

Supplementary File

SOMAscan Proteomics Identifies Novel Plasma Proteins in Amyotrophic Lateral Sclerosis Patients

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Table S1. Upregulated and Downregulated proteins between ALS patients and neurologically healthy controls using SAM Analysis of SOMAscan Results.

Increased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
SH21A	SH2D1A	1.689103	0.019616	
TIMP-3	TIMP3	1.292908	0.003370	
PDGF-BB	PDGFB	1.205864	0.005048	
TARC	CCL17	1.081839	0.003579	
PKC-D	PRKCD	1.004368	0.011062	
Gro-b/g	CXCL3 CXCL2	1.003084	0.009649	
NID2	NID2	0.515259	0.011365	
Nidogen	NID1	0.419618	0.012383	
Decreased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
Ephrin-A2	EFNA2	-0.281950	0.024822	
RGMB	RGMB	-0.313691	0.018977	
VCAM-1	VCAM1	-0.315918	0.007538	
sTie-1	TIE1	-0.334705	0.015354	
TLR4	TLR4	-0.371138	0.006458	
RGM-C	HJV	-0.388838	0.008869	
IGF-I sR	IGF1R	-0.415958	0.025886	
BMPR1A	BMPR1A	-0.449027	0.015265	
LYPD3	LYPD3	-0.456034	0.011876	
NCAM-L1	L1CAM	-0.482127	0.003169	
BCAM	BCAM	-0.485299	0.031243	
sTie-2	TEK	-0.488869	0.014479	
Ephrin-A3	EFNA3	-0.514959	0.002038	
MASP3	MASP1	-0.536970	0.011929	
Kallikrein 7	KLK7	-0.553409	0.024028	
Nectin-like protein 2	CADM1	-0.559252	0.045222	
RET	RET	-0.566935	0.019690	
IR	INSR	-0.594081	0.024830	
TLR4:MD-2 complex	TLR4 LY96	-0.597382	0.007250	
IL-13 Ra1	IL13RA1	-0.597611	0.014114	
CD23	FCER2	-0.602595	0.046311	
HAI-1	SPINT1	-0.620640	0.008536	
sE-Selectin	SELE	-0.625872	0.024278	
TECK	CCL25	-0.655796	0.041103	
Carbonic anhydrase I	CA1	-0.703275	0.003678	
ISLR2	ISLR2	-0.738526	0.000955	
IL-5 Ra	IL5RA	-0.742331	0.026171	
kallikrein 8	KLK8	-0.791363	0.021481	
Growth hormone receptor	GHR	-0.820837	0.004480	
BST1	BST1	-0.884313	0.037880	
CYTT	CST2	-0.905443	0.017786	
IL-18 BPa	IL18BP	-0.906512	0.001023	
TF	F3	-1.041923	0.001738	
ENPP7	ENPP7	-1.261712	0.003212	

Table S2. Upregulated and Downregulated proteins between ALS patients and neurologically healthy controls using limma analysis of SOMAscan Results.

Increased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
Troponin T	TNNT2	1.477579	0.000924	
TIMP-3	TIMP3	1.292908	0.003370	
PDGF-BB	PDGFB	1.205864	0.005048	
TARC	CCL17	1.081839	0.003579	
Gro-b/g	CXCL3 CXCL2	1.003084	0.009649	
DKK1	DKK1	0.826350	0.004899	
CSRP3	CSRP3	0.786139	0.002050	
Dkk-4	DKK4	0.685932	0.008682	
Decreased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
TLR4:MD-2 complex	TLR4 LY96	-0.597382	0.007250	
HAI-1	SPINT1	-0.620640	0.008536	
Carbonic anhydrase I	CA1	-0.703275	0.003678	
ISLR2	ISLR2	-0.738526	0.000955	
Growth hormone receptor	GHR	-0.820837	0.004480	
IL-1a	IL1A	-0.849595	0.003406	
IL-18 BPa	IL18BP	-0.906512	0.001023	
TF	F3	-1.041923	0.001738	
ENPP7	ENPP7	-1.261712	0.003212	

Table S3. Upregulated and Downregulated proteins between S-ALS patients and neurologically healthy controls using SAM Analysis of SOMAscan Results.

