

Supplementary Captions.

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**Table S1.** Normalized protein expression values for all samples and most important clinical variables.

Patient ID,POMP normalized,PSMB5 normalized,NRF2 normalized,XBP1 normalized,cMAF normalized,MAFB normalized,Refractoriness,ISS,Previous treatment,Progression,PFS,OS,Death,ASCT  
20,0.151273885,1.51050955,0.014996815,0.430254777,0.190796178,1398.08917,0,,0,0,19.4666667,19.4666667,1,1  
21,0.170496894,1.62360248,0.020068323,0.488819876,0.214192547,1751.5528,0,III,0,1,12.4666667,39.5,0,1  
25,0.21986755,2.32582781,0.014536424,0.42218543,0.204470199,2009.93377,0,I,0,0,35.0333333,35.0333333,0,1  
26,0.155095541,0.161783439,0.018904459,0.458917197,0.185127389,1668.78981,0,II,0,0,35.3666667,35.3666667,0,1  
27,0.14760479,1.46586826,0.014350299,0.432934132,0.190658683,2170.65868,1,I,1,1,4.7,31.2666667,1,0  
28,0.103492063,1.41650794,0.01404127,0.427301587,0.181968254,1311.11111,0,I,0,1,32.4,35.9666667,0,1  
30,0.200928793,1.7498452,0.018690403,0.460990712,0.20377709,1888.54489,0,III,0,1,7.6666667,8.0666667,1,0  
33,0.190909091,1.43039773,0.01615625,0.472443182,0.161477273,1806.81818,0,II,0,1,36.3,36.3,0,1  
34,0.151446945,1.35948553,0.017263666,0.433440514,0.17829582,1279.74277,0,II,1,1,10.8666667,18.6333333,1,1  
36,0.172755418,2.02352941,0.018306502,0.492879257,0.237832817,1622.29102,0,III,0,1,1.2,34.9666667,0,1  
37,0.143086817,1.71028939,0.015717042,0.464951768,0.221028939,1559.48553,0,III,0,1,4.6,35.6666667,0,1

38,0.155974843,1.85597484,0.015993711,0.435849057,0.224433962,1783.01887,0,II,0,1,14.2666667,44.3333333,0,1  
40,0.161587302,1.75809524,0.018876191,0.464126984,0.222031746,1850.79365,0,I,0,1,33.2,34.1333333,0,1  
41,0.142507645,1.53639144,0.017810398,0.455963303,0.217492355,1868.50153,0,III,0,1,11.7666667,13.7666667,1,0  
43,0.134202899,1.13942029,0.011295652,0.42,0.211768116,1321.73913,1,III,0,1,3.76666667,21.9333333,1,0  
52,0.205740181,2.00845921,0.015755287,0.479154079,0.177915408,1477.34139,1,III,0,1,1.03333333,12.4,1,0  
56,0.204746835,1.35727848,0.016237342,0.468987342,0.181297468,1462.02532,1,III,0,1,1.63333333,12.3666667,1,0  
58,0.153416149,1.71149068,0.015726708,0.464596273,0.19826087,1975.15528,0,II,1,1,12.5666667,42.0666667,0,0  
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73,0.139263804,1.45214724,0.017901841,0.432822086,0.189079755,1524.53988,0,I,0,1,29.5,56.7666667,0,1  
74,0.164240506,1.51297468,0.018012658,0.461392405,0.181044304,2212.02532,0,III,0,1,34.8,59.3333333,1,0  
78,0.175609756,1.50060976,0.016689024,0.418902439,0.211158537,1896.34146,1,I,1,1,1.03333333,1.26666667,1,0  
83,0.160909091,1.92818182,0.014227273,0.472424242,0.221090909,1466.66667,0,III,0,1,9.83333333,11.5333333,1,0  
84,0.142207792,1.71525974,0.018525974,0.474675325,0.244383117,1142.85714,0,III,1,1,13.2666667,16.0666667,1,1  
85,0.146290801,1.32967359,0.017186944,0.39851632,0.188071217,1703.26409,1,III,1,1,8.3,31.0333333,0,1  
86,0.205571031,1.72618384,0.01732312,0.369637883,0.228022284,1637.88301,0,III,0,1,20.3,27.0333333,1,1  
89,0.1678125,1.39125,0.0172375,0.4509375,0.1820625,2243.75,0,I,0,1,29.1666667,49.4666667,1,1  
90,0.173114754,1.9652459,0.020839344,0.423934426,0.225737705,2540.98361,0,II,0,1,8.16666667,21.0666667,1,0  
91,0.137650602,1.87138554,0.016409639,0.445481928,0.201174699,1939.75904,0,III,1,1,5.13333333,28.2333333,0,0  
92,0.135060976,1.47042683,0.016121951,0.430792683,0.177835366,1637.19512,0,I,0,1,10.5,29.7666667,1,0  
93,0.135504886,1.62801303,0.016276873,0.486644951,0.181335505,1657.98046,1,,0,1,12.0333333,33.9666667,0,0  
94,0.205475504,2.00864553,0.017507205,0.440345821,0.214005764,2285.30259,0,III,0,1,11.7666667,12.0333333,1,0  
101,0.214144737,2.13651316,0.016411184,0.422039474,0.226348684,1894.73684,1,I,0,1,22.6666667,23.2666667,0,0  
114,0.149262537,1.33362832,0.012386431,0.413274336,0.154100295,1336.28319,1,I,0,1,1.73333333,1.83333333,0,0  
115,0.113489736,1.39618768,0.017548387,0.378005865,0.218856305,973.607038,1,I,1,1,3.03333333,39.1333333,1,1  
118,0.129216867,1.61295181,0.013593374,0.402409639,0.172379518,1228.91566,0,,1,1,11.6333333,20.7,0,1  
119,0.206461538,1.972,0.019187692,0.426153846,0.203292308,2089.23077,1,I,0,1,10.5,29.7666667,1,0

