

Table S11

FOCUS 3x

n patients = 278	OS Univariate survival analysis			
Variable	HR	95% CI Low	95% CI High	p-value
CMS1 vs. CMS2	3.26	2.29	4.65	5 x 10 ⁻⁹
CMS3 vs. CMS2	1.18	0.78	1.76	0.435
CMS4 vs. CMS2	1.42	1.04	1.95	0.029
imCMS1 vs. imCMS2 (Slide A)	3.07	2.11	4.48	5 x 10 ⁻⁹
imCMS3 vs. imCMS2 (Slide A)	1.26	0.82	1.93	0.286
imCMS4 vs. imCMS2 (Slide A)	1.42	1.05	1.91	0.023
imCMS1 vs. imCMS2 (Slide B)	4.45	3	6.59	1 x 10 ⁻¹³
imCMS3 vs. imCMS2 (Slide B)	1.22	0.81	1.85	0.338
imCMS4 vs. imCMS2 (Slide B)	1.42	1.05	1.91	0.023

n patients = 267	OS Multivariable survival analysis (adjusted by gender, age, stage)			
Variable	HR	95% CI Low	95% CI High	p-value
CMS1 vs. CMS2	2.32	1.61	3.34	6 x 10 ⁻⁶
CMS3 vs. CMS2	1.04	0.68	1.58	0.867
CMS4 vs. CMS2	1.38	1	1.92	0.05
Age	1	0.98	1.01	0.677
Gender	1.14	0.87	1.5	0.343
Stage	2.37	1.9	2.95	2 x 10 ⁻¹⁴
imCMS1 vs. imCMS2 (Slide A)	2.55	1.73	3.78	3 x 10 ⁻⁶
imCMS3 vs. imCMS2 (Slide A)	1.15	0.74	1.77	0.537
imCMS4 vs. imCMS2 (Slide A)	1.38	1.02	1.89	0.039
Age	1	0.98	1.01	0.782
Gender	1.09	0.83	1.43	0.551
Stage	2.5	2	3.12	6 x 10 ⁻¹⁶
imCMS1 vs. imCMS2 (Slide B)	3.34	2.21	5.05	1 x 10 ⁻⁸
imCMS3 vs. imCMS2 (Slide B)	1.19	0.78	1.84	0.418
imCMS4 vs. imCMS2 (Slide B)	1.38	1.02	1.88	0.038
Age	1	0.99	1.02	0.916
Gender	1.11	0.84	1.46	0.452
Stage	2.45	1.95	3.07	7 x 10 ⁻¹⁵

OS: overall survival

TCGA 3x

n patients=395		OS Univariate survival analysis			
Variable	HR	95% CI Low	95% CI High	p-value	
CMS1 vs. CMS2	1.35	0.76	2.42	0.308	
CMS3 vs. CMS2	0.42	0.17	0.99	0.048	
CMS4 vs. CMS2	1.26	0.74	2.15	0.402	
imCMS1 vs. imCMS2	1.88	1.07	3.3	0.027	
imCMS3 vs. imCMS2	0.45	0.21	0.97	0.043	
imCMS4 vs. imCMS2	1.18	0.67	2.08	0.571	

n patients=380		OS Multivariate survival analysis (adjusted by age, gender, stage)			
Variable	HR	95% CI Low	95% CI High	p-value	
CMS1 vs. CMS2	1.41	0.75	2.67	0.285	
CMS3 vs. CMS2	0.6	0.24	1.48	0.267	
CMS4 vs. CMS2	1.04	0.59	1.82	0.894	
Age	1.04	1.02	1.06	0.001	
Gender	1.08	0.68	1.73	0.735	
Stage	2.15	1.63	2.83	6 x 10 ⁻⁸	
imCMS1 vs. imCMS2	1.78	0.95	3.32	0.07	
imCMS3 vs. imCMS2	0.57	0.25	1.28	0.172	
imCMS4 vs. imCMS2	1.03	0.57	1.88	0.91	
Age	1.04	1.02	1.06	0.001	
Gender	1.03	0.65	1.65	0.892	
Stage	2.13	1.61	2.81	9 x 10 ⁻⁸	

