

Supplementary Materials for

Single-cell morphology encodes metastatic potential

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- Fig. S8. Morphological features.
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Other Supplementary Material for this manuscript includes the following:

(available at advances.sciencemag.org/cgi/content/full/6/4/eaaw6938/DC1)

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Supplementary materials

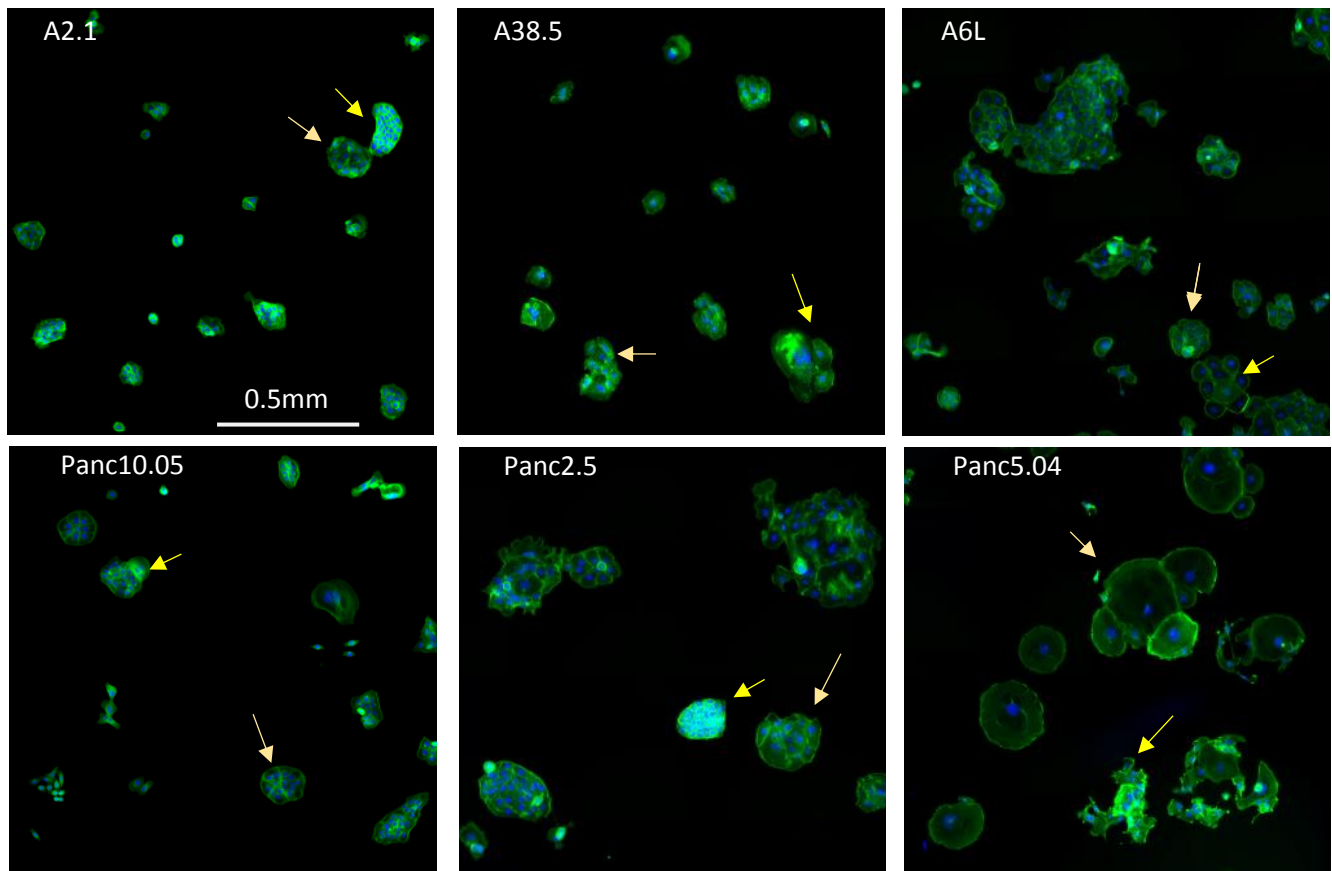


Fig. S1. Cell polymorphism in cancer cells derived from primary pancreatic tumors and metastases. Representative images of pancreatic cancer cells stained for their nucleus (blue) and F-actin (green) in the ultra low-density growth conditions. These include cancer cells derived from metastases that spread from pancreatic tumors (A2.1, A38.5 and A6L) and cells derived from primary pancreatic tumors (Panc10.05, Panc2.5 and Panc5.04). Cells formed several spatially and morphologically distinct progenies. Arrows highlight colonies with distinct cell morphology of the same type.

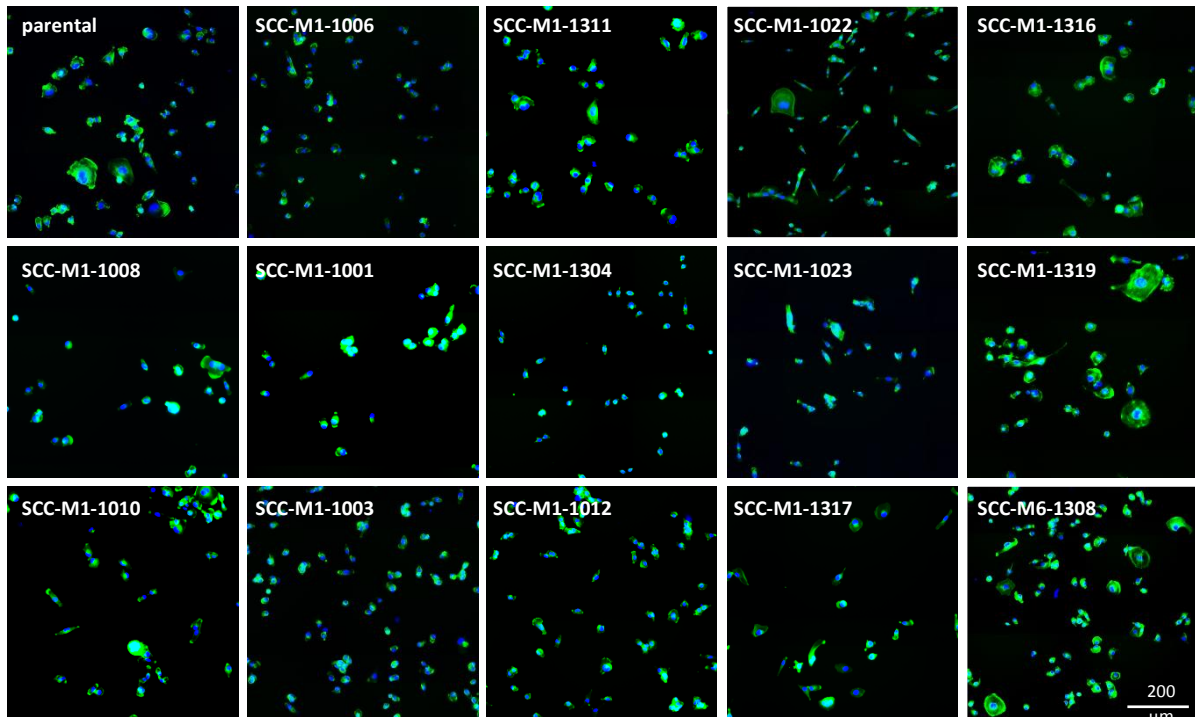


Fig. S2. Morphological spectrum of SCCs. Representative images of 14 different SCCs and their parental MDA-MB-231 breast cancer cells; nuclei and F-actin were stained and shown in blue and green, respectively.

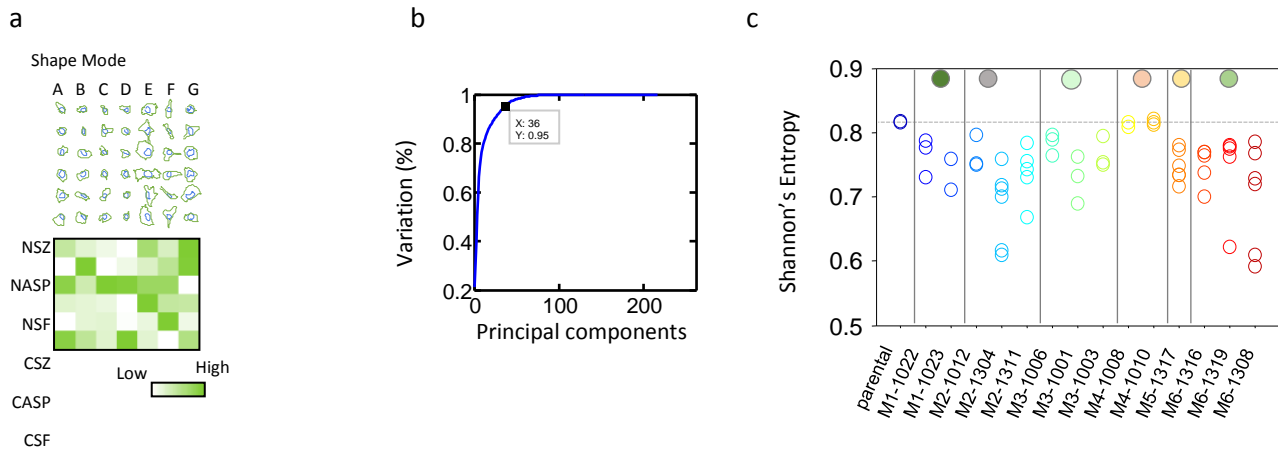


Fig. S3. Quantitative morphology analysis of metastasis of breast cancer cells. **a.** A heat map shows the average nucleus size (NSZ), the nucleus aspect ratio (NASP), the nucleus shape factor (NSF), the cell size (CSZ), the cell aspect ratio (CASP), and the cell shape factor (CSF) for the cell morph classes A-G. The aspect ratio is defined as the ratio between the lengths of the long axis and short axis. The shape factor was calculated using $4\pi \times (\text{Area}) / (\text{perimeter}^2)$. **b.** Principal component analysis for cell morphology feature matrix of SCC cells shows that 95% of variations were covered by the first 36 eigenvectors. **c.** Shannon's entropy of CM distribution for different SCCs. The numbers at the bottom indicate the corresponding morpho-types of the SCCs.

