

Table S01

FOCUS		All samples	CMS1	CMS2	CMS3	CMS4	Unclassified
Clinico-pathological features		Statistics	Statistics	Statistics	Statistics	Statistics	Statistics
<i>Age</i>	Median (range)	<i>n</i> =362 64 (31, 83)	<i>n</i> =52 60 (35, 82)	<i>n</i> =120 66 (39, 83)	<i>n</i> =36 65 (48, 79)	<i>n</i> =70 63 (32, 82)	<i>n</i> =84 65 (31, 81)
<i>Gender</i>	Male	<i>n</i> =362 229	<i>n</i> =52 29	<i>n</i> =120 79	<i>n</i> =36 22	<i>n</i> =70 42	<i>n</i> =84 57
	Female	133	23	41	14	28	27
<i>Site</i>	Right colon	<i>n</i> =326 130	<i>n</i> =45 27	<i>n</i> =106 28	<i>n</i> =33 17	<i>n</i> =64 21	<i>n</i> =78 37
	Left colon	196	18	78	16	43	41
<i>TNM Stage</i>	I	<i>n</i> =350 4	<i>n</i> =50 0	<i>n</i> =116 3	<i>n</i> =35 0	<i>n</i> =66 0	<i>n</i> =83 1
	II	39	2	22	6	5	4
	III	75	6	25	8	19	17
	IV	232	42	66	21	42	61
<i>Overall survival (months)</i>	Median (range)	<i>n</i> =362 23 (2, 172)	<i>n</i> =52 15 (4, 43)	<i>n</i> =120 30 (5, 119)	<i>n</i> =36 23 (5, 172)	<i>n</i> =70 23 (5, 107)	<i>n</i> =84 20 (2, 91)
Major molecular features							
<i>Microsatellite status</i>	Stable (MSS)	<i>n</i> =339 328	<i>n</i> =47 37	<i>n</i> =116 116	<i>n</i> =30 30	<i>n</i> =65 64	<i>n</i> =81 81
	Instable (MSI)	11	10	0	0	1	0
<i>CIMP</i>	CIMP.High	<i>n</i> =313 72	<i>n</i> =41 22	<i>n</i> =108 7	<i>n</i> =28 13	<i>n</i> =61 9	<i>n</i> =75 21
	CIMP.Low	83	11	29	5	13	25
	CIMP.Negative	158	8	72	10	39	29
Selected mutations							
<i>KRAS mut status</i>	Wild-type	<i>n</i> =339 171	<i>n</i> =47 31	<i>n</i> =116 68	<i>n</i> =30 8	<i>n</i> =65 33	<i>n</i> =81 31
	Mutated	168	16	48	22	32	50
<i>BRAF mut status</i>	Wild-type	<i>n</i> =339 303	<i>n</i> =47 27	<i>n</i> =116 116	<i>n</i> =30 27	<i>n</i> =65 59	<i>n</i> =81 74
	Mutated	36	20	0	3	6	7
<i>TP53 mut status</i>	Wild-type	<i>n</i> =339 101	<i>n</i> =47 18	<i>n</i> =116 21	<i>n</i> =30 17	<i>n</i> =65 20	<i>n</i> =81 25
	Mutated	238	29	95	13	45	56
<i>APC mut status</i>	Wild-type	<i>n</i> =339 50	<i>n</i> =47 18	<i>n</i> =116 7	<i>n</i> =30 3	<i>n</i> =65 14	<i>n</i> =81 8
	Mutated	289	29	109	27	51	73

