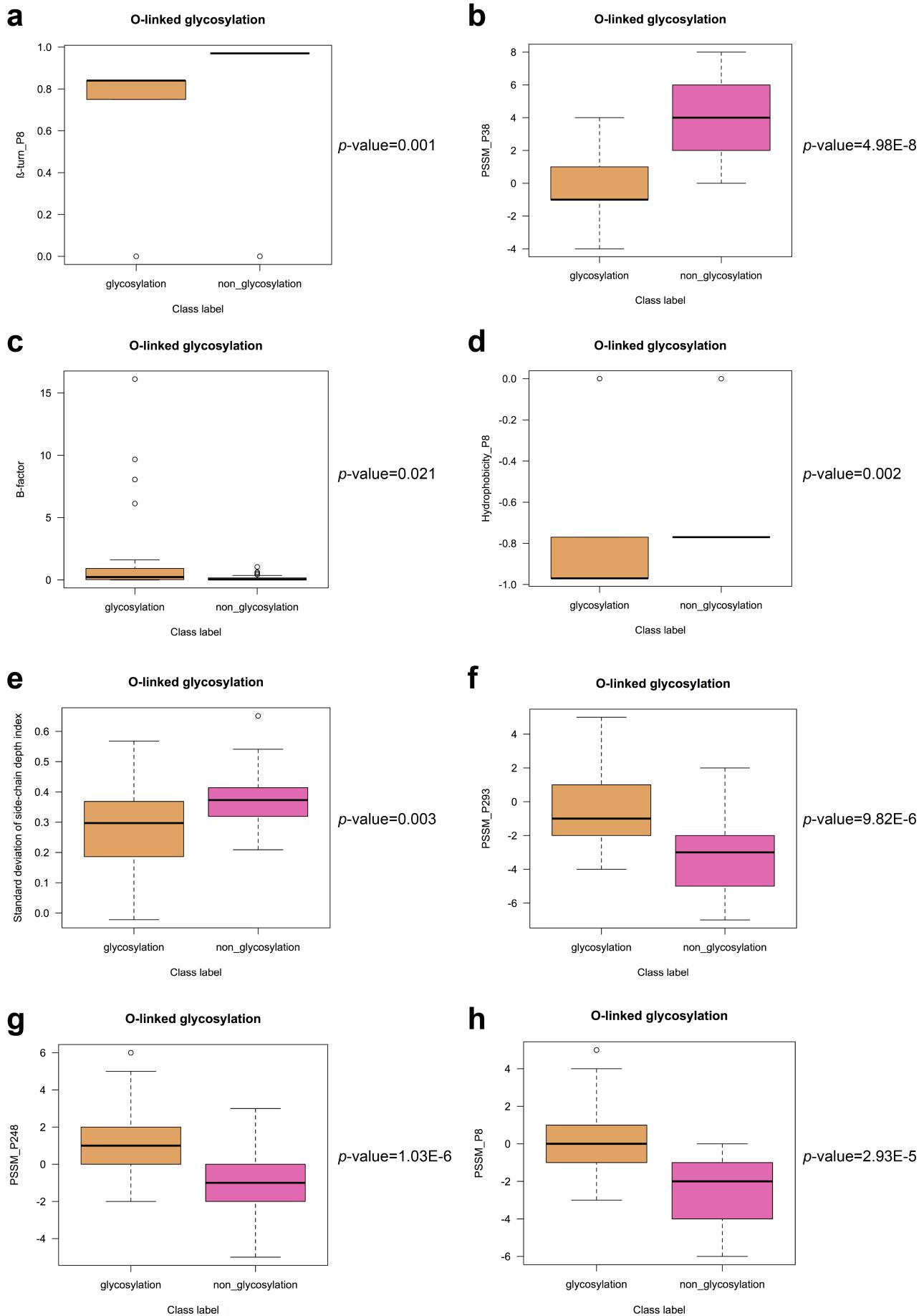


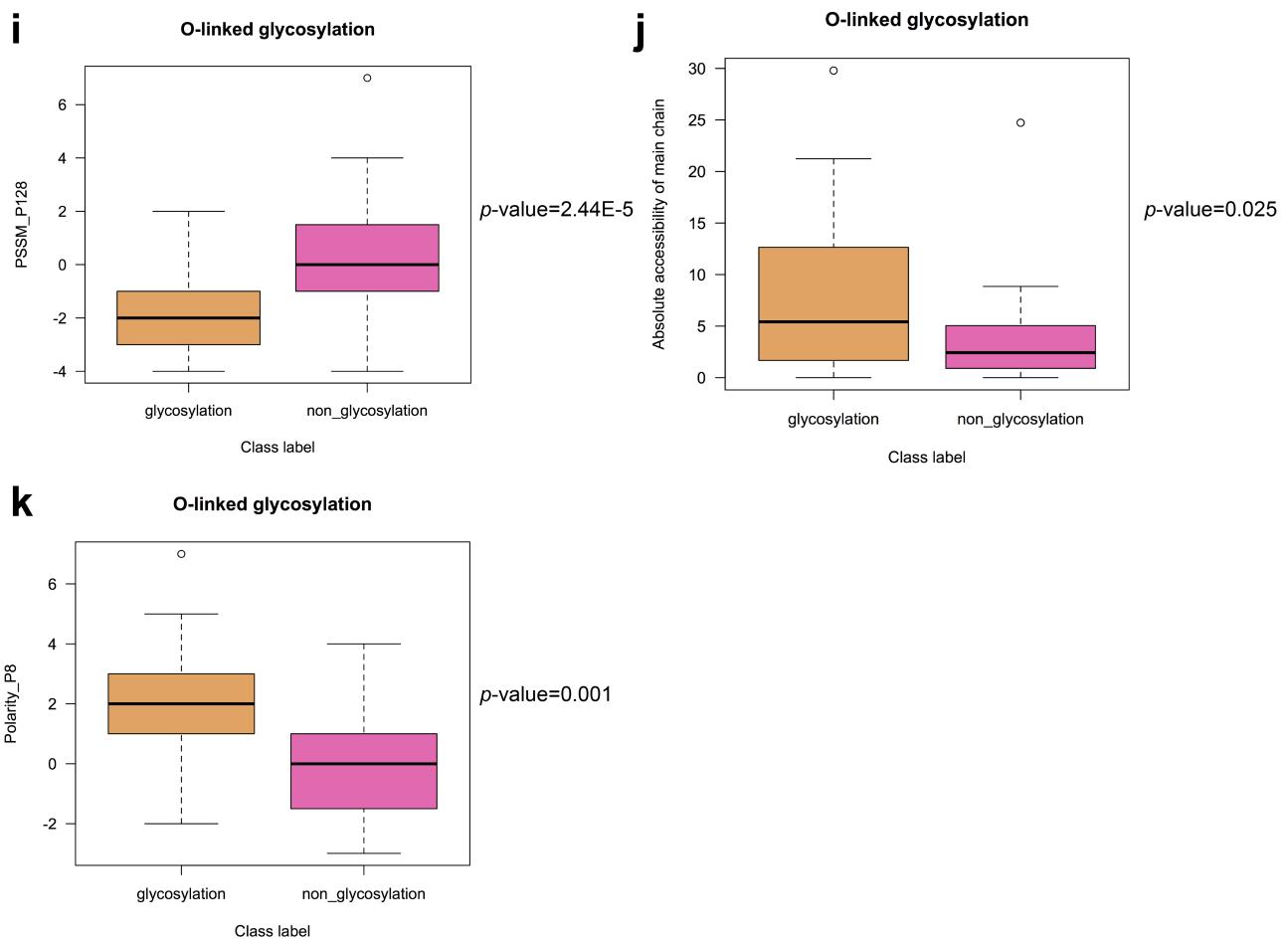
Supplementary Figure S1. IFS curves for the prediction of (a) N- and (b) O-linked glycosylation sites. The values highlighted in the graph correspond to the highest AUC scores obtained by IFS for N- and O-linked glycosylation, respectively.





Supplementary Figure S2. Boxplots showing the significant difference in feature value distributions of the optimal sequence and structural features listed in Table 1 for N-linked glycosylation prediction.





Supplementary Figure S3. Boxplots showing significant difference in feature value distributions of the optimal sequence and structural features listed in Table 2 for O-linked glycosylation prediction.

a

Introduction

Glycosylation is a ubiquitous type of protein post-translational modification (PTM) in eukaryotic cells, which plays vital roles in various biological processes such as cellular communication, ligand recognition, and subcellular recognition. It is estimated that >50% of the entire human proteome is glycosylated.