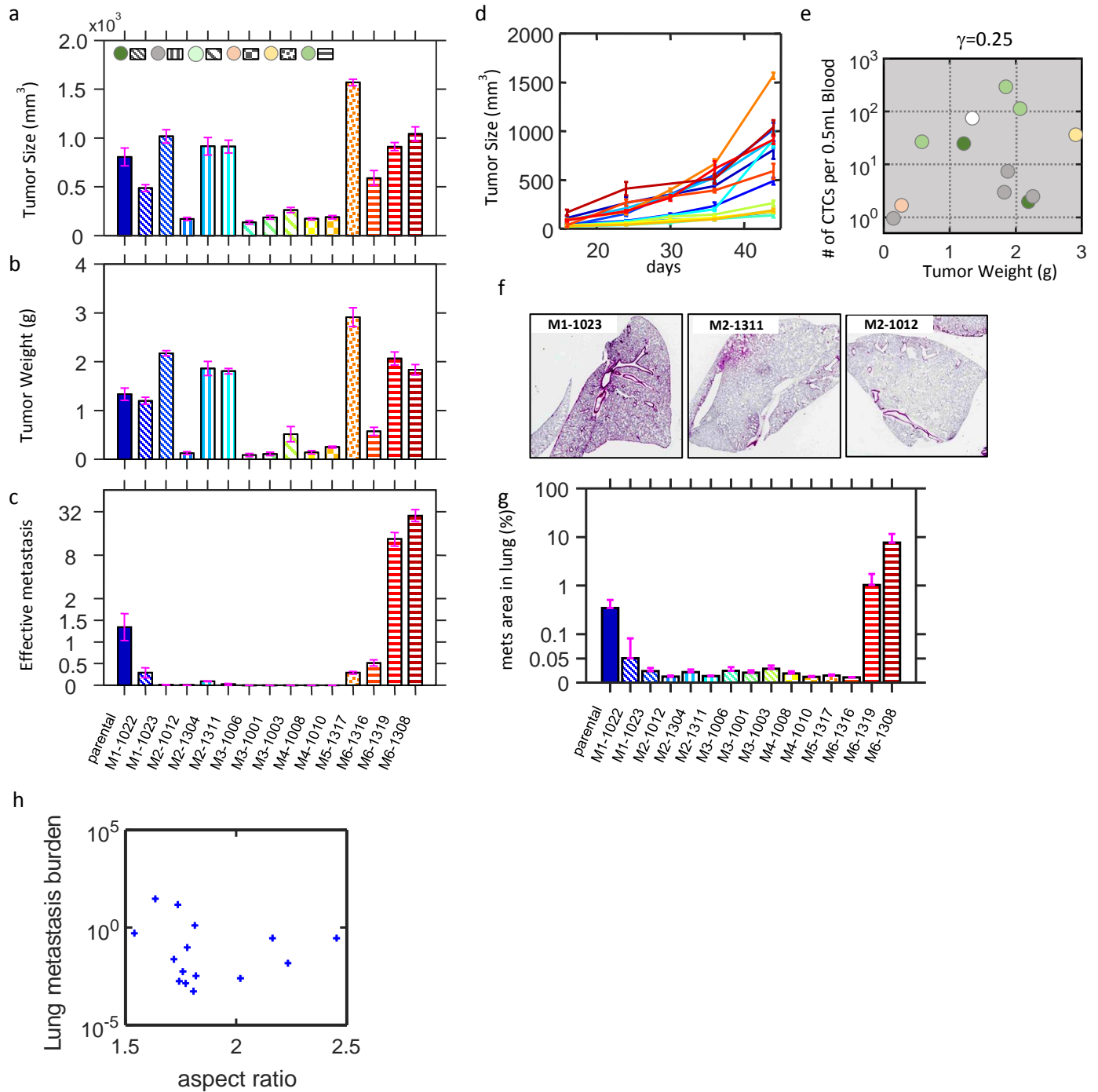


Fig. S4. SCCs derived from the same parental breast cancer cells show divergent invasive behavior in vivo. **a-c.** Bar graphs show the tumor size (a), tumor weight (b), and effective metastasis to the lung (c) after injecting different SCCs into the mammary pad of mice. **d.** Growth of tumors formed by different SCCs. **e.** Scatter plot shows that the CTC count and tumor weight of different SCCs are poorly correlated (Pearson correlation coefficient of 0.25). **f.** Histological sections of mice lung show metastatic lesion free are presented in SCC-1023, SCC-1311 and SCC-1012. At least four mice were tested for each SCC. **g.** Measurement of area ratio occupied by metastasized cancer cells in the tissue sections of lung of mice. **h.** A scatter plot shows the relationship between average cell aspect ratio and corresponding lung metastasis burden (measured by human DNA in the lung) among different SCCs. There is no clear correlation between the cell aspect ratio of SCCs and their metastatic potential, with a Pearson's correlation coefficient of 0.13. Error bars represent standard error of the mean.

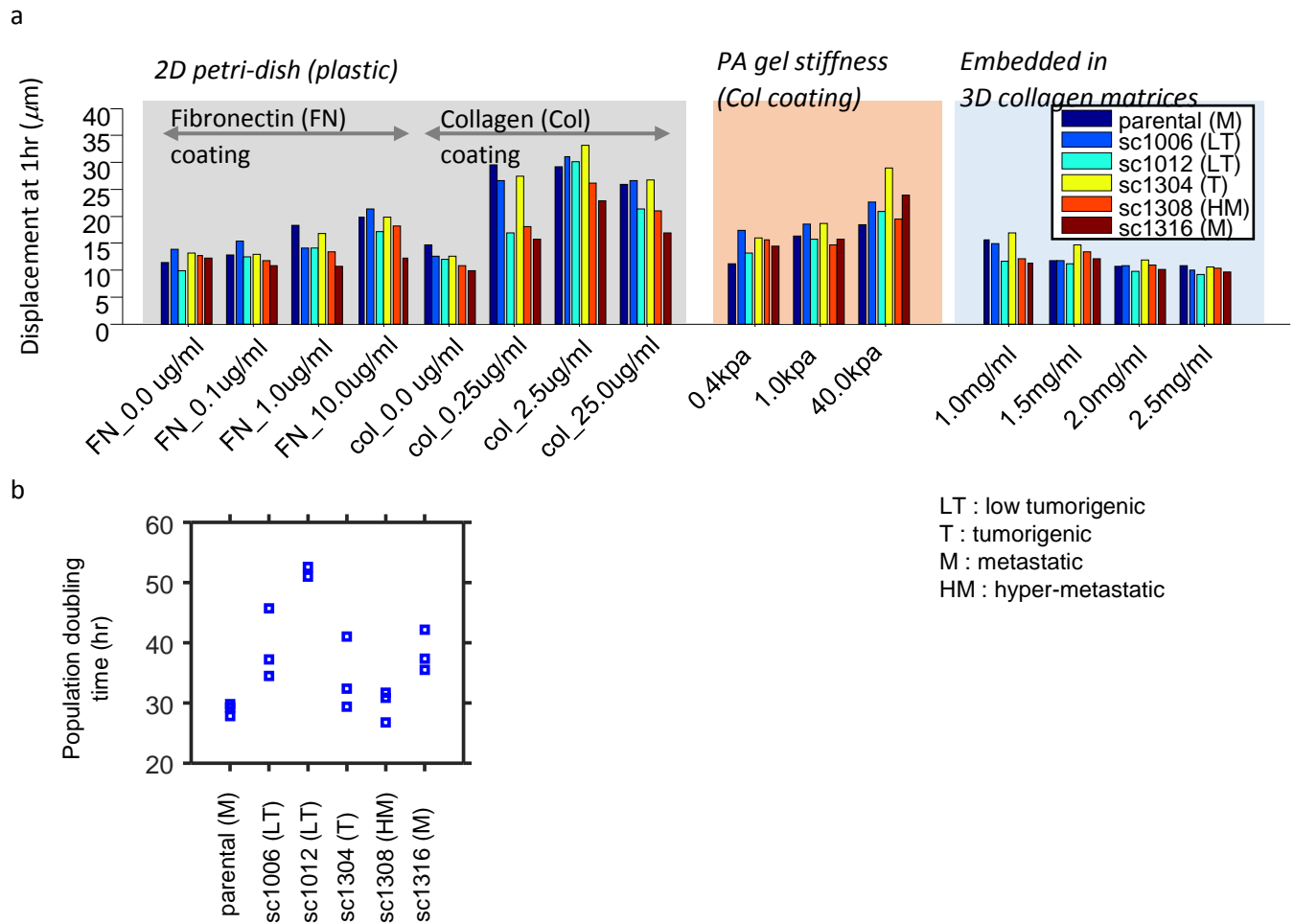
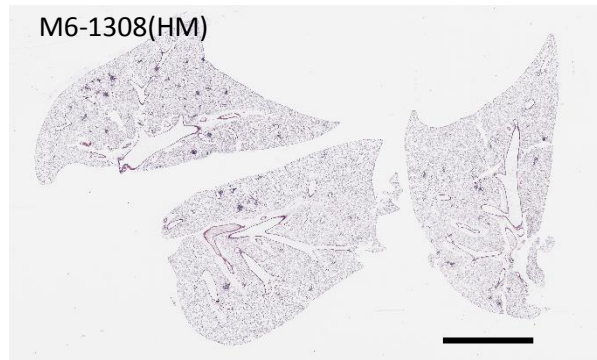
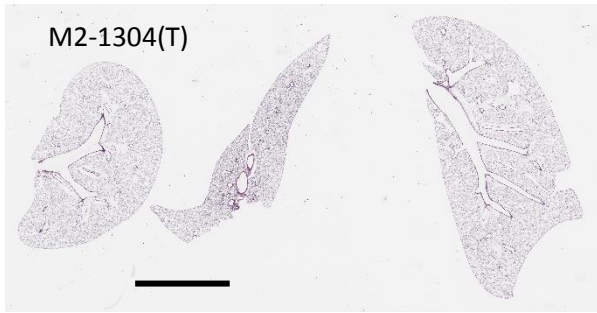


Fig. S5. Analysis of SCC with migration and proliferation assay. a. Migration speed of different SCCs under different culture models including 2D substrates with different concentration of fibronectin (FN) and collagen I (COL) coating on petri dishes, on 2D substrates of different stiffness (0.4kpa, 1.0kpa and 40.0kpa) and embedded in 3D collagen I matrices of four different collagen densities (1mg/ml, 1.5mg/ml, 2.0mg/ml and 2.5mg/ml). Overall the results show the metastatic SCCs, SC1308 and SC1316, do not display a more motile behavior than other SCCs in all *in vitro* models tested. **b.** Population doubling time of different SCCs. These results show little correlation between tumorigenicity (primary tumor growth) of SCCs *in vivo* and their doubling time *in vitro*.

a



b

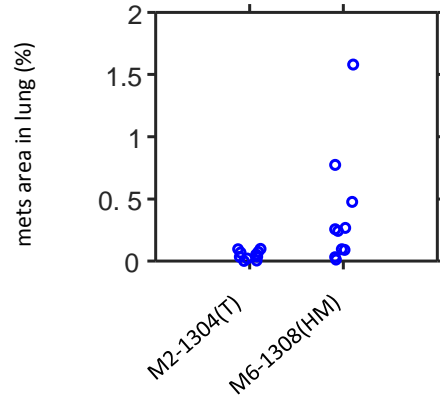


Fig. S6. Lung tissue sections in tail vein metastasis model for different SCCs. **a.** Histological sections of mice lung show there are multiple metastatic lesions in mice that were injected in the tail vein with SCC-1308 (HM) cells, but not with SCC-1304 (T) cells. **b.** Measurement of the area ratio occupied by metastasized tumor cells in the tissue sections of the lung of mice in these two samples.

