

## **SUPPLEMENTARY MATERIALS**

**Genomic characterization of small cell carcinomas of the uterine cervix**

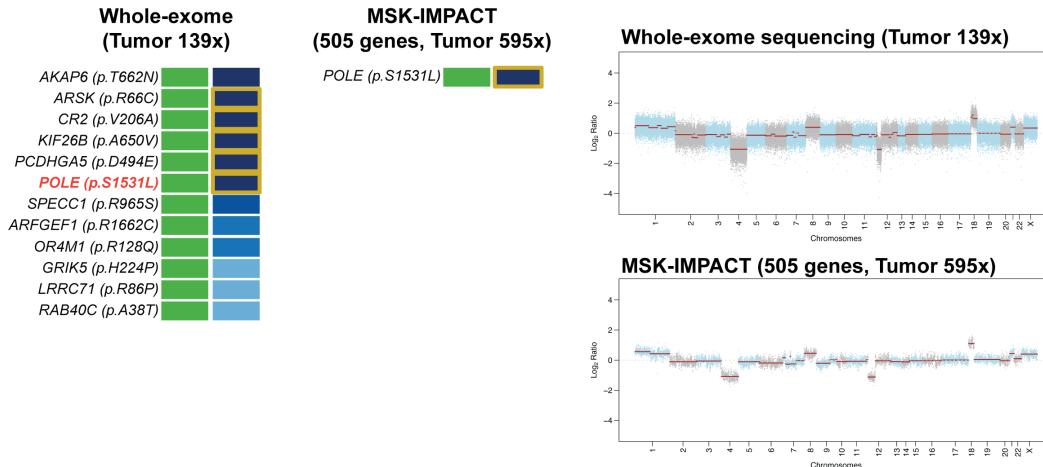
**Schultheis et al.**

**Supplementary Figures S1 and S2**

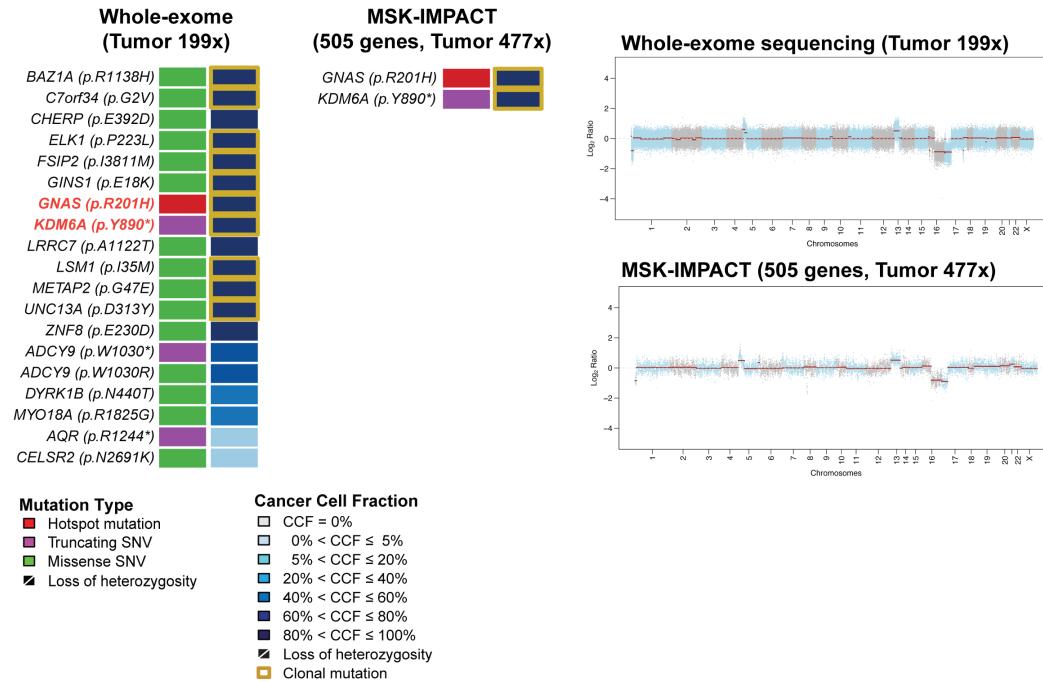
**Supplementary Tables S1 - S6**

## Supplementary Figure S1

### SCC1

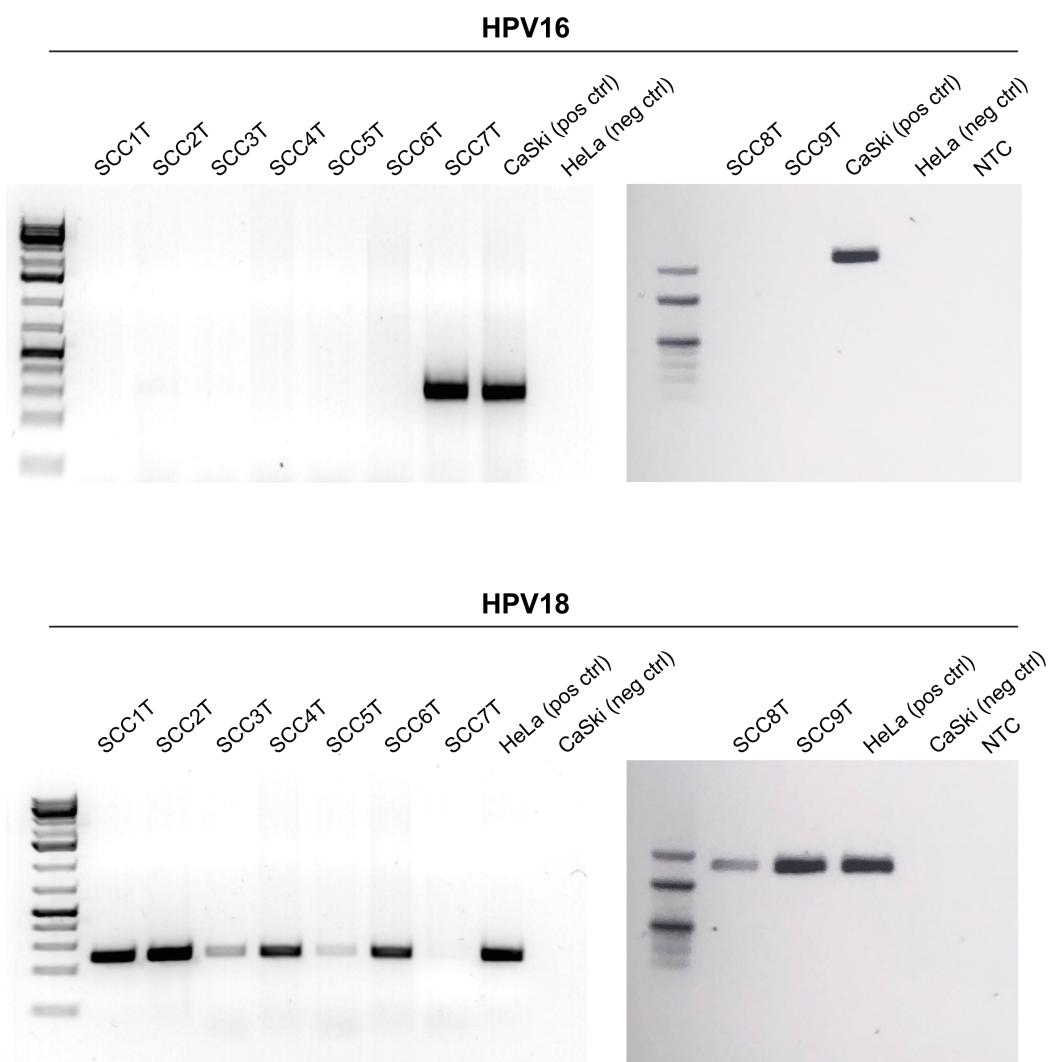


### SCC4



**Supplementary Figure S1: Comparison of whole-exome and high-depth targeted sequencing of uterine cervix small cell carcinomas SCC1 and SCC4.** The same tumor and normal derived DNA samples of SCC1 (top) and SCC4 (bottom) were subjected to whole-exome sequencing and MSK-IMPACT sequencing targeting 505 cancer-related genes. The mutations identified by whole-exome sequencing are shown on the left, and genes being part of the 505 MSK-IMPACT panel are labeled in an orange font. All mutations identified by whole-exome sequencing affecting genes that were part of the 505 targeted panel were identified by MSK-IMPACT sequencing (middle), and no additional mutations were found. The copy number profiles of SCC1 and SCC4 based on whole-exome sequencing and MSK-IMPACT sequencing are also comparable.

## Supplementary Figure S2



**Supplementary Figure S2. Detection of HPV16 and HPV18 DNA in uterine cervix small cell carcinomas.** DNA from uterine cervix small cell carcinomas was amplified with HPV16-specific primers (top) or with HPV18-specific primers (bottom). Agarose gel electrophoresis is shown. CaSki- and HeLa-derived cell line DNA, which are known to harbor HPV16 and HPV18, respectively, were included as positive controls. Ctrl, control. NTC, no template control.

**Supplementary Table S1.** Antibody clones, dilutions, antigen retrievals and scoring used for the immunohistochemical analyses performed.