Increased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
TIMP-3	TIMP3	1.488049	0.003717	
TARC	CCL17	1.451586	0.000576	
Protein disulfide isomerase A3	PDIA3	1.020243	0.031587	
AN32B	ANP32B	0.618600	0.002078	
NID2	NID2	0.602565	0.011215	
Decreased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
Ephrin-A2	EFNA2	-0.337748	0.021640	
sTie-1	TIE1	-0.346598	0.031437	
IL-16	IL16	-0.354345	0.054714	
VCAM-1	VCAM1	-0.368661	0.007593	
ULBP-2	ULBP2	-0.457169	0.010375	
VEGF sR3	FLT4	-0.469542	0.004093	
Ephrin-A3	EFNA3	-0.504992	0.008539	
BMPR1A	BMPR1A	-0.514741	0.017360	
BCAM	BCAM	-0.515190	0.050522	
IGF-I	IGF1	-0.516672	0.013849	
LYPD3	LYPD3	-0.532839	0.011828	
Cathepsin S	CTSS	-0.546092	0.033920	
sTie-2	TEK	-0.554973	0.017552	
NCAM-L1	L1CAM	-0.571303	0.002630	
IL-18 Ra	IL18R1	-0.602471	0.038568	
TLR4:MD-2 complex	TLR4 LY96	-0.661724	0.010767	
IL-13 Ra1	IL13RA1	-0.681839	0.016537	
Kallikrein 7	KLK7	-0.696381	0.014740	
granzyme A	GZMA	-0.710334	0.004400	
RET	RET	-0.749046	0.007686	
ISLR2	ISLR2	-0.787154	0.002363	
IR	INSR	-0.791866	0.009677	
IL-18 BPa	IL18BP	-0.844704	0.006945	
sE-Selectin	SELE	-0.860835	0.007021	
CD23	FCER2	-0.896461	0.009192	
Myostatin	MSTN	-0.897505	0.056884	
kallikrein 8	KLK8	-0.924181	0.021588	
Lactoferrin	LTF	-0.957086	0.031620	
Growth hormone receptor	GHR	-1.138476	0.000460	

Table S4. Downregulated proteins between F-ALS patients and neurologically healthy controls using SAM Analysis of SOMAscan Results. No significantly up-regulated proteins were identified.

Decreased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
RNF43	RNF43	-0.403979	0.027766	
TLR4	TLR4	-0.434531	0.006334	
RGM-C	HFE2	-0.481616	0.006041	
ISLR2	ISLR2	-0.689897	0.002233	
Carbonic anhydrase I	CA1	-0.812518	0.002971	
Discoidin domain receptor 2	DDR2	-1.138072	0.003942	
TF	F3	-1.466182	0.000018	
ENPP7	ENPP7	-1.498924	0.000856	

Table S5. Upregulated and Downregulated proteins between S-ALS patients and neurologically healthy controls using limma Analysis of SOMAscan Results.

Increased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
TIMP-3	TIMP3	1.488049	0.003717	
Troponin T	TNNT2	1.483767	0.003801	
TARC	CCL17	1.451586	0.000576	
G-CSF-R	CSF3R	1.193777	0.008583	
IGFBP-2	IGFBP2	1.114518	0.003953	
DKK1	DKK1	0.978497	0.004176	
Dkk-4	DKK4	0.863833	0.004338	
FGF-10	FGF10	0.754318	0.009419	
LIMP II	SCARB2	0.628880	0.004092	
AN32B	ANP32B	0.618600	0.002078	
SDF-1	CXCL12	0.616372	0.003589	
CD22	CD22	0.592353	0.009333	

Decreased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
Nogo Receptor	RTN4R	-0.590989	0.001815	
granzyme A	GZMA	-0.710334	0.004400	
RET	RET	-0.749046	0.007686	
ISLR2	ISLR2	-0.787154	0.002363	
IR	INSR	-0.791866	0.009677	
IL-18 BPa	IL18BP	-0.844704	0.006945	
sE-Selectin	SELE	-0.860835	0.007021	
CD23	FCER2	-0.896461	0.009192	
Growth hormone receptor	GHR	-1.138476	0.000460	

Table S6. Upregulated and Downregulated proteins between F-ALS patients and neurologically healthy controls using READAT Analysis of Somascan Results.