TCGA		All samples	CMS1	CMS2	CMS3	CMS4	Unclassified
Clinico-pathological features		Statistics	Statistics	Statistics	Statistics	Statistics	Statistics
<i>Age</i>	Median (range)	<i>n</i> =461 67 (31, 90)	<i>n</i> =73 77 (34, 90)	<i>n</i> =188 66 (34, 90)	<i>n</i> =58 67 (31, 88)	<i>n</i> =109 65 (37, 90)	<i>n</i> =33 66 (43, 89)
<i>Gender</i>	Male	<i>n</i> =461 245	<i>n</i> =73 32	<i>n</i> =188 105	<i>n</i> =58 30	<i>n</i> =109 59	<i>n</i> =33 19
	Female	216	41	83	28	50	14
<i>Site</i>	Right colon	<i>n</i> =447 194	<i>n</i> =70 63	<i>n</i> =182 48	<i>n</i> =57 30	<i>n</i> =106 37	<i>n</i> =32 16
	Left colon	253	7	134	27	69	16
<i>Stage at diagnosis</i>	I	<i>n</i> =444 99	<i>n</i> =72 18	<i>n</i> =181 43	<i>n</i> =55 20	<i>n</i> =104 11	<i>n</i> =32 7
	II	154	35	56	21	34	8
	III	128	15	50	11	38	14
	IV	63	4	32	3	21	3
<i>Overall survival</i>	Median (range)	<i>n</i> =460 22 (0, 150)	<i>n</i> =73 21 (0, 133)	<i>n</i> =187 23 (0, 141)	<i>n</i> =58 25 (0, 142)	<i>n</i> =109 22 (0, 150)	<i>n</i> =33 17 (0, 65)
<i>Progression-free interval</i>	Median (range)	<i>n</i> =460 19 (0, 150)	<i>n</i> =73 18 (0, 133)	<i>n</i> =187 21 (0, 141)	<i>n</i> =58 20 (0, 142)	<i>n</i> =109 17 (0, 150)	<i>n</i> =33 15 (0, 42)
Major molecular features							
<i>Microsatellite status</i>	Stable (MSS)	<i>n</i> =434 363	<i>n</i> =67 10	<i>n</i> =174 173	<i>n</i> =55 46	<i>n</i> =106 102	<i>n</i> =32 32
	Instable (MSI)	71	57	1	9	4	0
<i>CIMP</i>	CIMP.High	<i>n</i> =357 59	<i>n</i> =67 45	<i>n</i> =142 3	<i>n</i> =46 9	<i>n</i> =82 0	<i>n</i> =20 2
	CIMP.Low	70	11	22	19	14	4
	CIMP.Negative	228	11	117	18	68	14
Selected mutations							
<i>KRAS mut status</i>	Wild-type	<i>n</i> =392 228	<i>n</i> =60 44	<i>n</i> =152 101	<i>n</i> =49 11	<i>n</i> =102 61	<i>n</i> =29 11
	Mutated	164	16	51	38	41	18
<i>BRAF mut status</i>	Wild-type	<i>n</i> =392 351	<i>n</i> =60 26	<i>n</i> =152 152	<i>n</i> =49 47	<i>n</i> =102 97	<i>n</i> =29 29
	Mutated	41	34	0	2	5	0
<i>TP53 mut status</i>	Wild-type	<i>n</i> =392 166	<i>n</i> =60 39	<i>n</i> =152 48	<i>n</i> =49 32	<i>n</i> =102 34	<i>n</i> =29 13
	Mutated	226	21	104	17	68	16
<i>APC mut status</i>	Wild-type	<i>n</i> =392 115	<i>n</i> =60 44	<i>n</i> =152 21	<i>n</i> =49 15	<i>n</i> =102 30	<i>n</i> =29 5
	Mutated	277	16	131	34	72	24

GRAMPIAN		All samples	CMS1	CMS2	CMS3	CMS4	Unclassified
Clinico-pathological features		Statistics	Statistics	Statistics	Statistics	Statistics	Statistics
<i>Age</i>	Median (range)	<i>n</i> =223 67 (31, 91)	<i>n</i> =20 72 (45, 86)	<i>n</i> =63 67 (31, 89)	<i>n</i> =35 69 (36, 83)	<i>n</i> =26 60 (40, 79)	<i>n</i> =79 67 (43, 91)
<i>Gender</i>	Male	<i>n</i> =223 142	<i>n</i> =20 9	<i>n</i> =63 43	<i>n</i> =35 21	<i>n</i> =26 16	<i>n</i> =79 53
	Female	81	11	20	14	10	26
<i>Site</i>	Right colon	223 0	<i>n</i> =20 0	<i>n</i> =63 0	<i>n</i> =35 0	<i>n</i> =26 0	<i>n</i> =79 0
	Left colon	223	20	63	35	26	79
<i>Stage at diagnosis</i>	I	<i>n</i> =190 70	<i>n</i> =15 2	<i>n</i> =56 27	<i>n</i> =25 9	<i>n</i> =25 8	<i>n</i> =69 24
	II	61	5	14	9	9	24
	III	53	8	14	7	6	18
	IV	6	0	1	0	2	3
<i>Relapse-free survival</i>	Median (range)	<i>n</i> =125 36 (4, 113)	<i>n</i> =10 36 (24, 36)	<i>n</i> =35 36 (4, 41)	<i>n</i> =24 36 (6, 54)	<i>n</i> =14 36 (9, 36)	<i>n</i> =42 36 (6, 113)
Major molecular features							
<i>Microsatellite status</i>	Stable (MSS)	<i>n</i> =181 178	<i>n</i> =19 16	<i>n</i> =51 31	<i>n</i> =30 30	<i>n</i> =21 21	<i>n</i> =60 60
	Unstable (MSI)	3	3	0	0	0	0
<i>CIMP</i>	CIMP.High	<i>n</i> =118 21	<i>n</i> =11 5	<i>n</i> =40 2	<i>n</i> =24 8	<i>n</i> =11 0	<i>n</i> =32 6
	CIMP.Low	39	4	18	9	1	7
	CIMP.Negative	58	2	20	7	10	19
Selected mutations							
<i>KRAS mut status</i>	Wild-type	<i>n</i> =181 84	<i>n</i> =19 4	<i>n</i> =51 32	<i>n</i> =30 10	<i>n</i> =21 12	<i>n</i> =60 26
	Mutated	97	15	19	20	9	34
<i>BRAF mut status</i>	Wild-type	<i>n</i> =181 178	<i>n</i> =19 16	<i>n</i> =51 51	<i>n</i> =30 30	<i>n</i> =21 21	<i>n</i> =60 60
	Mutated	3	3	0	0	0	0
<i>TP53 mut status</i>	Wild-type	<i>n</i> =181 32	<i>n</i> =19 5	<i>n</i> =51 7	<i>n</i> =30 7	<i>n</i> =21 5	<i>n</i> =60 8
	Mutated	149	14	44	23	16	52
<i>APC mut status</i>	Wild-type	<i>n</i> =181 15	<i>n</i> =19 7	<i>n</i> =51 1	<i>n</i> =30 3	<i>n</i> =21 1	<i>n</i> =60 3
	Mutated	166	12	50	27	20	57