

Table S1. Distribution of SNP genotypes in cases and controls, by race

| Gene | SNP | Major/minor allele | Genotype | European-Americans | | | | African-Americans | | | |
|---------------------------------|------------|--------------------|----------|--------------------|---------|-------------------|---------|-------------------|---------|------------------|---------|
| | | | | Cases (n=922) | | Controls (n=1074) | | Cases (n=305) | | Controls (n=251) | |
| | | | | n | (col %) | n | (col %) | n | (col %) | n | (col %) |
| ALCOHOL METABOLISM GENES | | | | | | | | | | | |
| <i>ADH1B</i> | rs12507573 | C/A | AA | 188 | 20.4% | 234 | 21.8% | 56 | 18.4% | 40 | 15.9% |
| | | | AC | 462 | 50.1% | 536 | 49.9% | 149 | 48.9% | 127 | 50.6% |
| | | | CC | 272 | 29.5% | 304 | 28.3% | 100 | 32.8% | 83 | 33.1% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% |
| | rs1042026 | A/G | GG | 73 | 7.9% | 81 | 7.5% | 3 | 1.0% | 5 | 2.0% |
| | | | GA | 382 | 41.4% | 457 | 42.6% | 54 | 17.7% | 36 | 14.3% |
| | | | AA | 467 | 50.7% | 536 | 49.9% | 248 | 81.3% | 210 | 83.7% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs7673353 | C/T | TT | 0 | 0.0% | 0 | 0.0% | 7 | 2.3% | 12 | 4.8% |
| | | | TC | 3 | 0.3% | 6 | 0.6% | 89 | 29.2% | 65 | 25.9% |
| | | | CC | 919 | 99.7% | 1068 | 99.4% | 209 | 68.5% | 174 | 69.3% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs17028834 | T/C | CC | 1 | 0.1% | 0 | 0.0% | 2 | 0.7% | 5 | 2.0% |
| | | | CT | 3 | 0.3% | 2 | 0.2% | 80 | 26.2% | 50 | 19.9% |
| | | | TT | 918 | 99.6% | 1072 | 99.8% | 223 | 73.1% | 196 | 78.1% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| rs1693457 | T/C | CC | 27 | 2.9% | 27 | 2.5% | 9 | 3.0% | 16 | 6.4% | |
| | | CT | 297 | 32.2% | 333 | 31.0% | 88 | 28.9% | 86 | 34.3% | |
| | | TT | 597 | 64.8% | 714 | 66.5% | 208 | 68.2% | 148 | 59.0% | |
| | | missing | 1 | 0.1% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% | |
| rs1229984 | G/A | AA | 0 | 0.0% | 3 | 0.3% | 0 | 0.0% | 0 | 0.0% | |
| | | AG | 30 | 3.3% | 71 | 6.6% | 1 | 0.3% | 5 | 2.0% | |
| | | GG | 892 | 96.7% | 1000 | 93.1% | 304 | 99.7% | 246 | 98.0% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs1159918 | G/T | TT | 129 | 14.0% | 138 | 12.8% | 161 | 52.8% | 140 | 55.8% | |
| | | TG | 419 | 45.4% | 485 | 45.2% | 116 | 38.0% | 93 | 37.1% | |
| | | GG | 374 | 40.6% | 451 | 42.0% | 28 | 9.2% | 18 | 7.2% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs1229982 | G/T | TT | 30 | 3.3% | 45 | 4.2% | 69 | 22.6% | 51 | 20.3% | |
| | | TG | 282 | 30.6% | 327 | 30.4% | 146 | 47.9% | 125 | 49.8% | |
| | | GG | 610 | 66.2% | 702 | 65.4% | 90 | 29.5% | 75 | 29.9% | |
| | | missing | 0.0% | 0 | 0.0% | 0 | 0.0% | 0.0% | 0 | 0.0% | |
| <i>ADH1C</i> | rs2298753 | T/C | CC | 15 | 1.6% | 15 | 1.4% | 0 | 0.0% | 0 | 0.0% |
| | | | CT | 186 | 20.2% | 177 | 16.5% | 20 | 6.6% | 14 | 5.6% |
| | | | TT | 721 | 78.2% | 882 | 82.1% | 285 | 93.4% | 237 | 94.4% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs1614972 | C/T | TT | 73 | 7.9% | 96 | 8.9% | 67 | 22.0% | 67 | 26.7% |
| | | | TC | 403 | 43.7% | 446 | 41.5% | 144 | 47.2% | 129 | 51.4% |
| | | | CC | 446 | 48.4% | 532 | 49.5% | 94 | 30.8% | 55 | 21.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs1391088 | C/A | AA | 3 | 0.3% | 4 | 0.4% | 2 | 0.7% | 2 | 0.8% |
| | | | AC | 118 | 12.8% | 161 | 15.0% | 55 | 18.0% | 36 | 14.3% |
| | | | CC | 801 | 86.9% | 909 | 84.6% | 248 | 81.3% | 213 | 84.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs1693482 | C/T | TT | 158 | 17.1% | 172 | 16.0% | 6 | 2.0% | 4 | 1.6% |
| | | | TC | 437 | 47.4% | 497 | 46.3% | 94 | 30.8% | 64 | 25.5% |
| | | | CC | 325 | 35.2% | 403 | 37.5% | 205 | 67.2% | 183 | 72.9% |
| | | | missing | 2 | 0.2% | 2 | 0.2% | 0 | 0.0% | 0 | 0.0% |
| rs1631460 | C/G | GG | 160 | 17.4% | 173 | 16.1% | 6 | 2.0% | 4 | 1.6% | |
| | | GC | 443 | 48.0% | 508 | 47.3% | 95 | 31.1% | 65 | 25.9% | |
| | | CC | 318 | 34.5% | 393 | 36.6% | 204 | 66.9% | 182 | 72.5% | |
| | | missing | 1 | 0.1% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs11936869 | C/G | GG | 64 | 6.9% | 69 | 6.4% | 53 | 17.4% | 56 | 22.3% | |
| | | GC | 354 | 38.4% | 430 | 40.0% | 143 | 46.9% | 122 | 48.6% | |
| | | CC | 504 | 54.7% | 575 | 53.5% | 109 | 35.7% | 72 | 28.7% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% | |

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| Gene | SNP | Major/minor allele | Genotype | European-Americans | | | | African-Americans | | | |
|-----------|------------|--------------------|----------|--------------------|---------|-------------------|---------|-------------------|---------|------------------|---------|
| | | | | Cases (n=922) | | Controls (n=1074) | | Cases (n=305) | | Controls (n=251) | |
| | | | | n | (col %) | n | (col %) | n | (col %) | n | (col %) |
| ADH4 | rs29001227 | A/T | TT | 1 | 0.1% | 1 | 0.1% | 8 | 2.6% | 8 | 3.2% |
| | | | TA | 4 | 0.4% | 1 | 0.1% | 92 | 30.2% | 59 | 23.5% |
| | | | AA | 917 | 99.5% | 1072 | 99.8% | 205 | 67.2% | 184 | 73.3% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs1126672 | C/T | TT | 53 | 5.7% | 82 | 7.6% | 6 | 2.0% | 2 | 0.8% |
| | | | TC | 386 | 41.9% | 434 | 40.4% | 60 | 19.7% | 46 | 18.3% |
| | | | CC | 483 | 52.4% | 558 | 52.0% | 239 | 78.4% | 203 | 80.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs4699710 | T/C | CC | 70 | 7.6% | 98 | 9.1% | 13 | 4.3% | 5 | 2.0% |
| | | | CT | 412 | 44.7% | 458 | 42.6% | 90 | 29.5% | 78 | 31.1% |
| | | | TT | 440 | 47.7% | 518 | 48.2% | 202 | 66.2% | 168 | 66.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs10017466 | T/C | CC | 72 | 7.8% | 99 | 9.2% | 40 | 13.1% | 32 | 12.7% |
| | | | CT | 414 | 44.9% | 459 | 42.7% | 143 | 46.9% | 101 | 40.2% |
| | | | TT | 436 | 47.3% | 516 | 48.0% | 121 | 39.7% | 118 | 47.0% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 1 | 0.3% | 0 | 0.0% |
| rs1800759 | C/A | AA | 114 | 12.4% | 150 | 14.0% | 171 | 56.1% | 152 | 60.6% | |
| | | AC | 472 | 51.2% | 527 | 49.1% | 113 | 37.0% | 83 | 33.1% | |
| | | CC | 336 | 36.4% | 397 | 37.0% | 21 | 6.9% | 16 | 6.4% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs1800761 | G/A | AA | 31 | 3.4% | 28 | 2.6% | 26 | 8.5% | 22 | 8.8% | |
| | | AG | 293 | 31.8% | 343 | 31.9% | 124 | 40.7% | 100 | 39.8% | |
| | | GG | 598 | 64.9% | 703 | 65.5% | 155 | 50.8% | 129 | 51.4% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs3762894 | T/C | CC | 25 | 2.7% | 20 | 1.9% | 14 | 4.6% | 17 | 6.8% | |
| | | CT | 247 | 26.8% | 306 | 28.5% | 99 | 32.5% | 73 | 29.1% | |
| | | TT | 650 | 70.5% | 748 | 69.6% | 192 | 63.0% | 161 | 64.1% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| ADH7 | rs284787 | C/T | TT | 61 | 6.6% | 58 | 5.4% | 6 | 2.0% | 6 | 2.4% |
| | | | TC | 339 | 36.8% | 384 | 35.8% | 78 | 25.6% | 57 | 22.7% |
| | | | CC | 522 | 56.6% | 632 | 58.8% | 221 | 72.5% | 188 | 74.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs894369 | C/G | GG | 32 | 3.5% | 43 | 4.0% | 7 | 2.3% | 6 | 2.4% |
| | | | GC | 275 | 29.8% | 356 | 33.1% | 77 | 25.2% | 68 | 27.1% |
| | | | CC | 615 | 66.7% | 675 | 62.8% | 221 | 72.5% | 176 | 70.1% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% |
| | rs17588403 | T/A | AA | 44 | 4.8% | 31 | 2.9% | 4 | 1.3% | 3 | 1.2% |
| | | | AT | 274 | 29.7% | 344 | 32.0% | 66 | 21.6% | 58 | 23.1% |
| | | | TT | 604 | 65.5% | 699 | 65.1% | 235 | 77.0% | 190 | 75.7% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs1154454 | T/C | CC | 24 | 2.6% | 31 | 2.9% | 49 | 16.1% | 48 | 19.1% |
| | | | CT | 250 | 27.1% | 309 | 28.8% | 146 | 47.9% | 128 | 51.0% |
| | | | TT | 648 | 70.3% | 734 | 68.3% | 110 | 36.1% | 75 | 29.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| rs1154456 | T/C | CC | 116 | 12.6% | 127 | 11.8% | 7 | 2.3% | 7 | 2.8% | |
| | | CT | 438 | 47.5% | 494 | 46.0% | 87 | 28.5% | 68 | 27.1% | |
| | | TT | 368 | 39.9% | 453 | 42.2% | 211 | 69.2% | 176 | 70.1% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs1154460 | G/A | AA | 194 | 21.0% | 218 | 20.3% | 66 | 21.6% | 58 | 23.1% | |
| | | AG | 466 | 50.5% | 544 | 50.7% | 169 | 55.4% | 127 | 50.6% | |
| | | GG | 259 | 28.1% | 312 | 29.1% | 70 | 23.0% | 65 | 25.9% | |
| | | missing | 3 | 0.3% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% | |
| rs971074 | G/A | AA | 7 | 0.8% | 10 | 0.9% | 8 | 2.6% | 5 | 2.0% | |
| | | AG | 165 | 17.9% | 210 | 19.6% | 101 | 33.1% | 70 | 27.9% | |
| | | GG | 750 | 81.3% | 854 | 79.5% | 196 | 64.3% | 176 | 70.1% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |

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|------------|-----------|--------------------|----------|--------------------|---------|-------------------|---------|-------------------|---------|------------------|---------|
| | | | | Cases (n=922) | | Controls (n=1074) | | Cases (n=305) | | Controls (n=251) | |
| | | | | n | (col %) | n | (col %) | n | (col %) | n | (col %) |
| ALDH2 | rs1573496 | C/G | GG | 5 | 0.5% | 9 | 0.8% | 0 | 0.0% | 0 | 0.0% |
| | | | GC | 154 | 16.7% | 190 | 17.7% | 7 | 2.3% | 6 | 2.4% |
| | | | CC | 762 | 82.6% | 875 | 81.5% | 298 | 97.7% | 245 | 97.6% |
| | | | missing | 1 | 0.1% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs4767939 | A/G | GG | 35 | 3.8% | 39 | 3.6% | 52 | 17.0% | 38 | 15.1% |
| | | | GA | 297 | 32.2% | 315 | 29.3% | 145 | 47.5% | 125 | 49.8% |
| | | | AA | 589 | 63.9% | 720 | 67.0% | 108 | 35.4% | 88 | 35.1% |
| | | | missing | 1 | 0.1% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs2238151 | T/C | CC | 119 | 12.9% | 125 | 11.6% | 257 | 84.3% | 179 | 71.3% |
| | | | CT | 436 | 47.3% | 464 | 43.2% | 46 | 15.1% | 71 | 28.3% |
| | | | TT | 363 | 39.4% | 482 | 44.9% | 2 | 0.7% | 1 | 0.4% |
| | | | missing | 4 | 0.4% | 3 | 0.3% | 0 | 0.0% | 0 | 0.0% |
| | rs7312055 | G/A | AA | 0 | 0.0% | 0 | 0.0% | 28 | 9.2% | 12 | 4.8% |
| | | | AG | 5 | 0.5% | 8 | 0.7% | 111 | 36.4% | 101 | 40.2% |
| | | | GG | 917 | 99.5% | 1066 | 99.3% | 166 | 54.4% | 138 | 55.0% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| rs2158029 | G/A | AA | 0 | 0.0% | 0 | 0.0% | 6 | 2.0% | 1 | 0.4% | |
| | | AG | 6 | 0.7% | 11 | 1.0% | 45 | 14.8% | 41 | 16.3% | |
| | | GG | 916 | 99.3% | 1063 | 99.0% | 254 | 83.3% | 209 | 83.3% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs16941667 | C/T | TT | 6 | 0.7% | 5 | 0.5% | 6 | 2.0% | 4 | 1.6% | |
| | | TC | 153 | 16.6% | 158 | 14.7% | 74 | 24.3% | 58 | 23.1% | |
| | | CC | 763 | 82.8% | 911 | 84.8% | 225 | 73.8% | 188 | 74.9% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% | |
| rs16941669 | T/G | GG | 14 | 1.5% | 11 | 1.0% | 2 | 0.7% | 0 | 0.0% | |
| | | GT | 174 | 18.9% | 190 | 17.7% | 50 | 16.4% | 36 | 14.3% | |
| | | TT | 734 | 79.6% | 872 | 81.2% | 253 | 83.0% | 214 | 85.3% | |
| | | missing | 0 | 0.0% | 1 | 0.1% | 0 | 0.0% | 1 | 0.4% | |
| CYP2E1 | rs3813865 | G/C | CC | 0 | 0.0% | 1 | 0.1% | 4 | 1.3% | 3 | 1.2% |
| | | | CG | 45 | 4.9% | 49 | 4.6% | 76 | 24.9% | 53 | 21.1% |
| | | | GG | 877 | 95.1% | 1023 | 95.3% | 225 | 73.8% | 195 | 77.7% |
| | | | missing | 0 | 0.0% | 1 | 0.1% | 0 | 0.0% | 0 | 0.0% |
| | rs3813867 | G/C | CC | 1 | 0.1% | 0 | 0.0% | 1 | 0.3% | 1 | 0.4% |
| | | | CG | 51 | 5.5% | 58 | 5.4% | 30 | 9.8% | 25 | 10.0% |
| | | | GG | 870 | 94.4% | 1016 | 94.6% | 273 | 89.5% | 224 | 89.2% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 1 | 0.3% | 1 | 0.4% |
| | rs8192772 | T/C | CC | 6 | 0.7% | 7 | 0.7% | 3 | 1.0% | 2 | 0.8% |
| | | | CT | 127 | 13.8% | 146 | 13.6% | 64 | 21.0% | 53 | 21.1% |
| | | | TT | 789 | 85.6% | 921 | 85.8% | 237 | 77.7% | 196 | 78.1% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 1 | 0.3% | 0 | 0.0% |
| | rs915908 | G/A | AA | 21 | 2.3% | 27 | 2.5% | 0 | 0.0% | 1 | 0.4% |
| | | | AG | 211 | 22.9% | 266 | 24.8% | 18 | 5.9% | 22 | 8.8% |
| | | | GG | 690 | 74.8% | 781 | 72.7% | 287 | 94.1% | 228 | 90.8% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| rs915909 | C/T | TT | 0 | 0.0% | 0 | 0.0% | 5 | 1.6% | 2 | 0.8% | |
| | | TC | 1 | 0.1% | 4 | 0.4% | 53 | 17.4% | 40 | 15.9% | |
| | | CC | 921 | 99.9% | 1070 | 99.6% | 246 | 80.7% | 204 | 81.3% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 1 | 0.3% | 5 | 2.0% | |
| rs7092584 | C/T | TT | 11 | 1.2% | 9 | 0.8% | 4 | 1.3% | 5 | 2.0% | |
| | | TC | 168 | 18.2% | 197 | 18.3% | 80 | 26.2% | 58 | 23.1% | |
| | | CC | 743 | 80.6% | 868 | 80.8% | 219 | 71.8% | 187 | 74.5% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 2 | 0.7% | 1 | 0.4% | |
| rs743535 | C/T | TT | 11 | 1.2% | 6 | 0.6% | 7 | 2.3% | 4 | 1.6% | |
| | | CT | 149 | 16.2% | 176 | 16.4% | 82 | 26.9% | 57 | 22.7% | |
| | | CC | 757 | 82.1% | 886 | 82.5% | 212 | 69.5% | 188 | 74.9% | |
| | | missing | 5 | 0.5% | 6 | 0.6% | 4 | 1.3% | 2 | 0.8% | |