Antibody	Clone	Dilution	Antigen retrieval	Detection	Company	Scoring
Ki67	MIB1	1:400	ER2, 30'	Leica Bond-RX-BOND Polymer Refine Detection	DAKO	Nuclear positivity in tumor cells; percentage counting 500 tumor cell nuclei.
p63	4A4	RTU	CC1, 24'	Ventana - OptiView	Ventana	No nuclear positivity allowed in the tumor cells.
Chromogranin A	LK2H10	RTU	CC1, 8'	Ventana - Optiview	Ventana	Positive cytoplasmic staining in tumor cells at any percentage.
Synaptophysin	Snp88	1:2000	CC1, 32'	Ventana - Optiview	Biogenex	Positive cytoplasmic staining in tumor cells at any percentage.
p16	E6H4	RTU	CC1, 32'	Ventana - Optiview	Ventana	Nuclear positivity and cytoplasmic positivity in tumor cells.
Pan-cytokeratin	Clone AE1/AE3	1:600	ER2, 30'	Ventana - Optiview	DAKO	Cytoplasmic positivity in tumor cells at any percentage considered as positive

RTU, ready to use.

**Supplementary Table S2.** Primers for Sanger sequencing validation of mutations identified by whole-exome sequencing.

Sample	Gene	AA change	Chromosome	Position	Forward	Reverse
SCC1T	AKAP6	T662N	14	3208695	GAATACCTAGCACTGCCCTCTC	CCTGCCTAGCCTGTATGCT
SCC1T	ARSK	R66C	5	94901771	ATCAATGCTAATTAAATTCTTATTTC	TGTAGCCACTTCTCCCCATT
SCC1T	KIF26B	A650V	1	245775129	CCCAGGCGATGAGCAG	GAAGAACACGCTGTGAGTTGCG
SCC1T	POLE	S1531L	12	133219542	AATCGCTGGATGGCTCG	CAGTGGTGGAGTCTGCC
SCC1T	SPECC1	R965S	17	20163560	AGTATTAGCATCTCTCCTTACCATGT	CAGGACTACTACTGCCAGGG
SCC2T	AGA	E130Q	4	178360736	CTTAGATCGGTTTATCCATGATTT	TATGGATGTAGGAGCAGTAGGAGA
SCC2T	ARSF	C151*	X	3002330	TCACATGTTTTGCAATGTG	GAGGAGAGCTGACTCTCAA
SCC2T	BDP1	Q1353*	5	70806976	GGAAAACAGAGCTAGAGGAGACC	TAAGTGTGTTGGACTTAAGGACAAA
SCC2T	COL5A1	R498S	9	137630654	TTGGACCTTGGCCCTGC	TGAAGACACAAGCTCTGGG
SCC2T	GLI3	R905H	7	42005957	GTCTTCAGGCTCATCCTCTCC	GCTGGCCTACCTGAGCA
SCC2T	OR2D3	R138W	11	6942644	GAGGAAAACCATTCTTTATGG	ACTAGTGCCTTACTGGCC
SCC2T	PIK3CA	E545K	3	178936091	AGGGAAAAATATGACAAGAACAG	CTGCTTATTATCCATAAGGTATGG
SCC2T	SNRNP200	R1090W	2	96953698	TACATGCTGGTTCGATCATC	GAGCTGGAGGTAATGAAACCG
SCC2T	TP53	R175H	17	7578406	AGACCTAAAGAGCAATCAGTGAGGA	GCCAACCTGGCCAAGACCT
SCC2T	UTP20	L583S	12	101696298	GTGGTTCTCTGTGTTCTGC	GATTTACAACAAATAGAGGCTTT
SCC2T	ZNF584	E378K	19	58929017	CTGAAAGTCCACACTGGG	TCCTGACGCCCTCTGGA
SCC4T	CHERP	E392D	19	16639020	GTGGGATCTGGCTGTG	TAACTGCCTGCACTGGATTAG
SCC4T	MYO18A	R1825G	15	72208836	GACTGGTAATGCGATCTGTAGT	GAAACTGGCTGAGAATTGG
SCC4T	LRRC7	A1122T	1	70504985	TTCTCAGGACATCTGTGAATGAG	CGATCCAGCCCTCCCTATA
SCC4T	GINS1	E18K	20	25388508	CGCCATGTTCTGCGAAA	TGCCCCGTCCAAGTCTAC
SCC4T	BAZ1A	R1138H	14	35234363	GACATATCTGCAACGCC	TTCTCTGCTAGTCTATCCAAGTT
SCC4T	METAP2	G47E	12	95868095	GAAGAAGGAGCTGCCCTACG	TGAGCCAACAGTCAGCC
SCC4T	KDM6A	Y890*	X	44929570	AAACAGATCTGCTCTGGTAACC	TCAAACTTTCACTTGATGAACTT
SCC4T	ELK1	P223L	X	47497568	CTCTCCTGGGCCCTCTA	CCCGGGGAAATCACTG
SCC4T	ADCY9	W1030*	16	4016748	GGTCTGGAGTAGGTCTGG	ACTACCACGGAGACGTGAA
SCC4T	ADCY9	W1030R	16	4016748	GGTCTGGAGTAGGTCTGG	ACTACCACGGAGACGTGAA