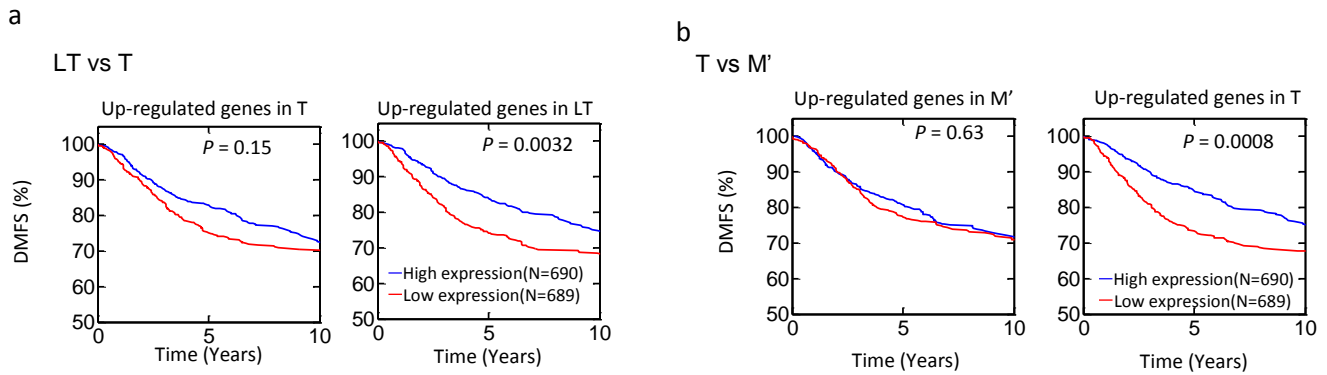


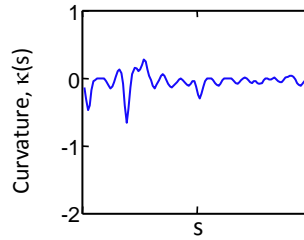
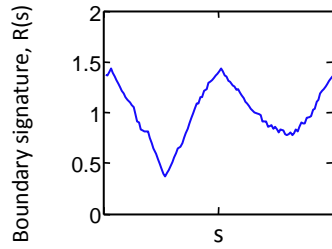
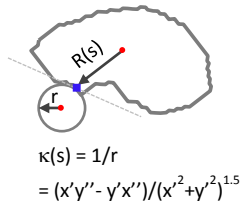
Fig. S7. Distinct gene expression profiles of SCCs reveal prognostic genes. a-b. Differentially expressed genes between LT-SCCs and T-SCCs (a) and between T-SCCs and M'-SCCs (b) were used to investigate the prognostication power. Patients with a higher expression of tumorigenic and metastatic suppressor genes (i.e. genes up-regulated in LT-SCCs compared to T-SCCs or genes up-regulated in T-SCCs compared to M'-SCCs) have a significantly longer survival time than patients with low expression ($P = 0.0032$ and $P = 0.0008$ for LT vs. T and T vs. M'). For the genes that are up-regulated in the T-SCCs compared to LT-SCCs or T-SCCs compared to M'-SCCs, no significant prognostic power was found ($P > 0.05$). P value was calculated using log-rank test.

Category I : Basic morphological features



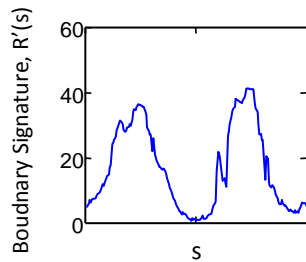
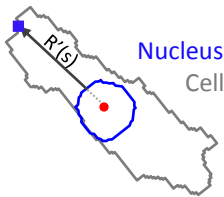
Features:
Area, perimeter, long axis,
aspect ratio,...etc.

Category II & III : Boundary signature and curvature



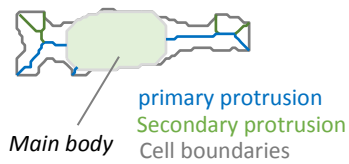
Features:
 $mean(R)$, $median(R)$, $std(R)$,
 $mean(\kappa)$, $median(\kappa)$, $std(\kappa)$...etc

Category IV: N-C positioning signature



Features:
 $mean(R')$, $median(R')$,
 $std(R')$,etc.

Category V: Protrusion



Features:
Protrusion numbers,
<protrusion length>
...etc.

Fig. S8. Morphological features. Schematic plots illustrating four major categories of morphological features that were extracted to quantitatively represent the morphology of individual cells.

Table S1. Summary of SCC information.

Cell name	Morphotype class	Gene expression profiles	Agressiveness grade	Tested mouse number	Turmor size(mm ³) (mean ± sem)	Turmor weight (g) (mean ± sem)	Xenograft mouse model		CTC counts (#/0.5ml blood) (mean ± sem)
							Effective metastasis (A.U.) (mean ± sem)		
parental	4	4	M	8	807.9 ± 257.2	1.34 ± 0.13	$1.3 \times 10^0 \pm 1.3 \times 10^{-1}$	71.8 ± 8.17	
SCC-1006	3	1	LT	4	137.0 ± 39.2	0.09 ± 0.03	$1.4 \times 10^{-3} \pm 4.5 \times 10^{-4}$	N.A.	
SCC-1001	3	1	LT	4	188.0 ± 39.7	0.11 ± 0.04	$3.4 \times 10^{-3} \pm 5.7 \times 10^{-4}$	N.A.	
SCC-1003	3	1	LT	4	263.2 ± 55.6	0.51 ± 0.16	$1.8 \times 10^{-3} \pm 4.2 \times 10^{-4}$	N.A.	
SCC-1008	4	1	LT	4	174.3 ± 26.1	0.14 ± 0.04	$2.5 \times 10^{-3} \pm 8.9 \times 10^{-4}$	N.A.	
SCC-1010	4	1	LT	4	189.6 ± 34.6	0.25 ± 0.02	$5.5 \times 10^{-4} \pm 1.2 \times 10^{-4}$	1.6 ± 0.4	
SCC-1012	2	1	LT	4	172.0 ± 29.6	0.13 ± 0.03	$5.6 \times 10^{-3} \pm 1.2 \times 10^{-3}$	1.0 ± 0.0	
SCC-1304	2	3	T	4	916.1 ± 180.4	1.86 ± 0.14	$9.6 \times 10^{-2} \pm 1.0 \times 10^{-2}$	7.4 ± 0.7	
SCC-1311	2	3	T	4	912.8 ± 133.0	1.81 ± 0.06	$2.4 \times 10^{-2} \pm 1.9 \times 10^{-2}$	3.0 ± 1.3	
SCC-1022	1	2	T	4	488.7 ± 71.4	1.20 ± 0.07	$2.9 \times 10^{-1} \pm 1.1 \times 10^{-1}$	24.6 ± 8.3	
SCC-1023	1	2	T	4	1017.9 ± 136.8	2.18 ± 0.05	$1.5 \times 10^{-2} \pm 5.6 \times 10^{-3}$	2.0 ± 0.4	
SCC-1317	5	3	T	4	1569.1 ± 65.3	2.92 ± 0.19	$2.9 \times 10^{-1} \pm 2.8 \times 10^{-2}$	35.8 ± 11.0	
SCC-1316	6	4	M	4	590.5 ± 171.8	0.58 ± 0.08	$5.2 \times 10^{-1} \pm 7.0 \times 10^{-2}$	26.8 ± 4.9	
SCC-1319	6	4	HM	4	912.5 ± 83.8	2.07 ± 0.13	$1.5 \times 10^1 \pm 3.0 \times 10^0$	112.6 ± 28.7	
SCC-1308	6	4	HM	4	1042.1 ± 145.0	1.84 ± 0.11	$3.0 \times 10^1 \pm 5.5 \times 10^0$	293.0 ± 55.4	

Table S2. Genes that are highly differentially expressed among SCP.