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| | | | | Cases (n=922) | | Controls (n=1074) | | Cases (n=305) | | Controls (n=251) | |
| | | | | n | (col %) | n | (col %) | n | (col %) | n | (col %) |
| | rs2249695 "tipsy" SNP | C/T | TT | 34 | 3.7% | 41 | 3.8% | 28 | 9.2% | 23 | 9.2% |
| | | | TC | 292 | 31.7% | 372 | 34.6% | 124 | 40.7% | 103 | 41.0% |
| | | | CC | 595 | 64.5% | 661 | 61.5% | 153 | 50.2% | 124 | 49.4% |
| | | | missing | 1 | 0.1% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% |
| | rs28969387 | A/T | TT | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | | | TA | 2 | 0.2% | 0 | 0.0% | 23 | 7.5% | 20 | 8.0% |
| | | | AA | 920 | 99.8% | 1074 | 100.0% | 282 | 92.5% | 231 | 92.0% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs11101812 | T/C | CC | 0 | 0.0% | 0 | 0.0% | 2 | 0.7% | 1 | 0.4% |
| | | | CT | 2 | 0.2% | 1 | 0.1% | 22 | 7.2% | 18 | 7.2% |
| | | | TT | 919 | 99.7% | 1073 | 99.9% | 279 | 91.5% | 230 | 91.6% |
| | | | missing | 1 | 0.1% | 0 | 0.0% | 2 | 0.7% | 2 | 0.8% |
| OXIDATIVE STRESS METABOLISM GENES | | | | | | | | | | | |
| <i>CAT</i> | rs1049982 | C/T | TT | 95 | 10.3% | 106 | 9.9% | 92 | 30.2% | 76 | 30.3% |
| | | | TC | 396 | 43.0% | 449 | 41.8% | 147 | 48.2% | 123 | 49.0% |
| | | | CC | 429 | 46.5% | 514 | 47.9% | 65 | 21.3% | 50 | 19.9% |
| | | | missing | 2 | 0.2% | 5 | 0.5% | 1 | 0.3% | 2 | 0.8% |
| <i>GPX1</i> | rs8179172 | T/A | AA | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 1 | 0.4% |
| | | | AT | 1 | 0.1% | 2 | 0.2% | 50 | 16.4% | 48 | 19.1% |
| | | | TT | 921 | 99.9% | 1072 | 99.8% | 255 | 83.6% | 202 | 80.5% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs1800668 | C/T | TT | 77 | 8.4% | 101 | 9.4% | 15 | 4.9% | 19 | 7.6% |
| | | | TC | 381 | 41.3% | 437 | 40.7% | 115 | 37.7% | 91 | 36.3% |
| | | | CC | 464 | 50.3% | 533 | 49.6% | 174 | 57.0% | 140 | 55.8% |
| | | | missing | 0 | 0.0% | 3 | 0.3% | 1 | 0.3% | 1 | 0.4% |
| | rs3811699 | A/G | GG | 79 | 8.6% | 100 | 9.3% | 23 | 7.5% | 28 | 11.2% |
| | | | GA | 379 | 41.1% | 442 | 41.2% | 133 | 43.6% | 105 | 41.8% |
| | | | AA | 464 | 50.3% | 532 | 49.5% | 149 | 48.9% | 118 | 47.0% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| rs3448 | C/T | TT | 84 | 9.1% | 83 | 7.7% | 29 | 9.5% | 17 | 6.8% | |
| | | TC | 350 | 38.0% | 397 | 37.0% | 115 | 37.7% | 96 | 38.2% | |
| | | CC | 488 | 52.9% | 594 | 55.3% | 161 | 52.8% | 138 | 55.0% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| <i>GPX2</i> | rs11623705 | G/T | TT | 13 | 1.4% | 15 | 1.4% | 0 | 0.0% | 0 | 0.0% |
| | | | TG | 199 | 21.6% | 220 | 20.5% | 15 | 4.9% | 19 | 7.6% |
| | | | GG | 710 | 77.0% | 839 | 78.1% | 290 | 95.1% | 232 | 92.4% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs2412065 | G/C | CC | 37 | 4.0% | 55 | 5.1% | 48 | 15.7% | 47 | 18.7% |
| | | | CG | 290 | 31.5% | 369 | 34.4% | 154 | 50.5% | 119 | 47.4% |
| | | | GG | 595 | 64.5% | 650 | 60.5% | 103 | 33.8% | 85 | 33.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs2737844 | C/T | TT | 76 | 8.2% | 109 | 10.1% | 209 | 68.5% | 162 | 64.5% |
| | | | TC | 361 | 39.2% | 462 | 43.0% | 82 | 26.9% | 75 | 29.9% |
| | | | CC | 481 | 52.2% | 498 | 46.4% | 13 | 4.3% | 14 | 5.6% |
| | | | missing | 4 | 0.4% | 5 | 0.5% | 1 | 0.3% | 0 | 0.0% |
| <i>GPX4</i> | rs757229 | G/C | CC | 192 | 20.8% | 233 | 21.7% | 104 | 34.1% | 86 | 34.3% |
| | | | CG | 484 | 52.5% | 522 | 48.6% | 145 | 47.5% | 123 | 49.0% |
| | | | GG | 246 | 26.7% | 318 | 29.6% | 55 | 18.0% | 42 | 16.7% |
| | | | missing | 0 | 0.0% | 1 | 0.1% | 1 | 0.3% | 0 | 0.0% |
| <i>SOD1</i> | rs11910115 | A/C | CC | 0 | 0.0% | 0 | 0.0% | 2 | 0.7% | 5 | 2.0% |
| | | | CA | 1 | 0.1% | 1 | 0.1% | 59 | 19.3% | 59 | 23.5% |
| | | | AA | 921 | 99.9% | 1073 | 99.9% | 244 | 80.0% | 187 | 74.5% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs4998557 | G/A | AA | 12 | 1.3% | 14 | 1.3% | 41 | 13.4% | 40 | 15.9% |
| | | | AG | 205 | 22.2% | 195 | 18.2% | 137 | 44.9% | 111 | 44.2% |
| | | | GG | 704 | 76.4% | 862 | 80.3% | 127 | 41.6% | 100 | 39.8% |
| | | | missing | 1 | 0.1% | 3 | 0.3% | 0 | 0.0% | 0 | 0.0% |

Table S1. Distribution of SNP genotypes in cases and controls, by race

| Gene | SNP | Major/minor allele | Genotype | European-Americans | | | | African-Americans | | | |
|-----------|------------|-----------------------|----------|--------------------|---------|-------------------|---------|-------------------|---------|------------------|---------|
| | | | | Cases (n=922) | | Controls (n=1074) | | Cases (n=305) | | Controls (n=251) | |
| | | | | n | (col %) | n | (col %) | n | (col %) | n | (col %) |
| | rs10432782 | T/G | GG | 12 | 1.3% | 14 | 1.3% | 20 | 6.6% | 20 | 8.0% |
| | | | GT | 205 | 22.2% | 194 | 18.1% | 113 | 37.0% | 101 | 40.2% |
| | | | TT | 704 | 76.4% | 865 | 80.5% | 172 | 56.4% | 130 | 51.8% |
| | | | missing | 1 | 0.1% | 1 | 0.1% | 0 | 0.0% | 0 | 0.0% |
| | rs2070424 | A/G | GG | 6 | 0.7% | 5 | 0.5% | 6 | 2.0% | 7 | 2.8% |
| | | | AG | 127 | 13.8% | 113 | 10.5% | 83 | 27.2% | 77 | 30.7% |
| | | | AA | 789 | 85.6% | 956 | 89.0% | 216 | 70.8% | 167 | 66.5% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs1041740 | C/T | TT | 88 | 9.5% | 101 | 9.4% | 6 | 2.0% | 3 | 1.2% |
| | | | TC | 387 | 42.0% | 473 | 44.0% | 62 | 20.3% | 43 | 17.1% |
| | | | CC | 447 | 48.5% | 500 | 46.6% | 237 | 77.7% | 205 | 81.7% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| SOD2 | rs4342445 | G/A | AA | 57 | 6.2% | 56 | 5.2% | 5 | 1.6% | 4 | 1.6% |
| | | | AG | 340 | 36.9% | 355 | 33.1% | 71 | 23.3% | 59 | 23.5% |
| | | | GG | 525 | 56.9% | 663 | 61.7% | 229 | 75.1% | 188 | 74.9% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs2842980 | A/T | TT | 42 | 4.6% | 61 | 5.7% | 30 | 9.8% | 24 | 9.6% |
| | | | TA | 299 | 32.4% | 338 | 31.5% | 139 | 45.6% | 92 | 36.7% |
| | | | AA | 581 | 63.0% | 675 | 62.8% | 136 | 44.6% | 135 | 53.8% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| | rs8031 | T/A | AA | 206 | 22.3% | 267 | 24.9% | 31 | 10.2% | 35 | 13.9% |
| | | | AT | 467 | 50.7% | 534 | 49.7% | 134 | 43.9% | 107 | 42.6% |
| | | | TT | 249 | 27.0% | 273 | 25.4% | 140 | 45.9% | 109 | 43.4% |
| | | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% |
| rs5746134 | C/T | TT | 0 | 0.0% | 0 | 0.0% | 7 | 2.3% | 3 | 1.2% | |
| | | TC | 0 | 0.0% | 4 | 0.4% | 93 | 30.5% | 58 | 23.1% | |
| | | CC | 922 | 100.0% | 1070 | 99.6% | 205 | 67.2% | 190 | 75.7% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |
| rs2758331 | C/A | AA | 204 | 22.1% | 264 | 24.6% | 18 | 5.9% | 18 | 7.2% | |
| | | AC | 468 | 50.8% | 537 | 50.0% | 113 | 37.0% | 87 | 34.7% | |
| | | CC | 250 | 27.1% | 273 | 25.4% | 174 | 57.0% | 146 | 58.2% | |
| | | missing | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | 0 | 0.0% | |

Table S2. SNP effects on odds of developing cancer (dominant genetic model)