SCC4T	GNAS	R201H	20	57484421	TATTACTGTTGGTGGCTTG	ACAGTTGGCTACTGAAAGTTGA
SCC4T	UNC13A	D313Y	19	17767038	TCCTCCCTCCAGGAAGTC	CTCCGACATGGAGGATGAG
SCC5T	ANKRD44	P945S	2	197858397	CTGAGCTGGTACTGGTACTTT	ATCCATGTAATTTGAGGCTAAAGTAC
SCC5T	ROBO2	W451C	3	77607216	TTTGACAGATAGACCTCACCTATAA	AGATGATAATTATGTACAAGTCAA
SCC5T	RQCD1	R259H	2	219457379	AGAATTGTTGTGTTGTGTT	GTTCAGCAGAGTATTCCCTAAAGTC
SCC5T	FAM120A	T996S	9	96324527	TTCATTCATTTAGCGCTCAA	TGCTGGTTTGTAAACCTCT
SCC5T	FLNC	R1008H	7	128484151	GTGAACACAGGAGGGCT	TAGCGCACAGCCTGGG
SCC5T	PCDH17	L48F	13	58206822	TGCCCCTACTCTCAAGAAC	CTGCTCGCTCCACGTCC
SCC5T	ONECUT1	P249L	15	53081336	ATTGCTGACCTGCGCG	AACGGCTTCGAAGGCC
SCC5T	CHAT	Q311P	10	50833698	GACAGCTGGTGGCTCAG	GTAATGGAGCTGAAACGCATG
SCC5T	FZR1	M27V	19	3525875	CTGGGGCACTCTCGG	GGGATGAAAGGGTCTCC
SCC5T	SCIN	V122M	7	12620694	GGCAGACACCTCATCAGTTT	CGACCCCTCACATGTAGGAG
SCC5T	ITSN1	P113S	21	35107500	CATAGCTATGAAACTTACAACTGAA	ATAATGTTAAACCGAGAGGGAAA
SCC5T	PLA2G4F	F318I	15	42442018	TGTCGAGGGCCTTG	CACCCAGAACCCCCA
SCC5T	CACNA1F	G985E	X	49074921	TGATCTGGCCCGG	ATCTCGTGGTGAAGATTCTG
SCC6T	IDH1	H309Y	2	209104653	GGGCTGTTGGAGAGCA	CTCTTCATGCAGTTGGACCC
SCC6T	LAMB4	R1252T	7	107696077	ATCAGTCCCTAAAGTCACATAGG	TCTGTGAGGAGACTTCAAAGA
SCC6T	METTL18	M11	1	169762834	AGAACATTCTGCCCCCTCTTCT	CCATGTATAAGCTGCTAAAGAGAAAATT
SCC6T	NIPBL	S2013I	5	37038770	GTTTCACTACCTTGTATTTAGTG	CTAAATTAAAGCACTTAAATGTTTATGT
SCC6T	NOTCH2	E1025K	1	120483288	AGTCATTAACCTGACAGTTTCTCC	TAGCTGATGTTGCTGTGTTGC
SCC6T	NPY5R	F78L	4	164271657	GATGACTTACAGTTTTCTGATTGGG	CATAATATGGCACATGACTTGC
SCC6T	ZCCHC11	D237N	1	52991244	GTAATTTAAAGGGTAAACGGGGG	TGAAAACACTCACCACGATCT
SCC6T	ZCCHC8	Q205*	12	122967884	AAGCCAAAAGTTATGATTTGGT	CTTCCGAAGGATGGAAA
SCC6T	RP11-1220K.2	R356C	7	141838447	TCTTGGGTTCCAGCTAGTCG	AAAAATGGGAATTGGTGTCAAC
SCC7T	C19orf57	E245*	19	14000936	CAGGGGACACCGCTCTCC	CCTCTGAGAACAGGGG
SCC7T	CEP97	Q566E	3	101477146	GAAGAACACATCTCAAGCAACTTC	TCCTTATTTCATCAGTTAACGACAA
SCC7T	CEP97	S330*	3	101476002	TAGACAGGTTATTGATTTAGGT	ATATGCACTTTCTGCTTATGA
SCC7T	CEP97	Q580*	3	101477188	ATTTAACCCAGAGATCTGTTGCT	CAACCAACAAAACAGTCATC
SCC7T	FGFR3	E140K	4	1801512	GGACCTGCCCATCT	CCTTAGCCCTCAGCTGCC
SCC7T	GBE1	H291Q	3	81692051	TGCTATCCAAAGATCATGAGTC	TCATTGTTAGCCGTTATGGAAC
SCC7T	IL24	E105K	1	207074845	GAAGATCCCTATCTGCTGTG	CCAGAGTAGAGAATGACTTCAGAGTC
SCC7T	KANK4	P795T	1	62728920	AGTGGGACACGCTGTAGTGA	GGAGGAACCTTGTGCTGCC
SCC7T	NLRP14	C505*	11	7064772	ATGAAAACGCTATGTTGACCC	CTTCATTCAAAGGGAAACAA
SCC7T	P2RX7	E175Q	12	121600313	TTTGACCCCTATAGGAATTAGAC	CTGAGAGGGGAAGGCC
SCC7T	RNF168	Q105*	3	196215543	ACTAAATTAAAATGAAACTAAAACACA	CAAAGAGGATATTGCTGCAGTT
SCC7T	SLC12A7	E989Q	5	1057647	AGGATCTGTTCCCCCAG	CAGGAACACCGCGCTCC
SCC7T	TMEM51	L213F	1	15546114	TCACTGACGGGGCTCG	CGTGGCTCAAGTGGGG
SCC7T	INPP5D	L46Q	2	234104059	CTGCAAATGAAACCCCTT	ACCCACGCCACTTAC
SCC7T	CACNA1A	R1977H	19	13320507	GTGGTACATGCAAGCCA	AAGATCTACGCGACCATGATG

