

Table S1 - Normalised Total Spectra Counts for Control, MCI and AD plasma using Scaffold v4.3.4 (significantly altered proteins in bold)

#	Identified Proteins (107)	Accession Number	AD plasma replicates				MCI Plasma replicates				Control plasma replicates			
			AD1	AD2	AD3	AD4	MCI1	MCI2	MCI3	MCI4	C1	C2	C3	C4
1	Alpha-2-macroglobulin precursor	gi 112911 (+2)	100.27	99.586	119.25	107.24	131.22	99.621	116.67	104.21	149.75	162.1	122.29	98.57
2	apolipoprotein B precursor	[Hor gi 105990532 (+1)	93.535	112.74	89.439	89.941	131.95	124.53	90.063	73.558	74.876	93.006	57.683	95.39
3	Chain A, Human Serum Albumin	gi 122920512 (+4)	86.052	115.24	83.051	65.726	137.05	132.62	75.735	67.428	81.562	90.349	73.834	71.543
4	complement component 3 precursor	gi 115298678	94.283	84.554	112.86	103.78	110.08	104.6	112.58	112.38	116.33	102.31	124.6	111.29
5	keratin 1 [Homo sapiens], gi 39	gi 11935049	53.876	57.622	83.051	53.619	35.721	39.226	96.204	149.16	52.815	47.832	101.52	71.543
6	fibrinogen, alpha polypeptide iso	gi 11761629 (+2)	39.659	30.69	40.461	32.863	32.076	29.886	47.078	34.736	31.421	30.559	41.532	41.336
7	proapolipoprotein	gi 178775	41.155	35.701	27.684	34.593	46.657	39.848	28.656	28.606	63.511	53.146	23.073	31.797
8	hCG2001591 [Homo sapiens]	gi 119623974 (+2)	41.155	35.074	29.813	39.782	39.367	43.584	30.703	32.692	36.77	39.195	29.995	42.926
9	Chain A, Trypsin In Complex Wit	gi 110590762 (+3)	23.945	17.537	10.648	17.296	29.889	29.264	12.281	6.1298	40.112	35.209	16.151	22.258
10	fibrinogen, beta chain prepropr	gi 70906435	36.666	30.69	63.885	41.511	34.263	31.754	34.797	42.909	32.758	27.902	50.761	47.695
11	keratin 10 (epidermolytic hyperl	gi 119581085 (+2)	26.938	40.711	53.238	13.837	18.225	15.566	47.078	63.342	28.079	20.594	64.605	28.617
12	similar to Keratin, type I cytoske	gi 114667176	19.455	21.295	27.684	20.756	13.851	17.434	32.75	67.428	20.725	21.923	34.61	12.719
13	fibrinogen gamma chain, isoform	gi 119625320 (+4)	29.183	23.174	12.777	13.837	28.431	25.528	20.469	12.26	28.079	30.559	18.459	25.437
14	keratin 2 [Homo sapiens], gi 64	gi 47132620	29.183	27.558	63.885	19.026	19.683	21.169	55.266	55.169	31.421	24.58	46.146	27.027
15	ceruloplasmin (ferroxidase), iso	gi 119599289 (+3)	23.945	21.921	17.036	13.837	23.328	20.547	18.422	12.26	16.713	17.937	13.844	17.488
16	Fibronectin 1 [Homo sapiens],	gi 109658664 (+11)	32.924	38.206	29.813	27.674	19.683	18.056	14.328	8.1731	18.719	19.266	18.459	19.078
17	Complement factor H precursor	gi 158517847 (+3)	20.204	13.779	23.425	22.485	19.683	7.4716	22.516	20.433	14.039	18.601	23.073	19.078
18	inter-alpha (globulin) inhibitor H	gi 119606782 (+3)	19.455	22.548	8.518	15.567	9.4771	10.585	12.281	8.1731	11.365	15.944	11.537	12.719