| Gene | SNP | major/ minor alleles | Oral cavity, oropharyngeal, hypopharyngeal cancer NOS | | | | Oropharyngeal cancer | | | |
|---------------------------------|------------|----------------------------|--|---|--------------------------------------|----------------------|-----------------------------------|---|--------------------------------------|----------------------|
| | | | n cases/controls | | Adjusted OR ^b (95% CI) | p-value ^c | n cases/controls | | Adjusted OR ^b (95% CI) | p-value ^c |
| | | | homozygous for major allele | one or two copies of minor allele | | | homozygous for major allele | one or two copies of minor allele | | |
| ALCOHOL METABOLISM GENES | | | | | | | | | | |
| <i>ADH1B</i> | rs12507573 | C/A | 71/386 | 152/935 | 0.94 (0.80-1.11) | 1.00 | 91/386 | 241/935 | 1.02 (0.88-1.17) | 1.00 |
| | rs1042026 | A/G | 125/744 | 98/578 | 1.04 (0.89-1.22) | 1.00 | 193/744 | 139/578 | 0.97 (0.84-1.11) | 1.00 |
| | rs7673353 | C/T | 204/1239 | 19/83 | 1.09 (0.79-1.50) | 1.00 | 309/1239 | 23/83 | 0.97 (0.72-1.29) | 1.00 |
| | rs17028834 | T/C | 211/1265 | 12/57 | 1.05 (0.72-1.53) | 1.00 | 312/1265 | 20/57 | 1.29 (0.94-1.77) | 1.00 |
| | rs1693457 | T/C | 136/860 | 86/461 | 1.13 (0.97-1.32) | 1.00 | 223/860 | 109/461 | 1.02 (0.89-1.18) | 1.00 |
| | rs1229984 | G/A | 219/1243 | 4/79 | 0.60 (0.36-1.01) | 1.00 | 322/1243 | 10/79 | 0.74 (0.52-1.06) | 1.00 |
| | rs1159918 | G/T | 65/468 | 158/854 | 1.16 (0.98-1.37) | 1.00 | 109/468 | 223/854 | 1.12 (0.97-1.29) | 1.00 |
| | rs1229982 | G/T | 129/775 | 94/547 | 0.99 (0.84-1.16) | 1.00 | 197/775 | 135/547 | 0.98 (0.85-1.12) | 1.00 |
| <i>ADH1C</i> | rs2298753 | T/C | 180/1117 | 43/205 | 1.18 (0.97-1.44) | 1.00 | 270/1117 | 62/205 | 1.19 (1.00-1.41) | 1.00 |
| | rs1614972 | C/T | 95/585 | 128/737 | 1.02 (0.87-1.19) | 1.00 | 141/585 | 191/737 | 1.02 (0.89-1.16) | 1.00 |
| | rs1391088 | C/A | 192/1119 | 31/203 | 0.98 (0.79-1.22) | 1.00 | 280/1119 | 52/203 | 1.03 (0.86-1.24) | 1.00 |
| | rs1693482 | C/T | 97/585 | 125/735 | 1.03 (0.88-1.20) | 1.00 | 145/585 | 186/735 | 1.03 (0.90-1.18) | 1.00 |
| | rs1631460 | C/G | 93/574 | 130/748 | 1.05 (0.90-1.23) | 1.00 | 144/574 | 188/748 | 1.02 (0.89-1.17) | 1.00 |
| | rs11936869 | C/G | 111/645 | 112/676 | 0.96 (0.83-1.12) | 1.00 | 161/645 | 171/676 | 0.99 (0.87-1.13) | 1.00 |
| <i>ADH4</i> | rs29001227 | A/T | 208/1253 | 15/69 | 1.00 (0.70-1.42) | 1.00 | 305/1253 | 27/69 | 1.33 (0.99-1.79) | 1.00 |
| | rs1126672 | C/T | 131/758 | 92/564 | 1.02 (0.87-1.19) | 1.00 | 201/758 | 131/564 | 0.96 (0.83-1.10) | 1.00 |
| | rs4699710 | T/C | 120/684 | 103/638 | 0.98 (0.84-1.13) | 1.00 | 179/684 | 153/638 | 0.96 (0.84-1.10) | 1.00 |
| | rs10017466 | T/C | 106/632 | 117/690 | 1.00 (0.87-1.16) | 1.00 | 157/632 | 174/690 | 1.02 (0.89-1.16) | 1.00 |
| | rs1800759 | C/A | 75/412 | 148/910 | 0.93 (0.79-1.10) | 1.00 | 101/412 | 231/910 | 1.00 (0.87-1.16) | 1.00 |
| | rs1800761 | G/A | 134/830 | 89/492 | 1.01 (0.87-1.18) | 1.00 | 193/830 | 139/492 | 1.07 (0.94-1.22) | 1.00 |
| | rs3762894 | T/C | 150/906 | 73/416 | 1.00 (0.86-1.18) | 1.00 | 218/906 | 114/416 | 1.04 (0.91-1.20) | 1.00 |
| <i>ADH7</i> | rs284787 | C/T | 126/817 | 97/505 | 1.14 (0.98-1.33) | 1.00 | 208/817 | 124/505 | 1.01 (0.88-1.16) | 1.00 |
| | rs894369 | C/G | 153/849 | 70/472 | 0.93 (0.80-1.09) | 1.00 | 217/849 | 115/472 | 0.96 (0.84-1.10) | 1.00 |
| | rs17588403 | T/A | 157/887 | 66/435 | 0.92 (0.78-1.08) | 1.00 | 230/887 | 102/435 | 0.94 (0.81-1.08) | 1.00 |
| | rs1154454 | T/C | 134/808 | 89/514 | 0.95 (0.81-1.11) | 1.00 | 208/808 | 124/514 | 0.94 (0.82-1.09) | 1.00 |
| | rs1154456 | T/C | 108/629 | 115/693 | 0.98 (0.84-1.14) | 1.00 | 150/629 | 182/693 | 1.08 (0.94-1.24) | 1.00 |
| | rs1154460 | G/A | 61/377 | 161/944 | 1.01 (0.86-1.20) | 1.00 | 88/377 | 244/944 | 1.06 (0.91-1.23) | 1.00 |
| | rs971074 | G/A | 168/1027 | 55/295 | 1.07 (0.90-1.27) | 1.00 | 257/1027 | 75/295 | 0.95 (0.81-1.12) | 1.00 |
| | rs1573496 | C/G | 186/1117 | 36/205 | 1.08 (0.88-1.33) | 1.00 | 292/1117 | 40/205 | 0.84 (0.69-1.02) | 1.00 |
| <i>ALDH2</i> | rs4767939 | A/G | 135/806 | 88/516 | 0.99 (0.85-1.16) | 1.00 | 191/806 | 140/516 | 1.08 (0.94-1.23) | 1.00 |
| | rs2238151 | T/C | 66/482 | 156/837 | 1.13 (0.95-1.34) | 1.00 | 114/482 | 217/837 | 1.02 (0.88-1.18) | 1.00 |
| | rs7312055 | G/A | 196/1201 | 27/121 | 0.88 (0.65-1.20) | 1.00 | 301/1201 | 31/121 | 0.83 (0.64-1.09) | 1.00 |
| | rs2158029 | G/A | 216/1269 | 7/53 | 0.74 (0.47-1.14) | 1.00 | 316/1269 | 16/53 | 1.06 (0.76-1.47) | 1.00 |
| | rs16941667 | C/T | 189/1097 | 34/224 | 0.96 (0.78-1.18) | 1.00 | 266/1097 | 66/224 | 1.11 (0.94-1.31) | 1.00 |
| | rs16941669 | T/G | 175/1084 | 48/236 | 1.14 (0.95-1.37) | 1.00 | 272/1084 | 60/236 | 0.98 (0.83-1.16) | 1.00 |
| <i>CYP2E1</i> | rs3813865 | G/C | 204/1215 | 19/106 | 1.02 (0.77-1.36) | 1.00 | 298/1215 | 34/106 | 1.11 (0.88-1.40) | 1.00 |
| | rs3813867 | G/C | 207/1237 | 16/84 | 1.02 (0.76-1.37) | 1.00 | 316/1237 | 16/84 | 0.81 (0.60-1.09) | 1.00 |
| | rs8192772 | T/C | 195/1114 | 28/208 | 0.92 (0.73-1.15) | 1.00 | 279/1114 | 53/208 | 1.07 (0.90-1.28) | 1.00 |
| | rs915908 | G/A | 164/1006 | 59/316 | 1.11 (0.93-1.32) | 1.00 | 263/1006 | 69/316 | 0.89 (0.75-1.04) | 1.00 |
| | rs915909 | C/T | 212/1271 | 11/46 | 1.11 (0.75-1.62) | 1.00 | 317/1271 | 15/46 | 1.02 (0.72-1.45) | 1.00 |
| | rs7092584 | C/T | 181/1052 | 42/269 | 0.98 (0.81-1.19) | 1.00 | 262/1052 | 70/269 | 1.05 (0.89-1.23) | 1.00 |
| | rs743535 | C/T | 182/1071 | 41/243 | 0.99 (0.82-1.20) | 1.00 | 268/1071 | 60/243 | 1.00 (0.85-1.19) | 1.00 |
| | rs2249695 | C/T | 108/682 | 115/639 | 1.01 (0.86-1.19) | 1.00 | 180/682 | 152/639 | 0.90 (0.78-1.04) | 1.00 |
| | rs28969387 | A/T | 219/1302 | 4/20 | 1.06 (0.59-1.90) | 1.00 | 326/1302 | 6/20 | 0.93 (0.56-1.53) | 1.00 |
| | rs11101812 | T/C | 219/1300 | 3/20 | 0.98 (0.52-1.85) | 1.00 | 327/1300 | 4/20 | 0.92 (0.52-1.63) | 1.00 |

Table S2, p. 2 of 2

| | | Oral cavity, oropharyngeal, hypopharyngeal cancer NOS | | | | | Oropharyngeal cancer | | | |
|--|------------|--|-----------------------------------|---|--------------------------------------|----------------------|-----------------------------------|---|--------------------------------------|----------------------|
| Gene | SNP | major/ minor alleles | n cases/controls | | Adjusted OR ^b (95% CI) | p-value ^c | n cases/controls | | Adjusted OR ^b (95% CI) | p-value ^c |
| | | | homozygous for major allele | one or two copies of minor allele | | | homozygous for major allele | one or two copies of minor allele | | |
| OXIDATIVE STRESS METABOLISM GENES | | | | | | | | | | |
| <i>CAT</i> | rs1049982 | C/T | 100/589 | 123/726 | 0.97 (0.831.13) | 1.00 | 134/589 | 197/726 | 1.07 (0.94-1.22) | 1.00 |
| <i>GPX1</i> | rs8179172 | T/A | 212/1271 | 11/51 | 0.97 (0.661.44) | 1.00 | 320/1271 | 12/51 | 0.90 (0.62-1.29) | 1.00 |
| | rs1800668 | C/T | 113/672 | 110/646 | 1.06 (0.911.23) | 1.00 | 172/672 | 160/646 | 1.02 (0.89-1.16) | 1.00 |
| | rs3811699 | A/G | 109/649 | 114/673 | 1.06 (0.911.23) | 1.00 | 167/649 | 165/673 | 1.01 (0.89-1.15) | 1.00 |
| | rs3448 | C/T | 113/729 | 110/593 | 1.12 (0.971.31) | 1.00 | 190/729 | 142/593 | 0.98 (0.86-1.12) | 1.00 |
| <i>GPX2</i> | rs11623705 | G/T | 185/1069 | 38/253 | 0.94 (0.771.14) | 1.00 | 268/1069 | 64/253 | 1.02 (0.86-1.21) | 1.00 |
| | rs2412065 | G/C | 122/733 | 101/589 | 0.98 (0.841.15) | 1.00 | 193/733 | 139/589 | 0.94 (0.82-1.08) | 1.00 |
| | rs2737844 | C/T | 88/511 | 134/806 | 0.95 (0.811.12) | 1.00 | 136/511 | 194/806 | 0.93 (0.81-1.08) | 1.00 |
| <i>GPX4</i> | rs757229 | G/C | 60/358 | 162/963 | 0.98 (0.831.16) | 1.00 | 85/358 | 247/963 | 1.02 (0.88-1.18) | 1.00 |
| <i>SOD1</i> | rs11910115 | A/C | 208/1257 | 15/65 | 1.01 (0.701.45) | 1.00 | 319/1257 | 13/65 | 0.86 (0.60-1.22) | 1.00 |
| | rs4998557 | G/A | 155/959 | 68/360 | 1.00 (0.841.20) | 1.00 | 232/959 | 99/360 | 1.07 (0.92-1.25) | 1.00 |
| | rs10432782 | T/G | 163/992 | 60/329 | 0.98 (0.821.17) | 1.00 | 240/992 | 91/329 | 1.07 (0.92-1.25) | 1.00 |
| | rs2070424 | A/G | 187/1120 | 36/202 | 0.95 (0.761.17) | 1.00 | 274/1120 | 58/202 | 1.05 (0.88-1.26) | 1.00 |
| | rs1041740 | C/T | 127/705 | 96/617 | 0.94 (0.801.10) | 1.00 | 174/705 | 158/617 | 0.99 (0.87-1.14) | 1.00 |
| <i>SOD2</i> | rs4342445 | G/A | 145/850 | 78/472 | 1.01 (0.861.18) | 1.00 | 202/850 | 130/472 | 1.08 (0.95-1.24) | 1.00 |
| | rs2842980 | A/T | 129/807 | 94/515 | 1.06 (0.911.23) | 1.00 | 189/807 | 143/515 | 1.09 (0.96-1.25) | 1.00 |
| | rs8031 | T/A | 77/382 | 146/940 | 0.91 (0.771.07) | 1.00 | 102/382 | 230/940 | 0.99 (0.86-1.14) | 1.00 |
| | rs5746134 | C/T | 207/1257 | 16/65 | 1.20 (0.851.68) | 1.00 | 304/1257 | 28/65 | 1.41 (1.05-1.89) | 1.00 |
| | rs2758331 | C/A | 86/419 | 137/903 | 0.90 (0.771.06) | 1.00 | 108/419 | 224/903 | 1.02 (0.88-1.17) | 1.00 |

^a Cases and controls do not sum to 1227 and 1325, respectively, because 4 cases and 3 controls are missing information on duration of cigarette smoking, and because a few subjects lack genotype information for some SNPs

^b Conditional logistic regression models for estimating main effect of each SNP were conditioned on sex, race, and age category, and adjusted for continuous duration of smoking in years (rounded to whole years). Odds ratios are for those with one or more copies of the minor allele versus the referent group of those homozygous for the major allele (dominant genetic model).

^c Bonferroni-corrected for 64 statistical tests

Table S3. Haplotype effects on risk of developing SCCHN (additive genetic model)