**Table S2.** Normalized protein levels according to ISS stage. Data are presented as mean and SD.

Variable	ISS I/II	SD	ISS III	SD.	p-value
POMP	0.16	0.03	0.17	0.03	0.6322
PSMB5	1.54	0.45	1.68	0.28	0.3641
NRF2	0.017	0.002	0.017	0.002	0.9069
XBPI	0.44	0.02	0.45	0.03	0.1094
cMAF	0.20	0.02	0.21	0.02	0.0383
MAFB	1757.25	390.04	1716.23	293.05	0.7262

**Table S3.** Normalized protein levels according to previous treatment. Data are presented as mean and SD.

Variable	No previous treatment	SD	Previous treatment	SD	<i>p</i> -value
<b>POMP</b>	0.17	0.03	0.15	0.02	0.0350
<b>PSMB5</b>	1.62	0.40	1.57	0.19	0.4121
<b>NRF2</b>	0.019	0.002	0.016	0.002	0.6760
<b>XBP1</b>	0.45	0.03	0.43	0.03	0.1113
<b>cMAF</b>	0.20	0.02	0.20	0.02	0.9843
<b>MAFB</b>	1757.41	310.81	1625.63	426.24	0.3009

**Table S4.** Normalized protein levels according to anemia at the diagnosis. Data are presented as mean and SD.

Variable	HB $\geq 10$ g/dL at diagnosis	SD	HB $< 10$ g/dL at diagnosis	SD	<i>p</i> -value
<b>POMP</b>	0.17	0.03	0.16	0.03	0.2114
<b>PSMB5</b>	1.61	0.42	1.61	0.25	0.8678
<b>NRF2</b>	0.017	0.002	0.017	0.002	0.5875
<b>XBP1</b>	0.44	0.03	0.44	0.03	0.9918
<b>cMAF</b>	0.199	0.02	0.21	0.02	0.2262
<b>MAFB</b>	1737.64	345.26	1734.92	336.98	0.9813

**Table S5.** Protein levels according to hypercalcemia at the diagnosis. Data are presented as mean and SD.

Variable	Calcium $\leq 2.5$ mmol/L at diagnosis	SD	Calcium $> 2.5$ mmol/L at diagnosis	SD	<i>p</i> -value
<b>POMP</b>	0.16	0.03	0.17	0.03	0.6292
<b>PSMB5</b>	1.60	0.41	1.65	0.21	0.6292
<b>NRF2</b>	0.016	0.002	0.018	0.002	0.5875
<b>XBP1</b>	0.44	0.03	0.45	0.02	0.9918
<b>cMAF</b>	0.20	0.02	0.20	0.02	0.2262
<b>MAFB</b>	1663.82	333.04	1915.38	289.76	0.9813