We present a novel bioinformatics tool called GlycoMine^{struct}, which is a comprehensive tool for the systematic *in silico* identification of N- and O-linked glycosylation sites in the human structural proteome. GlycoMine^{struct} was developed using the random forest algorithm and evaluated based on a well-prepared up-to-date benchmark dataset that encompasses all three types of glycosylation sites, which was curated from multiple public resources.

Online Web Server

b

The screenshot shows the "Online Web Server" page. The sidebar menu is identical to the homepage. The main area has a heading "Please upload a PDB File" with a file input field containing "选择文件 未选择任何文件". To its right are dropdown menus for "Type: N-linked" and "Email: lrtuy774@gmail.com". Below these are buttons for "Submit" and "Clear". A text input field contains "3WN4:A". The bottom part of the screen displays a table of prediction results:

Protein	Chain	Position	Residues	Probability
3WN4	A	247	ASN	0.944
3WN4	A	582	ASN	0.943
3WN4	A	511	ASN	0.938
3WN4	A	285	ASN	0.927
3WN4	A	293	ASN	0.916
3WN4	A	590	ASN	0.898
3WN4	A	362	ASN	0.887
3WN4	A	640	ASN	0.875
3WN4	A	680	ASN	0.866

Supplementary Figure S4. Screenshots of GlycoMine^{struct} web server. (a) The user interface and **(b)** Display of the prediction output.

Supplementary Table S1. The prediction performance of N- and O-linked glycosylation sites based on 5-fold cross-validation and independent tests. The performance was evaluated using six measures including AUC, MCC, ACC, Sensitivity, Specificity, and Precision.

			AUC	MCC	ACC	Sensitivity	Specificity	Precision
All OFS features	Benchmark	N-linked	0.949	0.838	0.919	0.936	0.901	0.912
		O-linked	0.923	0.814	0.906	0.871	0.939	0.931
	Independent	N-linked	0.941	0.830	0.912	0.972	0.853	0.867
		O-linked	0.922	0.805	0.895	0.778	0.979	0.972
Structural features only	Benchmark	N-linked	0.911	0.734	0.865	0.823	0.911	0.910
		O-linked	0.836	0.660	0.828	0.871	0.788	0.794
	Benchmark	N-linked	0.880	0.695	0.846	0.811	0.883	0.884
Sequence features only	Benchmark	O-linked	0.814	0.547	0.766	0.871	0.667	0.711
NGlycPred	Independent	N-linked	0.804	0.685	0.840	0.785	0.896	0.884

Supplementary Table S2. Statistical summary of the proteome-wide prediction of N- and O-linked glycosylation sites at the 99% specificity level. The prediction was performed for the whole human structural proteome with a total of 20538 proteins.

Glycosylation Type	Number of predicted glycosylated proteins	Number of predicted glycosylation sites
N-linked	3386	4996
O-linked	5298	10529

Supplementary Table S3. Lists of all initial features explored in this study. (a)

Sequence-derived features and **(b)** Structural features.

(a)

Feature type	Description	Software	Number of features
AAindex	The hydrophobicity score, flexibility score, polarity score and β -turn score	AAindex ²	$4 \times 15 = 60$
Physicochemical properties	14 physicochemical properties	BioJava ³	14
PSSMs	Position-specific scoring matrices	PSI-BLAST ⁴	$20 \times 15 = 300$
Residue conversation score	Residue conversation	PSI-BLAST ⁴	$1 \times 15 = 15$

(b)

Feature type	Description	Software	Number of features
Surface accessibility	All-atoms, total-side, main-chain, non-polar and all-polar accessibilities	NACCESS ⁵	5
Secondary structure	ACC: the number of water molecules in contact with corresponding residue; PHI and PSI: two IUPAC peptide backbone torsion angles.	DSSP ⁶	3
Log-odds ratio	Log-odds ratio is a statistical feature	DiscoTope ⁷	1
Depth Index	The average depth, standard deviation of depth index, side-chain average depth index and standard deviation of side-chain depth index	PSAIA ⁸	4
B-factor	The average B-factor of all the atoms in this residue	PDB file ⁹	1

Supplementary Table S4. Significantly enriched GO terms for N-linked glycosylated proteins.