19	hemopexin [Homo sapiens], gi :	gi 11321561 (+6)	11.224	12.527	10.648	5.1889	15.309	14.943	14.328	8.1731	16.045	16.608	11.537	9.539
20	inter-alpha (globulin) inhibitor H	gi 119585668 (+3)	20.952	10.648	17.036	13.837	14.58	11.83	14.328	10.216	13.371	12.622	9.2293	20.668
21	Chain A, Crystal Structure Of Th	gi 28373620 (+1)	14.217	11.9	8.518	10.378	11.664	13.075	8.1875	12.26	15.376	13.951	9.2293	15.898
22	inter-alpha (globulin) inhibitor H	gi 119585664 (+1)	13.469	20.042	12.777	10.378	10.206	12.453	16.375	16.346	7.3539	11.294	16.151	11.129
23	Chain A, Tertiary Structures Of T	gi 2098257 (+3)	15.714	12.527	17.036	15.567	17.496	13.698	10.234	10.216	16.713	15.28	18.459	17.488
24	Complement factor B [Homo sa	gi 13278732 (+2)	11.224	8.7686	4.259	5.1889	6.5611	9.9621	2.0469	4.0866	10.028	11.294	4.6146	6.3594
25	Chain I, 2.5a Crystal Structure O	gi 52695711 (+1)	12.721	11.274	8.518	12.107	13.122	11.207	6.1407	6.1298	10.028	7.972	6.922	17.488
26	apolipoprotein A-IV precursor [T	gi 178757 (+2)	11.224	9.3949	2.1295	1.7296	10.206	8.7168	4.0938	4.0866	9.3595	8.6363	2.3073	4.7695
27	gelsolin isoform 32	gi 114626427 (+2)	11.972	7.5159	4.259	10.378	10.206	9.9621	8.1875	10.216	8.691	9.9649	4.6146	6.3594

28	Chain A, Crystal Structure Of Cle	gi 443345	6.7345	5.6369	12.777	19.026	5.1031	6.8489	14.328	10.216	4.0112	6.6433	16.151	12.719
29	complement component 1 inhil	gi 114642584 (+8)	9.7276	7.5159	10.648	17.296	8.0191	7.4716	6.1407	8.1731	8.0225	9.9649	9.2293	12.719
30	Complement component 5 [Hor	gi 109731812 (+2)	5.9862	6.2633	4.259	8.6482	5.1031	5.6037	4.0938	6.1298	8.0225	8.6363	2.3073	11.129
31	Alpha-1B-glycoprotein precurs	gi 46577680 (+1)	6.7345	6.2633	10.648	6.9185	8.0191	6.8489	10.234	8.1731	6.6854	6.6433	4.6146	6.3594
32	plasminogen	gi 387026 (+1)	6.7345	8.7686	2.1295	0	4.3741	8.7168	4.0938	0	5.3483	8.6363	6.922	1.5898
33	Kininogen 1 [Homo sapiens]	gi 37748641 (+2)	8.9794	6.2633	6.3885	12.107	9.4771	4.981	2.0469	6.1298	4.6798	5.979	11.537	6.3594
34	apolipoprotein E mutant E3K	gi 364011 (+1)	6.7345	5.6369	10.648	5.1889	4.3741	5.6037	8.1875	6.1298	4.0112	5.3146	9.2293	9.539
35	alpha-1-microglobulin/bikunin	pgi 4502067 (+1)	6.7345	6.8896	4.259	5.1889	5.8321	6.2263	4.0938	2.0433	5.3483	4.6503	6.922	3.1797
36	coagulation factor II (thrombin)	gi 119588383 (+4)	4.4897	5.6369	6.3885	8.6482	5.8321	6.2263	4.0938	6.1298	8.691	9.3006	13.844	9.539
37	Angiotensinogen (serpin peptid	gi 15079348 (+10)	9.7276	8.1422	10.648	8.6482	8.0191	7.4716	10.234	8.1731	8.0225	8.6363	4.6146	7.9492
38	alpha-1-acid glycoprotein 1 prec	gi 1197209 (+2)	6.7345	6.2633	4.259	3.4593	7.2901	5.6037	4.0938	4.0866	2.0056	5.979	2.3073	6.3594
39	Alpha-2-HS-glycoprotein precur	gi 112910 (+2)	6.7345	3.758	6.3885	6.9185	8.0191	4.3584	8.1875	2.0433	6.6854	5.3146	11.537	11.129
