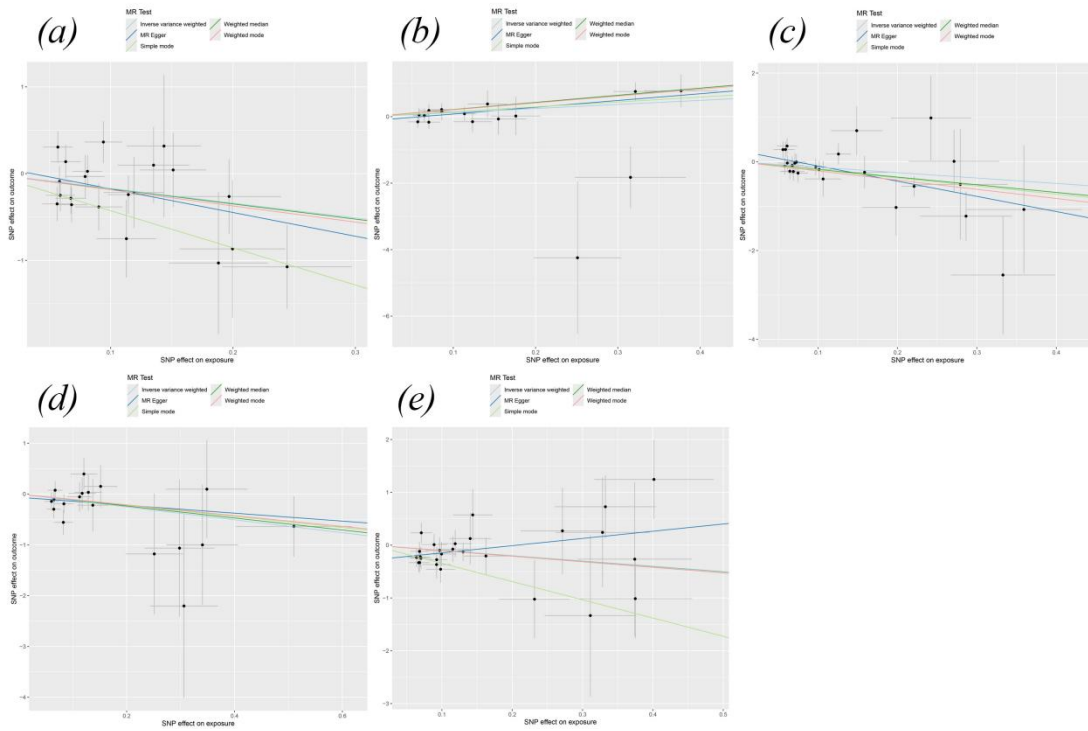


Supplementary figure

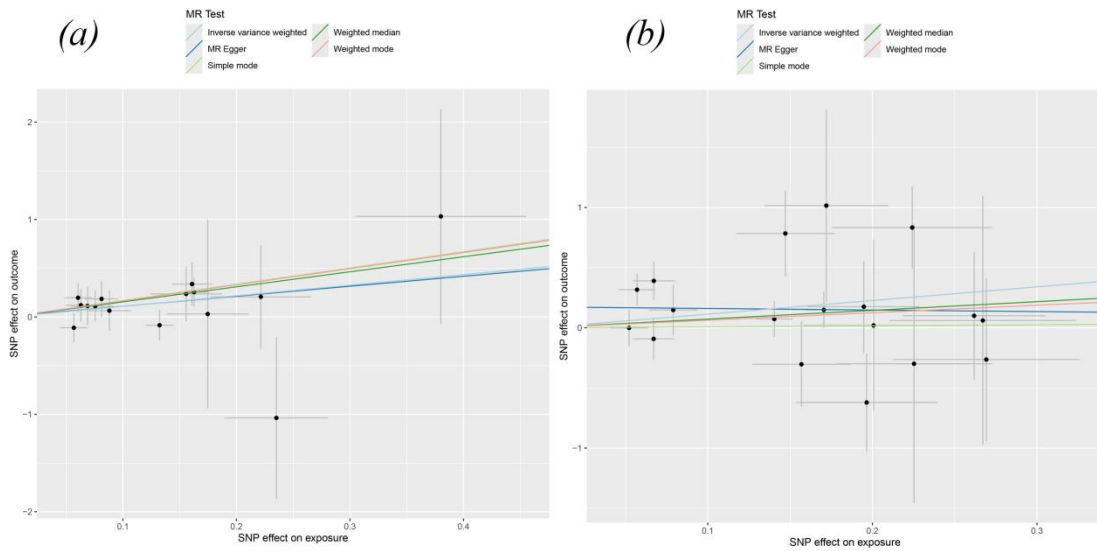
supplementary Figure S1-S4: Scatter plot of the Mendelian analysis.

supplementary Figure S5-S7: Funnel plot of Mendelian randomization analysis.

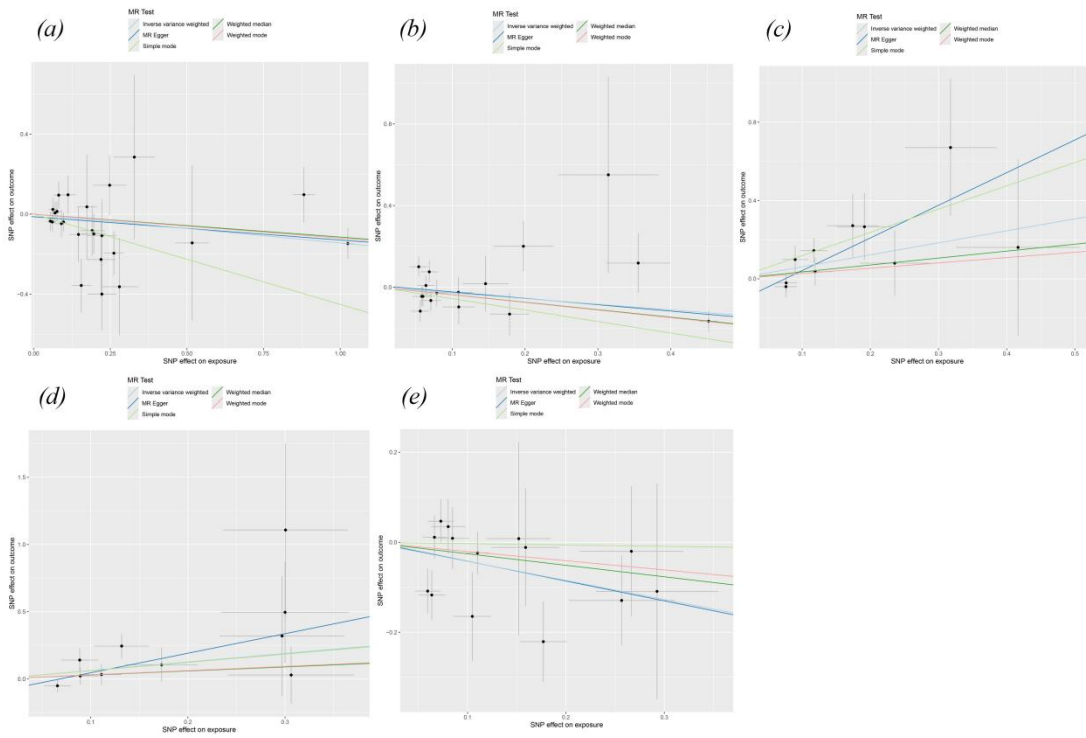
supplementary Figure S8-S13: Leave-one-out plot of the Mendelian analysis