| European-Americans | | | | | African-Americans | | | | | | | | | | | | |
|--------------------|-----------------------------------|------------|------------|---------------------|--------------------------|--|-----------|-----------------------------------|------------|------------|----------------------|--------------------------|--|----------------------|----------------------|----------------------|------|
| Gene | Haplotype definition ^a | | | Haplotype frequency | OR (95% CI) ^b | Bonferroni-adjusted p-value (13 tests) | Gene | Haplotype definition ^a | | | Haplotype frequency | OR (95% CI) ^b | Bonferroni-adjusted p-value (12 tests) | | | | |
| ADH1B | rs12507573 | rs1042026 | rs1693457 | | | | ADH1B | Haplotype 1: | | | | | | | | | |
| | A | A | T | 0.459 | 1.00 (ref) | | | rs12507573 | rs1042026 | | 0.483 | 1.00 (ref) | | | | | |
| | C | A | C | 0.183 | 1.17 (0.96 - 1.42) | 1.00 | | C | A | | 0.422 | 1.02 (0.75 - 1.39) | 1.00 | | | | |
| | C | A | T | 0.069 | 1.09 (0.83 - 1.44) | 1.00 | | A | A | | 0.096 | 0.80 (0.49 - 1.32) | 1.00 | | | | |
| | C | G | T | 0.287 | 1.05 (0.89 - 1.23) | 1.00 | | C | G | | | | | | | | |
| | * | * | * | 0.002 | 0.30 (0.06 - 1.55) | 1.00 | | | | | | | | | | | |
| | Haplotype 2: | | | | | | | | | | | | | | | | |
| | rs7673353 | rs17028834 | rs1693457 | rs1159918 | | | | C | T | T | G | 0.271 | 1.00 (ref) | | | | |
| | C | C | T | T | | | | C | C | T | T | 0.130 | 1.84 (1.09 - 3.11) | 0.28 | | | |
| | C | T | C | T | | | | C | T | C | T | 0.202 | 0.79 (0.52 - 1.22) | 1.00 | | | |
| | C | T | T | T | | | | C | T | T | T | 0.224 | 1.21 (0.80 - 1.83) | 1.00 | | | |
| | T | T | T | T | | | | T | T | T | T | 0.173 | 1.06 (0.67 - 1.66) | 1.00 | | | |
| ADH1C | Haplotype 1: | | | | | | ADH1C | rs1614972 | rs1391088 | rs1693482 | rs1631460 | rs11936869 | | | | | |
| | rs2298753 | rs1614972 | | 0.597 | 1.00 (ref) | | | T | C | C | C | G | 0.427 | 1.00 (ref) | | | |
| | T | C | | 0.106 | 1.24 (0.99 - 1.55) | 0.76 | | C | A | C | C | C | 0.080 | 1.45 (0.80 - 2.62) | 1.00 | | |
| | C | C | | 0.297 | 1.06 (0.91 - 1.23) | 1.00 | | C | C | C | C | C | 0.264 | 1.05 (0.74 - 1.51) | 1.00 | | |
| | T | T | | | | | | C | C | T | G | C | 0.160 | 1.40 (0.88 - 2.21) | 1.00 | | |
| | Haplotype 2: | | | | | | T | C | C | C | C | 0.059 | 1.29 (0.65 - 2.56) | 1.00 | | | |
| | rs1693482 | rs1631460 | rs11936869 | | | | * | * | * | * | * | 0.010 | 7.60 (0.99 - 58.20) | 0.62 | | | |
| | T | G | C | 0.400 | 1.00 (ref) | | | | | | | | | | | | |
| | C | C | C | 0.333 | 0.94 (0.80 - 1.10) | 1.00 | | | | | | | | | | | |
| | C | C | G | 0.263 | 1.00 (0.84 - 1.19) | 1.00 | | | | | | | | | | | |
| | C | G | C | 0.004 | 0.99 (0.33 - 2.94) | 1.00 | | | | | | | | | | | |
| ADH4 | Haplotype 1: | | | | | | ADH4 | Haplotype 1: | | | | | | | | | |
| | rs1126672 | rs4699710 | rs10017466 | rs1800759 | 0.617 | 1.00 (ref) | | rs29001227 | rs1126672 | rs4699710 | rs10017466 | rs1800759 | 0.408 | 1.00 (ref) | | | |
| | C | T | T | C | 0.029 | 1.32 (0.88 - 1.99) | | 1.00 | A | C | T | T | A | 0.074 | 0.94 (0.51 - 1.72) | 1.00 | |
| | C | C | C | A | 0.078 | 0.93 (0.72 - 1.22) | | 1.00 | A | C | C | C | A | 0.242 | 1.03 (0.70 - 1.50) | 1.00 | |
| | C | T | T | A | 0.273 | 0.96 (0.81 - 1.12) | | 1.00 | A | C | T | T | C | 0.110 | 1.50 (0.89 - 2.51) | 1.00 | |
| | * | * | * | * | 0.002 | 2.21 (0.61 - 7.97) | 1.00 | A | T | C | C | A | 0.164 | 1.61 (1.04 - 2.49) | 0.42 | | |
| | Haplotype 2: | | | | | | | Haplotype 2: | | | | | | | | | |
| | rs1800761 | rs3762894 | | 0.810 | 1.00 (ref) | | rs1800761 | rs3762894 | | 0.711 | 1.00 (ref) | | | | | | |
| | G | T | | 0.160 | 1.02 (0.84 - 1.23) | 1.00 | G | T | | 0.210 | 0.92 (0.65 - 1.31) | 1.00 | | | | | |
| | A | C | | 0.029 | 1.38 (0.91 - 2.07) | 1.00 | A | C | | 0.078 | 0.97 (0.55 - 1.70) | 1.00 | | | | | |
| | A | T | | | | | A | T | | | | | | | | | |
| ADH7 | Haplotype 1: | | | | | | ADH7 | Haplotype 1: | | | | | | | | | |
| | rs284787 | rs894369 | rs17588403 | 0.371 | 1.00 (ref) | | | rs284787 | rs894369 | rs17588403 | 0.578 | 1.00 (ref) | | | | | |
| | C | C | T | 0.192 | 1.03 (0.85 - 1.25) | 1.00 | | C | C | T | 0.124 | 0.73 (0.46 - 1.18) | 1.00 | | | | |
| | C | C | A | 0.196 | 0.88 (0.73 - 1.06) | 1.00 | | C | C | A | 0.154 | 0.90 (0.59 - 1.38) | 1.00 | | | | |
| | C | G | T | 0.241 | 1.13 (0.95 - 1.35) | 1.00 | | C | G | T | 0.143 | 0.85 (0.55 - 1.32) | 1.00 | | | | |
| | T | C | T | | | | T | C | T | | | | | | | | |
| | Haplotype 2: | | | | | | | Haplotype 2: | | | | | | | | | |
| | rs1154456 | rs1154460 | rs971074 | rs1573496 | 0.539 | 1.00 (ref) | rs1154456 | rs1154460 | rs971074 | 0.508 | 1.00 (ref) | | | | | | |
| | T | G | G | C | 0.353 | 1.08 (0.93 - 1.25) | 1.00 | T | G | G | 0.163 | 0.96 (0.63 - 1.48) | 1.00 | | | | |
| | C | A | G | C | 0.010 | 0.93 (0.47 - 1.84) | 1.00 | C | A | G | 0.177 | 1.18 (0.77 - 1.81) | 1.00 | | | | |
| | T | A | A | C | 0.092 | 0.98 (0.76 - 1.25) | 1.00 | T | A | A | 0.150 | 0.97 (0.63 - 1.48) | 1.00 | | | | |
| | T | A | G | C | 0.003 | 2.78 (0.90 - 8.63) | 1.00 | T | A | G | | | | | | | |
| | * | * | * | * | 0.002 | 0.97 (0.12 - 8.06) | 1.00 | | | | | | | | | | |
| ALDH2 | rs4767939 | rs2238151 | rs16941667 | rs16941669 | 0.650 | 1.00 (ref) | ALDH2 | rs4767939 | rs2238151 | rs7312055 | rs2158029 | rs16941667 | rs16941669 | 0.263 | 1.00 (ref) | | |
| | A | T | C | T | 0.102 | 1.12 (0.89 - 1.41) | | 1.00 | A | C | A | G | C | T | 0.081 | 1.50 (0.79 - 2.83) | 1.00 |
| | A | C | C | G | 0.057 | 1.18 (0.88 - 1.60) | | 1.00 | A | C | G | G | C | G | 0.141 | 1.06 (0.65 - 1.73) | 1.00 |
| | A | C | C | T | 0.106 | 1.13 (0.90 - 1.42) | | 1.00 | A | C | G | G | C | T | 0.111 | 0.48 (0.28 - 0.83) | 0.11 |
| | G | C | C | T | 0.083 | 1.22 (0.95 - 1.58) | | 1.00 | G | T | G | G | C | T | 0.090 | 1.02 (0.57 - 1.85) | 1.00 |
| | G | C | T | | | | G | C | G | A | C | T | 0.177 | 1.05 (0.67 - 1.67) | 1.00 | | |
| | G | C | T | | | | G | C | G | G | C | T | 0.137 | 1.23 (0.75 - 2.02) | 1.00 | | |
| CYP2E1 | rs915908 | rs7092584 | rs743535 | rs2249695 | 0.653 | 1.00 (ref) | CYP2E1 | rs915908 | rs7092584 | rs743535 | rs2249695 | 0.144 | 0.88 (0.72 - 1.07) | 1.00 | | | |
| | G | C | C | C | 0.102 | 0.73 (0.58 - 0.92) | | 0.11 | G | C | C | T | 0.010 | 0.80 (0.40 - 1.61) | 1.00 | | |
| | A | C | C | C | 0.010 | 0.80 (0.40 - 1.61) | | 1.00 | G | T | C | T | 0.089 | 1.03 (0.81 - 1.32) | 1.00 | | |
| | G | C | C | T | | | | | G | T | C | T | | | | | |
| | G | T | C | T | | | | | G | T | C | T | | | | | |
| | G | T | T | | | | | G | T | T | | | | | | | |
| GPX1 | rs1800668 | rs3811699 | rs3448 | | 0.434 | 1.00 (ref) | GPX1 | rs1800668 | rs3811699 | rs3448 | | 0.422 | 1.00 (ref) | | | | |
| | C | A | C | 0.271 | 1.05 (0.90 - 1.24) | 1.00 | | C | A | C | 0.271 | 1.14 (0.79 - 1.65) | 1.00 | | | | |
| | C | A | T | 0.294 | 1.02 (0.87 - 1.20) | 1.00 | | C | A | T | 0.060 | 0.87 (0.46 - 1.67) | 1.00 | | | | |
| | T | G | C | | | | | C | G | C | 0.247 | 1.04 (0.72 - 1.52) | 1.00 | | | | |
| | T | G | C | | | | T | G | C | | | | | | | | |
| GPX2 | rs11623705 | rs2412065 | rs2737844 | | 0.700 | 1.00 (ref) | GPX2 | rs2412065 | rs2737844 | | 0.417 | 1.00 (ref) | | | | | |
| | G | G | C | 0.092 | 0.69 (0.54 - 0.88) | 0.04 | | C | T | | 0.190 | 0.84 (0.56 - 1.25) | 1.00 | | | | |
| | G | C | T | 0.090 | 0.81 (0.63 - 1.03) | 1.00 | | G | C | | 0.393 | 1.07 (0.77 - 1.49) | 1.00 | | | | |
| | G | G | T | 0.119 | 0.98 (0.79 - 1.21) | 1.00 | | G | T | | | | | | | | |
| | T | C | T | | | | | | | | | | | | | | |
| SOD1 | rs4998557 | rs10432782 | rs2070424 | rs1041740 | 0.576 | 1.00 (ref) | SOD1 | rs4998557 | rs10432782 | rs2070424 | rs1041740 | 0.097 | 1.00 (0.59 - 1.71) | 1.00 | | | |
| | G | T | A | C | 0.048 | 1.10 (0.79 - 1.52) | | 1.00 | G | T | A | C | 0.167 | 0.56 (0.37 - 0.86) | 0.09 | | |
| | A | G | A | C | 0.065 | 1.42 (1.07 - 1.88) | | 0.18 | A | G | A | C | 0.104 | 1.13 (0.68 - 1.90) | 1.00 | | |
| | A | G | G | C | 0.311 | 0.98 (0.84 - 1.14) | | 1.00 | A | G | G | C | 0.111 | 1.15 (0.69 - 1.90) | 1.00 | | |
| | G | T | A | T | | | | A | T | A | C | | | | | | |
| | G | T | A | T | | | | G | T | A | T | | | | | | |

Table S3. Haplotype effects on risk of developing SCCHN (additive genetic model)

| European-Americans | | | | | African-Americans | | | | | | | | | | | |
|--------------------|-----------------------------------|-----------|--------|-----------|---------------------|--------------------------|--|------|-----------------------------------|-----------|--------|-----------|---------------------|--------------------------|--|------|
| Gene | Haplotype definition ^a | | | | Haplotype frequency | OR (95% CI) ^b | Bonferroni-adjusted p-value (13 tests) | Gene | Haplotype definition ^a | | | | Haplotype frequency | OR (95% CI) ^b | Bonferroni-adjusted p-value (12 tests) | |
| SOD2 | rs4342445 | rs2842980 | rs8031 | rs2758331 | | | | SOD2 | rs4342445 | rs2842980 | rs8031 | rs5746134 | rs2758331 | | | |
| | G | A | A | A | 0.486 | 1.00 (ref) | | | G | A | A | C | A | 0.245 | 1.00 (ref) | |
| | A | A | T | C | 0.230 | 1.18 (1.00 - 1.40) | 0.71 | | A | A | T | C | C | 0.133 | 1.00 (0.59 - 1.68) | 1.00 |
| | G | A | T | C | 0.071 | 0.96 (0.73 - 1.27) | 1.00 | | G | A | A | C | C | 0.091 | 0.68 (0.39 - 1.20) | 1.00 |
| | G | T | T | C | 0.211 | 1.00 (0.84 - 1.20) | 1.00 | | G | A | T | C | C | 0.227 | 0.65 (0.41 - 1.02) | 0.74 |
| | * | * | * | * | 0.002 | 3.01 (0.50 - 18.03) | 1.00 | | G | T | T | C | C | 0.151 | 0.91 (0.56 - 1.47) | 1.00 |
| | | | | | | | | | G | T | T | T | C | 0.154 | 1.46 (0.89 - 2.40) | 1.00 |

^a * indicates all remaining allele combinations

^b Odds ratios for main effects of each haplotype were calculated from unconditional logistic regression models, adjusted for smoking duration in whole years, and matching variables sex, age category, sex*age category.

Table S4. Odds ratios for developing SCCHN for SNP genotypes, at varying levels of lifetime alcohol consumption

| Gene | SNP | major/ minor alleles | lifetime ethanol consumption (ml) | Odds ratios in comparison to common referent group ^a | | | | Assessment of additive interaction between SNP and lifetime alcohol consumption |
|---------------------------------|------------|----------------------------|--------------------------------------|---|---|--|---|---|
| | | | | homozygous major allele | | at least one minor allele | | |
| | | | | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | |
| ALCOHOL METABOLISM GENES | | | | | | | | |
| ADH1B | rs12507573 | C/A | never drinkers | 40 / 72 | 1.00 (ref) | 77 / 208 | 0.64 (0.39 - 1.06) | -0.19 |
| | | | >0 to <134,699 | 68 / 142 | 0.62 (0.37 - 1.05) | 141 / 323 | 0.52 (0.32 - 0.85) | |
| | | | 134,699 to <757,550 | 84 / 118 | 0.74 (0.44 - 1.27) | 234 / 242 | 1.05 (0.65 - 1.69) | |
| | | | 757,550+ | 160 / 47 | 2.75 (1.56 - 4.86) | 345 / 126 | 2.21 (1.34 - 3.65) | |
| | rs1042026 | A/G | never drinkers | 59 / 157 | 1.00 (ref) | 58 / 123 | 1.20 (0.75 - 1.93) | -0.75 |
| | | | >0 to <134,699 | 112 / 253 | 0.80 (0.53 - 1.20) | 97 / 213 | 0.85 (0.56 - 1.30) | |
| | | | 134,699 to <757,550 | 188 / 206 | 1.48 (0.99 - 2.22) | 130 / 154 | 1.31 (0.86 - 2.01) | |
| | | | 757,550+ | 312 / 98 | 3.75 (2.42 - 5.81) | 193 / 75 | 3.20 (2.02 - 5.08) | |
| | rs7673353 | C/T | never drinkers | 115 / 261 | 1.00 (ref) | 2 / 19 | 0.29 (0.06 - 1.35) | 1.06 |
| | | | >0 to <134,699 | 199 / 438 | 0.72 (0.53 - 0.98) | 10 / 28 | 0.71 (0.31 - 1.62) | |
| | | | 134,699 to <757,550 | 289 / 346 | 1.17 (0.85 - 1.60) | 29 / 14 | 3.14 (1.44 - 6.84) | |
| | | | 757,550+ | 453 / 159 | 3.10 (2.19 - 4.39) | 52 / 14 | 3.44 (1.66 - 7.15) | |
| rs17028834 | T/C | never drinkers | 113 / 272 | 1.00 (ref) | 4 / 8 | 1.13 (0.28 - 4.49) | 3.55 | |
| | | >0 to <134,699 | 201 / 447 | 0.75 (0.55 - 1.01) | 8 / 19 | 1.01 (0.40 - 2.54) | | |
| | | 134,699 to <757,550 | 303 / 342 | 1.29 (0.95 - 1.77) | 15 / 18 | 1.39 (0.61 - 3.16) | | |
| | | 757,550+ | 453 / 163 | 3.06 (2.16 - 4.32) | 52 / 10 | 6.73 (3.06 - 14.81) | | |
| rs1693457 | T/C | never drinkers | 78 / 177 | 1.00 (ref) | 38 / 103 | 0.86 (0.52 - 1.41) | 1.02 | |
| | | >0 to <134,699 | 146 / 295 | 0.77 (0.54 - 1.11) | 63 / 170 | 0.64 (0.42 - 0.98) | | |
| | | 134,699 to <757,550 | 201 / 234 | 1.21 (0.84 - 1.75) | 117 / 126 | 1.31 (0.87 - 1.98) | | |
| | | 757,550+ | 327 / 122 | 2.85 (1.92 - 4.24) | 178 / 51 | 3.72 (2.34 - 5.93) | | |
| rs1229984 | G/A | never drinkers | 109 / 268 | 1.00 (ref) | 8 / 12 | 1.60 (0.59 - 4.32) | -1.85 | |
| | | >0 to <134,699 | 206 / 427 | 0.83 (0.61 - 1.13) | 3 / 39 | 0.15 (0.04 - 0.50) | | |
| | | 134,699 to <757,550 | 305 / 339 | 1.35 (0.98 - 1.85) | 13 / 21 | 1.07 (0.48 - 2.37) | | |
| | | 757,550+ | 499 / 169 | 3.36 (2.38 - 4.75) | 6 / 4 | 2.11 (0.49 - 9.10) | | |
| rs1159918 | G/T | never drinkers | 49 / 101 | 1.00 (ref) | 68 / 179 | 0.90 (0.55 - 1.46) | 1.03 | |
| | | >0 to <134,699 | 80 / 168 | 0.67 (0.41 - 1.07) | 129 / 298 | 0.73 (0.47 - 1.14) | | |
| | | 134,699 to <757,550 | 107 / 129 | 1.11 (0.68 - 1.80) | 211 / 231 | 1.26 (0.81 - 1.97) | | |
| | | 757,550+ | 140 / 58 | 2.42 (1.43 - 4.08) | 365 / 115 | 3.34 (2.08 - 5.36) | | |
| rs1229982 | G/T | never drinkers | 72 / 168 | 1.00 (ref) | 45 / 112 | 0.91 (0.56 - 1.47) | 0.52 | |
| | | >0 to <134,699 | 131 / 270 | 0.79 (0.54 - 1.16) | 78 / 196 | 0.64 (0.42 - 0.97) | | |
| | | 134,699 to <757,550 | 180 / 211 | 1.21 (0.83 - 1.79) | 138 / 149 | 1.28 (0.85 - 1.93) | | |
| | | 757,550+ | 274 / 98 | 2.91 (1.91 - 4.43) | 231 / 75 | 3.34 (2.14 - 5.21) | | |
| ADH1C | rs2298753 | T/C | never drinkers | 97 / 238 | 1.00 (ref) | 20 / 42 | 1.19 (0.63 - 2.27) | -0.56 |
| | | | >0 to <134,699 | 169 / 392 | 0.74 (0.53 - 1.03) | 40 / 74 | 0.96 (0.59 - 1.57) | |
| | | | 134,699 to <757,550 | 246 / 303 | 1.22 (0.87 - 1.71) | 72 / 57 | 1.84 (1.16 - 2.93) | |
| | | | 757,550+ | 429 / 144 | 3.34 (2.32 - 4.81) | 76 / 29 | 2.97 (1.71 - 5.16) | |
| | rs1614972 | C/T | never drinkers | 56 / 123 | 1.00 (ref) | 61 / 157 | 0.84 (0.53 - 1.34) | 0.02 |
| | | | >0 to <134,699 | 81 / 219 | 0.52 (0.33 - 0.82) | 128 / 247 | 0.84 (0.55 - 1.28) | |
| | | | 134,699 to <757,550 | 155 / 152 | 1.33 (0.86 - 2.06) | 163 / 208 | 1.05 (0.68 - 1.61) | |
| | | | 757,550+ | 216 / 75 | 2.99 (1.86 - 4.79) | 289 / 98 | 2.85 (1.80 - 4.50) | |
| | rs1391088 | C/A | never drinkers | 100 / 235 | 1.00 (ref) | 17 / 45 | 0.95 (0.50 - 1.82) | 0.12 |
| | | | >0 to <134,699 | 181 / 392 | 0.76 (0.55 - 1.06) | 28 / 74 | 0.66 (0.39 - 1.12) | |
| | | | 134,699 to <757,550 | 270 / 315 | 1.24 (0.89 - 1.72) | 48 / 45 | 1.59 (0.95 - 2.67) | |
| | | | 757,550+ | 427 / 145 | 3.18 (2.21 - 4.57) | 78 / 28 | 3.24 (1.87 - 5.62) | |
| rs1693482 | C/T | never drinkers | 44 / 123 | 1.00 (ref) | 73 / 157 | 1.24 (0.77 - 2.01) | -0.24 | |
| | | >0 to <134,699 | 84 / 199 | 0.87 (0.54 - 1.38) | 124 / 265 | 0.85 (0.55 - 1.33) | | |
| | | 134,699 to <757,550 | 135 / 164 | 1.37 (0.86 - 2.18) | 182 / 196 | 1.54 (0.98 - 2.41) | | |
| | | 757,550+ | 234 / 77 | 3.64 (2.22 - 5.96) | 271 / 96 | 3.65 (2.26 - 5.88) | | |
| rs1631460 | C/G | never drinkers | 43 / 119 | 1.00 (ref) | 74 / 161 | 1.23 (0.76 - 1.99) | -0.17 | |
| | | >0 to <134,699 | 81 / 195 | 0.86 (0.53 - 1.37) | 128 / 271 | 0.86 (0.55 - 1.34) | | |
| | | 134,699 to <757,550 | 134 / 162 | 1.38 (0.87 - 2.22) | 184 / 198 | 1.53 (0.97 - 2.41) | | |
| | | 757,550+ | 231 / 76 | 3.61 (2.19 - 5.93) | 273 / 97 | 3.66 (2.26 - 5.93) | | |
| rs11936869 | C/G | never drinkers | 61 / 135 | 1.00 (ref) | 56 / 145 | 0.90 (0.56 - 1.43) | -0.17 | |
| | | >0 to <134,699 | 102 / 237 | 0.64 (0.42 - 0.98) | 107 / 228 | 0.79 (0.52 - 1.20) | | |
| | | 134,699 to <757,550 | 170 / 174 | 1.37 (0.90 - 2.08) | 148 / 186 | 1.08 (0.71 - 1.65) | | |
| | | 757,550+ | 242 / 78 | 3.17 (2.00 - 5.02) | 263 / 95 | 2.90 (1.85 - 4.54) | | |