**Supplementary Table S3.** Sequencing statistics, number of somatic mutations and validation rates.

Case	Tissue Type	Sequencing Type	Mean Target Coverage (X)	Target Bases 2X	Target Bases 50X	Target Bases 100X	Total # Somatic Mutations	# Somatic Non-synonymous Mutations	Validation Rate Sanger Sequencing	Validation Rate MSK-IMPACT
SCC1	Normal	Whole-exome	88	100%	77%	34%	21	12	100% (5/5)	100% (1/1)
	Tumor		139	100%	92%	68%				
SCC2	Normal	Whole-exome	121	100%	98%	93%	34	24	100% (11/11)	Not performed
	Tumor		336	100%	90%	59%				
SCC3	Normal	MSK-IMPACT 505	122	95%	78%	56%	10	6	Not performed	Not performed
	Tumor		408	100%	99%	99%				
SCC4	Normal	Whole-exome	152	100%	96%	84%	32	19 (18 validated)	92% (11/12)	100% (2/2)
	Tumor		199	100%	95%	75%				
SCC5	Normal	Whole-exome	140	100%	97%	88%	41	26	100% (13/13)	Not performed
	Tumor		221	100%	93%	69%				
SCC6	Normal	Whole-exome	181	100%	93%	73%	42	26	100% (9/9)	Not performed
	Tumor		182	100%	96%	80%				
SCC7	Normal	Whole-exome	175	100%	91%	70%	84	59 (57 validated)	87% (13/15)	Not performed
	Tumor		183	100%	94%	75%				
SCC8	Normal Pool	MSK-IMPACT 505	115	100%	79%	55%	0	0	Not applicable	Not performed
	Tumor		278	99%	99%	97%				
SCC9	Normal	MSK-IMPACT 505	202	100%	99%	93%	10	8	Not performed	Not performed
	Tumor		405	100%	99%	99%				

**Supplementary Table S4.** Non-synonymous somatic mutations identified in small cell carcinomas of the cervix using whole-exome or MSK-IMPACT targeted sequencing.