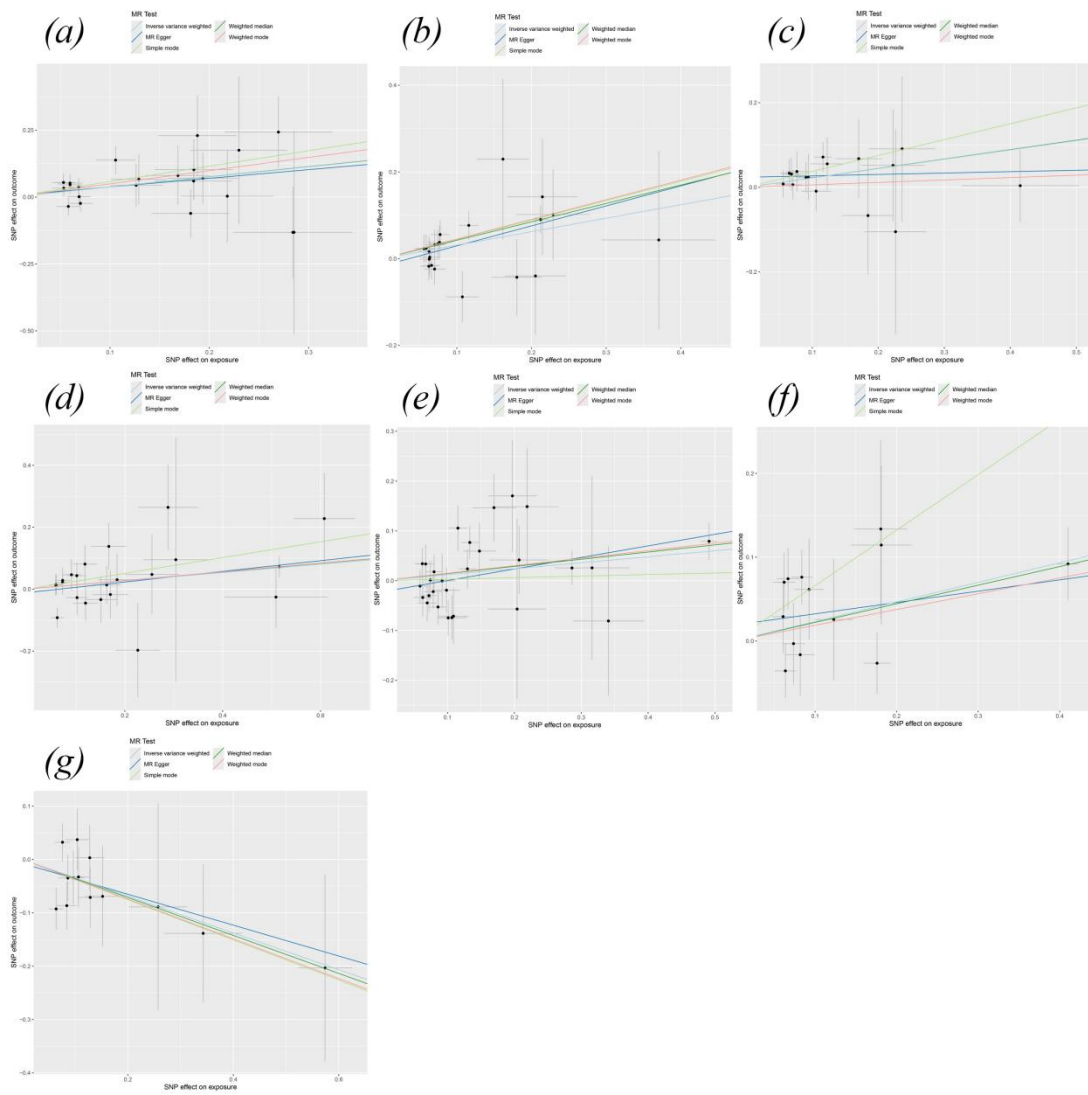
supplementary Figure S1: Scatter plot of the Mendelian analysis of inflammatory proteins for glossopharyngeal nerve disease. (a) CX3CL1. (b) MMP-1. (c) SLAMF1. (d) TNF. (e) TNFRSF9.



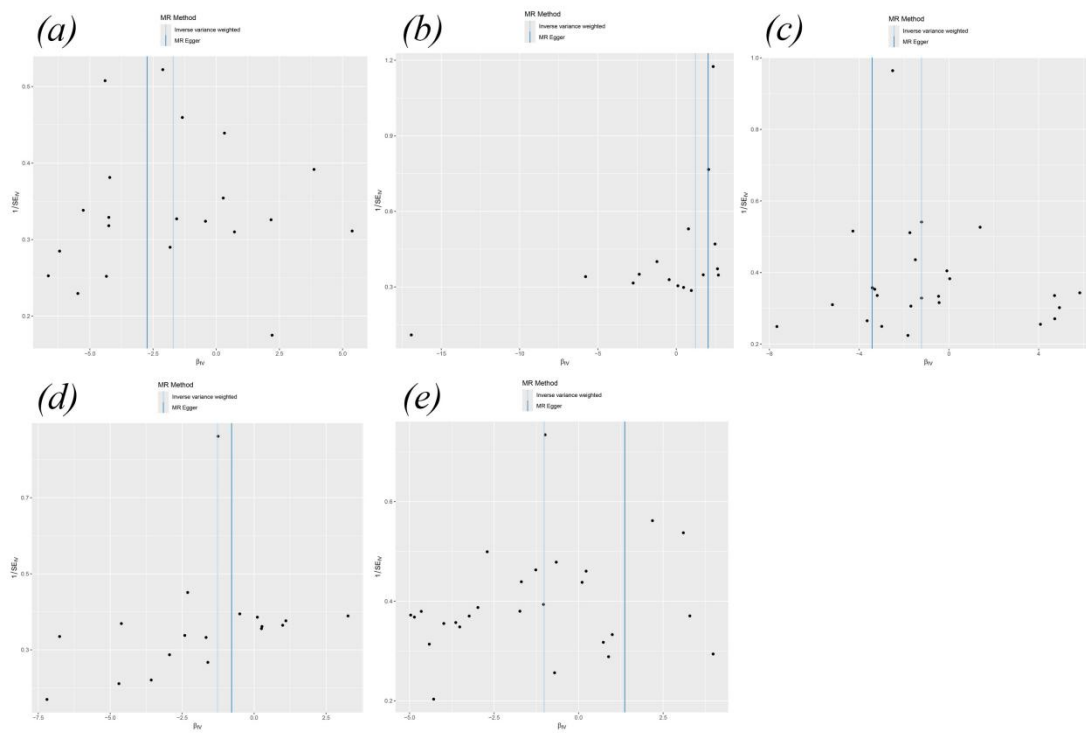
supplementary Figure S2. Scatter plot of the Mendelian analysis of inflammatory proteins for Phantom limb syndrome with pain. (a) FGF21. (b) Osteoprotegerin.



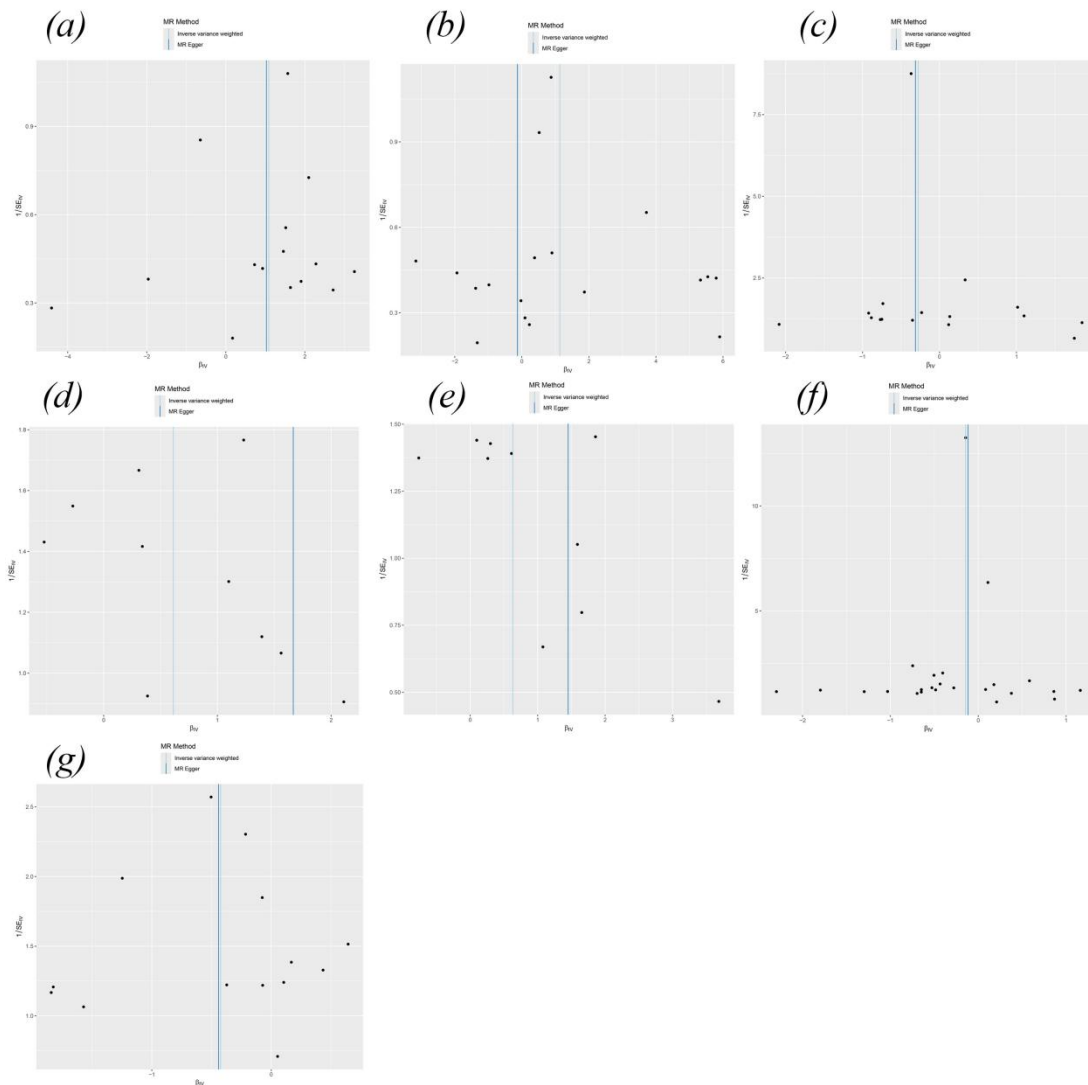
supplementary Figure S3: Scatter plot of the Mendelian analysis of inflammatory proteins for Small fiber neuropathy. (a) CD40LG. (b) IL-20RA. (c) IL-24. (d) CCL8. (e) OSM.



supplementary Figure S4: Scatter plot of the Mendelian analysis of inflammatory proteins for Unspecified neuralgia or neuritis. (a) CCL28. (b) CD244. (c) FGF23. (d) FGF5. (e) IL-12B. (f) CCL3. (g) SIRT2.