**Table S6.** Normalized protein levels according to renal failure at the diagnosis. Data are presented as mean and SD.

Variable	Creatinine $\leq 2$ mg/dL at diagnosis	SD	Creatinine $> 2$ mg/dL at diagnosis	SD	<i>p</i> -value
<b>POMP</b>	0.16	0.03	0.18	0.03	0.2784
<b>PSMB5</b>	1.60	0.38	1.68	0.11	0.4904
<b>NRF2</b>	0.017	0.002	0.018	0.0006	0.1488
<b>XPB1</b>	0.45	0.03	0.44	0.05	0.7213
<b>cMAF</b>	0.20	0.02	0.21	0.03	0.2588
<b>MAFB</b>	1738.56	330.58	1720.33	451.05	0.9203

**Table S7.** Normalized protein levels according to bone disease at the diagnosis. Data are presented as mean and SD.

Variable	Without bone disease	SD	Bone disease	SD	<i>p</i> -value
<b>POMP</b>	0.16	0.03	0.16	0.03	0.5273
<b>PSMB5</b>	1.62	0.25	1.60	0.44	0.8772
<b>NRF2</b>	0.017	0.002	0.017	0.002	0.8625
<b>XPB1</b>	0.45	0.02	0.44	0.04	0.7093
<b>cMAF</b>	0.20	0.02	0.20	0.02	0.4858
<b>MAFB</b>	1772.42	330.95	1677.26	356.10	0.3936

**Table S8.** Normalized protein levels according to the response to bortezomib-based chemotherapy according to the IMWG criteria. Data are presented as mean and SD.

Variable	$\geq$ VGPR	SD	$<$ VGPR	SD	<i>p</i> -value
<b>POMP</b>	0.16	0.03	0.17	0.03	0.3994
<b>PSMB5</b>	1.58	0.41	1.64	0.31	0.8107
<b>NRF2</b>	0.018	0.002	0.016	0.002	0.0167
<b>XPB1</b>	0.45	0.02	0.44	0.03	0.1234
<b>cMAF</b>	0.20	0.02	0.20	0.02	0.6520
<b>MAFB</b>	1761.34	374.04	1691.29	319.62	0.5320

**Table S9.** Normalized protein levels according to refractoriness to bortezomib. Data are presented as mean and SD.

Variable	Sensitive	SD	Refractory	SD	<i>p</i> -value
<b>POMP</b>	0.16	0.03	0.17	0.03	0.6819
<b>PSMB5</b>	1.63	0.38	1.56	0.31	0.1570
<b>NRF2</b>	0.017	0.002	0.016	0.002	0.0863
<b>XPB1</b>	0.45	0.03	0.44	0.03	0.1840
<b>cMAF</b>	0.20	0.02	0.20	0.02	0.3194
<b>MAFB</b>	1752.49	346.26	1665.87	342.37	0.4644

**Table S10.** The optimal cutpoints for protein levels determined by Cutoff Finder.

Protein	PFS			OS		
	Cut-off point	N below cut-off point	N above cut-off point	Cut-off point	N below cut-off point	N above cut-off point
<b>POMP</b>	0.1503	15	24	0.1729	28	11
<b>PSMB5</b>	1.525	17	22	1.713	24	15
<b>NRF2</b>	0.01786	28	11	0.01786	28	11
<b>XPB1</b>	0.4267	11	28	0.4612	26	13
<b>cMAF</b>	0.2078	22	17	0.1899	14	25
<b>MAFB</b>	1638	16	23	1501	11	28

**Table S11.** Univariate Cox regression analyses of dichotomized protein levels for progression-free survival for patients without previous treatment (N=29).

Variables	PFS				
	Coefficient	<i>p</i> -value	HR	95% CI	
<b>High POMP</b>	-0.233	0.2790	0.627	0.269	1.460
<b>High PSMB5</b>	0.308	0.1530	1.851	0.796	4.308
<b>High NRF2</b>	-0.157	0.4547	0.731	0.321	1.664
<b>High XPB1</b>	0.184	0.3799	1.45	0.635	3.287
<b>High cMAF</b>	0.352	0.0988	2.020	0.876	4.658
<b>High MAFB</b>	-0.403	0.0554	0.446	0.195	1.019

**Table S12.** The optimal cutpoints for protein levels determined by Cutoff Finder for PFS of patients without previous treatment (N=29).

<b>PFS</b>			
<b>Protein</b>	Cut-off point	N below	N above
<b>POMP</b>	0.1532	10	19
<b>PSMB5</b>	1.525	12	17
<b>NRF2</b>	0.01614	11	18
<b>XBP1</b>	0.4628	19	10
<b>cMAF</b>	0.2081	16	13
<b>MAFB</b>	1638	12	17