probe ID	Gene Symbol	Gene title	S.D. of SCP gene expression (in log2)	range of SCP gene expression (in log2)	max diff gene exp relative to parental (in log2)
11738785_at	SPANXB2	SPANX family, member B2	4.09	10.10	1.57
11738786_x_at	SPANXB2	SPANX family, member B2	3.95	9.99	2.08
11731994_s_at	SPANXE	SPANX family, member E	3.83	9.66	2.79
11726177_s_at	SPANXB2	SPANX family, member B2	3.78	9.49	2.34
11734286_a_at	COX7B2	cytochrome c oxidase subunit VIIb2	3.42	7.62	0.97
11726339_s_at	MAGEA3	melanoma antigen family A, 3	3.14	7.76	1.73
11749461_a_at	CDH11	cadherin 11, type 2, OB-cadherin (osteoblast)	3.03	7.98	2.13
11728829_at	INHBB	inhibin, beta B	2.89	7.97	2.13
11724322_a_at	CST6	cystatin E/M	2.80	6.99	5.07
11728190_s_at	CXCR4	chemokine (C-X-C motif) receptor 4	2.77	7.40	0.72
11754662_s_at	ADAMTS1	ADAM metalloproteinase with thrombospondin type 1 motif, 1	2.75	6.90	0.00
11720146_a_at	DAPK1	death-associated protein kinase 1	2.71	7.08	1.92
11744581_a_at	KHDRBS3	KH domain containing, RNA binding, signal transduction associated 3	2.68	6.95	0.00
11742710_a_at	SRGN	serglycin	2.63	7.03	0.13
11747952_a_at	TNFSF10	tumor necrosis factor (ligand) superfamily, member 10	2.59	6.84	6.19
11729544_at	MAGEC1	melanoma antigen family C, 1	2.53	6.76	0.72
11715455_x_at	KISS1	KiSS-1 metastasis-suppressor	2.50	7.75	1.06
11724885_at	CLIC3	chloride intracellular channel 3	2.47	7.24	5.61
11715454_at	KISS1	KiSS-1 metastasis-suppressor	2.46	7.44	0.89
11748907_a_at	RARRES3	retinoic acid receptor responder (tazarotene induced) 3	2.46	6.91	6.10
11729543_a_at	MAGEC1	melanoma antigen family C, 1	2.46	6.65	1.00
11743730_at	TNFSF10	tumor necrosis factor (ligand) superfamily, member 10	2.46	6.39	6.19
11719692_a_at	RARRES3	retinoic acid receptor responder (tazarotene induced) 3	2.43	6.83	5.93
11726250_at	AGR2	anterior gradient 2 homolog (Xenopus laevis)	2.41	6.61	2.70
11719172_x_at	AKR1C1	aldo-keto reductase family 1, member C1	2.36	7.51	4.97
11743731_a_at	TNFSF10	tumor necrosis factor (ligand) superfamily, member 10	2.36	6.53	5.90
11719414_a_at	PBX1	pre-B-cell leukemia homeobox 1	2.36	6.48	4.58
11729495_at	FAM133A	family with sequence similarity 133, member A	2.35	6.47	1.99
11727844_a_at	KRT81	keratin 81	2.35	6.17	0.86
11740312_at	HAS2	hyaluronan synthase 2	2.35	6.79	0.46
11732318_a_at	KHDRBS3	KH domain containing, RNA binding, signal transduction associated 3	2.34	6.17	0.06
11754189_s_at	AKR1C1	aldo-keto reductase family 1, member C1	2.32	8.07	4.50
11755928_s_at	MMP1	matrix metalloproteinase 1 (interstitial collagenase)	2.32	6.91	1.84
11719171_a_at	AKR1C1	aldo-keto reductase family 1, member C1	2.31	7.19	4.89
11757871_s_at	PBX1	pre-B-cell leukemia homeobox 1	2.31	5.97	4.14
11729496_at	FAM133A	family with sequence similarity 133, member A	2.29	5.97	1.59
11728189_a_at	CXCR4	chemokine (C-X-C motif) receptor 4	2.29	6.52	1.06
11739696_s_at	AFF3	AF4/FMR2 family, member 3	2.24	5.53	0.24
11746283_s_at	KRTAP2-3	keratin associated protein 2-3	2.22	5.87	1.35
11754879_a_at	SDR16C5	short chain dehydrogenase/reductase family 16C, member 5	2.21	5.98	1.94
11758031_s_at	BHLHE41	basic helix-loop-helix family, member e41	2.21	5.65	0.51
11747096_a_at	KCNK1	potassium channel, subfamily K, member 1	2.18	4.99	0.10
11731538_a_at	GPR110	G protein-coupled receptor 110	2.17	6.23	3.26
11741223_s_at	FAM198B	family with sequence similarity 198, member B	2.17	5.70	5.45
11758102_s_at	TMEM163	transmembrane protein 163	2.17	6.39	2.53
11743350_a_at	C15orf48	chromosome 15 open reading frame 48	2.17	6.63	5.25
11719562_at	SOX4	SRY (sex determining region Y)-box 4	2.16	5.54	0.93
11726812_a_at	TFF1	trefoil factor 1	2.14	6.39	2.88
11721984_at	C7orf29	chromosome 7 open reading frame 29	2.14	5.82	5.42
11727690_at	IL11	interleukin 11	2.13	6.14	1.22
11715711_a_at	AKR1C3	aldo-keto reductase family 1, member C3	2.13	6.67	5.00
11733461_at	SUSD5	sushi domain containing 5	2.12	5.59	0.16
11736687_a_at	FGF5	fibroblast growth factor 5	2.11	5.46	0.81
11737331_at	PDE1C	phosphodiesterase 1C, calmodulin-dependent 70kDa	2.11	5.66	1.12
11757480_x_at	IFI27	interferon, alpha-inducible protein 27	2.10	6.27	3.51
11722888_s_at	EMB	embigin	2.09	5.76	0.54
11727984_at	TENM3	teneurin transmembrane protein 3	2.08	5.57	0.01
11719120_a_at	KYNU	kynureninase	2.07	6.17	1.77
11758227_s_at	ROBO1	roundabout, axon guidance receptor, homolog 1 (Drosophila)	2.06	5.22	0.22
11751833_a_at	BCHE	butyrylcholinesterase	2.06	5.95	1.28
11719384_at	PRSS3	protease, serine, 3	2.06	5.76	1.07
11716329_s_at	GJA1	gap junction protein, alpha 1, 43kDa	2.06	6.71	0.37
11744195_at	FAT3	FAT tumor suppressor homolog 3 (Drosophila)	2.05	5.61	1.02
11724795_at	ZG16B	zymogen granule protein 16B	2.04	5.57	4.19
11736688_x_at	FGF5	fibroblast growth factor 5	2.04	5.33	0.62
11732569_at	SLCO1B3	solute carrier organic anion transporter family, member 1B3	2.04	5.84	0.12

11716017_at	MAL2	mal, T-cell differentiation protein 2 (gene/pseudogene)	2.03	5.95	3.06
11763394_a_at	LOC100505946	uncharacterized LOC100505946	2.00	5.50	2.82
11757532_s_at	RBM47	RNA binding motif protein 47	2.00	5.43	4.58
11725057_s_at	NEDD9	neural precursor cell expressed, developmentally down-regulated 9	1.99	5.70	4.21
11746830_a_at	AGR2	anterior gradient 2 homolog (Xenopus laevis)	1.99	5.84	2.34
11728191_x_at	CXCR4	chemokine (C-X-C motif) receptor 4	1.99	5.54	0.89
11716665_s_at	GDF15	growth differentiation factor 15	1.98	5.92	3.38
11731995_x_at	SPANXC	SPANX family, member C	1.98	6.61	1.21
11751225_s_at	CSF2RA	colony stimulating factor 2 receptor, alpha, low-affinity (granulocyte-macrophage)	1.97	5.75	3.77
11754862_a_at	RARRS2	retinoic acid receptor responder (tazarotene induced) 2	1.97	5.31	4.86
11756473_a_at	CEL2	CUGBP, Elav-like family member 2	1.97	5.53	0.96
11740390_a_at	RBM47	RNA binding motif protein 47	1.96	5.64	4.35
11758525_s_at	SOX4	SRY (sex determining region Y)-box 4	1.96	5.21	1.09
11753942_a_at	UCA1	urothelial cancer associated 1 (non-protein coding)	1.95	5.75	2.64
11730442_s_at	FAM155A	family with sequence similarity 155, member A	1.95	5.37	0.23
11741897_a_at	MMP1	matrix metalloproteinase 1 (interstitial collagenase)	1.95	6.33	2.06
11735833_a_at	KIAA1199	KIAA1199	1.95	6.34	5.55
11757534_s_at	FAM198B	family with sequence similarity 198, member B	1.94	5.34	5.08
11748362_s_at	ABCC3	ATP-binding cassette, sub-family C (CFTR/MRP), member 3	1.94	5.03	4.36
11725521_x_at	GLUL	glutamate-ammonia ligase	1.93	6.86	1.69
11735148_a_at	SOCS2	suppressor of cytokine signaling 2	1.92	5.40	0.00
11758070_s_at	PBX1	pre-B-cell leukemia homeobox 1	1.91	4.75	4.43
11736730_a_at	RIMS2	regulating synaptic membrane exocytosis 2	1.90	5.02	1.15
11739809_at	TMEM156	transmembrane protein 156	1.90	4.63	0.09
11740091_s_at	TMEM163	transmembrane protein 163	1.89	5.94	2.78
11762533_a_at	RAP1A	RAP1A, member of RAS oncogene family	1.89	7.66	7.66
11717762_a_at	CPVL	carboxypeptidase, vitellogenic-like	1.89	5.76	0.92
11717803_a_at	NTN4	netrin 4	1.89	5.25	0.53
11717305_a_at	ADIRF	adipogenesis regulatory factor	1.88	6.15	3.12
11736729_a_at	RIMS2	regulating synaptic membrane exocytosis 2	1.87	4.92	1.25
11726188_at	SHISA3	shisa homolog 3 (Xenopus laevis)	1.87	4.67	0.25
11729647_s_at	ESM1	endothelial cell-specific molecule 1	1.87	5.97	0.82
11757702_s_at	DSC2	desmocollin 2	1.87	5.28	1.39
11723751_at	TFF2	trefoil factor 2	1.86	6.03	3.49
11734808_s_at	DTNA	dystrobrevin, alpha	1.85	4.69	0.69
11740311_at	HAS2	hyaluronan synthase 2	1.85	5.65	0.50
11723464_s_at	CEL2	CUGBP, Elav-like family member 2	1.84	5.70	1.02
11756085_a_at	MAGEC1	melanoma antigen family C, 1	1.84	5.10	0.85
11725517_x_at	ABCG1	ATP-binding cassette, sub-family G (WHITE), member 1	1.84	5.22	2.66
11736685_a_at	FGF5	fibroblast growth factor 5	1.84	4.95	0.92
11718230_a_at	HLA-F	major histocompatibility complex, class I, F	1.83	4.63	4.20
11756072_s_at	SAA1	serum amyloid A1	1.83	6.58	5.83
11754856_a_at	CA12	carbonic anhydrase XII	1.82	6.34	3.87
11740528_a_at	AFF3	AF4/FMR2 family, member 3	1.82	4.90	0.49
11720947_x_at	CA12	carbonic anhydrase XII	1.82	6.42	3.84
11757046_a_at	LOC100127888	uncharacterized LOC100127888	1.82	5.01	2.13
11754601_s_at	SPON2	spondin 2, extracellular matrix protein	1.81	5.09	4.80
11717885_a_at	CCDC3	coiled-coil domain containing 3	1.81	5.05	5.05
11717742_a_at	GPRC5C	G protein-coupled receptor, family C, group 5, member C	1.80	5.52	4.25
11743129_at	SORL1	sortilin-related receptor, L(DLR class) A repeats containing	1.80	5.22	4.14
11755523_a_at	RBM47	RNA binding motif protein 47	1.78	4.83	4.24
11721625_s_at	GLUL	glutamate-ammonia ligase	1.78	6.76	1.88
11734271_a_at	CDH18	cadherin 18, type 2	1.77	4.41	1.01
11744484_s_at	HLA-DRA	major histocompatibility complex, class II, DR alpha	1.77	5.53	2.98
11729020_at	ZNF804A	zinc finger protein 804A	1.76	4.70	0.50
11746149_x_at	BCHE	butyrylcholinesterase	1.76	4.97	1.20
11726192_at	TFAP2C	transcription factor AP-2 gamma (activating enhancer binding protein 2 gamma)	1.76	4.91	0.85
11726333_s_at	LEF1	lymphoid enhancer-binding factor 1	1.75	5.14	0.04
11753915_a_at	SEL1L3	sel-1 suppressor of lin-12-like 3 (C. elegans)	1.74	5.69	1.15
11715893_s_at	IFI27	interferon, alpha-inducible protein 27	1.74	5.49	3.31
11758166_s_at	KAL1	Kallmann syndrome 1 sequence	1.73	6.31	2.30
11730106_a_at	PLCB1	phospholipase C, beta 1 (phosphoinositide-specific)	1.72	4.66	1.44
11719827_a_at	QPCT	glutamyl-peptide cyclotransferase	1.72	5.25	1.35
11759295_at	EHF	ets homologous factor	1.71	6.57	0.90
11755908_s_at	IGSF3	immunoglobulin superfamily, member 3	1.71	5.60	3.50
11732719_at	EREG	epiregulin	1.70	5.56	2.70