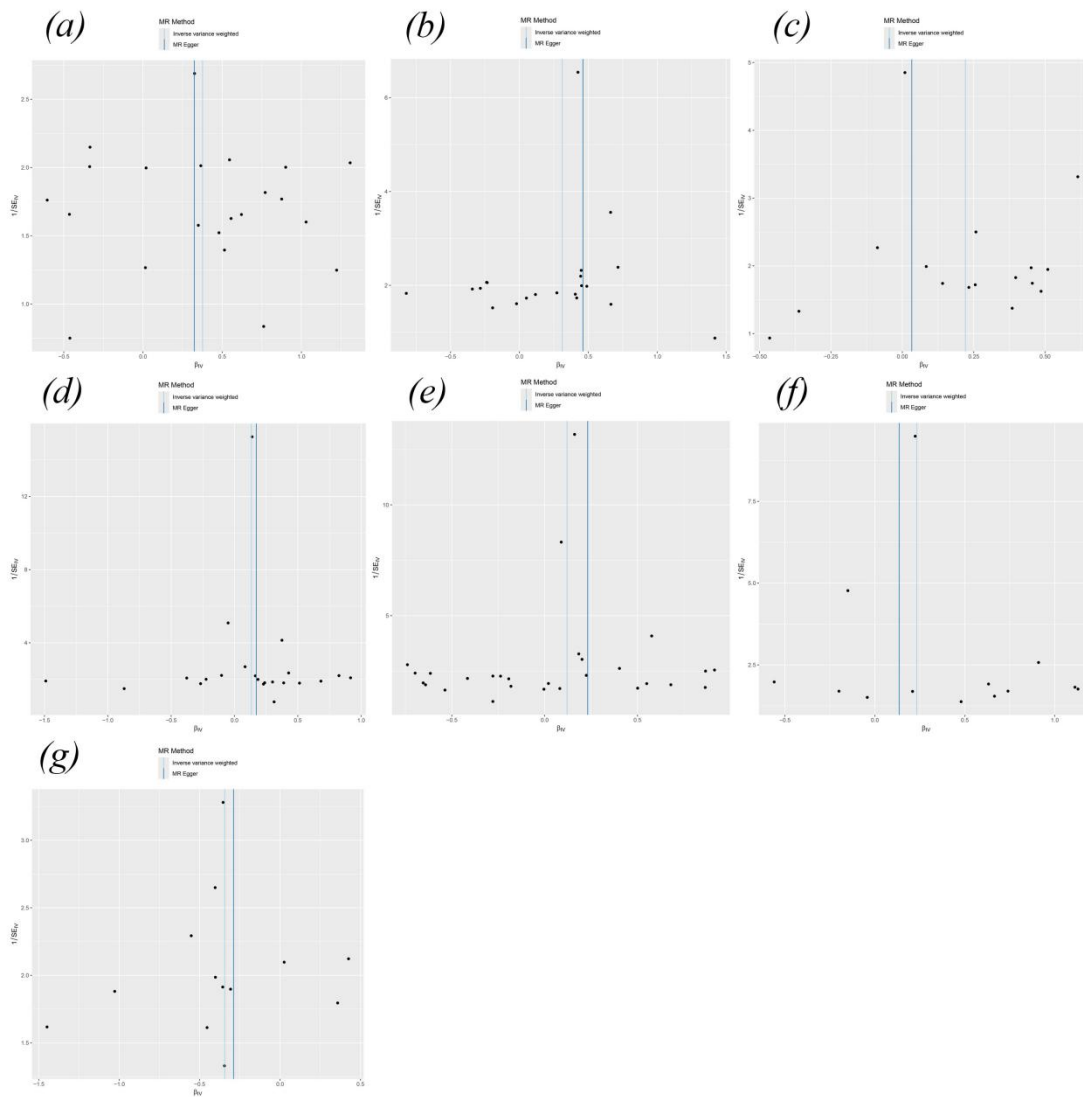
supplementary Figure S5: Funnel plot of Mendelian randomization analysis of inflammatory proteins for glossopharyngeal nerve disease. (a) CX3CL1. (b) MMP-1. (c) SLAMF1. (d) TNF. (e) TNFRSF9.



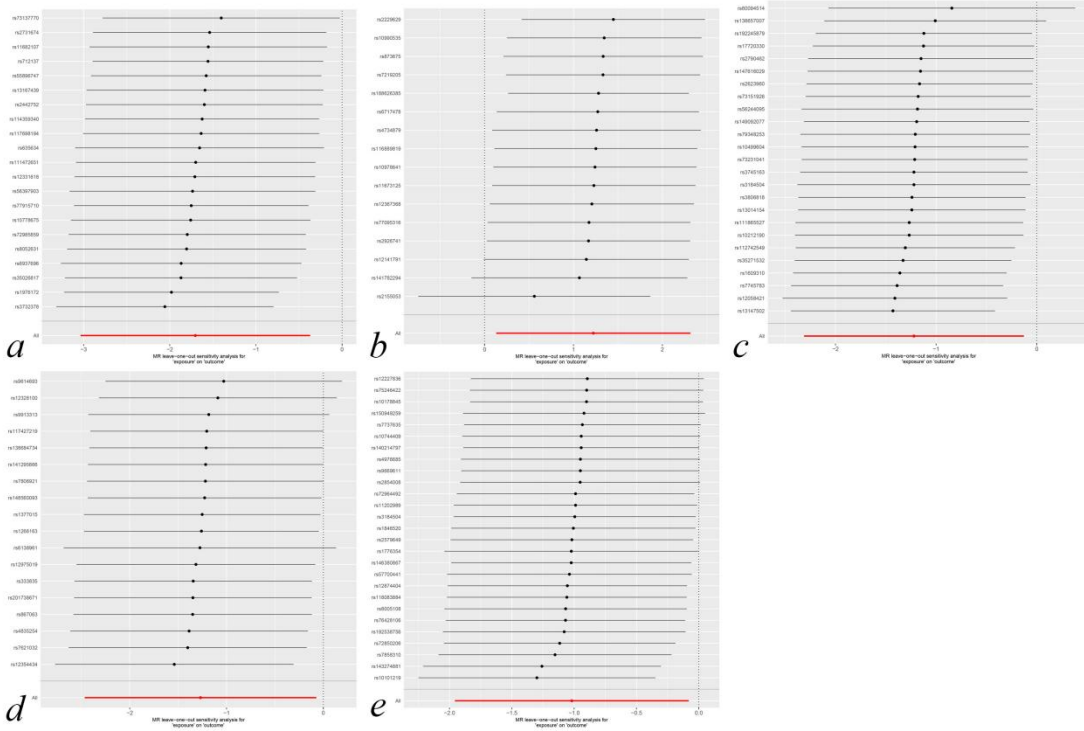
supplementary Figure S6:

(a),(b): Funnel plot of Mendelian randomization analysis of inflammatory proteins for Phantom limb syndrome with pain. (a) FGF21. (b) Osteoprotegerin.

