SOX2 activation predicts prognosis in patients with head and neck squamous cell carcinoma

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## Supplementary Figure Legends

Supplementary Fig. 1. SOX2 activation in HNSCC. (A) Cross-cancer summary of SOX2 alteration in 18 cancers using genome copy number data obtained from the TCGA cohort. HNSCC is the fourth cancer to have SOX2 amplification (21%). (B) Scatter plots of SOX2 mRNA expression and copy number alteration in patients in the TCGA cohort. SOX2 expression was substantially higher in samples lacking SOX2 amplification.

Supplementary Fig. 2. Association between SOX2 signature and other subtypes of HNSCC. Patients are shown in columns and are grouped according to SOX2 signature, tumour site, human papillomavirus (HPV) status, regional lymph node (LN) metastasis, T stage, and smoking status.

Supplementary Fig. 3. Frequency of molecular subtype between SOX2-high and SOX2-low subgroups.

Supplementary Fig. 4. Kaplan-Meier plots of overall survival (OS) of patients with HNSCC according to SOX2 mRNA expression in the TCGA cohort.



## Suppl. Figure 1



Suppl. Figure 2



SOX2-low

SOX2-high



Suppl. Figure 4

	correlation	p value of	correlation	p value of
TCGA RNA	coefficient with	correlation with	coefficient with	correlation with
sequencing gene	copy number of	copy number of	gene expression	gene expression
symbol	SOX2	SOX2	of SOX2	of SOX2
ABCC5	0.678	1.92523E-70	0.651	4.16604E-63
ACAP2	0.506	9.78107E-35	0.603	4.68376E-52
ACPL2	0.421	1.98927E-23	0.490	2.42246E-32
ACTL6A	0.738	1.74965E-89	0.471	1.00055E-29
ADRA2B	0.437	2.24913E-25	0.425	6.86321E-24
B4GALT4	0.437	2.39056E-25	0.529	2.56486E-38
BEX2	0.447	1.43944E-26	0.428	2.70066E-24
C3orf21	0.624	7.53622E-57	0.449	9.04895E-27
C3orf58	0.515	4.20488E-36	0.684	6.02487E-72
C6orf168	0.427	3.45506E-24	0.543	1.10478E-40
CACNA1B	0.446	2.13629E-26	0.506	8.68839E-35
CASP4	-0.413	1.63262E-22	-0.460	2.90495E-28
CEP70	0.449	8.2918E-27	0.521	5.32347E-37
CHST7	0.481	4.92114E-31	0.541	2.06789E-40
CLCN2	0.592	6.61921E-50	0.446	1.79782E-26
COCH	0.448	1.15867E-26	0.536	1.80041E-39
CYP26A1	0.404	1.28819E-21	0.601	1.07556E-51
DCUN1D1	0.635	2.30113E-59	0.406	8.46315E-22
DLG1	0.444	3.33888E-26	0.467	4.22562E-29
DLX6AS	0.405	1.0798E-21	0.511	1.50998E-35
DMRT3	0.405	1.02043E-21	0.505	1.23783E-34
DVL3	0.725	9.18162E-85	0.471	1.05205E-29

Table S1. Genes associated with SOX2

EIF4A2	0.499	1.0015E-33	0.423	1.02942E-23
EPCAM	0.415	8.05554E-23	0.537	1.12848E-39
FAM20B	0.434	5.36703E-25	0.427	4.14603E-24
FEZF1	0.426	5.3949E-24	0.492	1.39564E-32
GK5	0.412	2.02108E-22	0.446	1.81963E-26
GLT25D2	0.405	1.00949E-21	0.473	6.44051E-30
GPC3	0.404	1.41277E-21	0.472	6.70273E-30
GPR160	0.410	2.97186E-22	0.534	3.26647E-39
IFT80	0.503	3.13136E-34	0.422	1.32643E-23
IQCB1	0.418	4.21866E-23	0.506	1.11136E-34
KCNG3	0.428	2.91277E-24	0.514	6.0285E-36
KCNMB2	0.519	8.42383E-37	0.492	1.29582E-32
KCNMB3	0.631	2.39568E-58	0.441	7.58611E-26
KLHL24	0.524	1.39499E-37	0.455	1.54491E-27
LOC100130691	0.403	1.84918E-21	0.584	3.03542E-48
LSG1	0.676	6.02408E-70	0.465	5.82863E-29
MCCC1	0.686	1.03027E-72	0.607	5.70455E-53
MED12L	0.473	4.90775E-30	0.472	6.96423E-30
MRAP2	0.417	5.78204E-23	0.634	4.06282E-59
MSL2	0.485	1.12818E-31	0.503	2.92887E-34
MYNN	0.649	8.34691E-63	0.453	2.58657E-27
NCRNA00086	0.433	7.74741E-25	0.551	4.92812E-42
NMNAT3	0.410	2.95747E-22	0.520	5.88133E-37
NTRK2	0.415	8.83634E-23	0.764	1.2542E-99
NTS	0.424	7.56766E-24	0.619	1.0548E-55
OPA1	0.575	1.37104E-46	0.401	2.92893E-21
OR7A5	0.413	1.66114E-22	0.417	4.654E-23