11738858_x_at	CSF2RA	colony stimulating factor 2 receptor, alpha, low-affinity (granulocyte-macrophage)	1.70	5.15	3.40
11717721_s_at	ST6GAL1	ST6 beta-galactosamide alpha-2,6-sialyltransferase 1	1.70	4.93	2.00
11721454_a_at	SERPINA1	serpin peptidase inhibitor, clade A (alpha-1 antiproteinase, antitrypsin), member 1	1.69	4.88	4.06
11743856_a_at	CTHRC1	collagen triple helix repeat containing 1	1.69	4.77	2.34
11728203_a_at	PHGDH	phosphoglycerate dehydrogenase	1.69	4.41	0.75
11719828_x_at	QPCT	glutaminyl-peptide cyclotransferase	1.68	5.09	1.42
11716666_a_at	ID3	inhibitor of DNA binding 3, dominant negative helix-loop-helix protein	1.67	5.28	2.57
11734938_x_at	CSF2RA	colony stimulating factor 2 receptor, alpha, low-affinity (granulocyte-macrophage)	1.67	4.96	3.00
11744776_a_at	NNMT	nicotinamide N-methyltransferase	1.67	5.70	3.61
11716663_a_at	GDF15	growth differentiation factor 15	1.67	4.82	3.14
11757396_s_at	ANXA8L2	annexin A8-like 2	1.67	5.14	3.18
11751612_a_at	SOCS2	suppressor of cytokine signaling 2	1.67	4.81	0.34
11715918_s_at	CD24	CD24 molecule	1.67	5.31	1.57
11730048_at	SLITRK5	SLIT and NTRK-like family, member 5	1.67	4.24	0.62
11722991_a_at	DMD	dystrophin	1.66	4.29	1.49
11715467_s_at	ARHGDI3	Rho GDP dissociation inhibitor (GDI) beta	1.66	6.44	0.39
11754796_a_at	SLCO3A1	solute carrier organic anion transporter family, member 3A1	1.66	4.64	3.38
11720409_s_at	CYP1B1	cytochrome P450, family 1, subfamily B, polypeptide 1	1.66	5.00	2.94
11727583_a_at	EPHA3	EPH receptor A3	1.66	4.95	1.10
11718506_at	TFF3	trefoil factor 3 (intestinal)	1.66	4.99	3.82
11732158_a_at	ADAMTS1	ADAM metalloproteinase with thrombospondin type 1 motif, 1	1.65	4.38	0.01
11746930_a_at	BHLHE41	basic helix-loop-helix family, member e41	1.65	4.43	0.81
11719198_s_at	CCND2	cyclin D2	1.65	4.53	1.30
11750552_s_at	CARD16	caspace recruitment domain family, member 16	1.65	5.45	1.58
11764166_s_at	EHF	ets homologous factor	1.65	6.42	0.98
11725515_a_at	ABCG1	ATP-binding cassette, sub-family G (WHITE), member 1	1.65	4.68	2.19
11725516_s_at	ABCG1	ATP-binding cassette, sub-family G (WHITE), member 1	1.65	4.74	3.11
11744718_a_at	CTHRC1	collagen triple helix repeat containing 1	1.64	4.22	1.79
11742850_x_at	HLA-DRA	major histocompatibility complex, class II, DR alpha	1.64	4.90	3.08
11751745_x_at	PRSS3	protease, serine, 3	1.64	4.58	1.11
11746094_a_at	MPP4	membrane protein, palmitoylated 4 (MAGUK p55 subfamily member 4)	1.63	4.56	2.18
11739094_a_at	CXCR4	chemokine (C-X-C motif) receptor 4	1.62	4.92	0.97
11732870_a_at	CASP1	caspace 1, apoptosis-related cysteine peptidase	1.62	4.48	0.87
11719561_s_at	SOX4	SRY (sex determining region Y)-box 4	1.62	5.25	1.96
11720277_a_at	OLFML2A	olfactomedin-like 2A	1.62	5.06	4.09
11733091_a_at	TSPAN8	tetraspanin 8	1.62	5.10	1.98
11741453_at	SPINK4	serine peptidase inhibitor, Kazal type 4	1.61	4.46	3.97
11718620_a_at	CHRDL1	chordin-like 1	1.61	4.45	1.32
11759905_a_at	EXD3	exonuclease 3'-5' domain containing 3	1.60	4.66	4.39
11726396_x_at	KAL1	Kallmann syndrome 1 sequence	1.60	5.80	2.29
11719447_s_at	GBP2	guanylate binding protein 2, interferon-inducible	1.59	4.63	3.07
11740782_a_at	RNF128	ring finger protein 128, E3 ubiquitin protein ligase	1.59	4.36	3.69
11743197_at	TLR4	toll-like receptor 4	1.59	4.81	1.60
11745722_x_at	FHL1	four and a half LIM domains 1	1.59	5.55	2.43
11715624_at	SH3BGR1	SH3 domain binding glutamic acid-rich protein like	1.59	6.73	0.80
11725058_a_at	NEDD9	neural precursor cell expressed, developmentally down-regulated 9	1.58	4.32	3.70
11726458_s_at	GSPT2	G1 to S phase transition 2	1.58	4.75	2.95
11738935_a_at	KLRC3	killer cell lectin-like receptor subfamily C, member 3	1.58	5.01	4.65
11744079_s_at	PRSS2	protease, serine, 2 (trypsin 2)	1.58	5.79	4.28
11746463_a_at	IL6	interleukin 6 (interferon, beta 2)	1.58	5.53	4.82
11747104_s_at	CYP1B1	cytochrome P450, family 1, subfamily B, polypeptide 1	1.58	4.49	3.13
11720410_s_at	CYP1B1	cytochrome P450, family 1, subfamily B, polypeptide 1	1.58	4.90	2.66
11718904_s_at	FILIP1L	filamin A interacting protein 1-like	1.58	5.96	2.17
11755282_a_at	SCNN1A	sodium channel, non-voltage-gated 1 alpha subunit	1.58	4.51	4.20
11732993_a_at	RGS4	regulator of G-protein signaling 4	1.58	5.30	1.31
11726559_s_at	LCP1	lymphocyte cytosolic protein 1 (L-plastin)	1.58	5.31	2.47
11730184_at	CST1	cystatin SN	1.58	5.16	3.45
11750623_a_at	FILIP1L	filamin A interacting protein 1-like	1.57	5.31	2.67
11731465_a_at	CTSC	cathepsin C	1.57	4.41	0.00
11744848_x_at	PRR16	proline rich 16	1.57	4.79	0.28
11755141_a_at	CACNA2D4	calcium channel, voltage-dependent, alpha 2/delta subunit 4	1.57	4.72	3.24
11718347_a_at	S100P	S100 calcium binding protein P	1.57	4.87	3.26
11758952_at	AGR2	anterior gradient 2 homolog (Xenopus laevis)	1.56	4.68	3.30
11742873_a_at	CDH11	cadherin 11, type 2, OB-cadherin (osteoblast)	1.56	4.71	2.14
11729348_at	HMG5	high mobility group nucleosome binding domain 5	1.56	4.52	0.84
11742923_a_at	LAPTM5	lysosomal protein transmembrane 5	1.56	5.59	2.30
11726678_a_at	AFAP1L2	actin filament associated protein 1-like 2	1.56	5.53	2.95
11720411_s_at	CYP1B1	cytochrome P450, family 1, subfamily B, polypeptide 1	1.56	4.84	2.70