40	haptoglobin, isoform CRA_a [Ho	gi 119579598	9.7276	2.5053	2.1295	3.4593	6.5611	7.4716	4.0938	2.0433	3.3427	1.993	2.3073	1.5898
41	apolipoprotein H precursor [Hor	gi 153266841 (+2)	5.238	4.3843	0	1.7296	2.916	4.3584	2.0469	0	2.6742	5.3146	0	4.7695
42	apolipoprotein J precursor	gi 178855 (+6)	4.4897	1.879	2.1295	1.7296	3.645	2.4905	4.0938	4.0866	2.6742	1.993	0	7.9492
43	Chain A, The Structure Of Penta	gi 576259	4.4897	5.0106	4.259	3.4593	3.645	3.1132	6.1407	4.0866	4.0112	5.979	2.3073	3.1797
44	apolipoprotein D, apoD [human	gi 619383	2.9931	2.5053	4.259	3.4593	2.916	1.8679	4.0938	2.0433	2.6742	2.6573	4.6146	0
45	retinol binding protein 4, plasm	gi 119570453 (+6)	2.9931	2.5053	2.1295	1.7296	1.458	3.7358	0	0	2.6742	2.6573	2.3073	1.5898
46	paraoxonase 1 [Homo sapiens],	gi 19923106	4.4897	2.5053	4.259	1.7296	5.1031	2.4905	4.0938	4.0866	4.0112	3.3216	4.6146	1.5898
47	Vitronectin [Homo sapiens], gi	gi 13477169 (+2)	2.9931	3.1316	6.3885	3.4593	3.645	4.3584	4.0938	6.1298	3.3427	4.6503	6.922	3.1797
48	complement component 4 bind	gi 4502503	3.7414	3.1316	4.259	6.9185	3.645	3.7358	4.0938	2.0433	4.6798	2.6573	4.6146	4.7695
49	histidine-rich glycoprotein prec	gi 4504489	4.4897	6.8896	0	1.7296	1.458	1.8679	4.0938	4.0866	2.0056	1.993	6.922	3.1797
50	Keratin, type I microfibrillar	48 kgi 125090	0	0	0	17.296	0	6.2263	0	0	0.6685	3.986	0	0
51	similar to afamin	gi 114594313 (+1)	2.9931	3.758	2.1295	0	1.458	3.1132	2.0469	4.0866	1.3371	1.993	4.6146	3.1797
52	complement component 6, isof	gi 119576417 (+4)	4.4897	3.1316	0	6.9185	2.916	1.8679	4.0938	0	2.6742	2.6573	4.6146	1.5898
53	unnamed protein product [Hom	gi 158261613 (+1)	4.4897	3.1316	2.1295	1.7296	3.645	3.7358	2.0469	2.0433	3.3427	2.6573	0	7.9492
54	type II intermediate-filament	prgi 246276	0	0	0	19.026	0	4.981	0	0	0	0	0	0
55	complement component 7, isof	gi 119576411 (+4)	3.7414	4.3843	2.1295	3.4593	1.458	3.7358	4.0938	2.0433	2.6742	3.3216	0	1.5898
56	heparin cofactor II isoform 1 [P	agi 114685230 (+5)	1.4966	1.2527	2.1295	0	1.458	0.6226	0	2.0433	0.6685	1.3287	2.3073	4.7695
57	complement component 9, isof	gi 119576390 (+3)	5.238	3.758	2.1295	3.4593	4.3741	3.7358	4.0938	2.0433	2.6742	2.6573	4.6146	6.3594
58	complement component 1, q su	gi 150036344 (+5)	2.9931	1.879	2.1295	1.7296	2.916	1.8679	2.0469	2.0433	2.0056	2.6573	0	3.1797
59	serine (or cysteine) proteinase	igi 114665618 (+6)	2.2448	1.879	2.1295	3.4593	1.458	1.8679	4.0938	2.0433	2.0056	0.6643	6.922	7.9492
60	unnamed protein product [Hom	gi 16553735 (+2)	2.2448	1.2527	4.259	3.4593	1.458	1.8679	4.0938	2.0433	1.3371	2.6573	4.6146	4.7695

61	inter-alpha (globulin) inhibitor Hgi 119585666 (+5)	1.4966	1.2527	2.1295	1.7296	2.187	1.8679	2.0469	0	1.3371	1.3287	2.3073	3.1797