Increased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
PDE5A	PDE5A	2.515048	0.003031	
M2-PK	PKM	2.282772	0.004186	
AMPM2	METAP2	2.111687	0.008841	
FER	FER	1.846274	0.008930	
MAPK2	MAPKAPK2	1.693216	0.002242	
CPNE1	CPNE1	1.667009	0.007142	
DLRB1	DYNLRB1	1.655651	0.005362	
Troponin T	TNNT2	1.471392	0.002702	
PKC-D	PRKCD	1.426355	0.001771	
EGF	EGF	1.385597	0.002806	
Gro-b/g	CXCL3 CXCL2	1.213105	0.008527	
C5a	C5	1.196144	0.008846	
CAMK2A	CAMK2A	1.189491	0.003996	
cGMP-stimulated PDE	PDE2A	1.012715	0.003776	
C6	C6	0.941726	0.008187	
CSRP3	CSRP3	0.845542	0.000344	
Calcineurin B a	PPP3R1	0.807205	0.008612	
Decreased in ALS patients				
Target Fullname	Entrez Gene Sym	logFC	p-value	
calreticulin	CALR	-0.583783	0.004835	
ISLR2	ISLR2	-0.689897	0.002233	
TECK	CCL25	-0.808331	0.009408	
Carbonic anhydrase I	CA1	-0.812518	0.002971	
BMP-6	BMP6	-0.885880	0.000407	
IL-18 BPa	IL18BP	-0.968319	0.001700	
Discoidin domain receptor 2 DDR2		-1.138072	0.003942	
a2-Macroglobulin	A2M	-1.438991	0.004912	
TF	F3	-1.466182	0.000018	
ENPP7	ENPP7	-1.498924	0.000856	
Thrombin	F2	-1.509842	0.007796	
C3b	C3	-1.703522	0.003412	

Table S7. List of GO Annotation for TARC, TIMP-3, NID1 and NID2. Abbreviations: P-Pathway, F-Function; C- Cellular.

TARC				
Qualifier	GO Term	GO Name	Reference	GO Aspect
involved_in	GO:0043547	positive regulation of GTPase activity	PMID:21873635	P
enables	GO:0008009	chemokine activity	PMID:21873635	F
involved_in	GO:0048247	lymphocyte chemotaxis	PMID:21873635	P
is_active_in	GO:0005615	extracellular space	PMID:21873635	C