11722348_a_at	REP52	RALBP1 associated Eps domain containing 2	LT>T	2.01	1.12E-03 T>M'	0.32	5.89E-01 LT>M'	2.33	1.38E-06	TRUE
11732625_a_at	RIMS1	regulating synaptic membrane exocytosis 1	T>LT	-0.50	4.59E-02 M>T	-1.85	2.82E-04 M>LT	-2.35	3.51E-08	TRUE
11736729_a_at	RIMS2	regulating synaptic membrane exocytosis 2	T>LT	0.00	1.00E+00 M>T	-3.15	1.21E-02 M>LT	-3.15	7.87E-04	TRUE
11736730_a_at	RIMS2	regulating synaptic membrane exocytosis 2	LT>T	0.24	7.91E-01 M>T	-3.34	9.85E-03 M>LT	-3.10	9.81E-04	TRUE
11757347_s_at	RNF144A	ring finger protein 144A	LT>T	2.47	3.31E-03 T>M'	0.01	9.54E-01 LT>M'	2.48	6.37E-03	TRUE
11758227_s_at	ROBO1	roundabout, axon guidance receptor, homolog 1 (Drosophila)	T>LT	-3.04	3.23E-03 M>T	-0.93	2.16E-02 M>LT	-3.98	1.40E-03	TRUE
11718347_x_at	S100P	S100 calcium binding protein P	T>LT	-2.38	1.58E-02 T>M'	2.04	9.46E-02 M>LT	-0.35	2.59E-01	TRUE
11733442_x_at	SAA1	serum amyloid A1	LT>T	0.61	4.54E-01 T>M'	1.79	3.47E-02 LT>M'	2.40	6.18E-03	TRUE
11756072_s_at	SAA1	serum amyloid A1	LT>T	0.69	4.23E-01 T>M'	2.66	1.61E-02 LT>M'	3.35	1.31E-03	TRUE
11755282_a_at	SCNN1A	sodium channel, non-voltage-gated 1 alpha subunit	LT>T	1.33	9.16E-02 T>M'	1.73	3.95E-02 LT>M'	3.06	3.41E-04	TRUE
11754879_a_at	SDR16C5	short chain dehydrogenase/reductase family 16C, member 5	T>LT	-3.86	2.67E-03 T>M'	1.23	3.70E-01 M>LT	-2.63	4.36E-04	TRUE
11753915_a_at	SELLL3	sel-1 suppressor of lin-12-like 3 (C. elegans)	T>LT	-1.38	1.35E-01 M>T	-1.53	1.79E-01 M>LT	-2.91	3.50E-03	TRUE
11730390_at	SEMA3A	sema domain, immunoglobulin domain (Ig), short basic domain, secreted, (semaphorin) 3A	T>LT	-1.77	1.13E-02 M>T	-0.56	3.30E-01 M>LT	-2.33	3.63E-03	TRUE
11721454_a_at	SERPINA1	serpin peptidase inhibitor, clade A (alpha-1 antiproteinase, antitrypsin), member 1	LT>T	2.11	3.99E-02 T>M'	0.58	4.50E-01 LT>M'	2.69	8.76E-03	TRUE
11721455_s_at	SERPINA1	serpin peptidase inhibitor, clade A (alpha-1 antiproteinase, antitrypsin), member 1	LT>T	1.82	2.94E-02 T>M'	0.52	4.10E-01 LT>M'	2.34	5.20E-03	TRUE
11744315_at	SGK223	homolog of rat pragra of Rnd2	LT>T	0.27	2.86E-01 T>M'	2.21	2.82E-03 LT>M'	2.48	2.72E-04	TRUE
11718447_s_at	SH3BGR2	SH3 domain binding glutamic acid-rich protein like 2	T>LT	-1.60	2.96E-03 M>T	-1.23	3.56E-02 M>LT	-2.82	3.06E-07	FALSE
11726188_at	SHISA3	shisa homolog 3 (Xenopus laevis)	T>LT	-2.83	2.08E-03 M>T	-0.62	5.24E-01 M>LT	-3.45	1.17E-05	TRUE
11729140_at	SLAMF7	SLAM family member 7	T>LT	-3.20	1.38E-03 T>M'	2.13	2.46E-03 M>LT	-1.08	1.14E-02	TRUE
11749023_a_at	SLC16A4	solute carrier family 16, member 4 (monocarboxylic acid transporter 5)	LT>T	2.63	1.18E-03 M>T	0.04	6.66E-01 LT>M'	2.58	3.47E-03	TRUE
11749024_x_at	SLC16A4	solute carrier family 16, member 4 (monocarboxylic acid transporter 5)	LT>T	2.36	2.71E-03 T>M'	0.04	6.24E-01 LT>M'	2.40	6.11E-03	TRUE
11733491_a_at	SLC44A5	solute carrier family 44, member 5	T>LT	-2.12	5.61E-03 M>T	-0.54	5.10E-01 M>LT	-2.66	2.29E-06	TRUE
11733492_s_at	SLC44A5	solute carrier family 44, member 5	T>LT	-2.30	3.01E-03 M>T	-0.31	7.07E-01 M>LT	-2.62	7.93E-05	FALSE
11747964_a_at	SLC47A	solute carrier family 7, sodium bicarbonate cotransporter, member 7	T>LT	-1.72	1.56E-03 M>T	-0.98	1.04E-01 M>LT	-2.70	9.29E-04	TRUE
11733042_at	SLC7A2	solute carrier family 7 (cationic amino acid transporter, y+ system), member 2	T>LT	-1.11	4.80E-02 M>T	-1.24	7.87E-02 M>LT	-2.36	1.41E-04	TRUE
11732569_at	SLC10B3	solute carrier organic anion transporter family, member 1B3	T>LT	-2.88	3.19E-03 M>T	-1.21	3.29E-02 M>LT	-4.08	4.32E-04	TRUE
11754796_a_at	SLC30A1	solute carrier organic anion transporter family, member 3A1	T>LT	1.77	3.20E-02 T>M'	1.50	1.28E-01 LT>M'	3.27	4.59E-07	TRUE
11758172_s_at	SLC30A1	solute carrier organic anion transporter family, member 3A1	LT>T	1.18	1.52E-02 T>M'	1.25	2.82E-02 LT>M'	2.43	1.17E-06	TRUE
11730048_at	SLITRK5	SLIT and NTRK-like family, member 5	T>LT	-1.84	2.61E-02 M>T	-1.43	1.05E-01 M>LT	-3.28	3.99E-05	TRUE
11725808_x_at	SNAIP1	synuclein, alpha interacting protein	T>LT	-0.19	5.04E-01 M>T	-2.29	3.42E-03 M>LT	-2.48	1.47E-03	TRUE
11747095_x_at	SNAIP1	synuclein, alpha interacting protein	T>LT	-0.10	7.08E-01 M>T	-2.56	2.79E-03 M>LT	-2.66	1.12E-03	TRUE
11750730_x_at	SNAIP1	synuclein, alpha interacting protein	T>LT	-0.11	6.83E-01 M>T	-2.40	3.70E-03 M>LT	-2.51	1.11E-03	TRUE
11735148_a_at	SOC22	suppressor of cytokine signaling 2	T>LT	-3.02	8.86E-04 M>T	-0.76	8.50E-02 M>LT	-3.78	3.29E-04	TRUE
11751612_x_at	SOC22	suppressor of cytokine signaling 2	T>LT	-2.58	1.02E-03 M>T	-0.71	1.19E-01 M>LT	-3.29	2.86E-04	FALSE
11735570_a_at	SORBS2	sorbin and SH3 domain containing 2	LT>T	0.59	3.43E-01 T>M'	1.98	9.55E-03 LT>M'	2.57	5.67E-04	TRUE
11758588_x_at	SORCS2	sorilin-related VPS10 domain containing receptor 2	LT>T	2.85	2.87E-08 T>M'	0.35	8.22E-02 LT>M'	3.20	3.25E-09	FALSE
11743129_at	SORL1	sorilin-related receptor, (LDLR class) A repeats containing	LT>T	1.97	1.75E-04 T>M'	2.19	9.35E-04 LT>M'	4.16	6.25E-07	FALSE
11743130_at	SORL1	sorilin-related receptor, (LDLR class) A repeats containing	LT>T	1.77	9.93E-05 T>M'	1.80	1.10E-03 LT>M'	3.57	1.69E-07	TRUE
11719560_at	SOX4	SRY (sex determining region Y)-box 4	T>LT	-2.89	2.54E-04 T>M'	1.20	4.93E-02 M>LT	-1.69	4.13E-03	TRUE
11719561_s_at	SOX4	SRY (sex determining region Y)-box 4	T>LT	-3.16	2.96E-04 T>M'	1.12	5.35E-02 M>LT	-2.04	5.65E-03	TRUE
11719562_at	SOX4	SRY (sex determining region Y)-box 4	T>LT	-4.17	7.58E-05 T>M'	0.66	1.74E-01 M>LT	-3.51	6.32E-04	TRUE
11757741_s_at	SOX4	SRY (sex determining region Y)-box 4	T>LT	-2.40	4.98E-05 T>M'	0.63	1.01E-01 M>LT	-1.78	5.62E-04	TRUE
11758525_s_at	SOX4	SRY (sex determining region Y)-box 4	T>LT	-3.81	9.37E-05 T>M'	0.75	9.33E-02 M>LT	-3.06	1.05E-03	FALSE
11726177_s_at	SPANXB2	SPANX family, member B2	T>LT	-6.74	1.52E-05 T>M'	0.34	8.42E-01 M>LT	-6.40	7.02E-04	FALSE
11738785_at	SPANXB2	SPANX family, member B2	T>LT	-7.69	1.27E-06 T>M'	0.48	7.33E-01 M>LT	-7.20	2.46E-04	FALSE
11738786_x_at	SPANXB2	SPANX family, member B2	T>LT	-7.21	5.43E-06 T>M'	0.34	8.33E-01 M>LT	-6.87	3.69E-04	FALSE
11731995_x_at	SPANXC	SPANX family, member C	T>LT	-1.20	8.77E-04 M>T	-2.78	2.92E-02 M>LT	-3.98	1.98E-03	FALSE
11731130_at	SPANXE	SPANX family, member E	T>LT	-0.62	4.61E-02 M>T	-1.94	8.02E-02 M>LT	-2.57	1.41E-02	TRUE
11731994_s_at	SPANXE	SPANX family, member E	T>LT	-2.39	1.09E-01 M>T	-5.59	1.71E-02 M>LT	-7.97	6.16E-07	TRUE
11759051_x_at	SPANXE	SPANX family, member E	T>LT	-0.60	1.01E-01 M>T	-1.90	7.01E-02 M>LT	-2.49	9.32E-03	TRUE
11741453_x	SPINK4	serine peptidase inhibitor, Kazal type 4	LT>T	1.94	4.84E-03 T>M'	1.51	4.60E-02 LT>M'	3.45	4.54E-07	TRUE
11754601_s_at	SPON2	spondin 2, extracellular matrix protein	T>LT	2.92	3.07E-03 T>M'	0.18	4.67E-01 LT>M'	3.10	4.95E-03	TRUE
11742710_a_at	SRGN	serglycin	T>LT	-3.97	1.06E-03 M>T	-1.37	4.31E-02 M>LT	-5.34	1.64E-04	FALSE
11717721_s_at	ST6GAL1	ST6 beta-galactosidase alpha-2,6-sialyltransferase 1	T>LT	-2.94	1.55E-03 T>M'	0.73	4.08E-01 M>LT	-2.21	9.59E-03	TRUE
11716055_s_at	STMN3	stathmin-like 3	LT>T	1.33	1.67E-02 T>M'	1.10	3.92E-02 LT>M'	2.42	1.44E-04	FALSE
11733461_at	SUSO5	sushi domain containing 5	T>LT	-2.84	3.14E-03 M>T	-1.40	1.77E-01 M>LT	-4.24	2.68E-05	TRUE
11717301_at	TACSTD2	tumor-associated calcium signal transducer 2	LT>T	1.96	6.44E-04 T>M'	1.40	9.50E-03 LT>M'	3.36	2.79E-06	TRUE
11727984_at	TENM3	tenascin transmembrane protein 3	T>LT	-2.56	3.97E-02 M>T	-0.85	2.26E-01 M>LT	-3.41	1.27E-02	TRUE
11726192_at	TFA2P2C	transcription factor AP-2 gamma (activating enhancer binding protein 2 gamma)	T>LT	-1.59	8.94E-02 M>T	-1.68	1.27E-01 M>LT	-3.27	3.48E-05	TRUE
11726812_a_at	TFPI1	trefoil factor 1	T>LT	-3.19	1.49E-02 T>M'	2.39	1.53E-01 M>LT	-0.80	2.42E-01	TRUE
11723751_at	TFPI2	trefoil factor 2	T>LT	-2.44	2.64E-02 T>M'	2.86	4.47E-02 LT>M'	0.42	5.47E-01	TRUE
11718506_at	TFPI3	trefoil factor 3 (intestinal)	T>LT	-2.54	1.23E-02 T>M'	2.54	4.20E-02 M>LT	0.00	9.98E-01	TRUE
11728302_at	TGFA	transforming growth factor, alpha	LT>T	0.57	6.78E-02 T>M'	1.77	1.03E-02 LT>M'	2.34	8.76E-04	TRUE
11734548_a_at	TGFB1	transforming growth factor, beta-induced, 68kDa	LT>T	1.29	2.94E-02 T>M'	1.14	6.49E-02 LT>M'	2.44	1.11E-03	TRUE
11734549_s_at	TGFB1	transforming growth factor, beta-induced, 68kDa	LT>T	1.21	5.38E-02 T>M'	1.22	1.04E-01 LT>M'	2.42	2.80E-03	TRUE
11734550_x_at	TGFB1	transforming growth factor, beta-induced, 68kDa	LT>T	1.24	4.26E-02 T>M'	1.27	7.53E-02 LT>M'	2.51	1.47E-03	TRUE
11743197_at	TLR4	toll-like receptor 4	T>LT	-3.05	1.16E-04 T>M'	0.50	2.32E-01 M>LT	-2.55	7.18E-04	FALSE
11731603_at	TMEM154	transmembrane protein 154	T>LT	-1.31	1.03E-02 M>T	-1.24	1.59E-03 M>LT	-2.55	5.68E-04	TRUE
11758583_s_at	TMEM154	transmembrane protein 154	T>LT	-1.42	2.41E-02 M>T	-1.05	1.65E-02 M>LT	-2.48	2.89E-03	TRUE
11739809_at	TMEM156	transmembrane protein 156	T>LT	-3.40	2.57E-06 M>T	-0.48	2.51E-01 M>LT	-3.88	3.90E-08	TRUE
11740091_s_at	TMEM163	transmembrane protein 163	T>LT	-2.22	6.20E-02 M>T	-0.55	6.90E-01 M>LT	-2.77	1.48E-05	TRUE
11758102_s_at	TMEM163	transmembrane protein 163	T>LT	-2.72	3.90E-02 M>T	-0.27	6.32E-01 M>LT	-3.42	3.30E-06	TRUE
11743730_at	TNFSF10	tumor necrosis factor (ligand) superfamily, member 10	LT>T	3.31	8.67E-05 T>M'	0.71	4.90E-03 LT>M'	5.57	1.50E-08	TRUE
11743731_a_at	TNFSF10	tumor necrosis factor (ligand) superfamily, member 10	LT>T	3.17	1.75E-05 T>M'	2.26	5.61E-04 LT>M'	5.43	1.35E-07	FALSE
11747952_x_at	TNFSF10	tumor necrosis factor (ligand) superfamily, member 10	LT>T	3.41	1.06E-04 T>M'	2.48	4.95E-03 LT>M'	5.89	1.51E-08	FALSE
11733091_a_at	TSPAN8	tetraspanin 8	T>LT	-3.09	2.49E-04 T>M'	0.73	1.96E-01 M>LT	-2.35	1.54E-03	TRUE
11755851_a_at	UBR4	ubiquitin protein ligase E3 component n-recognin 4	T>LT	-1.04	8.16E-03 M>T	-1.33	4.35E-03 M>LT	-2.37	4.41E-05	TRUE
11758628_x_at	ULK2	unc-51-like kinase 2 (C. elegans)	LT>T	1.80	6.52E-03 T>M'	0.70	2.86E-01 LT>M'	2.50	1.27E-03	FALSE
11732465_a_at	USP44	ubiquitin specific peptidase 44	T>LT	-1.15	9.18E-03 M>T	-1.29	1.14E-02 M>LT	-2.44	4.19E-06	TRUE
11740301_a_at	USP44	ubiquitin specific peptidase 44	T>LT	-1.33	1.78E-03 M>T	-1.31	9.22E-03 M>LT	-2.64	4.26E-07	TRUE
11724795_at	ZG16B	zymogen granule protein 16B	T>LT	-3.58	3.74E-03 T>M'	2.74	5.15E-02 M>LT	-0.84	4.88E-04	TRUE
11729410_a_at	ZNF415	zinc finger protein 415	LT>T	0.80	7.29E-02 T>M'	1.61	4.06E-02 LT>M'	2.41	9.09E-04	FALSE
11729020_at	ZNF804A	zinc finger protein 804A	T>LT	-2.03	2.04E-04 M>T	-2.06	9.23E-04 M>LT	-4.09	6.38E-08	FALSE