Table S4. Odds ratios for developing SCCHN for SNP genotypes, at varying levels of lifetime alcohol consumption

| Gene | SNP | major/ minor alleles | lifetime ethanol consumption (ml) | Odds ratios in comparison to common referent group ^a | | | | Assessment of additive interaction between SNP and lifetime alcohol consumption | |
|-------------|------------|----------------------------|--------------------------------------|---|---|--|---|---|---|
| | | | | homozygous major allele | | at least one minor allele | | | ICR ^d (bold if statistically significant after Bonferroni correction) |
| | | | | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | | |
| <i>ADHA</i> | rs29001227 | A/T | never drinkers | 113 / 271 | 1.00 (ref) | 4 / 9 | 1.04 (0.27 - 4.02) | 4.55 | |
| | | | >0 to <134,699 | 199 / 445 | 0.74 (0.55 - 1.01) | 10 / 21 | 0.98 (0.41 - 2.31) | | |
| | | | 134,699 to <757,550 | 294 / 334 | 1.28 (0.94 - 1.76) | 24 / 26 | 1.47 (0.74 - 2.93) | | |
| | | | 757,550+ | 445 / 163 | 3.02 (2.13 - 4.27) | 60 / 10 | 7.61 (3.46 - 16.75) | | |
| | rs1126672 | C/T | never drinkers | 58 / 152 | 1.00 (ref) | 59 / 128 | 1.19 (0.74 - 1.92) | -0.14 | |
| | | | >0 to <134,699 | 120 / 263 | 0.84 (0.56 - 1.27) | 89 / 203 | 0.79 (0.51 - 1.22) | | |
| | | | 134,699 to <757,550 | 186 / 214 | 1.37 (0.91 - 2.06) | 132 / 146 | 1.46 (0.95 - 2.24) | | |
| | | | 757,550+ | 310 / 107 | 3.47 (2.24 - 5.37) | 195 / 66 | 3.53 (2.21 - 5.63) | | |
| | rs4699710 | T/C | never drinkers | 49 / 142 | 1.00 (ref) | 68 / 138 | 1.45 (0.90 - 2.32) | 0.09 | |
| | | | >0 to <134,699 | 108 / 239 | 0.93 (0.61 - 1.44) | 101 / 227 | 0.91 (0.58 - 1.41) | | |
| | | | 134,699 to <757,550 | 172 / 183 | 1.70 (1.10 - 2.63) | 146 / 177 | 1.45 (0.93 - 2.24) | | |
| | | | 757,550+ | 272 / 100 | 3.69 (2.33 - 5.85) | 233 / 73 | 4.23 (2.63 - 6.80) | | |
| | rs10017466 | T/C | never drinkers | 45 / 134 | 1.00 (ref) | 72 / 146 | 1.46 (0.91 - 2.35) | 1.08 | |
| | | | >0 to <134,699 | 100 / 224 | 0.93 (0.60 - 1.46) | 109 / 242 | 0.93 (0.60 - 1.45) | | |
| | | | 134,699 to <757,550 | 154 / 163 | 1.73 (1.11 - 2.72) | 164 / 197 | 1.49 (0.96 - 2.31) | | |
| | | | 757,550+ | 223 / 93 | 3.26 (2.03 - 5.24) | 281 / 80 | 4.80 (2.98 - 7.72) | | |
| | rs1800759 | C/A | never drinkers | 35 / 93 | 1.00 (ref) | 82 / 187 | 1.17 (0.70 - 1.95) | 0.66 | |
| | | | >0 to <134,699 | 70 / 153 | 0.80 (0.47 - 1.36) | 139 / 313 | 0.86 (0.53 - 1.39) | | |
| | | | 134,699 to <757,550 | 101 / 104 | 1.61 (0.95 - 2.73) | 217 / 256 | 1.37 (0.85 - 2.22) | | |
| | | | 757,550+ | 132 / 55 | 3.02 (1.72 - 5.31) | 373 / 118 | 3.85 (2.32 - 6.41) | | |
| | rs1800761 | G/A | never drinkers | 68 / 167 | 1.00 (ref) | 49 / 113 | 1.13 (0.70 - 1.81) | -0.63 | |
| | | | >0 to <134,699 | 121 / 300 | 0.70 (0.48 - 1.04) | 88 / 166 | 0.95 (0.63 - 1.44) | | |
| | | | 134,699 to <757,550 | 208 / 235 | 1.39 (0.95 - 2.04) | 110 / 125 | 1.29 (0.84 - 1.98) | | |
| | | | 757,550+ | 310 / 101 | 3.57 (2.35 - 5.45) | 195 / 72 | 3.07 (1.95 - 4.83) | | |
| rs3762894 | T/C | never drinkers | 82 / 178 | 1.00 (ref) | 35 / 102 | 0.75 (0.45 - 1.23) | -0.74 | | |
| | | >0 to <134,699 | 135 / 328 | 0.62 (0.43 - 0.89) | 74 / 138 | 0.83 (0.55 - 1.27) | | | |
| | | 134,699 to <757,550 | 226 / 263 | 1.15 (0.80 - 1.65) | 92 / 97 | 1.22 (0.79 - 1.88) | | | |
| | | 757,550+ | 348 / 108 | 3.28 (2.19 - 4.89) | 157 / 65 | 2.29 (1.45 - 3.60) | | | |
| <i>ADH7</i> | rs284787 | C/T | never drinkers | 67 / 166 | 1.00 (ref) | 50 / 114 | 1.05 (0.65 - 1.68) | 0.45 | |
| | | | >0 to <134,699 | 127 / 299 | 0.74 (0.50 - 1.08) | 82 / 167 | 0.83 (0.54 - 1.27) | | |
| | | | 134,699 to <757,550 | 195 / 210 | 1.36 (0.92 - 2.01) | 123 / 150 | 1.26 (0.83 - 1.92) | | |
| | | | 757,550+ | 308 / 112 | 3.11 (2.05 - 4.72) | 197 / 61 | 3.61 (2.26 - 5.74) | | |
| | rs894369 | C/G | never drinkers | 84 / 181 | 1.00 (ref) | 33 / 99 | 0.71 (0.42 - 1.17) | 0.80 | |
| | | | >0 to <134,699 | 140 / 303 | 0.70 (0.48 - 1.00) | 69 / 162 | 0.63 (0.42 - 0.96) | | |
| | | | 134,699 to <757,550 | 207 / 214 | 1.26 (0.87 - 1.83) | 111 / 146 | 1.00 (0.67 - 1.50) | | |
| | | | 757,550+ | 350 / 125 | 2.75 (1.85 - 4.07) | 155 / 48 | 3.25 (2.02 - 5.22) | | |
| | rs17588403 | T/A | never drinkers | 71 / 190 | 1.00 (ref) | 46 / 90 | 1.35 (0.83 - 2.19) | -1.33 | |
| | | | >0 to <134,699 | 142 / 307 | 0.87 (0.60 - 1.25) | 67 / 159 | 0.78 (0.51 - 1.21) | | |
| | | | 134,699 to <757,550 | 226 / 251 | 1.49 (1.03 - 2.16) | 92 / 109 | 1.32 (0.85 - 2.05) | | |
| | | | 757,550+ | 347 / 108 | 3.96 (2.63 - 5.96) | 158 / 65 | 2.97 (1.88 - 4.69) | | |
| | rs1154454 | T/C | never drinkers | 76 / 175 | 1.00 (ref) | 41 / 105 | 0.85 (0.52 - 1.39) | 0.13 | |
| | | | >0 to <134,699 | 143 / 276 | 0.82 (0.57 - 1.19) | 66 / 190 | 0.55 (0.36 - 0.84) | | |
| | | | 134,699 to <757,550 | 191 / 227 | 1.16 (0.80 - 1.70) | 127 / 133 | 1.31 (0.86 - 1.98) | | |
| | | | 757,550+ | 306 / 106 | 3.04 (2.02 - 4.58) | 199 / 67 | 3.02 (1.91 - 4.77) | | |
| | rs1154456 | T/C | never drinkers | 54 / 125 | 1.00 (ref) | 63 / 155 | 0.77 (0.48 - 1.22) | -0.03 | |
| | | | >0 to <134,699 | 91 / 218 | 0.62 (0.40 - 0.96) | 118 / 248 | 0.68 (0.44 - 1.04) | | |
| | | | 134,699 to <757,550 | 149 / 181 | 1.03 (0.67 - 1.59) | 169 / 179 | 1.20 (0.78 - 1.85) | | |
| | | | 757,550+ | 256 / 81 | 2.91 (1.82 - 4.65) | 249 / 92 | 2.65 (1.67 - 4.20) | | |
| | rs1154460 | G/A | never drinkers | 35 / 65 | 1.00 (ref) | 81 / 215 | 0.63 (0.37 - 1.07) | 0.92 | |
| | | | >0 to <134,699 | 57 / 131 | 0.53 (0.30 - 0.92) | 152 / 334 | 0.55 (0.34 - 0.90) | | |
| | | | 134,699 to <757,550 | 88 / 111 | 0.83 (0.48 - 1.45) | 229 / 249 | 0.97 (0.59 - 1.60) | | |
| | | | 757,550+ | 133 / 54 | 1.93 (1.07 - 3.46) | 371 / 119 | 2.48 (1.47 - 4.20) | | |
| rs971074 | G/A | never drinkers | 91 / 219 | 1.00 (ref) | 26 / 61 | 1.15 (0.65 - 2.03) | 0.86 | | |
| | | >0 to <134,699 | 166 / 357 | 0.79 (0.56 - 1.11) | 43 / 109 | 0.73 (0.46 - 1.17) | | | |
| | | 134,699 to <757,550 | 247 / 275 | 1.34 (0.95 - 1.89) | 71 / 85 | 1.31 (0.84 - 2.05) | | | |
| | | 757,550+ | 380 / 140 | 3.13 (2.15 - 4.55) | 125 / 33 | 4.14 (2.47 - 6.94) | | | |
| rs1573496 | C/G | never drinkers | 98 / 236 | 1.00 (ref) | 19 / 44 | 1.08 (0.57 - 2.05) | 0.46 | | |
| | | >0 to <134,699 | 175 / 386 | 0.76 (0.55 - 1.06) | 34 / 80 | 0.76 (0.46 - 1.27) | | | |
| | | 134,699 to <757,550 | 272 / 301 | 1.31 (0.94 - 1.83) | 46 / 59 | 1.29 (0.78 - 2.13) | | | |
| | | 757,550+ | 446 / 155 | 3.20 (2.23 - 4.59) | 58 / 18 | 3.73 (1.98 - 7.05) | | | |

Table S4. Odds ratios for developing SCCHN for SNP genotypes, at varying levels of lifetime alcohol consumption