Type	ID	Term in the		P-value	Term
		the positive	human proteome		
CC	GO:0005783	17	120	1.72E-3	Endoplasmic reticulum
	GO:0005829	97	1106	7.07E-4	Cytosol
	GO:0005739	32	271	6.50E-4	Mitochondrion
	GO:0005759	13	95	6.86E-3	Mitochondrial matrix
	GO:0005813	12	86	7.82E-3	Centrosome
	GO:0000922	6	27	6.56E-3	Spindle pole
	GO:0005794	20	176	7.63E-3	Golgi apparatus
	GO:0016020	17	122	2.03E-3	Membrane
	GO:0009986	16	132	8.82E-3	Cell surface
	GO:0005576	36	322	8.24E-4	Extracellular region
	GO:0016021	31	306	6.27E-3	Integral component of membrane
	GO:0030027	8	47	9.02E-3	Lamellipodium
	GO:0005615	33	327	5.35E-3	Extracellular space
	GO:0005789	14	109	8.58E-3	Endoplasmic reticulum membrane
	GO:0036020	2	2	4.62E-3	Endolysosome membrane
MF	GO:0005524	65	537	5.79E-7	ATP binding
	GO:0004674	17	129	3.46E-3	Protein serine/threonine kinase activity
	GO:0009055	12	68	1.28E-3	Electron carrier activity
	GO:0004252	10	63	6.24E-3	Serine-type endopeptidase activity
	GO:0008201	9	55	7.53E-3	Heparin binding
	GO:0008144	6	28	7.79E-3	Drug binding
	GO:0005506	10	66	8.37E-3	Iron ion binding
	GO:0016887	4	13	8.06E-3	Atpase activity
	GO:0004222	7	34	5.28E-3	Metalloendopeptidase activity
	GO:0004716	4	8	1.12E-3	Receptor signaling protein tyrosine kinase

					activity
GO:0004197	6	25	4.51E-3	Cysteine-type endopeptidase activity	
GO:0005149	2	2	4.62E-3	Interleukin-1 receptor binding	
GO:0005536	3	5	2.72E-3	Glucose binding	
GO:0043169	5	16	2.89E-3	Cation binding	
				Hydrolase activity, acting on	
GO:0016812	2	2	4.62E-3	Carbon-nitrogen (but not peptide) bonds, in cyclic amides	
GO:0008013	3	6	5.07E-3	Beta-catenin binding	
GO:0004029	3	6	5.07E-3	Aldehyde dehydrogenase (NAD) activity	
GO:0016303	3	4	1.17E-3	1-phosphatidylinositol-3-kinase activity	
GO:0015038	2	2	4.62E-3	Glutathione disulfide oxidoreductase activity	
GO:0046934	2	2	4.62E-3	Phosphatidylinositol-4,5-bisphosphate 3-kinase activity	
BP	GO:0045087	26	200	5.07E-4	Innate immune response
	GO:0043066	32	212	7.23E-6	Negative regulation of apoptotic process
	GO:0006508	28	170	5.21E-6	Proteolysis
	GO:0006006	7	37	8.21E-3	Glucose metabolic process
	GO:0070207	4	8	1.12E-3	Protein homotrimerization
	GO:0044281	26	198	4.38E-4	Small molecule metabolic process
	GO:0007409	6	18	7.69E-4	Axonogenesis
	GO:0046777	13	94	6.34E-3	Protein autophosphorylation
	GO:0002250	4	11	4.28E-3	Adaptive immune response
	GO:0001816	4	9	1.88E-3	Cytokine production
	GO:0071222	8	29	4.27E-4	Cellular response to lipopolysaccharide
	GO:0045080	3	7	8.28E-3	Positive regulation of chemokine biosynthetic process
	GO:0042493	16	133	9.36E-3	Response to drug
	GO:0030168	15	111	4.52E-3	Platelet activation
	GO:0002576	9	55	7.53E-3	Platelet degranulation
	GO:0006935	5	21	9.49E-3	Chemotaxis
	GO:0071260	7	33	4.50E-3	Cellular response to mechanical stimulus