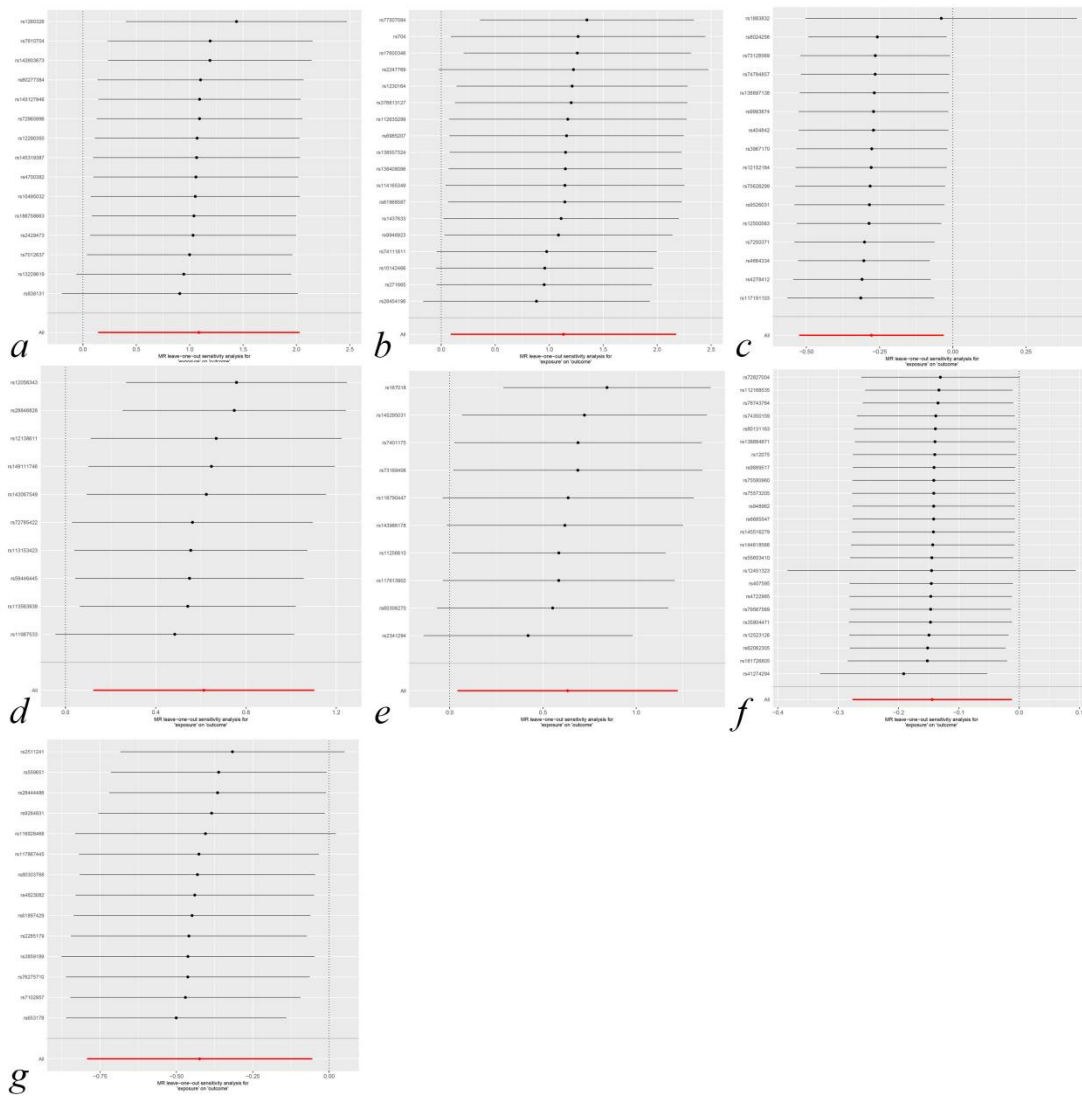
(c)-(g): Funnel plot of Mendelian randomization analysis of inflammatory proteins for Small fiber neuropathy. (c) CD40LG. (d) IL-20RA. (e) IL-24. (f) CCL8. (g) OSM



supplementary Figure S7: Funnel plot of Mendelian randomization analysis of inflammatory proteins for Unspecified neuralgia or neuritis. (a) CCL28. (b) CD244. (c) FGF23. (d) FGF5. (e) IL-12B. (f) CCL3. (g) SI RT2.

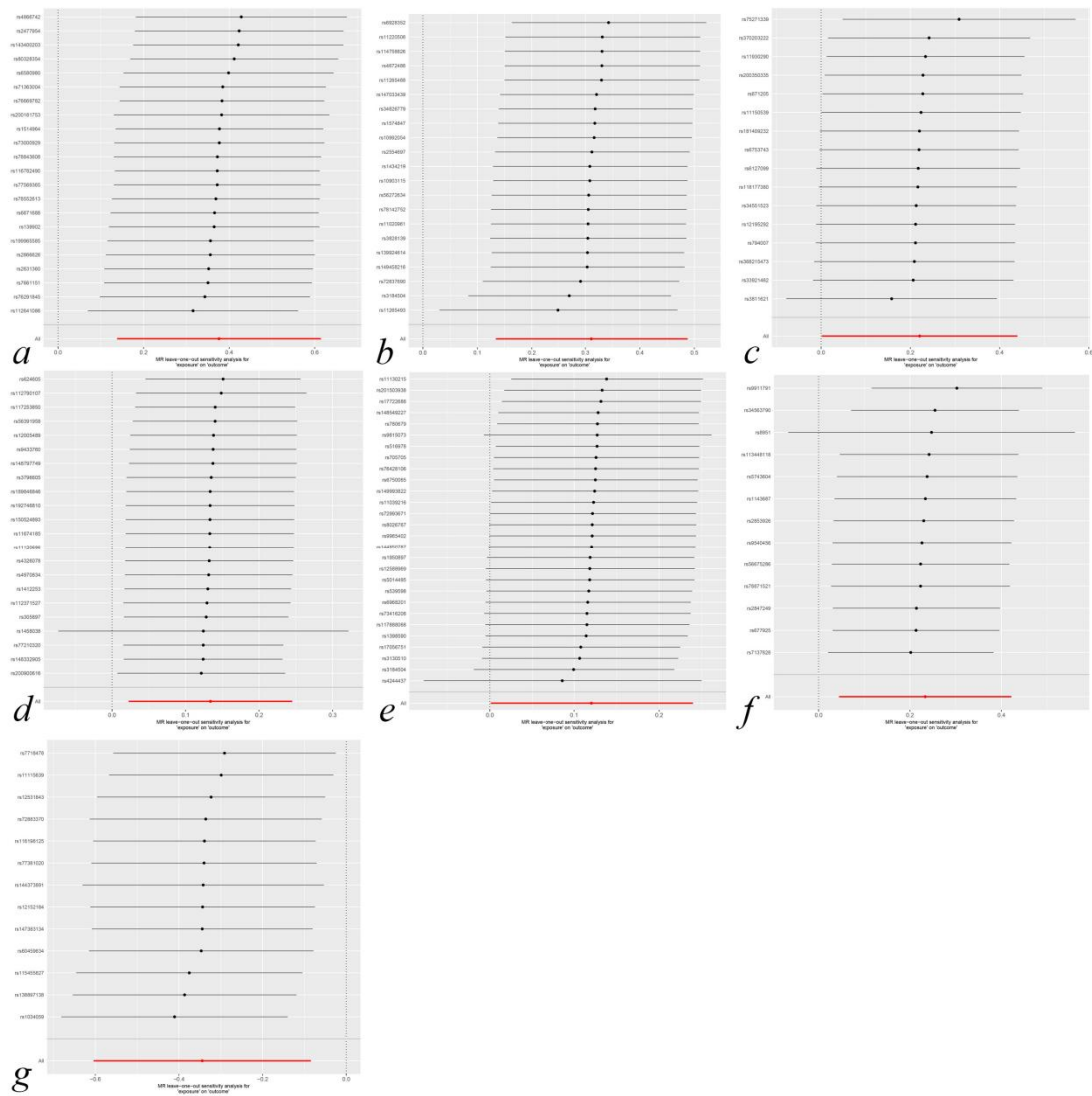


supplementary Figure S8: Leave-one-out plot of Mendelian randomization of inflammatory proteins to glossopharyngeal nerve disease. (a) CX3CL1. (b) MMP-1. (c) SLAMF1. (d) TNF. (e) TNFRSF9.

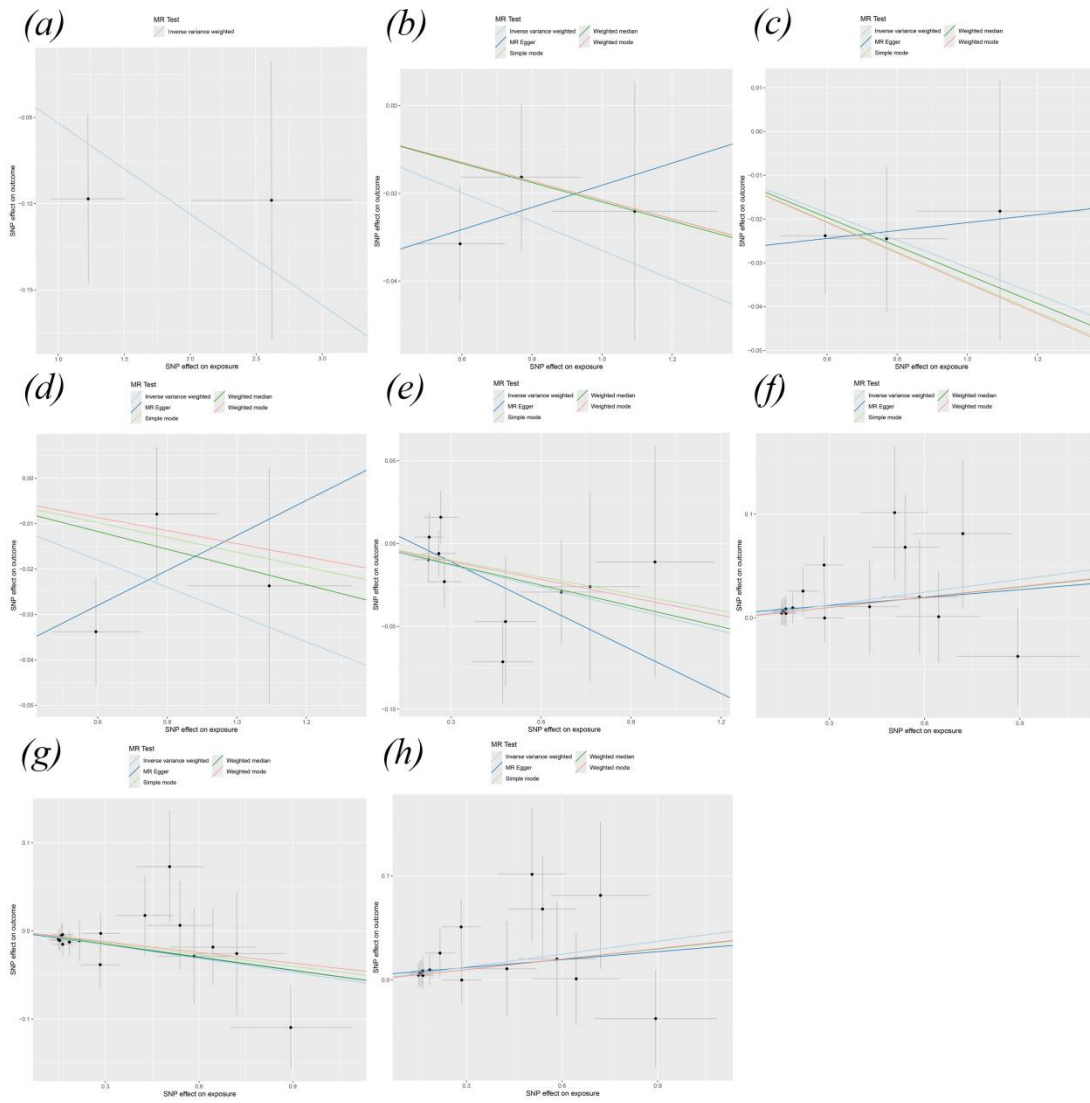


supplementary Figure S9:

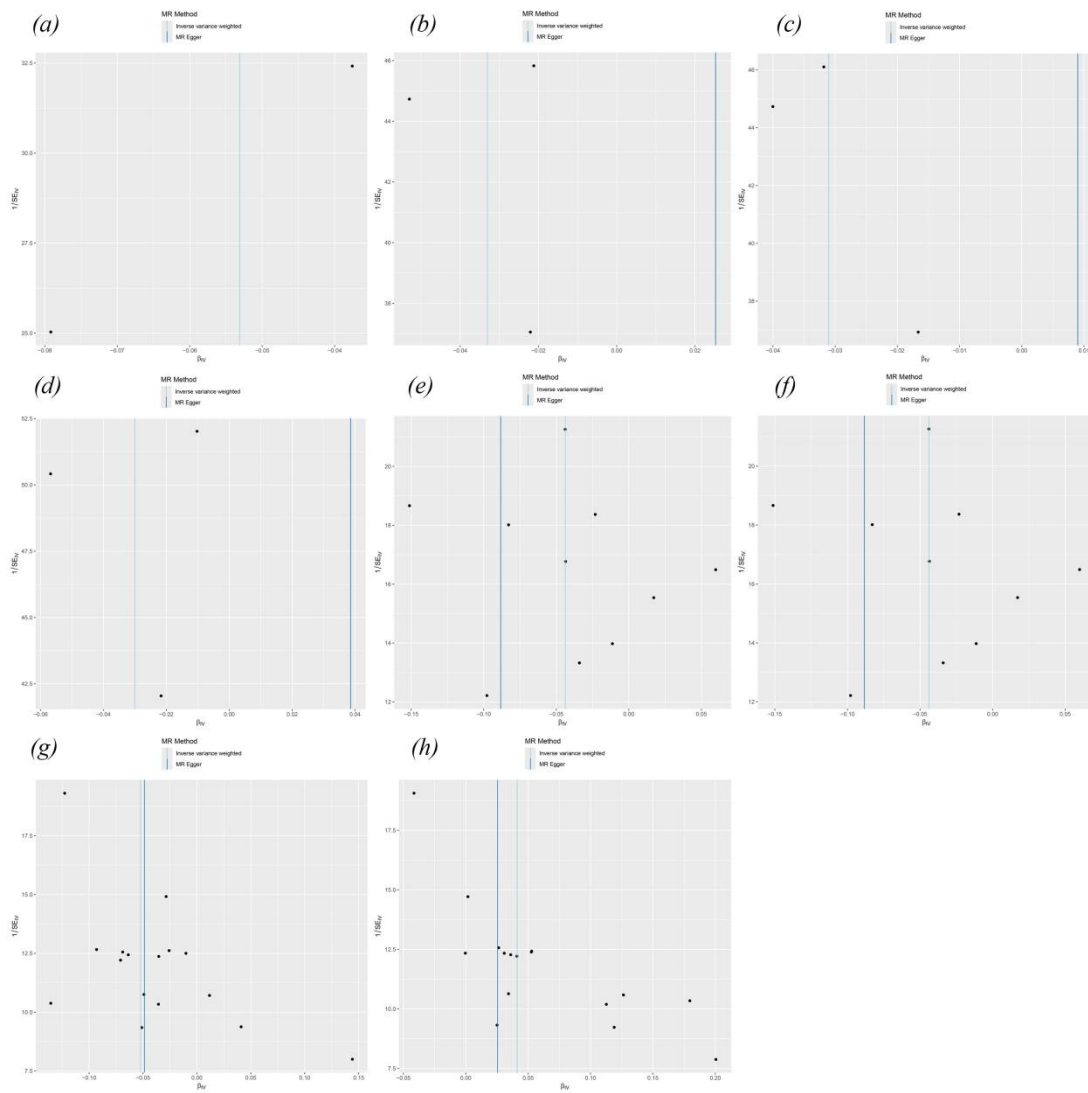
(a),(b): Leave-one-out plot of Mendelian randomization of inflammatory proteins to Phantom limb syndrome with pain. (a) FGF21. (b) Osteoprotegerin.
 (c)-(g): Leave-one-out plot of Mendelian randomization analysis of inflammatory proteins for Small fiber neuropathy. (c) CD40LG. (d) IL-20RA. (e) IL-24. (f) CCL8. (g) OSM



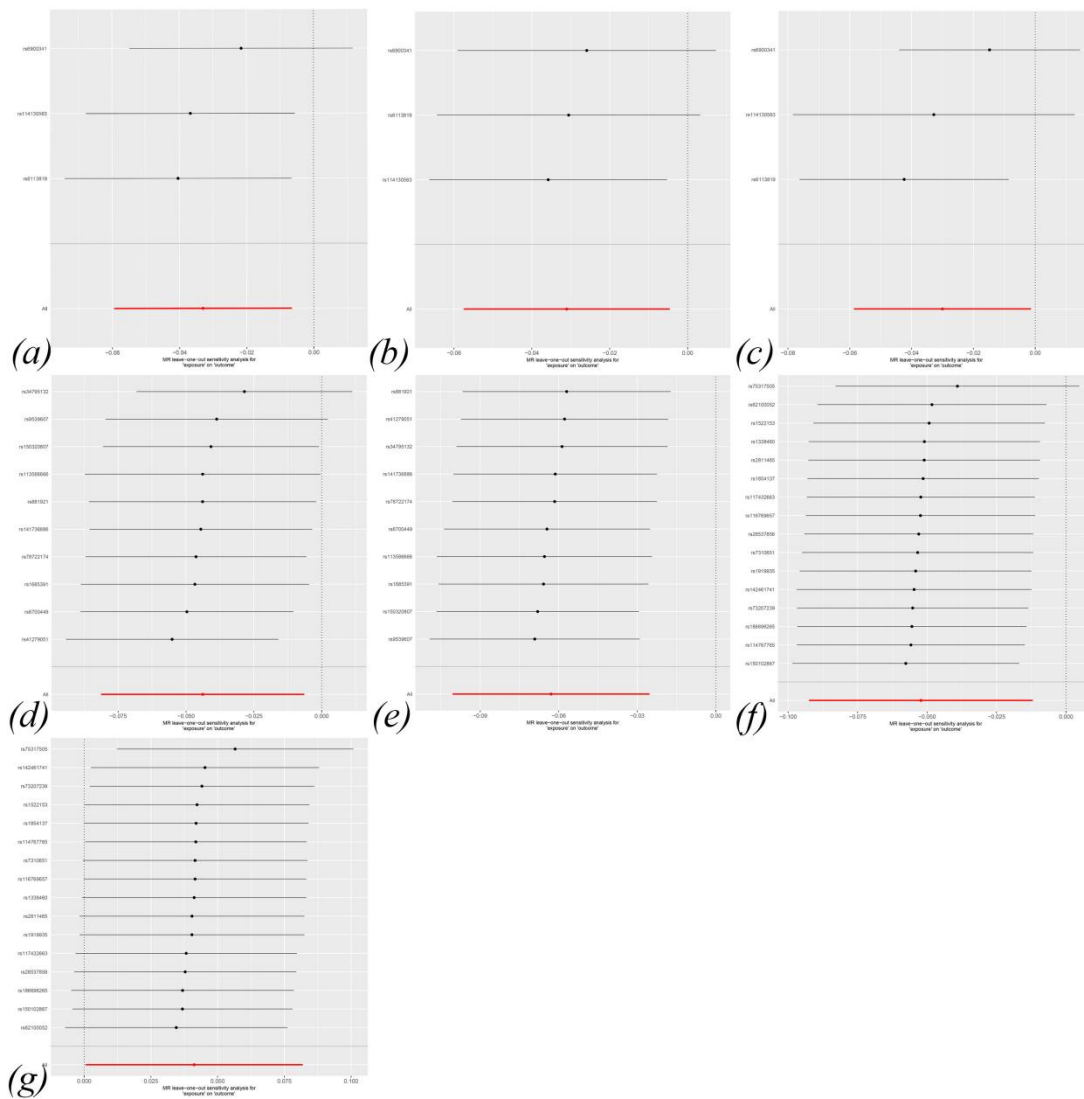
supplementary Figure S10: Leave-one-out plot of Mendelian randomization analysis of inflammatory proteins for Unspecified neuralgia or neuritis. (a) CCL28. (b) CD244. (c) FGF23. (d) FGF5. (e) IL-12B. (f) CCL3. (g) SIRT2.