involved_in	GO:0070098	chemokine-mediated signaling pathway	PMID:21873635	P
involved_in	GO:0070374	positive regulation of ERK1 and ERK2 cascade	PMID:21873635	P
involved_in	GO:0071346	cellular response to type II interferon	PMID:21873635	P
involved_in	GO:0071347	cellular response to interleukin-1	PMID:21873635	P
involved_in	GO:0071356	cellular response to tumor necrosis factor	PMID:21873635	P
involved_in	GO:0030593	neutrophil chemotaxis	PMID:21873635	P
involved_in	GO:0002548	monocyte chemotaxis	PMID:21873635	P
involved_in	GO:0006954	inflammatory response	PMID:21873635	P
involved_in	GO:0007186	G protein-coupled receptor signaling pathway	PMID:21873635	P
enables	GO:0048020	CCR chemokine receptor binding	PMID:21873635	F
enables	GO:0008009	chemokine activity	GO_REF:0000002	F
located_in	GO:0005576	extracellular region	GO_REF:0000002	C
involved_in	GO:0006955	immune response	GO_REF:0000002	P
located_in	GO:0005615	extracellular space	GO_REF:0000043	C
located_in	GO:0005576	extracellular region	GO_REF:0000043	C
enables	GO:0005125	cytokine activity	GO_REF:0000043	F
involved_in	GO:0006935	chemotaxis	GO_REF:0000043	P
involved_in	GO:0006954	inflammatory response	GO_REF:0000043	P
enables	GO:0008009	chemokine activity	PMID:9743322	F
enables	GO:0005102	signaling receptor binding	PMID:8702936	F
enables	GO:0005102	signaling receptor binding	PMID:9743322	F
involved_in	GO:0006935	chemotaxis	PMID:9743322	P
involved_in	GO:0007267	cell-cell signaling	PMID:8702936	P
located_in	GO:0005576	extracellular region	Reactome:R-HSA-443986	C
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
enables	GO:0005515	protein binding	PMID:28381538	F
involved_in	GO:0045662	negative regulation of myoblast differentiation	GO_REF:0000107	P
enables	GO:0031729	CCR4 chemokine receptor binding	GO_REF:0000107	F
located_in	GO:0005576	extracellular region	GO_REF:0000044	C
involved_in	GO:0061844	antimicrobial humoral immune response mediated by antimicrobial peptide	PMID:12949249	P
involved_in	GO:0031640	killing of cells of another organism	PMID:12949249	P
involved_in	GO:0031640	killing of cells of another organism	PMID:12949249	P

TIMP-3

Qualifier	GO Term	GO Name	Reference	GO Aspect
enables	GO:0002020	protease binding	PMID:21873635	F
enables	GO:0008191	metalloendopeptidase inhibitor activity	PMID:21873635	F
involved_in	GO:0051045	negative regulation of membrane protein	PMID:21873635	P

		ectodomain proteolysis		
is_active_in	GO:0005615	extracellular space	PMID:21873635	C
involved_in	GO:0009725	response to hormone	PMID:21873635	P
involved_in	GO:0010033	response to organic substance	PMID:21873635	P
involved_in	GO:0010951	negative regulation of endopeptidase activity	PMID:21873635	P
is_active_in	GO:0031012	extracellular matrix	PMID:21873635	C
involved_in	GO:0034097	response to cytokine	PMID:21873635	P
enables	GO:0008191	metalloendopeptidase inhibitor activity	GO_REF:0000002	F
enables	GO:0008191	metalloendopeptidase inhibitor activity	GO_REF:0000043	F
involved_in	GO:0050896	response to stimulus	GO_REF:0000043	P
enables	GO:0030414	peptidase inhibitor activity	GO_REF:0000043	F
involved_in	GO:0007601	visual perception	GO_REF:0000043	P
involved_in	GO:0010466	negative regulation of peptidase activity	GO_REF:0000043	P
located_in	GO:0005576	extracellular region	GO_REF:0000043	C
enables	GO:0004857	enzyme inhibitor activity	GO_REF:0000043	F
enables	GO:0046872	metal ion binding	GO_REF:0000043	F
located_in	GO:0005576	extracellular region	Reactome:R-HSA-481009	C
located_in	GO:0031089	platelet dense granule lumen	Reactome:R-HSA-481009	C
enables	GO:0005515	protein binding	PMID:18344519	F
involved_in	GO:0071310	cellular response to organic substance	GO_REF:0000107	P
located_in	GO:0005604	basement membrane	GO_REF:0000107	C
involved_in	GO:1904684	negative regulation of metalloendopeptidase activity	GO_REF:0000107	P
enables	GO:0008191	metalloendopeptidase inhibitor activity	PMID:18638486	F
involved_in	GO:1903984	positive regulation of TRAIL-activated apoptotic signaling pathway	PMID:19962668	P
part_of	GO:0062023	collagen-containing extracellular matrix	PMID:25037231	C
part_of	GO:0062023	collagen-containing extracellular matrix	PMID:28675934	C
part_of	GO:0062023	collagen-containing extracellular matrix	PMID:25037231	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:27559042	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:27068509	C
colocalizes_with	GO:0062023	collagen-containing extracellular matrix	PMID:20551380	C
involved_in	GO:0051045	negative regulation of membrane protein ectodomain proteolysis	PMID:18383040	P
involved_in	GO:0070373	negative regulation of ERK1 and ERK2 cascade	PMID:19962668	P
involved_in	GO:1904684	negative regulation of metalloendopeptidase activity	GO_REF:0000024	P
enables	GO:0005515	protein binding	PMID:18638486	F
located_in	GO:0005634	nucleus	PMID:21630459	C