GO:0045078	3	6	5.07E-3	Positive regulation of interferon-gamma biosynthetic process
GO:0006096	8	25	1.42E-4	Glycolytic process
GO:0005980	3	7	8.28E-3	Glycogen catabolic process
GO:0046686	5	14	1.52E-3	Response to cadmium ion
GO:0000281	3	7	8.28E-3	Mitotic cytokinesis
GO:0007160	7	26	1.12E-3	Cell-matrix adhesion
GO:0045780	3	7	8.28E-3	Positive regulation of bone resorption
GO:0050435	3	5	2.72E-3	Beta-amyloid metabolic process
GO:0007040	4	8	1.12E-3	Lysosome organization
GO:0007494	2	2	4.62E-3	Midgut development
GO:0060444	4	8	1.12E-3	Branching involved in mammary gland duct morphogenesis
GO:0030316	4	11	4.28E-3	Osteoclast differentiation
GO:0033628	3	7	8.28E-3	Regulation of cell adhesion mediated by integrin
GO:0002437	3	7	8.28E-3	Inflammatory response to antigenic stimulus
GO:0032611	2	2	4.62E-3	Interleukin-1 beta production
GO:0042346	4	11	4.28E-3	Positive regulation of NF-kappab import into nucleus
GO:0003073	2	2	4.62E-3	Regulation of systemic arterial blood pressure
GO:0030207	3	5	2.72E-3	Chondroitin sulfate catabolic process
GO:0045356	2	2	4.62E-3	Positive regulation of interferon-alpha biosynthetic process
GO:0060740	3	6	5.07E-3	Prostate gland epithelium morphogenesis
GO:0042733	4	9	1.88E-3	Embryonic digit morphogenesis
GO:0061198	2	2	4.62E-3	Fungiform papilla formation
GO:0060789	3	4	1.17E-3	Hair follicle placode formation
GO:0010042	2	2	4.62E-3	Response to manganese ion
GO:0071372	3	5	2.72E-3	Cellular response to follicle-stimulating hormone stimulus

GO:0003420	2	2	4.62E-3	Regulation of growth plate cartilage chondrocyte proliferation
GO:0070527	3	4	1.17E-3	Platelet aggregation
GO:0014909	3	4	1.17E-3	Smooth muscle cell migration
GO:0042117	2	2	4.62E-3	Monocyte activation
GO:0010641	2	2	4.62E-3	Positive regulation of platelet-derived growth factor receptor signaling pathway
GO:0050927	3	5	2.72E-3	Positive regulation of positive chemotaxis
GO:0035116	4	11	4.28E-3	Embryonic hindlimb morphogenesis
GO:0032331	3	6	5.07E-3	Negative regulation of chondrocyte differentiation
GO:0033146	2	2	4.62E-3	Regulation of intracellular estrogen receptor signaling pathway
GO:0050790	3	4	1.17E-3	Regulation of catalytic activity
GO:0051131	2	2	4.62E-3	Chaperone-mediated protein complex assembly
GO:0019953	2	2	4.62E-3	Sexual reproduction
GO:0055118	2	2	4.62E-3	Negative regulation of cardiac muscle contraction
GO:0048025	2	2	4.62E-3	Negative regulation of mrna splicing, via spliceosome
GO:0009966	3	4	1.17E-3	Regulation of signal transduction Canonical Wnt signaling pathway
GO:0061324	2	2	4.62E-3	involved in positive regulation of cardiac outflow tract cell proliferation
GO:0003344	2	2	4.62E-3	Pericardium morphogenesis

Supplementary Table S5. Significantly enriched GO terms for O-linked glycosylated proteins.

Type	ID	Term in the positive	Term in the human proteome	P-value	Term
CC	GO:0005783	17	120	1.72E-03	Endoplasmic reticulum
	GO:0005829	97	1106	7.07E-04	Cytosol
	GO:0005739	32	271	6.50E-04	Mitochondrion
	GO:0005759	13	95	6.86E-03	Mitochondrial matrix
	GO:0000922	6	27	6.56E-03	Spindle pole
	GO:0005794	20	176	7.63E-03	Golgi apparatus
	GO:0016020	17	122	2.03E-03	Membrane
	GO:0009986	16	132	8.82E-03	Cell surface
	GO:0005576	36	322	8.24E-04	Extracellular region
	GO:0016021	31	306	6.27E-03	Integral component of membrane
	GO:0030027	8	47	9.02E-03	Lamellipodium
	GO:0005615	33	327	5.35E-03	Extracellular space
	GO:0005789	14	109	8.58E-03	Endoplasmic reticulum membrane
	GO:0036020	2	2	4.62E-03	Endolysosome membrane
	GO:0030877	3	7	8.28E-03	Beta-catenin destruction complex
	GO:0036021	3	4	1.17E-03	Endolysosome lumen
	GO:0005524	65	537	5.79E-07	ATP binding
	GO:0004674	17	129	3.46E-03	Protein serine/threonine kinase activity
	GO:0009055	12	68	1.28E-03	Electron carrier activity
	GO:0004252	10	63	6.24E-03	Serine-type endopeptidase activity
	GO:0008201	9	55	7.53E-03	Heparin binding
	GO:0008144	6	28	7.79E-03	Drug binding
	GO:0005506	10	66	8.37E-03	Iron ion binding
	GO:0016887	4	13	8.06E-03	Atpase activity
	GO:0004222	7	34	5.28E-03	Metalloendopeptidase activity
	GO:0004716	4	8	1.12E-03	Receptor signaling protein tyrosine kinase activity