OS: overall survival

N patients=395		PFI Univariate survival analysis			
Variable	HR	95% CI Low	95% CI High	p-value	
CMS1 vs. CMS2	1.37	0.79	2.36	0.26	
CMS3 vs. CMS2	0.59	0.29	1.22	0.155	
CMS4 vs. CMS2	1.68	1.06	2.65	0.028	
imCMS1 vs. imCMS2	1.36	0.79	2.35	0.265	
imCMS3 vs. imCMS2	0.37	0.18	0.75	0.006	
imCMS4 vs. imCMS2	1.37	0.86	2.19	0.19	

n patients=380		PFI Multivariate survival analysis (adjusted by age, gender, stage)			
Variable	HR	95% CI Low	95% CI High	p-value	
CMS1 vs. CMS2	2	1.12	3.58	0.019	
CMS3 vs. CMS2	1	0.47	2.11	0.998	
CMS4 vs. CMS2	1.5	0.93	2.4	0.094	
Age	1	0.98	1.02	0.876	
Gender	0.83	0.55	1.25	0.376	
Stage	2.32	1.82	2.96	1 x 10 ⁻¹¹	
imCMS1 vs. imCMS2	1.6	0.89	2.87	0.115	
imCMS3 vs. imCMS2	0.52	0.25	1.08	0.081	
imCMS4 vs. imCMS2	1.22	0.75	1.99	0.417	
Age	1	0.99	1.02	0.797	
Gender	0.84	0.55	1.27	0.4	
Stage	2.19	1.72	2.78	1 x 10 ⁻¹⁰	

PFI: progression-free interval

GRAMPIAN 12x

n patients=83	RFS Univariate survival analysis			
Variable	HR	95% CI Low	95% CI High	p-value
CMS1 vs. CMS2	3.57	0.5	25.39	0.203
CMS3 vs. CMS2	2.14	0.36	12.83	0.403
CMS4 vs. CMS2	5.75	1.05	31.43	0.044
imCMS1 vs. imCMS2 (Slide A)	1.43	0.13	15.75	0.771
imCMS3 vs. imCMS2 (Slide A)	3.18	0.58	17.35	0.182
imCMS4 vs. imCMS2 (Slide A)	5.75	1.05	31.41	0.044
imCMS1 vs. imCMS2 (Slide B)	0	0	Inf	0.998
imCMS3 vs. imCMS2 (Slide B)	2.82	0.55	14.53	0.216
imCMS4 vs. imCMS2 (Slide B)	6.34	1.16	34.67	0.033

n patients=60	RFS Multivariate survival analysis (adjusted by age, gender, stage)			
Variable	HR	95% CI Low	95% CI High	p-value
CMS1 vs. CMS2	2.42	0.32	18.08	0.388
CMS3 vs. CMS2	1.8	0.23	14.38	0.579
CMS4 vs. CMS2	5.99	0.92	38.87	0.061
Age	0.92	0.86	0.99	0.031
Gender	4.79	0.92	25.12	0.064
Stage	18.47	3.39	100.64	0.001
imCMS1 vs. imCMS2 (Slide A)	1.36	0.12	15.61	0.804
imCMS3 vs. imCMS2 (Slide A)	3.18	0.41	24.4	0.267
imCMS4 vs. imCMS2 (Slide A)	6.63	1	44.12	0.05
Age	0.92	0.86	0.98	0.016
Gender	3.96	0.75	20.83	0.105
Stage	15.78	3.07	81.15	0.001
imCMS1 vs. imCMS2 (Slide B)	0	0	Inf	0.998
imCMS3 vs. imCMS2 (Slide B)	2.55	0.44	14.82	0.299
imCMS4 vs. imCMS2 (Slide B)	7.23	1.12	46.58	0.037
Age	0.93	0.87	1	0.043
Gender	3.52	0.68	18.25	0.134
Stage	12.73	2.71	59.79	0.001

RFS: relapse-free survival