NID1

Qualifier	GO Term	GO Name	Reference	GO Aspect
is_active_in	GO:0005886	plasma membrane	PMID:21873635	C
enables	GO:0005509	calcium ion binding	GO_REF:0000002	F
involved_in	GO:0007160	cell-matrix adhesion	GO_REF:0000002	P
located_in	GO:0005576	extracellular region	GO_REF:0000043	C

located_in	GO:0005604	basement membrane	GO_REF:0000043	C
involved_in	GO:0007155	cell adhesion	GO_REF:0000043	P
located_in	GO:0005576	extracellular region	Reactome:R-HSA-2426450	C
located_in	GO:0005576	extracellular region	Reactome:R-HSA-3791319	C
enables	GO:0005518	collagen binding	GO_REF:0000107	F
involved_in	GO:0010811	positive regulation of cell-substrate adhesion	GO_REF:0000107	P
located_in	GO:0071944	cell periphery	GO_REF:0000107	C
located_in	GO:0031012	extracellular matrix	GO_REF:0000107	C
located_in	GO:0005604	basement membrane	GO_REF:0000107	C
involved_in	GO:0030198	extracellular matrix organization	GO_REF:0000107	P
involved_in	GO:0032836	glomerular basement membrane development	GO_REF:0000107	P
involved_in	GO:0007160	cell-matrix adhesion	GO_REF:0000107	P
enables	GO:0043237	laminin-1 binding	GO_REF:0000107	F
enables	GO:0050840	extracellular matrix binding	GO_REF:0000107	F
located_in	GO:0005604	basement membrane	PMID:22952693	C
enables	GO:0005518	collagen binding	PMID:22952693	F
enables	GO:0043236	laminin binding	PMID:22952693	F
involved_in	GO:0071711	basement membrane organization	PMID:22952693	P
enables	GO:0005201	extracellular matrix structural constituent	PMID:25037231	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:25037231	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:28675934	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:20551380	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:27559042	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:27068509	F
part_of	GO:0062023	collagen-containing extracellular matrix	PMID:25037231	C
part_of	GO:0062023	collagen-containing extracellular matrix	PMID:25037231	C
part_of	GO:0062023	collagen-containing extracellular matrix	PMID:28675934	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:27559042	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:27068509	C
colocalizes_with	GO:0062023	collagen-containing extracellular matrix	PMID:20551380	C
colocalizes_with	GO:0062023	collagen-containing extracellular matrix	PMID:22261194	C
enables	GO:0005201	extracellular matrix structural constituent	GO_REF:0000024	F
enables	GO:0043394	proteoglycan binding	PMID:22952693	F
located_in	GO:0005604	basement membrane	GO_REF:0000044	C
involved_in	GO:0110011	regulation of basement membrane organization	PMID:2119632	P
involved_in	GO:0045785	positive regulation of cell adhesion	PMID:3794389	P
involved_in	GO:0051149	positive regulation of muscle cell differentiation	PMID:26555376	P
located_in	GO:0098637	protein complex involved in cell-matrix adhesion	PMID:3109910	C
involved_in	GO:2001046	positive regulation of integrin-mediated signaling pathway	PMID:23948589	P
located_in	GO:0005604	basement membrane	PMID:1678389	C
located_in	GO:0070062	extracellular exosome	PMID:23533145	C
located_in	GO:0070062	extracellular exosome	PMID:19056867	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:23658023	C

