**Appendix 3. Association of candidate genetic variants with number of injection**

|  |  |  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| **Gene** | **Variant** | **Major allele** | **Minor allele** | | **Time point** | | **N (major homo/hetero/minor homo)** | | **Genetic model** | **Number of injection** | | |
| **Mean\*** | | **Uncor-rected**  ***P* -value\*\*** |
| ***CFH*** | rs800292 | G | A | | 12 mo | | 201 / 156 / 37 | | Additive | 6.7 / 6.6 / 6.7 | | 0.7709 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.9686 |
|  |  |  |  | | 24 mo | | 187 / 145 / 34 | | Additive | 10.7 / 10.4 / 10.1 | | 0.4286 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.7429 |
| ***CFH*** | rs1061170† | T | C | | 12 mo | | 318 / 69 / 3 | | Additive | 6.7 / 6.5 / 7.7 | | 0.3124 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.1924 |
|  |  |  |  | | 24 mo | | 294 / 65 / 3 | | Additive | 12.7 / 10.2 / 10.6 | | 0.4915 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.3264 |
| ***CFH*** | rs1410996 | G | A | | 12 mo | | 186 / 160 / 47 | | Additive | 6.8 / 6.6 / 6.8 | | 0.6342 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.8463 |
|  |  |  |  | | 24 mo | | 173 / 149 / 44 | | Additive | 10.0 / 10.3 / 10.8 | | 0.1852 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.3442 |
| ***CFI*** | rs10033900 | T | C | | 12 mo | | 192 / 162 / 39 | | Additive | 6.7 / 6.7 / 6.6 | | 0.4398 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.6088 |
|  |  |  |  | | 24 mo | | 179 / 150 / 36 | | Additive | 10.7 / 10.5 / 9.6 | | 0.0884 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.1429 |
| ***C2*** | rs9332739† | G | C | | 12 mo | | 385 / 8 / 1 | | Additive | 6.7 / 6.6 / 6.0 | | 0.8792 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.896 |
|  |  |  |  | | 24 mo | | 358 / 7 / 1 | | Additive | 10.5 / 10.4 / 8.0 | | 0.7636 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.8682 |
| ***CFB*** | rs641153† | G | A | | 12 mo | | 343 / 48 / 1 | | Additive | 6.7 / 6.7 / 4.0 | | 0.797 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.6368 |
|  |  |  |  | | 24 mo | | 318 / 45 / 1 | | Additive | 10.5 / 10.7 / 9.8 | | 0.3764 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.3085 |
| ***SKIV2L*** | rs429608† | G | A | | 12 mo | | 344 / 44 / 4 | | Additive | 6.7 / 6.7 / 5.8 | | 0.931 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.7351 |
|  |  |  |  | | 24 mo | | 318 / 42 / 4 | | Additive | 10.5 / 10.7 / 9.8 | | 0.5354 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.4504 |
| ***VEGFA*** | rs699947 | C | A | | 12 mo | | 202 / 148 / 42 | | Additive | 6.6 / 6.8 / 7.1 | | 0.2701 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.3559 |
|  |  |  |  | | 24 mo | | 188 / 137 / 39 | | Additive | 10.4 / 10.5 / 11.0 | | 0.6564 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.5155 |
| ***VEGFA*** | rs3025039 | C | T | | 12 mo | | 236 / 133 / 25 | | Additive | 6.8 / 6.5 / 6.3 | | 0.0979 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.3211 |
|  |  |  |  | | 24 mo | | 219 / 124 / 23 | | Additive | 10.7 / 10.4 / 9.0 | | **0.0286** |
|  |  |  |  | |  | |  | | Recessive |  | | **0.015** |
| ***ARMS2*** | rs10490924 | T | G | | 12 mo | | 196 / 150 / 47 | | Additive | 6.8 / 6.8 / 6.1 | | 0.1589 |
|  |  |  |  | |  | |  | | Recessive |  | | **0.0344** |
|  |  |  |  | | 24 mo | | 183 / 139 / 44 | | Additive | 10.5 / 10.9 / 9.4 | | 0.4415 |
|  |  |  |  | |  | |  | | Recessive |  | | **0.0338** |
| ***HTRA1*** | rs11200638 | A | G | | 12 mo | | 202 / 144 / 48 | | Additive | 6.8 / 6.7 / 6.1 | | 0.1024 |
|  |  |  |  | |  | |  | | Recessive |  | | **0.0304** |
|  |  |  |  | | 24 mo | | 188 / 133 / 45 | | Additive | 10.5 / 10.9 / 9.4 | | 0.3129 |
|  |  |  |  | |  | |  | | Recessive |  | | **0.0275** |
| ***SCARB1*** | rs5888 | C | T | | 12 mo | | 216 / 146 / 31 | | Additive | 6.7 / 6.8 / 6.3 | | 0.9409 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.353 |
|  |  |  |  | | 24 mo | | 201 / 136 / 29 | | Additive | 10.5 / 10.7 / 10.0 | | 0.7495 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.4713 |
| ***PEDF*** | rs1136287 | C | T | | 12 mo | | 100 / 195 / 97 | | Additive | 6.5 / 6.6 / 7.0 | | 0.1205 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.1557 |
|  |  |  |  | | 24 mo | | 92 / 182 / 91 | | Additive | 10.0 / 10.5 / 10.9 | | 0.1307 |
|  |  |  |  | |  | |  | | Recessive |  | | 0.2968 |
| ***SYN3/***  ***TIMP3*** | rs9621532† | T | G | | 12 mo | | 372 / 20 / 1 | | Additive | 6.7 / 6.8 / 6.0 | | 0.588 |
|  |  |  | |  | |  | | Dominant |  | | 0.5294 |
|  |  |  |  | | 24 mo | | 346 / 19 / 1 | | Additive | 10.5 / 10.6 / 12.0 | | 0.5297 |
|  |  |  |  | |  | |  | | Dominant |  | | 0.6033 |
| ***APOE*** | apoE†† | - | - | 12 mo | | 42 / 274 / 70 | | ε2+ / ε3/ε3 / ε4+ | | | 6.3 / 6.8 / 6.7 | 0.2485 |
|  |  |  |  |  | |  | | others / ε2 carriers | | |  | 0.0985 |
|  |  |  |  |  | |  | | others / ε4 carriers | | |  | 0.3954 |
|  |  |  |  | 24 mo | | 40 / 253 / 66 | | ε2+ / ε3/ε3 / ε4+ | | | 10.3 / 10.6 / 10.5 | 0.7485 |
|  |  |  |  |  | |  | | others / ε2 carriers | | |  | 0.5963 |
|  |  |  |  |  | |  | | others / ε4 carriers | | |  | 0.6245 |

OR = Odds ratio; CI = Confidence interval

\*: Major allele homozygote / Heterozygote / Minor allele homozygote

\*\*: Uncorrected *P*-value from linear regression model. When applying Bonferroni correction, uncorrected *P*-value for significance should be less than 0.0031.

†: Analyzed under dominant model because of extremely low proportion of minor allele homozygotes

††: ε2+ = ε2 carrier, ε4+ = ε4 carrier