Sample ID	Sequencing platform	Chromosome	Position	Reference	Alternate	Gene symbol	A/A change	Mutation type	Tumor	Normal	LOH	Pathogenicity	Haplotype	Cancer cell	Clinical	RNASeq Validation	Sanger validation
SCC01	WES	14	33015044	C	A	AKAP6	p.T82R	Missense_Mutation	0.0473031	0.004	No	passenger	No	0.43	Biallelic	Mutation Expressed	Validation
SCC01	WES	9	88115642	G	A	ANPBP1	p.R196G	Missense_Mutation	0.0145452	0	No	passenger	No	0.44	Sicilian	Not Expressed	Not Tested
SCC01	WES	1	205940243	T	C	CSPG	p.V2058A	Missense_Mutation	0.288	0.0004830	No	passenger	No	0.05	Clinical	Not Expressed	Not Tested
SCC01	WES	18	42088173	T	G	GRIN2B	p.K224P	Missense_Mutation	0.1175965	0	No	heterozygote	No	0.25	Sicilian	NA	Not Tested
SCC01	WES	1	148685079	G	C	LRRK2	p.R898P	Missense_Mutation	0.125	0	No	passenger	No	0.36	Biallelic	NA	Not Tested
SCC01	WES	14	20470000	T	C	PTEN	p.L164F	Missense_Mutation	0.0004849	0	No	passenger	No	0.05	Sicilian	Not Expressed	Not Tested
SCC01	WES	14	140145129	G	A	PCDHGB8	p.D120Q	Missense_Mutation	0.544675	0	No	passenger	No	0.45	Sicilian	Not Expressed	Not Tested
SCC01	WES	13	130165943	G	A	POLR2E	p.S531L	Missense_Mutation	0.146874	0	No	passenger	No	0.1	Clinical	NA	Not Tested
SCC01	WES	17	20183668	G	A	SPES21	p.K68S	Missense_Mutation	0.0461505	0	No	passenger	No	0.69	Biallelic	Mutation Expressed	Validation
SCC01	WES	11	78872046	G	A	ACVR2A	p.R7Q	Missense_Mutation	0.0172438	0	No	passenger	No	0.1	Clinical	Not Expressed	Not Tested
SCC01	WES	25	30093130	G	A	ARID1A	p.C51Y	Missense_Mutation	0.4004948	0	No	passenger	No	0.1	Clinical	Not Expressed	Validation
SCC01	WES	16	73095349	A	G	ATXNL1	p.C55Y	Missense_Mutation	0.5460777	0	No	passenger	No	0.59	Clinical	NA	Not Tested
SCC01	WES	1	10000000	G	A	BRCA1	p.R100Q	Missense_Mutation	0.0004833	0	No	passenger	No	0.05	Sicilian	Mutation Expressed	Validation
SCC01	WES	6	3619441	A	C	BRPF2	p.K50R	Missense_Mutation	0.0073984	0	No	passenger	No	0.3	Biallelic	Mutation Expressed	Not Tested
SCC01	WES	12	41970000	T	TG	C10orf108	p.T145D	Missense_Mutation	0.1176983	0	No	passenger	No	0.47	Sicilian	NA	Not Tested
SCC01	WES	9	135036504	G	T	COLSAT1	p.Q88R	Missense_Mutation	0.4349508	0	No	heterozygote	No	0.01	Clinical	Not Expressed	Validation
SCC01	WES	10	8801253	G	A	CYP2C19	p.R40H	Missense_Mutation	0.00547423	0	No	passenger	No	0.18	Biallelic	NA	Not Tested
SCC01	WES	7	4090597	T	C	GLTMR	p.R99H	Missense_Mutation	0.4	0	No	heterozygote	No	0.88	Clinical	Not Expressed	Validation
SCC01	WES	11	880424	T	C	OSBPL2	p.L309P	Missense_Mutation	0.0004845	0	No	passenger	No	0.2	Clinical	Not Expressed	Validation
SCC01	WES	3	11093654	G	A	PRKCA	p.E54K	Missense_Mutation	0.4363108	0.0073264	No	heterozygote	No	0.1	Clinical	NA	Not Tested
SCC01	WES	6	30365468	T	C	PTN	p.E53Q	Missense_Mutation	0.1865	0	No	passenger	No	0.5	Biallelic	NA	Not Tested
SCC01	WES	2	9000000	G	A	RBBP4	p.R100Q	Missense_Mutation	0.4017674	0.0004833	No	passenger	No	0.05	Sicilian	NA	Not Tested
SCC01	WES	16	30748493	AGGG	A	SRCAF	p.E25Q	In_Frame_Del	0.0004856	0	No	passenger	No	0.05	Sicilian	NA	Not Tested
SCC01	WES	2	129364000	G	A	UGOT11	p.D150Q	Missense_Mutation	0.1173037	0.0120857	No	heterozygote	No	0.38	Biallelic	NA	Not Tested
SCC01	WES	12	101696298	T	C	UTP9X	p.L385K	Missense_Mutation	0.0327103	0	No	passenger	No	1	Clinical	NA	Not Tested
SCC01	WES	16	4017675	A	T	ADCP	p.W73R	Missense_Mutation	0.38205048	0	No	heterozygote	No	0.77	Biallelic	RNASeq not available	Validation
SCC01	WES	18	4016144	G	A	ADCP	p.W73R	Missense_Mutation	0.3820495	0	No	passenger	No	0.72	Biallelic	RNASeq not available	Validation
SCC01	WES	14	35243493	T	C	ASXL1	p.L104I	Missense_Mutation	0.0102043	0	No	heterozygote	No	0.1	Clinical	NA	Not Tested
SCC01	WES	1	10000000	G	A	CELP2	p.N207K	Missense_Mutation	0.000482138	0	No	passenger	No	0.14	Sicilian	NA	Not Tested
SCC01	WES	16	1983603	T	C	CHERP	p.E100Q	Missense_Mutation	0.00048005	0	No	passenger	No	0.02	Sicilian	NA	Not Tested
SCC01	WES	25	4197058	G	A	ELAVL1	p.P225L	Missense_Mutation	0.00740553	0	No	passenger	No	0.91	Clinical	RNASeq not available	Validation
SCC01	WES	2	1988170	A	G	F3MP1	p.L191M	Missense_Mutation	0.5108952	0	No	passenger	No	0.1	Clinical	RNASeq not available	Not Tested
SCC01	WES	20	57044401	G	A	GNAS	p.R61H	Missense_Mutation	0.111043	0	No	heterozygote	Yes	0.93	Clinical	RNASeq not available	Validation
SCC01	WES	23	44620530	G	A	KOMMT1	p.L122L	Missense_Mutation	0.486773	0.000482402	No	heterozygote	No	0.48	Clinical	NA	Not Tested