supplementary Figure S11: Scatter plot of the Mendelian analysis of Neuropathic pain for inflammatory proteins. (a)NT-3. (b) IL-10RA. (c) IL-17A. (d) TNFSF12. (e) IL-2RB. (f) IL-5. (g) CCL23. (h) CXCL5.



supplementary Figure S12: Funnel plot of the Mendelian analysis of Neuropathic pain for inflammatory proteins. (a)NT-3. (b) IL-10RA. (c) IL-17A. (d) TNFSF12. (e) IL-2RB. (f) IL-5. (g) CCL23. (h) CXCL5.



supplementary Figure S13: Leave-one-out plot of the Mendelian analysis of Neuropathic pain for inflammatory proteins. (a)IL-10RA. (b) IL-17A. (c) TNFSF12. (d) IL-2RB . (e) IL-5. (f) CCL23. (g) CXCL5.

Supplementary table

| MR estimates of Inflammatory proteins on Neuropathic pain | | | | | | | | | | | | | | | | | |
|---|-----------------------|-------|---------------------------|---------------|-------|---------------|----------------|-------|-----------------|----------------|-------|-------------|----------------|-------|---------------|----------------|-------|
| Outcome | Inflammatory proteins | nSNP | Inverse variance weighted | | | MR Egger | | | Weighted median | | | Simple mode | | | Weighted mode | | |
| | | | OR | 95CI | pval | OR | 95CI | pval | OR | 95CI | pval | OR | 95CI | pval | OR | 95CI | pval |
| Glossopharyngeal nerve disease | CX3CL1 | 21 | 0.182 | (0.048,0.691) | 0.012 | 0.065 | (0.002,2.013) | 0.135 | 0.174 | (0.027,1.14) | 0.068 | 0.014 | (0,0.457) | 0.026 | 0.155 | (0.008,3.144) | 0.239 |
| | MMP-1 | 16 | 3.399 | (1.14,10.133) | 0.028 | 7.635 | (1.171,49.794) | 0.052 | 8.536 | (2.267,32.142) | 0.002 | 4.341 | (0.481,39.189) | 0.211 | 7.904 | (2.092,29.869) | 0.008 |
| | SLAMF1 | 25 | 0.295 | (0.099,0.879) | 0.028 | 0.033 | (0.004,0.264) | 0.004 | 0.178 | (0.046,0.695) | 0.013 | 0.161 | (0.015,1.744) | 0.146 | 0.126 | (0.022,0.727) | 0.029 |
| | TNF | 18 | 0.281 | (0.085,0.931) | 0.038 | 0.457 | (0.053,3.967) | 0.488 | 0.307 | (0.051,1.834) | 0.195 | 0.333 | (0.029,3.806) | 0.389 | 0.346 | (0.049,2.419) | 0.3 |
| | TNFRSF9 | 27 | 0.361 | (0.141,0.923) | 0.033 | 3.921 | (0.506,30.361) | 0.203 | 0.365 | (0.085,1.555) | 0.173 | 0.032 | (0.002,0.478) | 0.019 | 0.349 | (0.035,3.493) | 0.379 |
| Phantom limb syndrome with pain | FGF21 | 15 | 2.961 | (1.152,7.614) | 0.024 | 2.787 | (0.287,27.032) | 0.393 | 4.684 | (1.242,17.659) | 0.023 | 5.33 | (0.785,36.187) | 0.109 | 5.214 | (1.027,26.479) | 0.066 |
| | Osteoprotegerin | 18 | 3.103 | (1.092,8.818) | 0.034 | 0.879 | (0.101,7.669) | 0.909 | 2.06 | (0.569,7.452) | 0.271 | 1.086 | (0.095,12.345) | 0.948 | 1.865 | (0.49,7.099) | 0.373 |
| Small fibre neuropathy | CD40LG | 16 | 0.758 | (0.592,0.97) | 0.027 | 0.731 | (0.513,1.042) | 0.105 | 0.693 | (0.56,0.858) | 0.001 | 0.571 | (0.274,1.19) | 0.155 | 0.689 | (0.545,0.87) | 0.007 |
| | IL-20RA | 10 | 1.846 | (1.131,3.013) | 0.014 | 5.299 | (1.579,17.777) | 0.027 | 1.425 | (0.723,2.808) | 0.307 | 3.279 | (0.98,10.972) | 0.086 | 1.312 | (0.484,3.56) | 0.607 |
| | IL-24 | 10 | 1.881 | (1.042,3.396) | 0.036 | 4.272 | (1.116,16.352) | 0.067 | 1.348 | (0.629,2.891) | 0.442 | 1.853 | (0.591,5.812) | 0.318 | 1.365 | (0.545,3.42) | 0.523 |
| | CCL8 | 24 | 0.866 | (0.758,0.988) | 0.033 | 0.89 | (0.757,1.047) | 0.173 | 0.892 | (0.771,1.031) | 0.121 | 0.637 | (0.372,1.091) | 0.114 | 0.887 | (0.772,1.018) | 0.102 |
| | OSM | 14 | 0.655 | (0.453,0.947) | 0.024 | 0.644 | (0.283,1.463) | 0.314 | 0.775 | (0.478,1.256) | 0.3 | 0.971 | (0.429,2.196) | 0.945 | 0.815 | (0.428,1.553) | 0.545 |
| Neuralgia and neuritis, unspecified | CCL28 | 22 | 1.457 | (1.148,1.849) | 0.002 | 1.383 | (0.837,2.284) | 0.22 | 1.461 | (1.035,2.063) | 0.031 | 1.781 | (0.932,3.404) | 0.095 | 1.639 | (0.881,3.049) | 0.134 |
| | CD244 | 21 | 1.365 | (1.143,1.629) | 0.001 | 1.586 | (1.089,2.309) | 0.026 | 1.529 | (1.179,1.983) | 0.001 | 1.562 | (1.003,2.433) | 0.062 | 1.573 | (1.195,2.071) | 0.004 |
| | FGF23 | 16 | 1.247 | (1.002,1.553) | 0.048 | 1.034 | (0.673,1.587) | 0.882 | 1.249 | (0.924,1.69) | 0.148 | 1.456 | (0.868,2.441) | 0.175 | 1.059 | (0.707,1.587) | 0.784 |
| | FGF5 | 22 | 1.143 | (1.023,1.278) | 0.018 | 1.188 | (1.013,1.392) | 0.047 | 1.149 | (1.018,1.298) | 0.025 | 1.29 | (0.935,1.78) | 0.135 | 1.151 | (1.023,1.295) | 0.029 |
| | IL-12B | 28 | 1.128 | (1.001,1.271) | 0.048 | 1.261 | (1.048,1.518) | 0.021 | 1.155 | (1.01,1.32) | 0.035 | 1.031 | (0.704,1.512) | 0.875 | 1.163 | (1.018,1.328) | 0.034 |
| | CCL3 | 13 | 1.263 | (1.046,1.525) | 0.015 | 1.145 | (0.849,1.544) | 0.395 | 1.249 | (1.021,1.528) | 0.031 | 1.938 | (1.063,3.531) | 0.052 | 1.207 | (0.978,1.489) | 0.105 |
| SIRT2 | 13 | 0.709 | (0.547,0.919) | 0.009 | 0.749 | (0.442,1.268) | 0.305 | 0.701 | (0.486,1.009) | 0.056 | 0.686 | (0.393,1.2) | 0.211 | 0.689 | (0.428,1.111) | 0.153 | |