62	Complement component 8, gar gi 109731764 (+4)	2.2448	1.2527	2.1295	1.7296	1.458	1.2453	4.0938	2.0433	2.6742	1.3287	0	3.1797
63	unnamed protein product [Hom gi 1335054	2.2448	0.6263	0	0	1.458	1.2453	8.1875	6.1298	2.0056	1.993	2.3073	3.1797
64	serum amyloid A4, constitutive gi 10835095 (+3)	2.9931	1.2527	2.1295	0	0	1.8679	0	2.0433	0.6685	0.6643	0	1.5898
65	keratin 5 [Homo sapiens], gi 14: gi 119395754 (+1)	0	10.021	4.259	0	0	0	6.1407	6.1298	5.3483	0	2.3073	6.3594
66	hCG2040284 [Homo sapiens] gi 119598461 (+2)	2.2448	2.5053	0	0	1.458	2.4905	0	0	2.0056	1.3287	0	0
67	alpha-1-acid glycoprotein 2 prec gi 29170378 (+2)	1.4966	1.879	4.259	1.7296	3.645	3.7358	2.0469	0	2.0056	3.986	0	1.5898
68	Complement C1r subcomponen gi 115204 (+6)	2.2448	2.5053	0	1.7296	0.729	1.8679	0	2.0433	1.3371	2.6573	0	1.5898
69	Chain A, Human Apolipoprotein gi 157831482	0	0	0	5.1889	0	0	2.0469	0	0	0	2.3073	0
70	alpha S1 casein gi 159793187 (+6)	0	0	4.259	0	0	0	6.1407	8.1731	0	0	6.922	0
71	similar to Ficolin (collagen/fibrin gi 114555009 (+4)	1.4966	1.879	2.1295	1.7296	1.458	1.2453	2.0469	0	1.3371	1.3287	2.3073	1.5898
72	C-type lectin domain family 3, r gi 156627579 (+2)	0.7483	1.879	0	1.7296	0.729	1.2453	2.0469	0	2.0056	1.3287	2.3073	3.1797
73	protein S (alpha) gi 114588068 (+5)	0	0.6263	0	0	0	1.2453	4.0938	0	0.6685	1.3287	0	1.5898
74	Kallikrein B, plasma (Fletcher fac gi 109659056 (+4)	0	3.1316	0	0	0	1.2453	0	0	2.0056	3.3216	0	0
75	hair type II keratin intermediate gi 1308	0	0	0	20.756	0	8.0942	0	0	0	2.6573	0	0
76	alpha-2-glycoprotein 1, zinc gi 114614917 (+6)	2.2448	1.2527	0	0	1.458	0.6226	0	2.0433	0.6685	0.6643	0	1.5898
77	hypothetical protein LOC507421 gi 134085840	0	0	0	17.296	0	7.4716	0	0	0	0.6643	0	0
78	alpha-2-plasmin inhibitor [Homo c gi 115583663 (+5)	0	1.2527	0	0	1.458	0.6226	0	0	1.3371	0.6643	2.3073	0
79	apolipoprotein M [Homo sapien gi 22091452 (+1)	0.7483	0.6263	0	0	0	1.8679	0	0	0.6685	0	0	0
80	Rubber elongation factor protei gi 132270	0.7483	0	0	1.7296	0	0	4.0938	2.0433	0	1.3287	2.3073	1.5898
81	complement factor B isoform 1 gi 109070536	7.4828	6.8896	2.1295	3.4593	0	6.8489	0	0	6.6854	6.6433	0	0
82	coagulation factor XII (Hagemangi 119605410 (+6)	1.4966	0.6263	0	1.7296	0	0	0	0	0	0	0	1.5898
83	hypothetical protein LOC539595 gi 149642659	0	0	0	17.296	0	7.4716	0	0	0	0	0	0
84	Chain A, Thyroxine-Binding Glob gi 114793604 (+5)	0.7483	1.2527	0	0	0.729	1.2453	0	0	0.6685	0.6643	0	0
85	complement component 2 prec gi 14550407 (+4)	2.9931	2.5053	2.1295	3.4593	0	1.2453	0	2.0433	0	0	0	0
86	keratin 6A [Homo sapiens] gi 114644568 (+5)	6.7345	11.274	0	0	0	0	0	6.1298	8.0225	0	0	4.7695
87	pregnancy-zone protein isoform gi 114643429 (+4)	0	0	0	10.378	14.58	12.453	14.328	12.26	18.051	17.273	0	9.539