РССВ	0.460	2.8295E-28	0.446	2.0147E-26
РНС3	0.552	2.56483E-42	0.442	5.69698E-26
PIR	0.405	1.20088E-21	0.593	5.22962E-50
PNCK	0.437	2.44554E-25	0.439	1.30532E-25
PRKCI	0.614	1.3106E-54	0.454	1.83606E-27
RAB6B	0.488	5.16015E-32	0.431	1.17188E-24
RASL11A	0.425	5.8441E-24	0.509	3.16502E-35
RIMKLA	0.405	1.22257E-21	0.557	3.96621E-43
RSRC1	0.501	5.75908E-34	0.467	4.26433E-29
SENP2	0.708	3.30702E-79	0.406	7.83144E-22
SGEF	0.439	1.42469E-25	0.712	1.4375E-80
SLC25A36	0.538	7.76435E-40	0.420	2.71419E-23
SOX2OT	0.607	4.3003E-53	0.797	6.2569E-114
STRBP	0.427	3.25372E-24	0.644	1.6076E-61
TBCCD1	0.644	1.99628E-61	0.458	5.55391E-28
TBL1XR1	0.726	3.81792E-85	0.467	3.42099E-29
TMEM20	0.415	8.64226E-23	0.493	9.52065E-33
TRA2B	0.502	3.50901E-34	0.443	4.48054E-26
TRIM59	0.480	6.78894E-31	0.423	9.85771E-24
TSEN15	0.443	4.26773E-26	0.449	8.30335E-27
ULBP1	0.440	9.56135E-26	0.401	2.7856E-21
VWA5B2	0.551	4.86855E-42	0.525	1.23116E-37
WNK2	0.414	1.03158E-22	0.603	3.61689E-52
ZNF639	0.787	1.4867E-109	0.419	3.40964E-23
ZXDC	0.420	2.18478E-23	0.419	3.09197E-23

 Table S2. Comparison of somatic mutation frequencies between SOX2-high and SOX2-low

 subtypes.

		SOX-high	SOX-low	Fisher's p value
AKT1	MUT	1	1	0.054
	WT	215	276	0.054
APC	MUT	13	8	0 115
	WT	203	269	0.115
ATM	MUT	5	10	0.442
	WT	211	267	0.445
CDH1	MUT	4	1	0 174
	WT	212	276	0.174
CDKN2A	MUT	34	77	1 505 07
	MUT	182	200	1.39E-07
CSF1R	WT	1	0	0.429
	MUT	215	277	0.438
EGFR	WT	6	13	0.249
	MUT	210	264	0.348
ERBB2	WT	6	2	0 146
	MUT	210	275	0.140
ERBB4	MUT	10	11	0.922
	WT	206	266	0.825
FBXW7	MUT	14	18	1
	WT	202	259	1
FGFR2	MUT	0	3	0.26
	WT	216	274	0.20
FGFR3	MUT	7	5	2 11E 06
	MUT	209	272	5.11E-00
FLT3	WT	3	3	0 115
	MUT	212	274	0.115
GNA11	WT	0	3	0.26
	MUT	216	274	0.20
HRAS	WT	1	26	3.11E-06
	MUT	215	251	
IDH1	MUT	1	3	0.635
	WT	215	274	
JAK3	MUT	2	3	1
	WT	214	274	

KIT	MUT	2	3	1
	WT	214	274	1
NOTCH1	MUT	16	69	1 505 07
	MUT	200	208	1.39E-07
NRAS	WT	1	0	0 429
	MUT	215	277	0.438
PIK3CA	WT	47	43	0.070
	MUT	169	234	0.079
PTEN	WT	10	4	0.052
	MUT	206	273	0.055
RB1	MUT	12	6	0.054
	WT	204	271	0.034
RET	MUT	8	4	0.142
	WT	208	273	0.142
SMAD4	MUT	5	7	1
	WT	211	270	1
SMO	MUT	1	0	0 429
	MUT	215	277	0.438
SRC	WT	0	0	1
	MUT	216	277	1
STK11	WT	2	3	1
	MUT	214	274	1
TP53	WT	141	210	0.012
	MUT	75	67	0.012
VHL	MUT	0	0	0 429
	WT	216	277	0.438