| Gene | SNP | major/ minor alleles | lifetime ethanol consumption (ml) | Odds ratios in comparison to common referent group ^a | | | | Assessment of additive interaction between SNP and lifetime alcohol consumption |
|---------------|------------|----------------------------|--------------------------------------|---|---|--|---|---|
| | | | | homozygous major allele | | at least one minor allele | | |
| | | | | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | |
| <i>ALDH2</i> | rs4767939 | A/G | never drinkers | 76 / 174 | 1.00 (ref) | 41 / 106 | 0.88 (0.54 - 1.44) | 1.09 |
| | | | >0 to <134,699 | 120 / 289 | 0.65 (0.44 - 0.94) | 89 / 177 | 0.84 (0.56 - 1.27) | |
| | | | 134,699 to <757,550 | 194 / 218 | 1.21 (0.83 - 1.77) | 124 / 142 | 1.28 (0.85 - 1.93) | |
| | | | 757,550+ | 261 / 103 | 2.69 (1.78 - 4.07) | 243 / 70 | 3.66 (2.35 - 5.71) | |
| | rs2238151 | T/C | never drinkers | 50 / 95 | 1.00 (ref) | 67 / 184 | 0.70 (0.43 - 1.14) | 1.91 |
| | | | >0 to <134,699 | 75 / 173 | 0.54 (0.34 - 0.88) | 133 / 293 | 0.64 (0.41 - 0.99) | |
| | | | 134,699 to <757,550 | 97 / 138 | 0.81 (0.50 - 1.31) | 220 / 222 | 1.19 (0.76 - 1.86) | |
| | | | 757,550+ | 122 / 65 | 1.66 (0.99 - 2.80) | 381 / 106 | 3.27 (2.02 - 5.28) | |
| | rs7312055 | G/A | never drinkers | 111 / 254 | 1.00 (ref) | 6 / 26 | 0.41 (0.15 - 1.12) | 0.97 |
| | | | >0 to <134,699 | 201 / 427 | 0.74 (0.54 - 1.01) | 8 / 39 | 0.32 (0.13 - 0.78) | |
| | | | 134,699 to <757,550 | 282 / 328 | 1.19 (0.87 - 1.64) | 36 / 32 | 1.30 (0.69 - 2.46) | |
| | | | 757,550+ | 422 / 152 | 2.93 (2.06 - 4.17) | 83 / 21 | 3.31 (1.76 - 6.23) | |
| | rs2158029 | G/A | never drinkers | 116 / 270 | 1.00 (ref) | 1 / 10 | 0.28 (0.03 - 2.30) | 2.73 |
| | | | >0 to <134,699 | 207 / 443 | 0.77 (0.57 - 1.04) | 2 / 23 | 0.13 (0.03 - 0.60) | |
| | | | 134,699 to <757,550 | 302 / 349 | 1.25 (0.91 - 1.70) | 16 / 11 | 1.79 (0.72 - 4.45) | |
| | | | 757,550+ | 470 / 166 | 3.09 (2.19 - 4.35) | 35 / 7 | 5.10 (2.04 - 12.73) | |
| | rs16941667 | C/T | never drinkers | 98 / 233 | 1.00 (ref) | 19 / 47 | 1.06 (0.57 - 1.99) | 0.99 |
| | | | >0 to <134,699 | 163 / 390 | 0.70 (0.50 - 0.98) | 46 / 75 | 1.11 (0.69 - 1.79) | |
| | | | 134,699 to <757,550 | 259 / 294 | 1.29 (0.92 - 1.80) | 59 / 66 | 1.42 (0.89 - 2.29) | |
| | | | 757,550+ | 405 / 148 | 3.12 (2.17 - 4.49) | 100 / 25 | 4.17 (2.39 - 7.28) | |
| rs16941669 | T/G | never drinkers | 97 / 218 | 1.00 (ref) | 20 / 61 | 0.74 (0.41 - 1.34) | 1.45 | |
| | | >0 to <134,699 | 162 / 380 | 0.67 (0.48 - 0.94) | 47 / 86 | 0.88 (0.55 - 1.39) | | |
| | | 134,699 to <757,550 | 254 / 302 | 1.17 (0.84 - 1.64) | 64 / 58 | 1.46 (0.90 - 2.35) | | |
| | | 757,550+ | 415 / 149 | 2.90 (2.01 - 4.18) | 90 / 23 | 4.09 (2.30 - 7.26) | | |
| <i>CYP2E1</i> | rs3813865 | G/C | never drinkers | 108 / 259 | 1.00 (ref) | 9 / 20 | 1.24 (0.51 - 3.00) | 1.95 |
| | | | >0 to <134,699 | 196 / 432 | 0.76 (0.56 - 1.04) | 13 / 34 | 0.75 (0.37 - 1.54) | |
| | | | 134,699 to <757,550 | 289 / 324 | 1.32 (0.96 - 1.82) | 29 / 36 | 1.27 (0.69 - 2.31) | |
| | | | 757,550+ | 437 / 160 | 3.11 (2.19 - 4.42) | 68 / 13 | 5.31 (2.65 - 10.65) | |
| | rs3813867 | G/C | never drinkers | 110 / 263 | 1.00 (ref) | 7 / 16 | 0.94 (0.35 - 2.47) | 3.88 |
| | | | >0 to <134,699 | 197 / 441 | 0.74 (0.54 - 1.01) | 12 / 25 | 0.80 (0.37 - 1.76) | |
| | | | 134,699 to <757,550 | 301 / 329 | 1.35 (0.98 - 1.85) | 17 / 31 | 0.65 (0.33 - 1.28) | |
| | | | 757,550+ | 460 / 165 | 3.00 (2.11 - 4.25) | 44 / 8 | 6.82 (2.90 - 16.02) | |
| | rs8192772 | T/C | never drinkers | 99 / 237 | 1.00 (ref) | 18 / 43 | 1.07 (0.55 - 2.05) | 1.89 |
| | | | >0 to <134,699 | 179 / 394 | 0.75 (0.54 - 1.03) | 30 / 72 | 0.84 (0.50 - 1.41) | |
| | | | 134,699 to <757,550 | 264 / 294 | 1.30 (0.94 - 1.81) | 53 / 66 | 1.29 (0.80 - 2.08) | |
| | | | 757,550+ | 413 / 152 | 3.02 (2.11 - 4.33) | 92 / 21 | 4.97 (2.78 - 8.91) | |
| | rs915908 | G/A | never drinkers | 93 / 216 | 1.00 (ref) | 24 / 64 | 1.04 (0.59 - 1.85) | -1.29 |
| | | | >0 to <134,699 | 152 / 352 | 0.74 (0.53 - 1.04) | 57 / 114 | 0.80 (0.52 - 1.24) | |
| | | | 134,699 to <757,550 | 254 / 277 | 1.35 (0.96 - 1.91) | 64 / 83 | 1.12 (0.71 - 1.77) | |
| | | | 757,550+ | 417 / 131 | 3.56 (2.45 - 5.17) | 88 / 42 | 2.31 (1.39 - 3.82) | |
| | rs915909 | C/T | never drinkers | 115 / 270 | 1.00 (ref) | 2 / 9 | 0.62 (0.12 - 3.24) | 0.80 |
| | | | >0 to <134,699 | 206 / 449 | 0.76 (0.56 - 1.03) | 3 / 15 | 0.35 (0.09 - 1.30) | |
| | | | 134,699 to <757,550 | 302 / 348 | 1.26 (0.92 - 1.71) | 16 / 12 | 1.74 (0.71 - 4.24) | |
| | | | 757,550+ | 468 / 163 | 3.15 (2.23 - 4.44) | 36 / 9 | 3.56 (1.56 - 8.17) | |
| rs7092584 | C/T | never drinkers | 92 / 223 | 1.00 (ref) | 25 / 57 | 1.17 (0.66 - 2.07) | 1.06 | |
| | | >0 to <134,699 | 167 / 375 | 0.75 (0.54 - 1.05) | 42 / 91 | 0.89 (0.56 - 1.44) | | |
| | | 134,699 to <757,550 | 252 / 279 | 1.36 (0.97 - 1.92) | 65 / 80 | 1.24 (0.79 - 1.94) | | |
| | | 757,550+ | 384 / 142 | 3.10 (2.14 - 4.49) | 120 / 31 | 4.33 (2.58 - 7.26) | | |
| rs743535 | C/T | never drinkers | 95 / 223 | 1.00 (ref) | 22 / 53 | 0.93 (0.51 - 1.69) | 2.26 | |
| | | >0 to <134,699 | 169 / 386 | 0.70 (0.50 - 0.98) | 40 / 78 | 0.91 (0.56 - 1.49) | | |
| | | 134,699 to <757,550 | 252 / 279 | 1.29 (0.92 - 1.81) | 60 / 80 | 1.04 (0.66 - 1.65) | | |
| | | 757,550+ | 386 / 147 | 2.79 (1.93 - 4.04) | 116 / 26 | 4.98 (2.89 - 8.59) | | |
| rs2249695 | C/T | never drinkers | 73 / 136 | 1.00 (ref) | 44 / 144 | 0.56 (0.35 - 0.91) | 1.21 | |
| | | >0 to <134,699 | 127 / 253 | 0.63 (0.43 - 0.94) | 82 / 213 | 0.52 (0.34 - 0.80) | | |
| | | 134,699 to <757,550 | 160 / 178 | 1.06 (0.71 - 1.59) | 158 / 181 | 0.95 (0.63 - 1.43) | | |
| | | 757,550+ | 218 / 92 | 2.12 (1.37 - 3.28) | 286 / 81 | 2.89 (1.83 - 4.55) | | |
| rs28969387 | A/T | never drinkers | 115 / 275 | 1.00 (ref) | 2 / 5 | 1.03 (0.16 - 6.58) | 0.05 | |
| | | >0 to <134,699 | 206 / 464 | 0.75 (0.55 - 1.01) | 3 / 2 | 2.25 (0.33 - 15.13) | | |
| | | 134,699 to <757,550 | 311 / 353 | 1.30 (0.95 - 1.77) | 7 / 7 | 1.06 (0.33 - 3.38) | | |
| | | 757,550+ | 493 / 170 | 3.22 (2.29 - 4.54) | 12 / 3 | 3.30 (0.88 - 12.40) | | |

Table S4. Odds ratios for developing SCCHN for SNP genotypes, at varying levels of lifetime alcohol consumption

| Gene | SNP | major/ minor alleles | lifetime ethanol consumption (ml) | Odds ratios in comparison to common referent group ^a | | | | Assessment of additive interaction between SNP and lifetime alcohol consumption |
|--|------------|----------------------------|--------------------------------------|---|---|--|---|---|
| | | | | homozygous major allele | | at least one minor allele | | |
| | | | | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | |
| | rs11101812 | T/C | never drinkers | 115 / 275 | 1.00 (ref) | 1 / 5 | 0.41 (0.04 - 3.90) | 19.34 |
| | | | >0 to <134,699 | 205 / 463 | 0.74 (0.55 - 1.00) | 4 / 3 | 2.70 (0.52 - 14.05) | |
| | | | 134,699 to <757,550 | 312 / 348 | 1.31 (0.96 - 1.79) | 5 / 10 | 0.67 (0.20 - 2.23) | |
| | | | 757,550+ | 489 / 172 | 3.10 (2.20 - 4.36) | 15 / 1 | 21.85 (2.60 - 183.30) | |
| OXIDATIVE STRESS METABOLISM GENES | | | | | | | | |
| CAT | rs1049982 | C/T | never drinkers | 60 / 137 | 1.00 (ref) | 57 / 140 | 1.10 (0.69 - 1.77) | -0.15 |
| | | | >0 to <134,699 | 95 / 216 | 0.75 (0.49 - 1.15) | 114 / 247 | 0.81 (0.54 - 1.22) | |
| | | | 134,699 to <757,550 | 135 / 150 | 1.41 (0.92 - 2.16) | 183 / 209 | 1.29 (0.85 - 1.94) | |
| | | | 757,550+ | 200 / 71 | 3.33 (2.10 - 5.28) | 303 / 102 | 3.28 (2.12 - 5.07) | |
| GPX1 | rs8179172 | T/A | never drinkers | 116 / 270 | 1.00 (ref) | 1 / 10 | 0.14 (0.02 - 1.25) | 0.16 |
| | | | >0 to <134,699 | 205 / 450 | 0.73 (0.54 - 0.99) | 4 / 16 | 0.44 (0.13 - 1.46) | |
| | | | 134,699 to <757,550 | 305 / 349 | 1.21 (0.88 - 1.65) | 13 / 11 | 2.15 (0.85 - 5.44) | |
| | | | 757,550+ | 475 / 160 | 3.10 (2.19 - 4.38) | 30 / 13 | 2.40 (1.10 - 5.27) | |
| | rs1800668 | C/T | never drinkers | 56 / 146 | 1.00 (ref) | 61 / 132 | 1.16 (0.73 - 1.85) | 0.37 |
| | | | >0 to <134,699 | 97 / 222 | 0.76 (0.50 - 1.16) | 112 / 243 | 0.85 (0.56 - 1.29) | |
| | | | 134,699 to <757,550 | 178 / 185 | 1.43 (0.94 - 2.16) | 140 / 175 | 1.33 (0.87 - 2.04) | |
| | | | 757,550+ | 272 / 95 | 3.23 (2.07 - 5.05) | 233 / 77 | 3.75 (2.37 - 5.95) | |
| | rs3811699 | A/G | never drinkers | 55 / 142 | 1.00 (ref) | 62 / 138 | 1.14 (0.72 - 1.82) | 0.37 |
| | | | >0 to <134,699 | 95 / 216 | 0.76 (0.50 - 1.17) | 114 / 250 | 0.85 (0.56 - 1.29) | |
| | | | 134,699 to <757,550 | 171 / 176 | 1.45 (0.95 - 2.21) | 147 / 184 | 1.31 (0.86 - 2.01) | |
| | | | 757,550+ | 258 / 91 | 3.21 (2.04 - 5.04) | 247 / 82 | 3.71 (2.35 - 5.88) | |
| | rs3448 | C/T | never drinkers | 60 / 158 | 1.00 (ref) | 57 / 122 | 1.27 (0.80 - 2.03) | 0.32 |
| | | | >0 to <134,699 | 116 / 256 | 0.84 (0.56 - 1.26) | 93 / 210 | 0.84 (0.55 - 1.28) | |
| | | | 134,699 to <757,550 | 157 / 192 | 1.35 (0.89 - 2.03) | 161 / 168 | 1.55 (1.03 - 2.34) | |
| | | | 757,550+ | 273 / 105 | 3.34 (2.17 - 5.14) | 232 / 68 | 3.93 (2.49 - 6.22) | |
| GPX2 | rs11623705 | G/T | never drinkers | 86 / 232 | 1.00 (ref) | 31 / 48 | 1.58 (0.90 - 2.77) | -0.76 |
| | | | >0 to <134,699 | 166 / 367 | 0.83 (0.59 - 1.17) | 43 / 99 | 0.83 (0.52 - 1.32) | |
| | | | 134,699 to <757,550 | 263 / 292 | 1.47 (1.05 - 2.08) | 55 / 68 | 1.24 (0.77 - 2.01) | |
| | | | 757,550+ | 421 / 142 | 3.60 (2.47 - 5.24) | 84 / 31 | 3.41 (2.00 - 5.84) | |
| | rs2412065 | G/C | never drinkers | 67 / 158 | 1.00 (ref) | 50 / 122 | 1.00 (0.62 - 1.60) | 0.06 |
| | | | >0 to <134,699 | 126 / 251 | 0.85 (0.57 - 1.26) | 83 / 215 | 0.65 (0.43 - 0.99) | |
| | | | 134,699 to <757,550 | 186 / 204 | 1.32 (0.89 - 1.96) | 132 / 156 | 1.25 (0.82 - 1.90) | |
| | | | 757,550+ | 278 / 99 | 3.18 (2.07 - 4.88) | 227 / 74 | 3.24 (2.06 - 5.10) | |
| | rs2737844 | C/T | never drinkers | 58 / 117 | 1.00 (ref) | 59 / 161 | 0.76 (0.48 - 1.22) | 0.03 |
| | | | >0 to <134,699 | 100 / 173 | 0.84 (0.54 - 1.30) | 108 / 292 | 0.54 (0.35 - 0.82) | |
| | | | 134,699 to <757,550 | 129 / 147 | 1.09 (0.70 - 1.70) | 188 / 211 | 1.12 (0.73 - 1.72) | |
| | | | 757,550+ | 180 / 62 | 2.84 (1.73 - 4.65) | 322 / 111 | 2.63 (1.66 - 4.16) | |
| GPX4 | rs757229 | G/C | never drinkers | 35 / 83 | 1.00 (ref) | 82 / 197 | 0.92 (0.55 - 1.52) | 0.09 |
| | | | >0 to <134,699 | 58 / 114 | 0.74 (0.43 - 1.28) | 150 / 352 | 0.69 (0.43 - 1.12) | |
| | | | 134,699 to <757,550 | 67 / 109 | 0.90 (0.52 - 1.56) | 251 / 251 | 1.34 (0.83 - 2.17) | |
| | | | 757,550+ | 116 / 40 | 3.05 (1.68 - 5.54) | 389 / 132 | 3.05 (1.84 - 5.06) | |
| SOD1 | rs11910115 | A/C | never drinkers | 116 / 271 | 1.00 (ref) | 1 / 9 | 0.31 (0.03 - 2.91) | 2.22 |
| | | | >0 to <134,699 | 205 / 441 | 0.76 (0.56 - 1.04) | 4 / 25 | 0.27 (0.08 - 0.92) | |
| | | | 134,699 to <757,550 | 299 / 340 | 1.27 (0.93 - 1.73) | 19 / 20 | 1.33 (0.62 - 2.85) | |
| | | | 757,550+ | 469 / 165 | 3.09 (2.19 - 4.35) | 36 / 8 | 4.63 (1.94 - 11.06) | |
| | rs4998557 | G/A | never drinkers | 85 / 202 | 1.00 (ref) | 32 / 78 | 0.91 (0.54 - 1.53) | 1.51 |
| | | | >0 to <134,699 | 157 / 341 | 0.75 (0.53 - 1.07) | 52 / 122 | 0.69 (0.44 - 1.09) | |
| | | | 134,699 to <757,550 | 208 / 258 | 1.15 (0.80 - 1.64) | 110 / 102 | 1.53 (1.00 - 2.34) | |
| | | | 757,550+ | 328 / 124 | 2.76 (1.87 - 4.08) | 176 / 49 | 4.18 (2.60 - 6.72) | |
| | rs10432782 | T/G | never drinkers | 86 / 208 | 1.00 (ref) | 31 / 72 | 0.95 (0.56 - 1.61) | 1.15 |
| | | | >0 to <134,699 | 162 / 351 | 0.77 (0.55 - 1.09) | 47 / 115 | 0.65 (0.41 - 1.04) | |
| | | | 134,699 to <757,550 | 222 / 267 | 1.20 (0.84 - 1.70) | 96 / 93 | 1.48 (0.96 - 2.27) | |
| | | | 757,550+ | 350 / 128 | 2.91 (1.98 - 4.27) | 154 / 44 | 4.00 (2.48 - 6.46) | |
| | rs2070424 | A/G | never drinkers | 95 / 238 | 1.00 (ref) | 22 / 42 | 1.17 (0.63 - 2.17) | 0.01 |
| | | | >0 to <134,699 | 181 / 402 | 0.78 (0.56 - 1.08) | 28 / 64 | 0.76 (0.44 - 1.30) | |
| | | | 134,699 to <757,550 | 260 / 302 | 1.31 (0.94 - 1.83) | 58 / 58 | 1.41 (0.87 - 2.30) | |
| | | | 757,550+ | 407 / 139 | 3.27 (2.27 - 4.72) | 98 / 34 | 3.45 (2.06 - 5.79) | |

