

Supplemental Data section:

The following tables provide a tissue-site breakdown of oral preneoplastic (OPLs) site at baseline and of the oral cancer site in patients who developed oral cancer during the follow-up.

Table 1: Side, anatomical site, EGFR expression by immunohistochemistry (IHC, semi-quantitative evaluation) and EGFR gene copy number (FISH) of oral preneoplastic lesions (OPLs) at baseline in 17 patients who developed same site oral cancer during follow-up.

Pts	OPL histology	OPL and oral cancer site	EGFR IHC	EGFR FISH
1	moderate dysplasia	ventral tongue	high	+
2	moderate dysplasia	left lateral tongue	high	+
3	hyperplasia	left lateral tongue	high	+
4	mild dysplasia	base of tongue	low	NA
5	hyperplasia	left lateral tongue	low	NA
6	mild dysplasia	left lateral tongue	high	+
7	moderate dysplasia	right ventral tongue	high	+
8	hyperplasia	left lateral tongue	high	+
9	hyperplasia	left lateral tongue	low	NA
10	hyperplasia	left lateral tongue	high	+
11	hyperplasia	left retromolar area	high	+
12	mild dysplasia	right buccal mucosa	low	NA
13	hyperplasia	right lateral tongue	high	NA
14	mild hyperplasia	left floor of mouth	high	+
15	mild dysplasia	right lateral tongue	high	+
16	hyperplasia	right lateral tongue	high	NA
17	hyperplasia	right mandibular gingiva	high	-

Table 2: Side, anatomical site, EGFR expression by immunohistochemistry (IHC, semi-quantitative evaluation) and EGFR gene copy number (FISH) of oral preneoplastic lesions (OPLs) at baseline and oral cancer site in 18 patients who developed oral cancer during follow-up at a site different from OPL site at inclusion.

Pts	OPL histology	OPL site	Oral cancer site	EGFR IHC	EGFR FISH
1	hyperplasia	right tonsillar pillar	right cheek	high	-
2	hyperplasia	left anterior floor	right lateral tongue	high	-
3	hyperplasia	left tongue	right mandibular alveolar ridge	high	-
4	mild dysplasia	right lateral tongue	left posterior tongue	low	NA
5	hyperplasia	left floor of mouth	right alveolar ridge	high	+
6	hyperplasia	right mandibula	mid lower lip	high	-
7	hyperplasia	right buccal mucosa	posterior tongue	high	-
8	hyperplasia	anterior mandibula	left lateral tongue	high	+
9	mild dysplasia	left cheek	left posterior maxilla	high	-
10	mild dysplasia	right lateral tongue	left ventral tongue	high	+
11	hyperplasia	left cheek	right ventral tongue	high	+
12	hyperplasia	anterior mandibula	left maxilla	high	-
13	severe dysplasia	right maxilla	anterior hard palate	high	NA
14	hyperplasia	right cheek	posterior tongue	high	NA
15	hyperplasia	left tongue	left floor of mouth	low	NA
16	mild dysplasia	anterior floor of mouth	left lateral tongue	high	NA
17	mild dysplasia	left lateral tongue	left retromolar trigone	high	+
18	moderate dysplasia	right floor of mouth	anterior floor of mouth	high	+

Table 3: Distribution of EGFR expression by immunohistochemistry(semi-quantitative evaluation) in different subgroups of OPLs, based on histology, in all the samples 145 samples analyzed in the study

Histology	EGFR expression		Total
	Low	High	
Hyperplasia	29	68	97
	20.00	46.90	66.90
	29.90	70.10	
	69.05	66.02	
Mild dysplasia	12	25	37
	8.28	17.24	25.52
	32.43	67.57	
	28.57	24.27	
Moderate dysplasia	1	9	10
	0.69	6.21	6.90
	10.00	90.00	
	2.38	8.74	
Severe dysplasia	0	1	1
	0.00	0.69	0.69
	0.00	100.00	
	0.00	0.97	
Total	42	103	145
	28.97	71.03	100.00

Table 4: Distribution of EGFR gene copy number by FISH (defined as FISH positive or negative) in different subgroups of OPLs, based on histology, in the 49 samples with high EGFR expression (IHC)

Histology	FISH status		Total
	Negative	Positive	
Hyperplasia	24	9	33
	48.98	18.37	67.35
	72.73	27.27	
	82.76	45.00	
Mild dysplasia	4	6	10
	8.16	12.24	20.41
	40.00	60.00	
	13.79	30.00	
Moderate dysplasia	1	4	5
	2.04	8.16	10.20
	20.00	80.00	
	3.45	20.00	
Severe dysplasia	0	1	1
	0.00	2.04	2.04
	0.00	100.00	
	0.00	5.00	
Total	29	20	49
	59.18	40.82	100.00

No association was found between FISH positivity and the degree of dysplasia (Fisher's Exact Test, P=0.72).