SCC01	WES	3	44620530	G	A	KOMMT1	p.L122L	Missense_Mutation	0.486773	0.000482402	No	heterozygote	No	0.48	Clinical	NA	Not Tested
SCC01	WES	6	38054043	A	C	LRRK2	p.S284W	Missense_Mutation	0.00740553	0	No	passenger	No	0.96	Clinical	RNASeq not available	Validation
SCC01	WES	17	2517602	G	C	MYO18A	p.R192Q	Missense_Mutation	0.1017648	0	No	heterozygote	No	0.41	Biallelic	RNASeq not available	Validation
SCC01	WES	19	17767038	G	A	UNC13A	p.D101Y	Missense_Mutation	0.0207304	0	No	heterozygote	No	0.92	Clinical	RNASeq not available	Validation
SCC01	WES	2	1602875	G	A	PTEN	p.R207Q	Missense_Mutation	0.00740553	0	No	heterozygote	No	0.97	Clinical	RNASeq not available	Validation
SCC01	WES	19	2000000	G	A	NPAS6	p.E240K	Missense_Mutation	0.0004804	0	No	passenger	No	0.86	Clinical	RNASeq not available	Validation
SCC01	WES	15	43081326	A	G	NEIL2	p.R45Q	Missense_Mutation	0.00740551	0	No	passenger	No	0.94	Clinical	RNASeq not available	Validation
SCC01	WES	18	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.95	Clinical	NA	Not Tested
SCC01	WES	18	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.95	Clinical	NA	Not Tested
SCC01	WES	2	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.95	Clinical	NA	Not Tested
SCC01	WES	23	40974021	G	T	CACHA1	p.D98R	Missense_Mutation	0.02016903	0	No	heterozygote	No	0.97	Clinical	NA	Not Tested
SCC01	WES	12	10000000	G	A	CHD1	p.D211P	Missense_Mutation	0.0141463	0	No	heterozygote	No	0.1	Clinical	NA	Not Tested
SCC01	WES	9	19951447	G	C	FAM129A	p.D102V	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.9	Clinical	NA	Not Tested
SCC01	WES	2	21045739	G	A	ROBO2	p.R202Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	2	21045739	G	A	ROBO2	p.R202Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	16	10000000	G	A	TCF4	p.E237K	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.97	Clinical	NA	Not Tested
SCC01	WES	19	202875	A	G	PTEN	p.M27Y	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.91	Clinical	NA	Not Tested
SCC01	WES	1	2000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.91	Clinical	NA	Not Tested
SCC01	WES	16	8805000	T	C	ZFYVE2	p.T254Y	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.95	Clinical	NA	Not Tested
SCC01	WES	12	10000000	G	A	ZFYVE2	p.T254Y	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.95	Clinical	NA	Not Tested
SCC01	WES	16	8805000	T	C	ZFYVE2	p.T254Y	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.95	Clinical	NA	Not Tested
SCC01	WES	19	39195703	G	C	ACTN4	p.K103N	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.82	Clinical	RNASeq not available	Validation
SCC01	WES	20	39195703	G	T	APBL1	p.A73T	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.92	Clinical	RNASeq not available	Validation
SCC01	WES	3	40951571	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.97	Clinical	NA	Not Tested
SCC01	WES	15	40974021	G	A	PLA2G4F	p.T219M	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.85	Clinical	NA	Not Tested
SCC01	WES	8	27312109	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.91	Clinical	NA	Not Tested
SCC01	WES	23	40974021	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.97	Clinical	NA	Not Tested
SCC01	WES	7	2000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	16	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	1	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	16	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	12	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	2	10000000	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.99	Clinical	NA	Not Tested
SCC01	WES	19	39195703	G	C	ACTN4	p.K103N	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.82	Clinical	RNASeq not available	Validation
SCC01	WES	20	39195703	G	T	APBL1	p.A73T	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.92	Clinical	RNASeq not available	Validation
SCC01	WES	3	40951571	G	A	PTEN	p.R207Q	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.95	Clinical	NA	Not Tested
SCC01	WES	15	40974021	G	A	PLA2G4F	p.T219M	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.85	Clinical	NA	Not Tested
SCC01	WES	8	24840504	G	T	DHODH	p.L251H	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.75	Clinical	NA	Not Tested
SCC01	WES	4	10000000	G	A	FGFR2	p.E149K	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.89	Clinical	NA	Not Tested
SCC01	WES	23	40974021	G	A	PLA2G4F	p.T219M	Missense_Mutation	0.0004804	0	No	heterozygote	No	0.84	Clinical	NA	Not Tested
SCC01	WES	3	40974021	G	A	PLA2G4F	p.T219M	Missense_Mutation									