Table S4. Odds ratios for developing SCCHN for SNP genotypes, at varying levels of lifetime alcohol consumption

| Gene | SNP | major/ minor alleles | lifetime ethanol consumption (ml) | Odds ratios in comparison to common referent group ^a | | | | Assessment of additive interaction between SNP and lifetime alcohol consumption | |
|-------------|-----------|----------------------------|--------------------------------------|---|---|--|---|---|---|
| | | | | homozygous major allele | | at least one minor allele | | | ICR ^d (bold if statistically significant after Bonferroni correction) |
| | | | | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | n (cases/ controls) ^b | Adjusted odds ratio ^c (95% CI) | | |
| | rs1041740 | C/T | never drinkers | 56 / 156 | 1.00 (ref) | 61 / 124 | 1.43 (0.89 - 2.28) | -1.09 | |
| | | | >0 to <134,699 | 104 / 239 | 0.89 (0.59 - 1.35) | 105 / 227 | 0.90 (0.59 - 1.37) | | |
| | | | 134,699 to <757,550 | 193 / 191 | 1.71 (1.13 - 2.59) | 125 / 169 | 1.33 (0.86 - 2.05) | | |
| | | | 757,550+ | 291 / 94 | 4.15 (2.65 - 6.49) | 214 / 79 | 3.49 (2.21 - 5.52) | | |
| SOD2 | rs4342445 | G/A | never drinkers | 55 / 177 | 1.00 (ref) | 62 / 103 | 2.14 (1.33 - 3.44) | 0.27 | |
| | | | >0 to <134,699 | 135 / 296 | 1.10 (0.74 - 1.63) | 74 / 170 | 0.96 (0.62 - 1.49) | | |
| | | | 134,699 to <757,550 | 184 / 227 | 1.61 (1.07 - 2.40) | 134 / 133 | 2.11 (1.38 - 3.24) | | |
| | | | 757,550+ | 331 / 123 | 4.04 (2.65 - 6.17) | 174 / 50 | 5.45 (3.33 - 8.90) | | |
| | rs2842980 | A/T | never drinkers | 74 / 169 | 1.00 (ref) | 43 / 111 | 0.87 (0.54 - 1.41) | 0.89 | |
| | | | >0 to <134,699 | 121 / 282 | 0.69 (0.47 - 1.01) | 88 / 184 | 0.76 (0.50 - 1.14) | | |
| | | | 134,699 to <757,550 | 196 / 222 | 1.27 (0.87 - 1.87) | 122 / 138 | 1.15 (0.76 - 1.74) | | |
| | | | 757,550+ | 291 / 108 | 2.77 (1.84 - 4.19) | 214 / 65 | 3.54 (2.25 - 5.56) | | |
| | rs8031 | T/A | never drinkers | 38 / 75 | 1.00 (ref) | 79 / 205 | 0.84 (0.51 - 1.40) | -0.97 | |
| | | | >0 to <134,699 | 59 / 139 | 0.58 (0.34 - 0.98) | 150 / 327 | 0.71 (0.44 - 1.15) | | |
| | | | 134,699 to <757,550 | 101 / 113 | 1.14 (0.68 - 1.93) | 217 / 247 | 1.15 (0.71 - 1.86) | | |
| | | | 757,550+ | 155 / 40 | 3.73 (2.08 - 6.71) | 350 / 133 | 2.61 (1.59 - 4.30) | | |
| rs5746134 | C/T | never drinkers | 113 / 264 | 1.00 (ref) | 4 / 16 | 0.55 (0.16 - 1.87) | 4.29 | | |
| | | >0 to <134,699 | 201 / 447 | 0.73 (0.54 - 0.99) | 8 / 19 | 0.80 (0.31 - 2.08) | | | |
| | | 134,699 to <757,550 | 299 / 342 | 1.25 (0.91 - 1.71) | 19 / 18 | 1.47 (0.68 - 3.18) | | | |
| | | 757,550+ | 446 / 163 | 2.96 (2.09 - 4.19) | 59 / 10 | 6.80 (3.03 - 15.26) | | | |
| rs2758331 | C/A | never drinkers | 38 / 82 | 1.00 (ref) | 79 / 198 | 0.92 (0.55 - 1.52) | -1.51 | | |
| | | >0 to <134,699 | 60 / 151 | 0.58 (0.34 - 0.98) | 149 / 315 | 0.78 (0.48 - 1.24) | | | |
| | | 134,699 to <757,550 | 108 / 127 | 1.14 (0.68 - 1.90) | 210 / 233 | 1.26 (0.78 - 2.03) | | | |
| | | 757,550+ | 179 / 44 | 4.23 (2.39 - 7.50) | 326 / 129 | 2.64 (1.61 - 4.34) | | | |

^a Conditional logistic regression models conditioned on sex, race, and age category (age in years: 20-49, 50-54, 55-59, 60-64, 65-69, 70-74, 75-80), and adjusted for continuous duration of smoking in years (rounded to nearest whole year)

^b Cases and controls do not sum to 1227 and 1325, respectively, because (1) 4 cases and 3 controls are missing information on duration of cigarette smoking, (2) 77 cases and 45 controls are missing information on lifetime alcohol consumption, and (3) a few subjects are missing genotype information for some SNPs

^c Odds ratios (ORs) for each SNP*drinking category were calculated from conditional logistic regression models including one SNP coded for dominant genetic model, categorized lifetime ethanol consumption, conditioned on sex, race, and age category, and adjusted for continuous smoking duration rounded to whole years.

^d ICRs were calculated using cancer odds ratios of subjects in three categories: (1) the highest drinking category and no minor allele (i.e., those singly exposed to drinking only -- OR₀₁); (2) never-drinkers with at least one minor allele (i.e., those singly exposed to only the variant allele -- OR₁₀); and (3) subjects in the highest drinking category and at least one minor allele (i.e., those doubly exposed to both alcohol and the variant allele -- OR₁₁), compared to never-drinkers homozygous for the major allele (i.e., the referent group that was not exposed to either the variant allele or to drinking -- OR₀₀ = 1.0). ICRs statistically different from 0 indicate departure from additive interaction. p-values for each ICR (not shown) were Bonferroni-adjusted for 64 statistical tests.

Table S5. SNP effects (dominant genetic model) on odds of developing SCCHN, overall and by race

(Odds ratios (ORs) for carriers of the minor allele compared to homozygotes for the major allele)

| Gene | SNP ¹ | major/ minor alleles | Genotype | ALL SUBJECTS COMBINED | | | AFRICAN-AMERICANS ONLY | | | EUROPEAN-AMERICANS ONLY | | |
|---------------------------------|------------------|-------------------------|-----------|-----------------------------------|--------------------------------------|---|-----------------------------------|--------------------------------------|---|-----------------------------------|--------------------------------------|---|
| | | | | n cases/ controls ² | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) | n cases/ controls ⁴ | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) | n cases/ controls ⁵ | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) |
| ALCOHOL METABOLISM GENES | | | | | | | | | | | | |
| ADH1B | rs12507573 | C/A | CC | 371/386 | | | 99/83 | | | 272/303 | | |
| | | | AA or CA | 852/935 | 0.96 (0.87-1.06) | 1.00 | 204/167 | 1.05 (0.67-1.64) | 1.00 | 648/768 | 0.89 (0.72-1.11) | 1.00 |
| | rs1042026 | A/G | AA | 712/744 | | | 246/210 | | | 466/534 | | |
| | | | GG or AG | 511/578 | 1.00 (0.91-1.10) | 1.00 | 57/41 | 0.90 (0.51-1.58) | 1.00 | 454/537 | 0.99 (0.82-1.20) | 1.00 |
| | rs7673353 | C/T | CC | 1124/1239 | | | 207/174 | | | 917/1065 | | |
| | | | TT or CT | 99/83 | 1.01 (0.83-1.23) | 1.00 | 96/77 | 1.16 (0.73-1.85) | 1.00 | 3/6 | 0.35 (0.08-1.48) | 1.00 |
| | rs17028834 | T/C | TT | 1138/1265 | | | 222/196 | | | 916/1069 | | |
| | | | CC or TC | 85/57 | 1.28 (1.03-1.59) | 1.00 | 81/55 | 1.92 (1.14-3.21) | 0.88 | 4/2 | 3.02 (0.48-18.98) | 1.00 |
| | rs1693457 | T/C | TT | 803/860 | | | 208/148 | | | 595/712 | | |
| | | | CC or TC | 419/461 | 1.01 (0.92-1.11) | 1.00 | 95/102 | 0.64 (0.41-0.99) | 1.00 | 324/359 | 1.15 (0.94-1.41) | 1.00 |
| tag SNP, intron 4 | rs1229984 | G/A | GG | 1192/1243 | | | 302/246 | | | 890/997 | | |
| | | | AA or GA | 31/79 | 0.72 (0.57-0.91) | 0.35 | 1/5 | 0.12 (0.01-1.24) | 1.00 | 30/74 | 0.54 (0.34-0.87) | 0.69 |
| rs1159918 | G/T | GG | 402/468 | | | 28/18 | | | 374/450 | | | |
| | | TT or GT | 821/854 | 1.07 (0.97-1.18) | 1.00 | 275/233 | 1.39 (0.66-2.92) | 1.00 | 546/621 | 1.14 (0.94-1.39) | 1.00 | |
| rs1229982 | G/T | GG | 699/775 | | | 89/75 | | | 610/700 | | | |
| | | TT or GT | 524/547 | 1.00 (0.91-1.10) | 1.00 | 214/176 | 1.33 (0.84-2.10) | 1.00 | 310/371 | 0.96 (0.78-1.17) | 1.00 | |
| ADH1C | rs2298753 | T/C | TT | 1002/1117 | | | 283/237 | | | 719/880 | | |
| | | | CC or TC | 221/205 | 1.12 (1.00-1.27) | 1.00 | 20/14 | 1.13 (0.46-2.77) | 1.00 | 201/191 | 1.28 (1.00-1.63) | 1.00 |
| | rs1614972 | C/T | CC | 539/585 | | | 94/55 | | | 445/530 | | |
| | | | TT or CT | 684/737 | 1.00 (0.92-1.09) | 1.00 | 209/196 | 0.74 (0.46-1.20) | 1.00 | 475/541 | 1.07 (0.88-1.29) | 1.00 |
| | rs1391088 | C/A | CC | 1045/1119 | | | 246/213 | | | 799/906 | | |
| | | | AA or CA | 178/203 | 1.01 (0.89-1.14) | 1.00 | 57/38 | 1.55 (0.87-2.77) | 1.00 | 121/165 | 0.91 (0.69-1.21) | 1.00 |
| rs1693482 | C/T | CC | 527/585 | | | 203/183 | | | 324/402 | | | |
| | | TT or CT | 694/735 | 1.05 (0.95-1.15) | 1.00 | 100/68 | 1.32 (0.83-2.12) | 1.00 | 594/667 | 1.05 (0.86-1.29) | 1.00 | |
| rs1631460 | C/G | CC | 519/574 | | | 202/182 | | | 317/392 | | | |
| | | GG or CG | 703/748 | 1.04 (0.95-1.14) | 1.00 | 101/69 | 1.31 (0.82-2.09) | 1.00 | 602/679 | 1.05 (0.86-1.28) | 1.00 | |
| rs11936869 | C/G | CC | 611/645 | | | 109/72 | | | 502/573 | | | |
| | | GG or CG | 612/676 | 0.98 (0.90-1.07) | 1.00 | 194/178 | 0.76 (0.48-1.19) | 1.00 | 418/498 | 1.00 (0.83-1.22) | 1.00 | |
| ADH4 | rs29001227 | A/T | AA | 1119/1253 | | | 204/184 | | | 915/1069 | | |
| | | | TT or AT | 104/69 | 1.25 (1.02-1.53) | 1.00 | 99/67 | 1.70 (1.05-2.73) | 1.00 | 5/2 | 3.42 (0.58-20.22) | 1.00 |
| | rs1126672 | C/T | CC | 720/758 | | | 237/203 | | | 483/555 | | |
| | | | TT or CT | 503/564 | 1.01 (0.92-1.10) | 1.00 | 66/48 | 1.45 (0.85-2.49) | 1.00 | 437/516 | 0.98 (0.81-1.19) | 1.00 |
| | rs4699710 | T/C | TT | 640/684 | | | 200/168 | | | 440/516 | | |
| | | | CC or TC | 583/638 | 1.01 (0.93-1.11) | 1.00 | 103/83 | 1.10 (0.70-1.74) | 1.00 | 480/555 | 1.02 (0.84-1.24) | 1.00 |
| | rs10017466 | T/C | TT | 556/632 | | | 120/118 | | | 436/514 | | |
| | | | CC or TC | 666/690 | 1.05 (0.97-1.15) | 1.00 | 182/133 | 1.65 (1.07-2.55) | 1.00 | 484/557 | 1.04 (0.85-1.26) | 1.00 |
| tag SNP, intron 5 | rs1800759 | C/A | CC | 357/412 | | | 21/16 | | | 336/396 | | |
| | | | AA or CA | 866/910 | 1.02 (0.92-1.12) | 1.00 | 282/235 | 1.57 (0.70-3.52) | 1.00 | 584/675 | 1.02 (0.83-1.24) | 1.00 |
| rs1800761 | G/A | GG | 750/830 | | | 153/129 | | | 597/701 | | | |
| | | AA or GA | 473/492 | 1.02 (0.93-1.12) | 1.00 | 150/122 | 0.99 (0.64-1.51) | 1.00 | 323/370 | 1.06 (0.86-1.29) | 1.00 | |
| rs3762894 | T/C | TT | 839/906 | | | 190/161 | | | 649/745 | | | |
| | | CC or TC | 384/416 | 0.98 (0.89-1.08) | 1.00 | 113/90 | 1.06 (0.68-1.66) | 1.00 | 271/326 | 0.94 (0.76-1.16) | 1.00 | |
| ADH7 | rs284787 | C/T | CC | 739/817 | | | 219/188 | | | 520/629 | | |
| | | | TT or CT | 484/505 | 1.06 (0.96-1.16) | 1.00 | 84/63 | 0.86 (0.54-1.39) | 1.00 | 400/442 | 1.14 (0.94-1.39) | 1.00 |
| | rs894369 | C/G | CC | 832/849 | | | 219/176 | | | 613/673 | | |
| | | | GG or CG | 391/472 | 0.92 (0.84-1.01) | 1.00 | 84/74 | 1.09 (0.67-1.76) | 1.00 | 307/398 | 0.82 (0.67-1.00) | 1.00 |
| | rs17588403 | T/A | TT | 837/887 | | | 233/190 | | | 604/697 | | |
| | | | AA or TA | 386/435 | 0.96 (0.87-1.05) | 1.00 | 70/61 | 0.78 (0.48-1.29) | 1.00 | 316/374 | 0.95 (0.77-1.16) | 1.00 |
| | rs1154454 | T/C | TT | 756/808 | | | 110/75 | | | 646/733 | | |
| | | | CC or TC | 467/514 | 0.94 (0.86-1.04) | 1.00 | 193/176 | 0.68 (0.43-1.08) | 1.00 | 274/338 | 0.94 (0.76-1.16) | 1.00 |
| | rs1154456 | T/C | TT | 577/629 | | | 209/176 | | | 368/453 | | |
| | | | CC or TC | 646/693 | 1.03 (0.94-1.13) | 1.00 | 94/75 | 0.90 (0.57-1.43) | 1.00 | 552/618 | 1.10 (0.90-1.34) | 1.00 |
| | rs1154460 | G/A | GG | 329/377 | | | 70/65 | | | 259/312 | | |
| | | | AA or GA | 891/944 | 1.04 (0.95-1.15) | 1.00 | 233/185 | 1.11 (0.67-1.82) | 1.00 | 658/759 | 1.08 (0.87-1.34) | 1.00 |
| rs971074 | G/A | GG | 943/1027 | | | 195/176 | | | 748/851 | | | |
| | | AA or GA | 280/295 | 1.01 (0.91-1.12) | 1.00 | 108/75 | 1.21 (0.77-1.90) | 1.00 | 172/220 | 0.94 (0.74-1.20) | 1.00 | |
| rs1573496 | C/G | CC | 1056/1117 | | | 296/245 | | | 760/872 | | | |
| | | GG or CG | 166/205 | 0.99 (0.87-1.12) | 1.00 | 7/6 | 0.78 (0.20-3.03) | 1.00 | 159/199 | 0.97 (0.75-1.24) | 1.00 | |
| ALDH2 | rs4767939 | A/G | AA | 694/806 | | | 107/88 | | | 587/718 | | |
| | | | GG or AG | 528/516 | 1.08 (0.98-1.18) | 1.00 | 196/163 | 1.07 (0.69-1.67) | 1.00 | 332/353 | 1.19 (0.97-1.45) | 1.00 |
| | rs2238151 | T/C | TT | 364/482 | | | 2/1 | | | 362/481 | | |
| | | | CC or TC | 855/837 | 1.13 (1.03-1.25) | 0.84 | 301/250 | 0.91 (0.06-14.48) | 1.00 | 554/587 | 1.28 (1.05-1.56) | 0.88 |