* Threshold value for Fold difference is 2.32 in log2 and P < 0.05; Genes that have significant difference between Groups are highlighted in red

Table S4. List of genes affiliated with regulation of cell shape.

CLU(LT>M')	clusterin
COCH(LT>M')	coagulation factor C homolog, cochlin (Limulus polyphemus)
DCLK1(M'>T,M'>LT)	doublecortin-like kinase 1
DOCK10(M'>LT)	dedicator of cytokinesis 10
GJA1(M'>LT)	gap junction protein, alpha 1, 43kDa
HAS2(M'>LT)	hyaluronan synthase 2
ICAM1(LT>M')	intercellular adhesion molecule 1
IL6(LT>M')	interleukin 6 (interferon, beta 2)
IL7R(M'>LT)	interleukin 7 receptor
LEF1(T>LT,M'>LT)	lymphoid enhancer-binding factor 1
MAP2(LT>T)	microtubule-associated protein 2
NRXN3(M'>LT)	neurexin 3
NTN4(T>LT,M'>LT)	netrin 4
PARD6B(LT>M')	par-6 partitioning defective 6 homolog beta (C. elegans)
RIMS1(M'>LT)	regulating synaptic membrane exocytosis 1
RIMS2(M'>T,M'>LT)	regulating synaptic membrane exocytosis 2
ROBO1(T>LT,M'>LT)	roundabout, axon guidance receptor, homolog 1 (Drosophila)
SEMA3A(M'>LT)	sema domain, immunoglobulin domain (Ig), short basic domain, secr
SLITRK5(M'>LT)	SLIT and NTRK-like family, member 5
SPON2(LT>T,LT>M')	spondin 2, extracellular matrix protein
TACSTD2(LT>M')	tumor-associated calcium signal transducer 2
ULK2(LT>M')	unc-51-like kinase 2 (C. elegans)

Table S5. Summary of CM features.

Feature name	Feature category	shape type	Feature description
f1	Basic	Nucleus	area
f2	Basic	Nucleus	perimeter
f3	Basic	Nucleus	long axis length
f4	Basic	Nucleus	short axis length
f5	Basic	Nucleus	angle
f6	Basic	Nucleus	solidity
f7	Basic	Nucleus	equi diameter
f8	Basic	Nucleus	aspect ratio
f9	Basic	Nucleus	circularity
f10	Basic	Nucleus	roundness
f11	Basic	Cell	area
f12	Basic	Cell	perimeter
f13	Basic	Cell	long axis length
f14	Basic	Cell	short axis length
f15	Basic	Cell	angle
f16	Basic	Cell	solidity
f17	Basic	Cell	equi diameter
f18	Basic	Cell	aspect ratio
f19	Basic	Cell	circularity
f20	Basic	Cell	roundness
f21	Boundary signature	Nucleus	mean(R) ($\equiv \langle R \rangle$)
f22	Boundary signature	Nucleus	median(R)
f23	Boundary signature	Nucleus	mode(R)
f24	Boundary signature	Nucleus	maR(R)
f25	Boundary signature	Nucleus	min(R)
f26	Boundary signature	Nucleus	25% percentile (R)
f27	Boundary signature	Nucleus	75% percentile (R)
f28	Boundary signature	Nucleus	mean(R(R > f27));
f29	Boundary signature	Nucleus	mean(R(R < f26));
f30	Boundary signature	Nucleus	$\Sigma(R)$
f31	Boundary signature	Nucleus	harmonic mean(R)
f32	Boundary signature	Nucleus	3% trimmed mean(R)
f33	Boundary signature	Nucleus	5% trimmed mean(R)
f34	Boundary signature	Nucleus	15% trimmed mean(R)
f35	Boundary signature	Nucleus	25% trimmed mean(R)
f36	Boundary signature	Nucleus	standard deviation(R) ($\equiv \sigma_R$)
f37	Boundary signature	Nucleus	$\sigma_R / \langle R \rangle$
f38	Boundary signature	Nucleus	$\sigma_R / \text{median}(R) $
f39	Boundary signature	Nucleus	$\sigma_R / \text{mode}(R) $
f40	Boundary signature	Nucleus	skewness(R)
f41	Boundary signature	Nucleus	kurtosis(R)
f42	Boundary signature	Nucleus	mean(R - $\langle R \rangle$)
f43	Boundary signature	Nucleus	range(X)
f44	Boundary signature	Nucleus	interquartile range (X)
f45	Boundary signature	Nucleus	$\Sigma(R^2)$
f46	Boundary signature	Nucleus	$\Sigma(R^3)$
f47	Boundary signature	Nucleus	mean(R ²)
f48	Boundary signature	Nucleus	mean(R ³)
f49	Boundary signature	Nucleus	mean(R ⁴)
f50	Boundary signature	Nucleus	mean(R ⁵)
f51	Boundary signature	Nucleus	$\Sigma(R - \langle R \rangle ^2)$
f52	Boundary signature	Nucleus	$\Sigma(R - \langle R \rangle ^3)$
f53	Boundary signature	Nucleus	mean(R - $\langle R \rangle$ ²)
f54	Boundary signature	Nucleus	mean(R - $\langle R \rangle$ ³)
f55	Boundary signature	Nucleus	mean(R - $\langle R \rangle$ ⁴)
f56	Boundary signature	Nucleus	mean(R - $\langle R \rangle$ ⁵)
f57	Boundary signature	Nucleus	number of peaks
f58	Boundary signature	Nucleus	gini coefficient (R)
f59	Boundary signature	Cell	mean(R) ($\equiv \langle R \rangle$)
f60	Boundary signature	Cell	median(R)
f61	Boundary signature	Cell	mode(R)
f62	Boundary signature	Cell	maR(R)
f63	Boundary signature	Cell	min(R)
f64	Boundary signature	Cell	25% percentile (R)
f65	Boundary signature	Cell	75% percentile (R)
f66	Boundary signature	Cell	mean(R(R > f65));
f67	Boundary signature	Cell	mean(R(R < f64));
f68	Boundary signature	Cell	$\Sigma(R)$

