

Table S1. Characteristics of the cohort. data not available (d.n.a.), Primary Tumor (PT), Lymph node metastasis (LN), Distant metastasis (DM), Local recurrence (LR), Cancer of unknown primary (CUP), positive (pos.), negative (neg.), without (w/o), progression (prog), overall survival (OS), Progrssion-free survival (PFS)

			LSD1 staining n=	SNAIL staining n=
Patients analyzed			339	339
	d.n.a.		5	7
Tissue Type	PT		334	332
	LN		163	165
	DM		22	21
	LR		59	57
Sub-Site	Hypopharynx		46	46
	Larynx		98	95
	Oral Cavity		76	76
	Oropharynx		103	104
	CUP / d.n.a.		11	11
Sex	male		259	258
	female		74	73
	d.n.a.		1	1
Age	≤ 61		54	52
	> 61		178	178
	d.n.a.		2	2
T-Stage	T1		68	69
	T2		98	98
	T3		96	96
	T4		70	67
	d.n.a.		2	2
N-Stage	N0		147	146
	N1		51	51
	N2		83	83
	N3		50	50
	d.n.a.		3	3
M-Stage	M0		317	315
	M1		14	14
	d.n.a.		3	3
UICC	I		66	69
	II		59	57
	III		59	59
	IV		149	146
	d.n.a.		1	1
p16	pos.		85	88
	neg.		249	244
	d.n.a.		0	0
Alcohol cons.	Yes		139	139
	No		183	180
	d.n.a.		2	3
Nicotine cons.	Yes		281	279
	No		36	35
	d.n.a.		17	18
5 y. PFS	dead or prog.		148	147
	alive w/o prog		178	177
	d.n.a.		8	8
5 y. OS	dead		119	117
	alive		207	107
	d.n.a.		8	8

Table S2. Univariate and multivariate Cox regression for PFS in primary hypopharyngeal HNSCC. Survival analysis of 5-year PFS in primary hypopharyngeal head and neck squamous cell carcinoma (HNSCC) using the Cox proportional hazards regression model. Only tumor and patient characteristics showing a significant correlation with 5-year PFS in the univariate analysis were included in the multivariate analysis. Characteristics and corresponding p-values showing a significant correlation in the multivariate analysis are bold.

Variable	Category	Univariate HR (95% -Konf. Int., p=)	Multivariate HR (95%- conf. int., p=)
LSD1-Expression	LSD1 low	2.53	3.72
	LSD high	(1.05-6.09, p=0.038)	(1.24-10.94, p=0.028)
T- Stage	T1,2,3	3.25	1.03
	T4	(1.29-8.21, p=0.013)	(0.35-3.01, p=0.924)
N-Stage	N0,1	4.18	0.15
	N2,3	(1.53-11.43, p=0.005)	(0.02-1.02, p=0.052)
M-Stage	M0	4.21	
	M1	(0.53-33.28, p=0.173)	
UICC-Stage	UICC I,II,III	7.37	44.56
	UICC IV	(2.16-25.15, p=0.001)	(4.18-475.45, p=0.002)
Sex	Male	0.39	
	Female	(0.09-1.66, p=0.202)	
Age	≤ 61 y.	0.97	
	> 61 y.	(0.42-2.22, p=0.943)	
p16	Positive	2.76	
	Negative	(0.37-20.51, p=0.321)	
Alcohol	Yes	0.71	
	No	(0.28-1.75, p=0.453)	
Nicotine	Yes	2.63	
	No	(0.61-11.41, p=0.197)	

Table S3. Univariate and multivariate Cox regression for in primary HNSCC. 5-year OS analysis using the Cox proportional hazards regression model. Multivariate analysis was performed with all tumor and patient characteristics showing a significant correlation with patient survival in the univariate analysis. Characteristics and corresponding p-values showing a significant correlation in the multivariate analysis are bold.

Variable	Category	Univariate HR (95% -Konf. Int., p=)	Multivariate HR (95%- conf. int., p=)
SNAIL-Expression	SNAIL low	1.45	1.57
	SNAIL high	(1.01-2.10, p=0.046)	(1.07-2.31, p=0.021)
T- Stage	T1,2	2.14	1.79
	T3,4	(1.47-3.11, p<0,001)	(1.08-2.99, p=0.025)
N-Stage	N0	1.67	1.49
	N1,2,3	(1.14-2.44, p=0.009)	(0.98-2.36, p=0.088)
M-Stage	M0	5.15	5.15
	M1	(2.74-9.67, p<0.001)	(2.64-10.03, p<0.001)
UICC-Stage	UICC I,II	2.32	1.19
	UICC III,IV	(1.52-3.53, p<0,001)	(0.61-2.34, p=0.606)
Sex	Male	0.80	
	Female	(0.51-1.27, p=0.345)	
Age	≤ 61 y.	1.25	
	> 61 y.	(0.86-1.81, p=0.236)	
p16	Positive	1.93	1.70
	Negative	(1.20-3.13, p=0.007)	(0.96-3.01, p=0.071)
Alcohol	Yes	0.57	0.74
	No	(0.40-0.84, p=0.004)	(0.50-1.09, p=0.125)
Nicotine	Yes	0.60	
	No	(0.29-1.22, p=0.158)	

Table S4. Univariate and multivariate cox regression of PFS in primary HNSCC. Survival analysis of 5-year PFS in primary HNSCC using the Cox proportional hazards regression model. All tumor and patient characteristics showing a significant correlation with 5-year survival in the univariate analysis were included in the multivariate analysis. Characteristics and corresponding p-values showing a significant correlation in the multivariate analysis are bold.

Variable	Category	Univariate HR (95% -Konf. Int., p=)	Multivariate HR (95%- conf. int., p=)
SNAIL-Expression	SNAIL low	1.63	1.82
	SNAIL high	(1.17-2.26, p=0.004)	(1.29-2.57, p=0.001)
T- Stage	T1,2	2.10	1.79
	T3,4	(1.51-2.94, p<0.001)	(1.12-2.85, p=0.014)
N-Stage	N0	1.40	
	N1,2,3	(1.00-1.95, p=0.051)	
M-Stage	M0	3.41	3.45
	M1	(1.83-6.32, p<0.001)	(1.80-6.63, p<0.001)
UICC-Stage	UICC I,II	2.19	1.38
	UICC III,IV	(1.52-3.16, p<0.001)	(0.81-2.34, p=0.238)
Sex	Male	0.84	
	Female	(0.56-1.26, p=0.398)	
Age	≤ 61 y.	1.22	
	> 61 y.	(0.88-1.69, p=0.241)	
p16	Positive	2.20	1.79
	Negative	(1.42-3.41, p<0.001)	(1.10-2.90, p=0.018)
Alcohol	Yes	0.63	0.81
	No	(0.45-0.87, p=0.006)	(0.57-1.14, p=0.224)
Nicotine	Yes	0.76	
	No	(0.43-1.35, p=0.354)	

Figure S1. Exemplary HE stained TMA Cores. Exemplary Tissue Micro Array (TMA) cores of the cohort stained with standard hematoxylin and eosin (HE) staining.