Table S5. SNP effects (dominant genetic model) on odds of developing SCCHN, overall and by race

(Odds ratios (ORs) for carriers of the minor allele compared to homozygotes for the major allele)

| Gene | SNP ¹ | major/ minor alleles | Genotype | ALL SUBJECTS COMBINED | | | AFRICAN-AMERICANS ONLY | | | EUROPEAN-AMERICANS ONLY | | |
|--|------------------|-------------------------|-----------|-----------------------------------|--------------------------------------|---|-----------------------------------|--------------------------------------|---|-----------------------------------|--------------------------------------|---|
| | | | | n cases/ controls ² | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) | n cases/ controls ⁴ | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) | n cases/ controls ⁵ | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) |
| CYP2E1 | rs7312055 | G/A | GG | 1081/1201 | | | 166/138 | | | 915/1063 | | |
| | | | AA or GA | 142/121 | 0.93 (0.78-1.12) | 1.00 | 137/113 | 0.82 (0.53-1.26) | 1.00 | 5/8 | 0.74 (0.22-2.54) | 1.00 |
| | rs2158029 | G/A | GG | 1166/1269 | | | 252/209 | | | 914/1060 | | |
| | | | AA or GA | 57/53 | 0.95 (0.75-1.19) | 1.00 | 51/42 | 0.95 (0.53-1.69) | 1.00 | 6/11 | 0.55 (0.18-1.66) | 1.00 |
| | rs16941667 | C/T | CC | 985/1097 | | | 224/188 | | | 761/909 | | |
| | | | TT or CT | 238/224 | 1.10 (0.98-1.23) | 1.00 | 79/62 | 1.21 (0.75-1.97) | 1.00 | 159/162 | 1.21 (0.93-1.58) | 1.00 |
| | rs16941669 | T/G | TT | 984/1084 | | | 251/214 | | | 733/870 | | |
| | | | GG or TG | 239/236 | 1.06 (0.95-1.19) | 1.00 | 52/36 | 1.52 (0.83-2.77) | 1.00 | 187/200 | 1.08 (0.85-1.38) | 1.00 |
| | rs3813865 | G/C | GG | 1098/1215 | | | 223/195 | | | 875/1020 | | |
| | | | CC or GC | 125/106 | 1.08 (0.92-1.27) | 1.00 | 80/56 | 1.22 (0.75-2.00) | 1.00 | 45/50 | 1.15 (0.73-1.82) | 1.00 |
| | rs3813867 | G/C | GG | 1139/1237 | | | 271/224 | | | 868/1013 | | |
| | | | CC or GC | 83/84 | 0.97 (0.82-1.16) | 1.00 | 31/26 | 1.13 (0.56-2.29) | 1.00 | 52/58 | 0.92 (0.60-1.39) | 1.00 |
| | rs8192772 | T/C | TT | 1023/1114 | | | 235/196 | | | 788/918 | | |
| | | | CC or TC | 199/208 | 1.07 (0.95-1.21) | 1.00 | 67/55 | 1.30 (0.77-2.19) | 1.00 | 132/153 | 1.14 (0.86-1.50) | 1.00 |
| | rs915908 | G/A | GG | 974/1006 | | | 286/228 | | | 688/778 | | |
| | | | AA or GA | 249/316 | 0.93 (0.84-1.04) | 1.00 | 17/23 | 0.55 (0.24-1.25) | 1.00 | 232/293 | 0.90 (0.72-1.12) | 1.00 |
| | rs915909 | C/T | CC | 1164/1271 | | | 245/204 | | | 919/1067 | | |
| | | | TT or CT | 58/46 | 1.00 (0.79-1.27) | 1.00 | 57/42 | 1.08 (0.62-1.88) | 1.00 | 1/4 | 0.22 (0.02-2.17) | 1.00 |
| rs7092584 | C/T | CC | 959/1052 | | | 217/187 | | | 742/865 | | | |
| | | TT or CT | 262/269 | 1.05 (0.95-1.17) | 1.00 | 84/63 | 1.45 (0.89-2.37) | 1.00 | 178/206 | 1.05 (0.82-1.35) | 1.00 | |
| rs743535 | C/T | CC | 966/1071 | | | 210/188 | | | 756/883 | | | |
| | | TT or CT | 248/243 | 1.06 (0.95-1.19) | 1.00 | 89/61 | 1.46 (0.90-2.37) | 1.00 | 159/182 | 1.05 (0.81-1.36) | 1.00 | |
| rs2249695 | C/T | CC | 622/682 | | | 28/23 | | | 594/659 | | | |
| | | TT or CT | 600/639 | 0.93 (0.85-1.03) | 1.00 | 275/227 | 1.16 (0.55-2.44) | 1.00 | 325/412 | 0.86 (0.70-1.05) | 1.00 | |
| rs28969387 | A/T | AA | 1198/1302 | | | 280/231 | | | 918/1071 | | | |
| | | TT or AT | 25/20 | 0.98 (0.70-1.37) | 1.00 | 23/20 | 0.82 (0.37-1.78) | 1.00 | 2/0 | N/A | 1.00 | |
| rs11101812 | T/C | TT | 1194/1300 | | | 277/230 | | | 917/1070 | | | |
| | | CC or TC | 26/20 | 1.11 (0.79-1.56) | 1.00 | 24/19 | 1.19 (0.52-2.72) | 1.00 | 2/1 | 2.97 (0.20-44.70) | 1.00 | |
| OXIDATIVE STRESS METABOLISM GENES | | | | | | | | | | | | |
| CAT | rs1049982 | C/T | CC | 521/589 | | | 92/76 | | | 429/513 | | |
| | | | TT or CT | 700/726 | 1.02 (0.93-1.11) | 1.00 | 211/173 | 0.93 (0.59-1.49) | 1.00 | 489/553 | 1.05 (0.86-1.27) | 1.00 |
| GPX1 | rs8179172 | T/A | TT | 1172/1271 | | | 253/202 | | | 919/1069 | | |
| | | | AA or TA | 51/51 | 0.92 (0.73-1.17) | 1.00 | 50/49 | 0.88 (0.51-1.52) | 1.00 | 1/2 | 0.27 (0.02-3.00) | 1.00 |
| | rs1800668 | C/T | CC | 635/672 | | | 173/140 | | | 462/532 | | |
| | | | TT or CT | 588/646 | 1.04 (0.95-1.14) | 1.00 | 130/110 | 1.13 (0.74-1.74) | 1.00 | 458/536 | 1.06 (0.88-1.29) | 1.00 |
| rs3811699 | A/G | AA | 609/649 | | | 147/118 | | | 462/531 | | | |
| | | GG or AG | 614/673 | 1.04 (0.95-1.13) | 1.00 | 156/133 | 1.19 (0.78-1.82) | 1.00 | 458/540 | 1.05 (0.87-1.28) | 1.00 | |
| rs3448 | C/T | CC | 648/729 | | | 160/138 | | | 488/591 | | | |
| | | TT or CT | 575/593 | 1.03 (0.95-1.13) | 1.00 | 143/113 | 1.07 (0.70-1.64) | 1.00 | 432/480 | 1.06 (0.87-1.29) | 1.00 | |
| GPX2 | rs11623705 | G/T | GG | 996/1069 | | | 288/232 | | | 708/837 | | |
| | | | TT or GT | 227/253 | 1.00 (0.89-1.12) | 1.00 | 15/19 | 0.70 (0.30-1.64) | 1.00 | 212/234 | 1.04 (0.83-1.31) | 1.00 |
| | rs2412065 | G/C | GG | 694/733 | | | 101/85 | | | 593/648 | | |
| | | | CC or GC | 529/589 | 0.94 (0.86-1.03) | 1.00 | 202/166 | 1.15 (0.73-1.80) | 1.00 | 327/423 | 0.85 (0.69-1.03) | 1.00 |
| rs2737844 | C/T | CC | 493/511 | | | 13/14 | | | 480/497 | | | |
| tag SNP, intron 1 | | TT or CT | 725/806 | 0.89 (0.81-0.99) | 1.00 | 289/237 | 2.15 (0.85-5.46) | 1.00 | 436/569 | 0.78 (0.64-0.94) | 0.73 | |
| GPX4 | rs757229 | G/C | GG | 300/358 | | | 55/42 | | | 245/316 | | |
| | | | CC or GC | 922/963 | 1.03 (0.93-1.14) | 1.00 | 247/209 | 0.71 (0.41-1.26) | 1.00 | 675/754 | 1.13 (0.91-1.40) | 1.00 |
| SOD1 | rs11910115 | A/C | AA | 1161/1257 | | | 242/187 | | | 919/1070 | | |
| | | | CC or AC | 62/65 | 0.93 (0.74-1.17) | 1.00 | 61/64 | 0.93 (0.55-1.56) | 1.00 | 1/1 | 0.76 (0.03-17.42) | 1.00 |
| | rs4998557 | G/A | GG | 829/959 | | | 126/100 | | | 703/859 | | |
| | | | AA or GA | 393/360 | 1.09 (0.99-1.21) | 1.00 | 177/151 | 0.80 (0.52-1.24) | 1.00 | 216/209 | 1.33 (1.05-1.68) | 1.00 |
| | rs10432782 | T/G | TT | 874/992 | | | 171/130 | | | 703/862 | | |
| | | | GG or TG | 348/329 | 1.07 (0.97-1.18) | 1.00 | 132/121 | 0.65 (0.42-1.00) | 1.00 | 216/208 | 1.35 (1.07-1.71) | 0.82 |
| rs2070424 | A/G | AA | 1003/1120 | | | 215/167 | | | 788/953 | | | |
| tag SNP, intron 3 | | GG or AG | 220/202 | 1.06 (0.94-1.20) | 1.00 | 88/84 | 0.52 (0.33-0.83) | 0.36 | 132/118 | 1.47 (1.10-1.97) | 0.58 | |
| SOD2 | rs1041740 | C/T | CC | 682/705 | | | 236/205 | | | 446/500 | | |
| | | | TT or CT | 541/617 | 0.98 (0.90-1.07) | 1.00 | 67/46 | 1.19 (0.70-2.02) | 1.00 | 474/571 | 0.92 (0.76-1.12) | 1.00 |
| | rs4342445 | G/A | GG | 752/850 | | | 229/188 | | | 523/662 | | |
| | | | AA or GA | 471/472 | 1.11 (1.02-1.22) | 1.00 | 74/63 | 1.11 (0.68-1.82) | 1.00 | 397/409 | 1.27 (1.04-1.54) | 1.00 |
| rs2842980 | A/T | AA | 715/807 | | | 136/135 | | | 579/672 | | | |
| | | TT or AT | 508/515 | 1.02 (0.94-1.12) | 1.00 | 167/116 | 1.53 (1.00-2.35) | 1.00 | 341/399 | 0.96 (0.79-1.18) | 1.00 | |
| rs8031 | T/A | TT | 387/382 | | | 138/109 | | | 249/273 | | | |
| | | AA or TA | 836/940 | 0.98 (0.89-1.08) | 1.00 | 165/142 | 1.01 (0.66-1.55) | 1.00 | 671/798 | 0.96 (0.77-1.20) | 1.00 | |

Table S5. SNP effects (dominant genetic model) on odds of developing SCCHN, overall and by race

(Odds ratios (ORs) for carriers of the minor allele compared to homozygotes for the major allele)

| Gene | SNP ¹ | major/ minor alleles | Genotype | ALL SUBJECTS COMBINED | | | AFRICAN-AMERICANS ONLY | | | EUROPEAN-AMERICANS ONLY | | |
|------|------------------|-------------------------|----------|-----------------------------------|--------------------------------------|---|-----------------------------------|--------------------------------------|---|-----------------------------------|--------------------------------------|---|
| | | | | n cases/ controls ² | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) | n cases/ controls ⁴ | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) | n cases/ controls ⁵ | Adjusted OR ³ (95% CI) | p-value (Bonferroni- corrected for 64 tests) |
| | rs5746134 | C/T | CC | 1125/1257 | | | 205/190 | | | 920/1067 | | |
| | | | TT or CT | 98/65 | 1.20 (0.98-1.48) | 1.00 | 98/61 | 1.66 (1.02-2.70) | 1.00 | 0/4 | N/A | 1.00 |
| | <i>rs2758331</i> | C/A | CC | 422/419 | | | 172/146 | | | 250/273 | | |
| | | | AA or CA | 801/903 | 0.99 (0.90-1.09) | 1.00 | 131/105 | 1.13 (0.74-1.73) | 1.00 | 670/798 | 0.95 (0.76-1.18) | 1.00 |

¹ Two SNP names highlighted in **bold** show strong evidence of OR differences between African-Americans and European-Americans using the criterion of non-overlapping 95% CIs. Four SNP names highlighted in *italics* show some evidence of OR differences between African- and European-Americans, but 95% CIs overlap somewhat

² Cases and controls do not sum to 1227 and 1325, respectively, because 4 cases and 3 controls are missing information on duration of cigarette smoking, and a few subjects lack genotype information for some SNPs

³ Conditional logistic regression models for calculating main effects of each SNP were conditioned on sex, race, and age category, and adjusted for continuous duration of smoking in years (rounded to whole years)

⁴ Numbers of African-American subjects do not sum to 305 cases and 251 controls, respectively, because 2 cases are missing information on duration of cigarette smoking, and a few subjects lack genotype information for some SNPs.

⁵ Numbers of European-American subjects do not sum to 922 cases and 1074 controls, respectively, because 2 cases and 3 controls are missing information on duration of cigarette smoking, and a few subjects lack genotype information for some SNPs.