GO:0004197	6	25	4.51E-03	Cysteine-type endopeptidase activity
GO:0005149	2	2	4.62E-03	Interleukin-1 receptor binding
GO:0005536	3	5	2.72E-03	Glucose binding
GO:0043169	5	16	2.89E-03	Cation binding
				Hydrolase activity, acting on
GO:0016812	2	2	4.62E-03	Carbon-nitrogen (but not peptide) bonds, in cyclic amides
GO:0008013	3	6	5.07E-03	Beta-catenin binding
GO:0004029	3	6	5.07E-03	Aldehyde dehydrogenase (NAD) activity
GO:0016303	3	4	1.17E-03	1-phosphatidylinositol-3-kinase activity
GO:0015038	2	2	4.62E-03	Glutathione disulfide oxidoreductase activity
GO:0046934	2	2	4.62E-03	Phosphatidylinositol-4,5-bisphosphate 3-kinase activity
GO:0045087	26	200	5.07E-04	Innate immune response
GO:0043066	32	212	7.23E-06	Negative regulation of apoptotic process
GO:0006508	28	170	5.21E-06	Proteolysis
GO:0006006	7	37	8.21E-03	Glucose metabolic process
GO:0070207	4	8	1.12E-03	Protein homotrimerization
GO:0044281	26	198	4.38E-04	Small molecule metabolic process
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GO:0045078	3	6	5.07E-03	Positive regulation of interferon-gamma biosynthetic process
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GO:0000281	3	7	8.28E-03	Mitotic cytokinesis
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GO:0045780	3	7	8.28E-03	Positive regulation of bone resorption
GO:0050435	3	5	2.72E-03	Beta-amyloid metabolic process
GO:0007040	4	8	1.12E-03	Lysosome organization
GO:0007494	2	2	4.62E-03	Midgut development
GO:0060444	4	8	1.12E-03	Branching involved in mammary gland duct morphogenesis
GO:0030316	4	11	4.28E-03	Osteoclast differentiation
GO:0033628	3	7	8.28E-03	Regulation of cell adhesion mediated by integrin
GO:0002437	3	7	8.28E-03	Inflammatory response to antigenic stimulus
GO:0032611	2	2	4.62E-03	Interleukin-1 beta production
GO:0042346	4	11	4.28E-03	Positive regulation of NF-kappab import into nucleus
GO:0003073	2	2	4.62E-03	Regulation of systemic arterial blood pressure
GO:0030207	3	5	2.72E-03	Chondroitin sulfate catabolic process
GO:0045356	2	2	4.62E-03	Positive regulation of interferon-alpha biosynthetic process
GO:0060740	3	6	5.07E-03	Prostate gland epithelium morphogenesis
GO:0042733	4	9	1.88E-03	Embryonic digit morphogenesis
GO:0061198	2	2	4.62E-03	Fungiform papilla formation
GO:0060789	3	4	1.17E-03	Hair follicle placode formation
GO:0010042	2	2	4.62E-03	Response to manganese ion
GO:0071372	3	5	2.72E-03	Cellular response to follicle-stimulating

