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| **Supplementary file Table 1a: Comparison of HIF-1α with demographic and clinicopathologic characteristics of OSCC** |
|  | **Category** | **Mean IRS HIF-1α**  | **P Value** |
|  | Age  |  ≥ 60 yrs(N=20) | 4.50+1.57 | 0.788 |
| <60 yrs (N=30) | 4.63+ 1.88 |
|  | Gender  | Male (N=33) | 4.27+1.85 | 0.059 |
| Female (N=17) | 5.18+1.38 |
|  | Positive family history of cancer | No(N=47) | 4.62+1.75 | 0.650 |
| Yes(N=3) | 4+2 |
|  | Positive family history of OSCC  | No(N=49) | 4.59+1.76 | 0.742 |
| Yes(N=1) | 4+ 0 |
|  | Tobacco habit  | No(N=7) | 5.57+1.61 | 0.12 |
| Yes(N=43) | 4.42+1.73 |
|  | Drinking habit | No(N=26) | 4.81+1.67 | 0.34 |
| Yes(N=24) | 4.33+1.83 |
|  | Systemic conditions  | No(N=37) | 4.62+1.86 | 0.750 |
| Yes(N=13) | 4.46+1.45 |
|  | Dietary habits  | Veg (N=1) | 4+0 | 0.742 |
| Non veg(N=49)  | 4.59+1.76 |
|  | Site of lesion  | Buccal mucosa (N=17) | 4.35+1.65 | 0.548 |
| Tongue (N=17) | 4.53+1.73 |
| Floor of mouth (N=2) | 5+4.24 |
| Palate(N=3) | 5.33+3.05 |
| Alveolar mucosa/gingiva(N=7) | 5.43+0.976 |
| Retromolar region (N=4) | 3.50+1.0 |
|  | Size of lesion | Less than or equal to 2cm (N=18) | 4.44+1.46 | 0.114 |
| > 2 to < 4/ equal (N=24) | 4.29+1.78 |
| > 4 cm (N=8) | 5.75+1.98 |
|  | lymph node involvement  | No(N=25) | 4.64+1.63 | 0.811 |
| Yes(N=25) | 4.52+1.89 |
|  | Type of lesion | Ulcer (N=11) | 4+1.78 | 0.315 |
| Ulceroproliferative (N=35) | 4.83+1.74 |
| White/red patch (N=4) | 4+1.63 |
| Swelling(N=0) | 0 |
|  | Histologic grading | WDSCC (N=9) | 4.56+1.13 | 0.143 |
| MDSCC (N=38) | 4.74+1.84 |
| PDSCC (N=3) | 2.67+1.15 |
|  | Presence of Necrosis | No(N=37) | 4.51+1.83 | 0.628 |
| Yes(N=13) | 4.77+1.53 |
|  | Inflammation | Mild (N=1) | 2+0 | 0.259 |
| Moderate(N=20)  | 4.85+1.56 |
| Severe (N=29) | 4.48+1.84 |
|  | Invasion of adjacent structures  | No(N=30) | 4.67+1.70 | 0.678 |
| Yes(N=20) | 4.45+1.84 |
|  | Tumor budding | Low(N=18)  | 4.89+1.41 | 0.314 |
| High (N=32) | 4.41+1.91 |
| **Supplementary file Table 1b: Comparison of HIF-2α with demographic and clinicopathologic characteristics of OSCC** |
|  | Category | Mean IRS HF2 | P Value |
|  | Age  |  ≥ 60 yrs(N=20) | 3.35+1.18 | 0.501 |
| <60 yrs (N=30) | 3.36+1.40 |
|  | Gender  | Male (N=33) | 3.67+1.53 | 0.119 |
| Female (N=17) | 3.18+0.63 |
|  | Positive family history of cancer | No(N=47) | 3.51+1.26 | 0.907 |
| Yes(N=3) | 3.33+2.30 |
|  | Positive family history of OSCC  | No(N=49) | 3.49+1.32 | 0.705 |
| Yes(N=1) | 4+0 |
|  | Tobacco habit  | No(N=7) | 3.57+1.39 | 0.887 |
| Yes(N=43) | 3.49+1.31 |
|  | Drinking habit | No(N=26) | 3.35+1.09 | 0.401 |
| Yes(N=24) | 3.67+1.52 |
|  | Systemic conditions  | No(N=37) | 3.35+1.18 | 0.256 |
|  | Yes(N=13) | 3.92+1.60 |
|  | Dietary habits  | Veg (N=1) | 4+0 | 0.705 |
| Non veg(N=49)  | 3.49+1.325 |
|  | Site of lesion  | Buccal mucosa (N=17) | 3.47+1.37 | 0.849 |
| Tongue (N=17) | 3.65+1.32 |
| Floor of mouth (N=2) | 3+1.41 |
| Palate(N=3) | 4+2 |
| Alveolar mucosa/gingiva(N=7) | 3+0.81 |
| Retromolar region (N=4) | 3.75+1.70 |
|  | Size of lesion | Less than or equal to 2cm (N=18) | 3.78+1.39 | 0.504 |
| > 2 to < 4/ equal (N=24) | 3.29+1.26 |
| > 4 cm (N=8) | 3.50+1.31 |
|  | lymph node involvement  | No(N=25) | 3.44+1.35 | 0.750 |
| Yes(N=25) | 3.56+1.29 |
|  | Type of lesion | Ulcer (N=11) | 3.18+1.25 | 0.230 |
| Ulceroproliferative (N=35) | 3.49+1.26 |
| White/red patch (N=4) | 4.50+1.73 |
| Swelling(N=0) | 0 |
|  | Histologic grading | WDSCC (N=9) | 3.56+1.42 | 0.289 |
| MDSCC (N=38) | 3.58+1.30 |
| PDSCC (N=3) | 2.23+0.577 |
|  | Presence of Necrosis | No(N=37) | 3.54+1.32 | 0.719 |
| Yes(N=13) | 3.38+1.32 |
|  | Inflammation | Mild (N=1) | 4+0 | 0.538 |
| Moderate(N=20)  | 3.25+1.20 |
| Severe (N=29) | 3.66+1.396 |
|  | Invasion of adjacent structures  | No(N=30) | 3.57+1.45 | 0.647 |
| Yes(N=20) | 3.40+1.09 |
|  | Tumor budding | Low(N=18)  | 3.83+1.38 | 0.196 |
| High (N=32) | 3.31+1.25 |
|  | (\* Chi-square test, two-sided, P.value<0.05)  |