**Supplementary Table S5.** Putative in-frame fusion transcripts with driver probability >0.1 identified in SCC1 and SCC2 using RNA-sequencing.

Sample ID	Genes	Genomic coordinates	Spanning and split reads (n)	Split reads (n)	Driver probability (Oncofuse)	Algorithm	5' location	3' location	P-value
SCC2T	VMP1-RPS6KB1	chr17:57842500>chr17:57987922	3	3	0.997438	chimerascan	Intron	Intron	0.04913752
SCC2T	SLC35G2-NCK1	chr3:136538057>chr3:136646825	2	1	0.9961532	chimerascan	Exon	Intron	0.05754801
SCC1T	DMC1-DDX17	chr22:38917612>chr22:38897286	3	2	0.9858879	chimerascan	Intron	Intron	0.14232601
SCC1T	ABCA10-ABC46	chr17:67145173>chr17:67136890	2	2	0.9736921	chimerascan	Intron	Intron	0.19960184
SCC1T	ARFIP2-TRIM3	chr11:6498947>chr11:6486963	2	2	0.9446882	chimerascan	Intron	Intron	0.34136487
SCC2T	PDLIM2-C8orf58	chr8:22447255>chr8:22458394	2	2	0.8289519	chimerascan	Intron	Intron	0.63653723
SCC2T	GKAP1-KIF27	chr9:86431910>chr9:86530594	7	2	0.5981402	chimerascan	Intron	Intron	1
SCC2T	GKAP1-KIF27	chr9:86431910>chr9:86530594	28	17	0.5981402	defuse	Intron	Intron	1
SCC1T	POLA2-CDC42EP2	chr11:65063462>chr11:65088014	4	3	0.5912114	chimerascan	Intron	Intron	1
SCC2T	POLA2-CDC42EP2	chr11:65063462>chr11:65088014	23	10	0.5912114	chimerascan	Intron	Intron	1
SCC2T	POLA2-CDC42EP2	chr11:65063462>chr11:65088014	41	21	0.5912114	defuse	Intron	Intron	0.97330609
SCC2T	PIBF1-KLF5	chr13:73573134>chr13:73635998	2	2	0.5481406	chimerascan	Intron	Intron	1
SCC1T	TLK2-FAM157A	chr17:60637488>chr3:197907625	33	33	0.2722771	chimerascan	Intron	Intron	1
SCC2T	TLK2-FAM157A	chr17:60637488>chr3:197907625	44	29	0.2722771	chimerascan	Intron	Intron	1
SCC1T	TLK2-FAM157B	chr17:60637488>chr9:141134069	33	33	0.2722771	chimerascan	Intron	Intron	1
SCC2T	HOXB6-HOXB3	chr17:46681841>chr17:46651377	3	2	0.1747878	chimerascan	Intron	Exon	1
SCC1T	ZNF716-ZNF678	chr7:57522293>chr1:227842036	4	4	0.1540229	chimerascan	Intron	Intron	1
SCC1T	F8-TMLHE	chrX:154114408>chrX:154718952	2	1	0.1096824	chimerascan	Intron	Exon	1
SCC1T	MBD1-CCDC11	chr18:47799193>chr18:47788590	3	2	0.1056514	chimerascan	Intron	Intron	1

**Supplementary Table S6.** Putative HPV integration sites based on RNA-sequencing.

Sample ID	hg19 locus	hg19 annotation	Mate 1 HPV18 Position	Mate 2 hg19 Chromosome 18 Position
SCC1T	18q12.3	-	882	39411292
SCC1T	18q12.3	-	2280	39497187
SCC1T	18q12.3	-	3669	39401088
SCC2T	8p22	RP11-89M16.1-002 (lncRNA)	2603	129517175
SCC2T	8p22	RP11-89M16.1-002 (lncRNA)	4908	129518183
SCC2T	8p22	RP11-89M16.1-002 (lncRNA)	5996	129517009
SCC2T	8p22	RP11-89M16.1-002 (lncRNA)	5998	129509523