88	hypothetical protein isoform 2 gi 114654573 (+4)	1.4966	1.2527	0	0	2.187	1.8679	0	0	0.6685	0	0	1.5898
89	Chain A, Structural Changes Of T gi 1065033 (+143)	0.7483	0.6263	2.1295	0	0.729	1.2453	0	0	0.6685	0.6643	2.3073	0
90	hemoglobin alpha-1 globin chair gi 13650074 (+22)	0.7483	0.6263	0	1.7296	0	1.2453	2.0469	0	0	0	0	3.1797
91	lumican isoform 1 gi 114646184 (+4)	0.7483	1.879	0	0	0	0	0	0	0	0	2.3073	0
92	unnamed protein product [Hom gi 158257796 (+3)	0	2.5053	0	0	0	0	0	0	1.3371	2.6573	0	0
93	complement component 8, beta; gi 119627047 (+4)	1.4966	1.2527	0	1.7296	0.729	0	2.0469	0	0	0	0	1.5898

94	similar to Fibrinogen beta chain	gi 119908847	2.9931	3.1316	0	0	0	3.1132	0	0	3.3427	1.993	6.922	0
95	similar to apolipoprotein F	gi 114644115 (+3)	0.7483	0	0	0	0	1.2453	2.0469	0	0	0.6643	0	1.5898
96	lipoprotein CIII	gi 224917	0	0.6263	4.259	3.4593	0	0	0	0	0	0	4.6146	0
97	Human Vitamin D Binding Prote	gi 18655424 (+1)	13.469	11.9	8.518	10.378	0	13.075	0	12.26	14.039	0	0	15.898
98	complement component 8, alph	gi 114556800 (+5)	0.7483	1.2527	0	1.7296	0	0.6226	0	0	0.6685	0	0	0
99	attractin, isoform CRA_b [Homo	gi 119630936 (+6)	0	0	0	0	0	0	2.0469	2.0433	0	1.3287	2.3073	0
100	keratin associated protein 13.1	gi 41350563	0	0	0	6.9185	0	0.6226	0	0	0	0	0	0
101	hair keratin A1	gi 115495167	0	0	0	13.837	0	6.8489	0	0	0	0	0	0
102	fibulin 1 isoform 3	gi 109094533 (+5)	0.7483	0.6263	0	0	0	0.6226	0	0	1.3371	0	0	0
103	similar to fibrinogen, gamma ch	gi 73977992 (+2)	0	8.1422	0	6.9185	0	0	0	0	0	0	0	12.719
104	hypothetical protein	gi 119922176 (+2)	0	0	0	6.9185	0	0.6226	0	0	0	0	0	0
105	Chain A, Apo-Human Serum Tra	gi 110590597 (+6)	0	1.879	0	0	0	0	0	0	0	0	0	0
106	Chain A, Yeast Alcohol Dehydro	gi 112491285 (+4)	0	1.2527	0	0	0	0	0	0	0	0	0	0
107	Keratin-associated protein 11-1	gi 52783049	0	0	0	5.1889	0	0	0	0	0	0	0	0

Charge state deconvolution and deisotoping were not performed. All MS/MS samples were analysed using Mascot (Matrix Science, London, UK). Mascot was set up to search the NCBI nr database (33055681 entries). Mascot was searched with a fragment ion mass tolerance of 0.20Da and a parent tolerance of 0.25Da. Oxidation of methionine and iodoacetamide derivative of cysteine were specified as variable modifications.

Scaffold was used to validate MS/MS based peptide and protein identifications. Peptide identifications were accepted if they could be established at greater than 95.0% probability by the Peptide Prophet algorithm. Protein identifications were accepted if they could be established at greater than 99.0% probability and contained at least 2 identified peptides. Protein probabilities were assigned by the Protein Prophet algorithm.