f69	Boundary signature	Cell	harmonic mean(R)
f70	Boundary signature	Cell	3% trimmed mean(R)
f71	Boundary signature	Cell	5% trimmed mean(R)
f72	Boundary signature	Cell	15% trimmed mean(R)
f73	Boundary signature	Cell	25% trimmed mean(R)
f74	Boundary signature	Cell	standard deviation(R) ($\equiv \sigma_R$)
f75	Boundary signature	Cell	$\sigma_R/ <R> $
f76	Boundary signature	Cell	$\sigma_R/ \text{median}(R) $
f77	Boundary signature	Cell	$\sigma_R/ \text{mode}(R) $
f78	Boundary signature	Cell	skewness(R)
f79	Boundary signature	Cell	kurtosis(R)
f80	Boundary signature	Cell	mean(R-<R>)
f81	Boundary signature	Cell	range(X)
f82	Boundary signature	Cell	interquartile range (X)
f83	Boundary signature	Cell	$\Sigma(R^2)$
f84	Boundary signature	Cell	$\Sigma(R^3)$
f85	Boundary signature	Cell	mean(R^2)
f86	Boundary signature	Cell	mean(R^3)
f87	Boundary signature	Cell	mean(R^4)
f88	Boundary signature	Cell	mean(R^5)
f89	Boundary signature	Cell	$\Sigma(R-<R> ^2)$
f90	Boundary signature	Cell	$\Sigma(R-<R> ^3)$
f91	Boundary signature	Cell	mean(R-<R> ^2)
f92	Boundary signature	Cell	mean(R-<R> ^3)
f93	Boundary signature	Cell	mean(R-<R> ^4)
f94	Boundary signature	Cell	mean(R-<R> ^5)
f95	Boundary signature	Cell	number of peaks
f96	Boundary signature	Cell	gini coefficient (R)
f97	Curvature	Nucleus	mean(κ) ($\equiv \langle \kappa \rangle$)
f98	Curvature	Nucleus	median(κ)
f99	Curvature	Nucleus	mode(κ)
f100	Curvature	Nucleus	max(κ)
f101	Curvature	Nucleus	min(κ)
f102	Curvature	Nucleus	25% percentile (κ)
f103	Curvature	Nucleus	75% percentile (κ)
f104	Curvature	Nucleus	mean($\kappa(\kappa > f103)$);
f105	Curvature	Nucleus	mean($\kappa(\kappa < f102)$);
f106	Curvature	Nucleus	$\Sigma(\kappa)$
f107	Curvature	Nucleus	harmonic mean(κ)
f108	Curvature	Nucleus	3% trimmed mean(κ)
f109	Curvature	Nucleus	5% trimmed mean(κ)
f110	Curvature	Nucleus	15% trimmed mean(κ)
f111	Curvature	Nucleus	25% trimmed mean(κ)
f112	Curvature	Nucleus	standard deviation(κ) ($\equiv \sigma_\kappa$)
f113	Curvature	Nucleus	$\sigma_\kappa/ \langle \kappa \rangle $
f114	Curvature	Nucleus	$\sigma_\kappa/ \text{median}(\kappa) $
f115	Curvature	Nucleus	$\sigma_\kappa/ \text{mode}(\kappa) $
f116	Curvature	Nucleus	skewness(κ)
f117	Curvature	Nucleus	kurtosis(κ)
f118	Curvature	Nucleus	mean($\kappa-\langle \kappa \rangle$)
f119	Curvature	Nucleus	range(κ)
f120	Curvature	Nucleus	interquartile range (κ)
f121	Curvature	Nucleus	$\Sigma(\kappa^2)$
f122	Curvature	Nucleus	$\Sigma(\kappa^3)$
f123	Curvature	Nucleus	mean(κ^2)
f124	Curvature	Nucleus	mean(κ^3)
f125	Curvature	Nucleus	mean(κ^4)
f126	Curvature	Nucleus	mean(κ^5)
f127	Curvature	Nucleus	$\Sigma(\kappa-\langle \kappa \rangle ^2)$
f128	Curvature	Nucleus	$\Sigma(\kappa-\langle \kappa \rangle ^3)$
f129	Curvature	Nucleus	mean($\kappa-\langle \kappa \rangle$ ^2)
f130	Curvature	Nucleus	mean($\kappa-\langle \kappa \rangle$ ^3)
f131	Curvature	Nucleus	mean($\kappa-\langle \kappa \rangle$ ^4)
f132	Curvature	Nucleus	mean($\kappa-\langle \kappa \rangle$ ^5)
f133	Curvature	Nucleus	number of peaks
f134	Curvature	Nucleus	gini coefficient (κ)
f135	Curvature	Cell	mean(κ) ($\equiv \langle \kappa \rangle$)

f136	Curvature	Cell	median(κ)
f137	Curvature	Cell	mode(κ)
f138	Curvature	Cell	max(κ)
f139	Curvature	Cell	min(κ)
f140	Curvature	Cell	25% percentile (κ)
f141	Curvature	Cell	75% percentile (κ)
f142	Curvature	Cell	mean($\kappa(\kappa > f141)$);
f143	Curvature	Cell	mean($\kappa(\kappa < f140)$);
f144	Curvature	Cell	$\Sigma(\kappa)$
f145	Curvature	Cell	harmonic mean(κ)
f146	Curvature	Cell	3% trimmed mean(κ)
f147	Curvature	Cell	5% trimmed mean(κ)
f148	Curvature	Cell	15% trimmed mean(κ)
f149	Curvature	Cell	25% trimmed mean(κ)
f150	Curvature	Cell	standard deviation(κ) ($\equiv \sigma_{\kappa}$)
f151	Curvature	Cell	$\sigma_{\kappa}/ \langle \kappa \rangle $
f152	Curvature	Cell	$\sigma_{\kappa}/ \text{median}(\kappa) $
f153	Curvature	Cell	$\sigma_{\kappa}/ \text{mode}(\kappa) $
f154	Curvature	Cell	skewness(κ)
f155	Curvature	Cell	kurtosis(κ)
f156	Curvature	Cell	mean($ \kappa - \langle \kappa \rangle $)
f157	Curvature	Cell	range(κ)
f158	Curvature	Cell	interquartile range (κ)
f159	Curvature	Cell	$\Sigma(\kappa^2)$
f160	Curvature	Cell	$\Sigma(\kappa^3)$
f161	Curvature	Cell	mean(κ^2)
f162	Curvature	Cell	mean(κ^3)
f163	Curvature	Cell	mean(κ^4)
f164	Curvature	Cell	mean(κ^5)
f165	Curvature	Cell	$\Sigma(\kappa - \langle \kappa \rangle ^2)$
f166	Curvature	Cell	$\Sigma(\kappa - \langle \kappa \rangle ^3)$
f167	Curvature	Cell	mean($ \kappa - \langle \kappa \rangle ^2$)
f168	Curvature	Cell	mean($ \kappa - \langle \kappa \rangle ^3$)
f169	Curvature	Cell	mean($ \kappa - \langle \kappa \rangle ^4$)
f170	Curvature	Cell	mean($ \kappa - \langle \kappa \rangle ^5$)
f171	Curvature	Cell	number of peaks
f172	Curvature	Cell	gini coefficient (κ)
f173	NC-positioning signature	Nucleus & Cell	mean(R') ($\equiv \langle R' \rangle$)
f174	NC-positioning signature	Nucleus & Cell	median(R')
f175	NC-positioning signature	Nucleus & Cell	mode(R')
f176	NC-positioning signature	Nucleus & Cell	ma R' (R')
f177	NC-positioning signature	Nucleus & Cell	min(R')
f178	NC-positioning signature	Nucleus & Cell	25% percentile (R')
f179	NC-positioning signature	Nucleus & Cell	75% percentile (R')
f180	NC-positioning signature	Nucleus & Cell	$\Sigma(R')$
f181	NC-positioning signature	Nucleus & Cell	harmonic mean(R')
f182	NC-positioning signature	Nucleus & Cell	3% trimmed mean(R')
f183	NC-positioning signature	Nucleus & Cell	5% trimmed mean(R')
f184	NC-positioning signature	Nucleus & Cell	15% trimmed mean(R')
f185	NC-positioning signature	Nucleus & Cell	25% trimmed mean(R')
f186	NC-positioning signature	Nucleus & Cell	standard deviation(R') ($\equiv \sigma_{R'}$)
f187	NC-positioning signature	Nucleus & Cell	mean($ R' - \langle R' \rangle $)
f188	NC-positioning signature	Nucleus & Cell	range(R')
f189	NC-positioning signature	Nucleus & Cell	interquartile range (R')
f190	NC-positioning signature	Nucleus & Cell	$\Sigma(R'^2)$
f191	NC-positioning signature	Nucleus & Cell	$\Sigma(R'^3)$
f192	NC-positioning signature	Nucleus & Cell	mean(R'^2)
f193	NC-positioning signature	Nucleus & Cell	mean(R'^3)
f194	NC-positioning signature	Nucleus & Cell	mean(R'^4)
f195	NC-positioning signature	Nucleus & Cell	mean(R'^5)
f196	NC-positioning signature	Nucleus & Cell	$\Sigma(R' - \langle R' \rangle ^2)$
f197	NC-positioning signature	Nucleus & Cell	$\Sigma(R' - \langle R' \rangle ^3)$
f198	NC-positioning signature	Nucleus & Cell	mean($ R' - \langle R' \rangle ^2$)
f199	NC-positioning signature	Nucleus & Cell	mean($ R' - \langle R' \rangle ^3$)
f200	NC-positioning signature	Nucleus & Cell	mean($ R' - \langle R' \rangle ^4$)
f201	NC-positioning signature	Nucleus & Cell	mean($ R' - \langle R' \rangle ^5$)
f202	NC-positioning signature	Nucleus & Cell	number of peaks
f203	NC-positioning signature	Nucleus & Cell	correlation coefficient between nucleus boundary signature and cell boundary signature

f204	Protrusion	Cell	mean protrusion length
f205	Protrusion	Cell	max protrusion length
f206	Protrusion	Cell	number of protrusion
f207	Protrusion	Cell	number of primary protrusion
f208	Protrusion	Cell	number of secondary protrusion
f209	Protrusion	Cell	Branching index (=f208/f207)
f210	Protrusion	Cell	mean primary protrusion length
f211	Protrusion	Cell	max primary protrusion length
f212	Protrusion	Cell	standard deviation of primary protrusion length
f213	Protrusion	Cell	mean secondary protrusion length
f214	Protrusion	Cell	max secondary protrusion length
f215	Protrusion	Cell	standard deviation of secondary protrusion length