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| **Table S2. Association of a burden of rare variants in *LDLR* with plasma LDL-C levels and MI-risk for variants classified as non-disruptive and unclear.** |
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| **pheno-****type** | **variants analyzed** | **variant count** | **allele count** | **allele freq.** | **LDL-C >190mg/dl****(n=251)** | **LDL-C <190mg/dl****(n=1,901)** | **P-value** | **OR** | **95% CI** |
| **plasma****LDL-C** | non-disruptive | 46 | 99 | 0.046 | 19 | 80 | 0.023 | 1.8 | 1.0-3.1 |
| unclear | 10 | 10 | 0.005 | 2 | 8 | 0.330 | 1.9 | 0.2-9.6 |
| non-disruptive + unclear | 56 | 109 | 0.051 | 21 | 88 | 0.038 | 1.8 | 1.0-3.1 |
| **pheno-****type** | **variants analyzed** | **variant count** | **allele count** | **allele freq.** | **MI case****(n=1,716)** | **MI control****(n=1,519)** | **P-value** | **OR** | **95% CI** |
| **MI** | non-disruptive | 46 | 136 | 0.042 | 82 | 54 | 0.035 | 1.3 | 0.9-1.9 |
| unclear | 10 | 12 | 0.004 | 10 | 2 | 0.043 | 4.4 | 0.9-41.6 |
| non-disruptive + unclear | 56 | 148 | 0.046 | 92 | 56 | 0.114 | 1.5 | 1.0-2.1 |