Supplementary Table S1: MR estimates of Inflammatory proteins on Neuropathic pain

| MR estimates of Neuropathic pain on Inflammatory proteins | | | | | | | | | | | | | | | | | | |
|---|-----------------------|--------|---------------------------|-----------------|-------|--------|----------------|-------|--------|-----------------|-------|--------|-----------------|-------|--------|----------------|-------|--|
| Exposure | Inflammatory proteins | nSNP | Inverse variance weighted | | | | MR Egger | | | Weighted median | | | Simple mode | | | Weighted mode | | |
| | | | beta | 95CI | pval | beta | 95CI | pval | beta | 95CI | pval | beta | 95CI | pval | beta | 95CI | pval | |
| Glossopharyngeal nerve disease | NT-3 | 2.000 | -0.053 | (-0.101,-0.005) | 0.030 | \ | \ | \ | \ | \ | \ | \ | \ | \ | \ | \ | \ | |
| Phantom limb syndrome with pain | IL-10RA | 3.000 | -0.033 | (-0.059,-0.007) | 0.015 | 0.025 | (-0.097,0.148) | 0.755 | -0.022 | (-0.057,0.013) | 0.217 | -0.022 | (-0.061,0.018) | 0.397 | -0.022 | (-0.062,0.019) | 0.410 | |
| | IL-17A | 3.000 | -0.031 | (-0.057,-0.005) | 0.021 | 0.009 | (-0.114,0.132) | 0.909 | -0.033 | (-0.065,0) | 0.047 | -0.034 | (-0.071,0.002) | 0.208 | -0.035 | (-0.075,0.006) | 0.236 | |
| | TNFSF12 | 3.000 | -0.030 | (-0.059,-0.001) | 0.040 | 0.039 | (-0.089,0.166) | 0.659 | -0.020 | (-0.052,0.013) | 0.240 | -0.016 | (-0.057,0.024) | 0.514 | -0.014 | (-0.054,0.025) | 0.548 | |
| Small fibre neuropathy | IL-2RB | 10.000 | -0.044 | (-0.081,-0.006) | 0.022 | -0.088 | (-0.183,0.006) | 0.105 | -0.042 | (-0.093,0.01) | 0.111 | -0.034 | (-0.123,0.055) | 0.475 | -0.036 | (-0.11,0.038) | 0.366 | |
| | IL-5 | 10.000 | -0.063 | (-0.101,-0.025) | 0.001 | -0.057 | (-0.152,0.038) | 0.273 | -0.056 | (-0.105,-0.006) | 0.027 | -0.090 | (-0.164,-0.016) | 0.042 | -0.048 | (-0.118,0.022) | 0.215 | |
| Neuralgia and neuritis, unspecified | CCL23 | 16.000 | -0.052 | (-0.092,-0.012) | 0.011 | -0.049 | (-0.128,0.03) | 0.248 | -0.049 | (-0.103,0.004) | 0.069 | -0.045 | (-0.132,0.043) | 0.336 | -0.041 | (-0.12,0.039) | 0.333 | |
| | CXCL5 | 16.000 | 0.041 | (0,0.082) | 0.047 | 0.025 | (-0.055,0.106) | 0.544 | 0.033 | (-0.023,0.089) | 0.243 | 0.033 | (-0.06,0.127) | 0.493 | 0.033 | (-0.048,0.115) | 0.433 | |

Supplementary Table S2: MR estimates of Neuropathic pain on Inflammatory proteins