located_in	GO:0110165	cellular anatomical entity	GO_REF:0000117	C
NID2				
Qualifier	GO Term	GO Name	Reference	GO Aspect
is_active_in	GO:0005615	extracellular space	PMID:21873635	C
is_active_in	GO:0005604	basement membrane	PMID:21873635	C
involved_in	GO:0007160	cell-matrix adhesion	PMID:21873635	P
enables	GO:0005509	calcium ion binding	GO_REF:0000002	F
involved_in	GO:0007160	cell-matrix adhesion	GO_REF:0000002	P
located_in	GO:0005576	extracellular region	GO_REF:0000043	C
located_in	GO:0005604	basement membrane	GO_REF:0000043	C
involved_in	GO:0007155	cell adhesion	GO_REF:0000043	P
enables	GO:0005518	collagen binding	PMID:9733643	F
located_in	GO:0005604	basement membrane	PMID:9733643	C
involved_in	GO:0007155	cell adhesion	PMID:9733643	P
located_in	GO:0005576	extracellular region	Reactome:R-HSA-2426450	C
located_in	GO:0031012	extracellular matrix	GO_REF:0000107	C
located_in	GO:0005604	basement membrane	GO_REF:0000107	C
involved_in	GO:0007160	cell-matrix adhesion	GO_REF:0000107	P
located_in	GO:0005604	basement membrane	PMID:22952693	C
involved_in	GO:0071711	basement membrane organization	PMID:22952693	P
enables	GO:0005201	extracellular matrix structural constituent	PMID:28675934	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:20551380	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:27559042	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:27068509	F
enables	GO:0005201	extracellular matrix structural constituent	PMID:28327460	F
part_of	GO:0062023	collagen-containing extracellular matrix	PMID:28675934	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:28327460	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:27559042	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:27068509	C
colocalizes_with	GO:0062023	collagen-containing extracellular matrix	PMID:20551380	C
enables	GO:0005201	extracellular matrix structural constituent	GO_REF:0000024	F
located_in	GO:0005604	basement membrane	GO_REF:0000044	C
located_in	GO:0005886	plasma membrane	GO_REF:0000052	C
located_in	GO:0070062	extracellular exosome	PMID:23533145	C
located_in	GO:0062023	collagen-containing extracellular matrix	PMID:23658023	C
enables	GO:0005515	protein binding	PMID:11956183	F
located_in	GO:0110165	cellular anatomical entity	GO_REF:0000117	C

Table S8. ROC curve data of ALS, S-ALS and F-ALS versus controls (H-CTR and NoH-CTR).

ALS versus controls

Protein Name	AUC	P value	Z score	Sensitivity	Specificity	Optimal cut-off point
TARC	0.68	0.0034	2.93	0.56	0.72	57.22
TIMP3	0.60	0.14194	1.47	-	-	-
NID1	0.52	0.78816	0.27	-	-	-
NID2	0.58	0.22565	1.27	-	-	-

S-ALS versus controls

Protein Name	AUC	P value	Z score	Sensitivity	Specificity	Optimal cut-off point
TARC	0.67	0.00915	2.60	0.562	0.722	57.22
TIMP3	0.67	0.00827	2.64	0.656	0.694	49.4
NID1	0.54	0.55898	0.58	-	-	-
NID2	0.51	0.89388	0.13	-	-	-

F-ALS versus controls

Protein Name	AUC	P value	Z score	Sensitivity	Specificity	Optimal cut-off point
TARC	0.72	0.00856	2.63	1	0.375	31.49
TIMP3	0.65	0.1389	1.48.			
NID1	0.53	0.79092	0.26.			
NID2	0.81	0.00002	4.32.	0.583	0.938	22.74

Table S9. Stepwise logistic regression analysis of ALS, S-ALS and F-ALS versus controls (H-CTR and NoH-CTR). Abbreviations: VIF-Variance Inflation Factor

ALS versus controls

Protein Name	P value	VIF
TARC	0.00836	1.360
NID1	0.03857	1.360

S-ALS versus controls

Protein Name	P value	VIF
TARC	0.00240	1.428
NID1	0.05980	1.428

F-ALS versus controls

Protein Name	P value	VIF
TARC	0.07890	1.113
TIMP3	0.02540	1.087
NID2	0.02120	1.026