				hormone stimulus
GO:0003420	2	2	4.62E-03	Regulation of growth plate cartilage chondrocyte proliferation
GO:0070527	3	4	1.17E-03	Platelet aggregation
GO:0014909	3	4	1.17E-03	Smooth muscle cell migration
GO:0042117	2	2	4.62E-03	Monocyte activation
GO:0010641	2	2	4.62E-03	Positive regulation of platelet-derived growth factor receptor signaling pathway
GO:0050927	3	5	2.72E-03	Positive regulation of positive chemotaxis
GO:0035116	4	11	4.28E-03	Embryonic hindlimb morphogenesis
GO:0032331	3	6	5.07E-03	Negative regulation of chondrocyte differentiation
GO:0050790	3	4	1.17E-03	Regulation of catalytic activity
GO:0051131	2	2	4.62E-03	Chaperone-mediated protein complex assembly
GO:0019953	2	2	4.62E-03	Sexual reproduction
GO:0055118	2	2	4.62E-03	Negative regulation of cardiac muscle contraction
GO:0048025	2	2	4.62E-03	Negative regulation of mrna splicing, via spliceosome
GO:0009966	3	4	1.17E-03	Regulation of signal transduction
GO:0061324	2	2	4.62E-03	Canonical Wnt signaling pathway involved in positive regulation of cardiac outflow tract cell proliferation
GO:0003344	2	2	4.62E-03	Pericardium morphogenesis
GO:0005783	17	120	1.72E-03	Endoplasmic reticulum
GO:0005829	97	1106	7.07E-04	Cytosol
GO:0005739	32	271	6.50E-04	Mitochondrion
GO:0005759	13	95	6.86E-03	Mitochondrial matrix
GO:0000922	6	27	6.56E-03	Spindle pole
GO:0005794	20	176	7.63E-03	Golgi apparatus

	GO:0016020	17	122	2.03E-03	Membrane
	GO:0009986	16	132	8.82E-03	Cell surface
	GO:0005576	36	322	8.24E-04	Extracellular region
	GO:0016021	31	306	6.27E-03	Integral component of membrane
	GO:0030027	8	47	9.02E-03	Lamellipodium
	GO:0005615	33	327	5.35E-03	Extracellular space
	GO:0005789	14	109	8.58E-03	Endoplasmic reticulum membrane
	GO:0036020	2	2	4.62E-03	Endolysosome membrane
	GO:0030877	3	7	8.28E-03	Beta-catenin destruction complex
	GO:0036021	3	4	1.17E-03	Endolysosome lumen
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MF	GO:0005524	65	537	5.79E-07	ATP binding
	GO:0004674	17	129	3.46E-03	Protein serine/threonine kinase activity
	GO:0009055	12	68	1.28E-03	Electron carrier activity
	GO:0004252	10	63	6.24E-03	Serine-type endopeptidase activity
	GO:0008201	9	55	7.53E-03	Heparin binding
	GO:0008144	6	28	7.79E-03	Drug binding
	GO:0005506	10	66	8.37E-03	Iron ion binding
	GO:0016887	4	13	8.06E-03	Atpase activity
	GO:0004222	7	34	5.28E-03	Metalloendopeptidase activity
	GO:0004716	4	8	1.12E-03	Receptor signaling protein tyrosine kinase activity
	GO:0004197	6	25	4.51E-03	Cysteine-type endopeptidase activity
	GO:0005149	2	2	4.62E-03	Interleukin-1 receptor binding
	GO:0005536	3	5	2.72E-03	Glucose binding
	GO:0043169	5	16	2.89E-03	Cation binding
					Hydrolase activity, acting on carbon-nitrogen (but not peptide) bonds, in cyclic amides
	GO:0016812	2	2	4.62E-03	
	GO:0008013	3	6	5.07E-03	Beta-catenin binding
	GO:0004029	3	6	5.07E-03	Aldehyde dehydrogenase (NAD) activity
	GO:0016303	3	4	1.17E-03	1-phosphatidylinositol-3-kinase activity
	GO:0015038	2	2	4.62E-03	Glutathione disulfide oxidoreductase

				activity
GO:0046934	2	2	4.62E-03	Phosphatidylinositol-4,5-bisphosphate 3-kinase activity
BP	GO:0045087	26	200	5.07E-04 Innate immune response
	GO:0043066	32	212	7.23E-06 Negative regulation of apoptotic process
	GO:0006508	28	170	5.21E-06 Proteolysis
	GO:0006006	7	37	8.21E-03 Glucose metabolic process
	GO:0070207	4	8	1.12E-03 Protein homotrimerization
	GO:0044281	26	198	4.38E-04 Small molecule metabolic process
	GO:0007409	6	18	7.69E-04 Axonogenesis
	GO:0046777	13	94	6.34E-03 Protein autophosphorylation
	GO:0002250	4	11	4.28E-03 Adaptive immune response
	GO:0001816	4	9	1.88E-03 Cytokine production
	GO:0071222	8	29	4.27E-04 Cellular response to lipopolysaccharide
	GO:0045080	3	7	8.28E-03 Positive regulation of chemokine biosynthetic process
	GO:0042493	16	133	9.36E-03 Response to drug
	GO:0030168	15	111	4.52E-03 Platelet activation
	GO:0002576	9	55	7.53E-03 Platelet degranulation
	GO:0006935	5	21	9.49E-03 Chemotaxis
	GO:0071260	7	33	4.50E-03 Cellular response to mechanical stimulus
	GO:0045078	3	6	5.07E-03 Positive regulation of interferon-gamma biosynthetic process
	GO:0006096	8	25	1.42E-04 Glycolytic process
	GO:0005980	3	7	8.28E-03 Glycogen catabolic process
	GO:0046686	5	14	1.52E-03 Response to cadmium ion
	GO:0000281	3	7	8.28E-03 Mitotic cytokinesis
	GO:0007160	7	26	1.12E-03 Cell-matrix adhesion
	GO:0045780	3	7	8.28E-03 Positive regulation of bone resorption
	GO:0050435	3	5	2.72E-03 Beta-amyloid metabolic process
	GO:0007040	4	8	1.12E-03 Lysosome organization
	GO:0007494	2	2	4.62E-03 Midgut development

GO:0060444	4	8	1.12E-03	Branching involved in mammary gland duct morphogenesis
GO:0030316	4	11	4.28E-03	Osteoclast differentiation
GO:0033628	3	7	8.28E-03	Regulation of cell adhesion mediated by integrin
GO:0002437	3	7	8.28E-03	Inflammatory response to antigenic stimulus
GO:0032611	2	2	4.62E-03	Interleukin-1 beta production
GO:0042346	4	11	4.28E-03	Positive regulation of NF-kappab import into nucleus
GO:0003073	2	2	4.62E-03	Regulation of systemic arterial blood pressure
GO:0030207	3	5	2.72E-03	Chondroitin sulfate catabolic process
GO:0045356	2	2	4.62E-03	Positive regulation of interferon-alpha biosynthetic process
GO:0060740	3	6	5.07E-03	Prostate gland epithelium morphogenesis
GO:0042733	4	9	1.88E-03	Embryonic digit morphogenesis
GO:0061198	2	2	4.62E-03	Fungiform papilla formation
GO:0060789	3	4	1.17E-03	Hair follicle placode formation
GO:0010042	2	2	4.62E-03	Response to manganese ion
GO:0071372	3	5	2.72E-03	Cellular response to follicle-stimulating hormone stimulus
GO:0003420	2	2	4.62E-03	Regulation of growth plate cartilage chondrocyte proliferation
GO:0070527	3	4	1.17E-03	Platelet aggregation
GO:0014909	3	4	1.17E-03	Smooth muscle cell migration
GO:0042117	2	2	4.62E-03	Monocyte activation
GO:0010641	2	2	4.62E-03	Positive regulation of platelet-derived growth factor receptor signaling pathway
GO:0050927	3	5	2.72E-03	Positive regulation of positive chemotaxis
GO:0035116	4	11	4.28E-03	Embryonic hindlimb morphogenesis

GO:0032331	3	6	5.07E-03	Negative regulation of chondrocyte differentiation
GO:0050790	3	4	1.17E-03	Regulation of catalytic activity
GO:0051131	2	2	4.62E-03	Chaperone-mediated protein complex assembly
GO:0019953	2	2	4.62E-03	Sexual reproduction
GO:0055118	2	2	4.62E-03	Negative regulation of cardiac muscle contraction
GO:0048025	2	2	4.62E-03	Negative regulation of mrna splicing, via spliceosome
GO:0009966	3	4	1.17E-03	Regulation of signal transduction
GO:0061324	2	2	4.62E-03	Canonical Wnt signaling pathway involved in positive regulation of cardiac outflow tract cell proliferation
GO:0003344	2	2	4.62E-03	Pericardium morphogenesis

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