| Exposure | SNP | effect_allele | other_allele | SNPs information of inflammatory proteins with Glossopharyngeal nerve disease | | | | | | | | | | R2 | f |
|-------------|-------------|---------------|--------------|---|--------------|-------------|------------|---------------|--------------|-------------|---------------|---------------------|----------|----------|---|
| | | | | beta_exposure | beta_outcome | se_exposure | se_outcome | pval_exposure | pval_outcome | se_exposure | pval_exposure | samplesize_exposure | | | |
| CX3CL1 | rs10778675 | A | C | 0.079 | -0.034 | 0.169 | 0.152 | 8.90E-01 | 0.244 | 0.016 | 5.37E-07 | 14295 | 1.75E-03 | 2.51E+01 | |
| | rs11472651 | C | G | -0.119 | 0.217 | 0.936 | 0.950 | 5.96E-01 | 0.411 | 0.025 | 1.57E-06 | 14728 | 1.56E-03 | 2.31E+01 | |
| | rs114359340 | A | G | -0.188 | 1.030 | 0.975 | 0.988 | 2.09E-01 | 0.819 | 0.041 | 3.41E-06 | 13651 | 1.58E-03 | 2.16E+01 | |
| | rs11682107 | A | G | -0.068 | 0.284 | 0.557 | 0.544 | 1.08E-01 | 0.177 | 0.013 | 8.45E-08 | 12412 | 2.31E-03 | 2.87E+01 | |
| | rs117898194 | A | G | -0.200 | 0.867 | 0.022 | 0.013 | 2.73E-01 | 0.792 | 0.043 | 3.83E-06 | 13859 | 1.54E-03 | 2.13E+01 | |
| | rs12331618 | A | G | -0.058 | 0.091 | 0.491 | 0.426 | 6.09E-01 | 0.178 | 0.011 | 3.30E-07 | 14729 | 1.77E-03 | 2.61E+01 | |
| | rs13167439 | T | C | -0.090 | 0.384 | 0.890 | 0.882 | 1.62E-01 | 0.274 | 0.019 | 2.89E-06 | 14295 | 1.53E-03 | 2.19E+01 | |
| | rs1976172 | C | G | -0.057 | -0.306 | 0.353 | 0.380 | 9.42E-02 | 0.183 | 0.012 | 2.12E-06 | 14732 | 1.52E-03 | 2.25E+01 | |
| | rs2442752 | T | C | -0.059 | 0.250 | 0.618 | 0.655 | 1.76E-01 | 0.185 | 0.012 | 4.80E-07 | 14743 | 1.72E-03 | 2.53E+01 | |
| | rs2731674 | T | G | -0.068 | 0.358 | 0.274 | 0.260 | 7.50E-02 | 0.201 | 0.013 | 1.04E-07 | 14740 | 1.92E-03 | 2.83E+01 | |
| | rs35026817 | T | G | 0.063 | 0.137 | 0.333 | 0.302 | 4.80E-01 | 0.194 | 0.012 | 2.12E-07 | 14740 | 1.82E-03 | 2.69E+01 | |
| | rs3732378 | A | G | 0.094 | 0.364 | 0.167 | 0.159 | 1.30E-01 | 0.240 | 0.015 | 9.94E-10 | 14740 | 2.53E-03 | 3.73E+01 | |
| | rs5589674 | T | C | 0.113 | -0.750 | 0.937 | 0.961 | 9.32E-02 | 0.447 | 0.024 | 3.71E-06 | 14734 | 1.45E-03 | 2.14E+01 | |
| | rs56397903 | A | G | 0.197 | -0.264 | 0.974 | 0.955 | 5.38E-01 | 0.428 | 0.042 | 3.46E-06 | 14743 | 1.46E-03 | 2.15E+01 | |
| | rs835634 | T | C | -0.115 | 0.242 | 0.188 | 0.202 | 2.69E-01 | 0.219 | 0.017 | 5.29E-12 | 11792 | 4.02E-03 | 4.70E+01 | |
| | rs6937696 | T | C | -0.081 | -0.026 | 0.651 | 0.644 | 8.86E-01 | 0.185 | 0.012 | 1.40E-11 | 14742 | 3.09E-03 | 4.57E+01 | |
| | rs712137 | A | C | 0.056 | -0.349 | 0.660 | 0.728 | 7.74E-02 | 0.198 | 0.012 | 2.71E-06 | 14737 | 1.49E-03 | 2.20E+01 | |
| | rs72985859 | T | C | 0.135 | 0.097 | 0.081 | 0.044 | 8.23E-01 | 0.436 | 0.029 | 3.72E-06 | 8297 | 2.57E-03 | 2.14E+01 | |
| | rs73137770 | C | G | 0.244 | -1.074 | 0.981 | 0.988 | 2.55E-02 | 0.481 | 0.053 | 3.91E-06 | 13808 | 1.54E-03 | 2.13E+01 | |
| | rs77915710 | A | G | -0.144 | -0.317 | 0.040 | 0.012 | 6.99E-01 | 0.820 | 0.030 | 2.32E-06 | 14742 | 1.51E-03 | 2.23E+01 | |
| | rs8052631 | A | G | -0.151 | -0.042 | 0.946 | 0.954 | 9.22E-01 | 0.426 | 0.027 | 2.19E-08 | 14295 | 2.19E-03 | 3.13E+01 | |
| | rs10978641 | A | T | 0.065 | 0.030 | 0.746 | 0.792 | 8.90E-01 | 0.218 | 0.014 | 3.56E-06 | 14296 | 1.50E-03 | 2.15E+01 | |
| | rs10990535 | T | C | 0.070 | -0.166 | 0.253 | 0.265 | 4.07E-01 | 0.201 | 0.014 | 1.91E-07 | 14734 | 1.84E-03 | 2.71E+01 | |
| | rs11673125 | T | C | -0.059 | -0.056 | 0.296 | 0.248 | 7.84E-01 | 0.205 | 0.013 | 4.10E-06 | 14741 | 1.44E-03 | 2.12E+01 | |
| | rs116889819 | A | G | -0.176 | -0.018 | 0.046 | 0.025 | 9.75E-01 | 0.578 | 0.030 | 4.45E-09 | 14744 | 2.33E-03 | 3.44E+01 | |
| rs12141791 | A | G | 0.086 | 0.212 | 0.698 | 0.634 | 2.45E-01 | 0.183 | 0.013 | 3.90E-11 | 14296 | 3.04E-03 | 4.37E+01 | | |
| rs12367368 | T | G | 0.086 | 0.148 | 0.115 | 0.150 | 5.49E-01 | 0.247 | 0.019 | 3.67E-06 | 14744 | 1.45E-03 | 2.14E+01 | | |
| rs141782294 | A | G | 0.377 | 0.777 | 0.978 | 0.968 | 1.14E-01 | 0.491 | 0.045 | 2.56E-17 | 14731 | 4.84E-03 | 7.16E+01 | | |
| rs188626385 | T | C | -0.251 | 4.244 | 0.018 | 0.001 | 6.42E-02 | 2.293 | 0.053 | 2.20E-06 | 12935 | 1.73E-03 | 2.24E+01 | | |
| rs2155053 | T | C | -0.321 | -0.756 | 0.946 | 0.951 | 5.67E-03 | 0.273 | 0.026 | 4.68E-34 | 14741 | 9.94E-03 | 1.48E+02 | | |
| rs2229629 | A | G | -0.315 | 1.825 | 0.014 | 0.011 | 4.85E-02 | 0.925 | 0.067 | 2.80E-06 | 10887 | 2.01E-03 | 2.19E+01 | | |
| rs2926791 | T | C | -0.070 | -0.185 | 0.344 | 0.310 | 3.28E-01 | 0.189 | 0.013 | 2.31E-08 | 14296 | 2.18E-03 | 3.12E+01 | | |
| rs4734879 | A | G | 0.114 | 0.038 | 0.742 | 0.779 | 6.80E-01 | 0.214 | 0.014 | 1.11E-16 | 14296 | 4.79E-03 | 6.87E+01 | | |
| rs6717478 | A | T | 0.155 | -0.070 | 0.966 | 0.962 | 8.81E-01 | 0.471 | 0.034 | 4.65E-06 | 14742 | 1.42E-03 | 2.10E+01 | | |
| rs7219205 | A | G | 0.057 | -0.157 | 0.502 | 0.579 | 3.85E-01 | 0.180 | 0.012 | 2.57E-06 | 14296 | 1.54E-03 | 2.21E+01 | | |
| rs77095316 | T | G | -0.142 | -0.380 | 0.071 | 0.050 | 3.51E-01 | 0.408 | 0.024 | 2.69E-09 | 14744 | 2.40E-03 | 3.54E+01 | | |
| rs873675 | T | C | 0.123 | -0.153 | 0.068 | 0.092 | 6.21E-01 | 0.308 | 0.023 | 1.34E-07 | 14742 | 1.88E-03 | 2.78E+01 | | |
| rs10212190 | A | T | 0.073 | -0.007 | 0.615 | 0.604 | 9.71E-01 | 0.180 | 0.012 | 1.88E-09 | 14285 | 2.52E-03 | 3.61E+01 | | |
| rs10499604 | T | C | 0.058 | -0.098 | 0.327 | 0.313 | 6.04E-01 | 0.189 | 0.013 | 3.48E-06 | 14730 | 1.46E-03 | 2.15E+01 | | |
| rs111865527 | T | C | 0.271 | 0.010 | 0.986 | 0.985 | 9.89E-01 | 0.709 | 0.057 | 1.88E-06 | 14244 | 1.59E-03 | 2.27E+01 | | |
| rs112742549 | T | C | 0.242 | 0.988 | 0.979 | 0.992 | 2.97E-01 | 0.948 | 0.051 | 1.71E-06 | 13499 | 1.69E-03 | 2.29E+01 | | |
| rs12058421 | A | C | -0.126 | -0.175 | 0.162 | 0.166 | 4.64E-01 | 0.238 | 0.016 | 2.80E-15 | 14731 | 4.22E-03 | 6.24E+01 | | |
| rs13014154 | A | G | -0.061 | 0.028 | 0.370 | 0.357 | 8.78E-01 | 0.183 | 0.012 | 6.50E-07 | 14285 | 1.73E-03 | 2.48E+01 | | |
| rs13147502 | A | G | -0.061 | -0.355 | 0.596 | 0.537 | 4.52E-02 | 0.177 | 0.013 | 1.62E-06 | 12843 | 1.79E-03 | 2.30E+01 | | |
| rs138657007 | A | G | -0.286 | 1.225 | 0.018 | 0.008 | 2.73E-02 | 0.555 | 0.058 | 6.98E-07 | 12790 | 1.92E-03 | 2.46E+01 | | |
| rs147616029 | T | C | -0.064 | 0.213 | 0.377 | 0.370 | 2.43E-01 | 0.182 | 0.013 | 4.87E-07 | 12850 | 1.97E-03 | 2.53E+01 | | |
| rs149092077 | A | G | -0.359 | 1.075 | 0.989 | 0.996 | 4.55E-01 | 1.440 | 0.073 | 8.63E-07 | 11137 | 2.17E-03 | 2.42E+01 | | |
| rs1609310 | T | G | -0.056 | -0.274 | 0.514 | 0.362 | 1.37E-01 | 0.184 | 0.012 | 3.46E-06 | 14286 | 1.51E-03 | 2.15E+01 | | |
| rs17720330 | C | G | -0.198 | 1.029 | 0.021 | 0.018 | 1.07E-01 | 0.639 | 0.043 | 4.04E-06 | 14727 | 1.44E-03 | 2.12E+01 | | |
| rs19245879 | A | G | 0.333 | -2.551 | 0.989 | 0.996 | 5.62E-02 | 1.536 | 0.066 | 3.76E-07 | 13799 | 1.87E-03 | 2.58E+01 | | |
| rs2623960 | T | C | 0.069 | -0.220 | 0.220 | 0.243 | 2.83E-01 | 0.205 | 0.014 | 1.50E-06 | 14286 | 1.62E-03 | 2.31E+01 | | |
| rs2790462 | T | C | -0.075 | 0.255 | 0.298 | 0.228 | 2.22E-01 | 0.209 | 0.013 | 3.06E-09 | 14733 | 2.38E-03 | 3.51E+01 | | |
| rs3184504 | T | C | 0.097 | -0.118 | 0.476 | 0.407 | 5.09E-01 | 0.179 | 0.013 | 8.55E-14 | 11783 | 4.70E-03 | 5.57E+01 | | |
| rs35271532 | A | G | -0.149 | -0.701 | 0.045 | 0.026 | 2.01E-01 | 0.549 | 0.032 | 4.51E-06 | 14286 | 1.47E-03 | 2.10E+01 | | |
| rs3745163 | A | C | 0.067 | -0.082 | 0.230 | 0.242 | 6.88E-01 | 0.205 | 0.015 | 3.35E-06 | 13669 | 1.58E-03 | 2.16E+01 | | |
| rs3806818 | A | G | 0.071 | -0.031 | 0.191 | 0.194 | 8.90E-01 | 0.224 | 0.015 | 3.82E-06 | 13853 | 1.54E-03 | 2.13E+01 | | |
| rs56244095 | A | C | 0.101 | -0.176 | 0.212 | 0.282 | 3.74E-01 | 0.198 | 0.016 | 8.38E-11 | 14286 | 2.94E-03 | 4.22E+01 | | |
| rs60094514 | T | C | 0.221 | -0.554 | 0.851 | 0.817 | 1.56E-02 | 0.229 | 0.016 | 2.45E-42 | 14732 | 1.25E-02 | 1.86E+02 | | |
| rs73151926 | T | C | 0.106 | -0.387 | 0.081 | 0.052 | 3.34E-01 | 0.401 | 0.023 | 3.45E-06 | 14286 | 1.51E-03 | 2.15E+01 | | |
| rs73231041 | C | G | -0.279 | 0.510 | 0.015 | 0.006 | 6.82E-01 | 1.245 | 0.060 | 3.05E-06 | 12735 | 1.71E-03 | 2.18E+01 | | |
| rs7745783 | T | C | -0.059 | -0.277 | 0.497 | 0.506 | 1.14E-01 | 0.175 | 0.012 | 7.44E-07 | 14286 | 1.71E-03 | 2.45E+01 | | |
| rs79348253 | A | G | -0.159 | 0.237 | 0.045 | 0.063 | 5.14E-01 | 0.364 | 0.030 | 7.75E-08 | 14719 | 1.96E-03 | 2.89E+01 | | |
| rs117427219 | A | G | -0.306 | 2.203 | 0.017 | 0.002 | 2.22E-01 | 1.806 | 0.063 | 1.13E-06 | 10850 | 2.18E-03 | 2.37E+01 | | |
| rs12328100 | A | G | -0.065 | 0.299 | 0.478 | 0.475 | 8.88E-02 | 0.176 | 0.013 | 5.97E-07 | 11784 | 2.11E-03 | 2.49E+01 | | |
| rs12354434 | A | G | -0.121 | -0.394 | 0.089 | 0.092 | 2.04E-01 | 0.310 | 0.025 | 1.03E-06 | 11337 | 2.10E-03 | 2.39E+01 | | |
| rs1266163 | A | T | 0.137 | -0.221 | 0.062 | 0.030 | 6.68E-01 | 0.514 | 0.029 | 1.90E-06 | 11337 | 2.00E-03 | 2.27E+01 | | |
| rs12975019 | C | G | -0.113 | 0.056 | 0.894 | 0.892 | 8.44E-01 | 0.286 | 0.023 | 6.14E-07 | 10278 | 2.41E-03 | 2.49E+01 | | |
| rs1377015 | A | G | -0.065 | 0.109 | 0.286 | 0.292 | 5.79E-01 | 0.196 | 0.014 | 4.25E-06 | 11783 | 1.79E-03 | 2.11E+01 | | |
| rs138684734 | A | G | 0.251 | -1.180 | 0.980 | 0.994 | 3.22E-01 | 1.190 | 0.053 | 2.44E-06 | 10850 | 2.04E-03 | 2.22E+01 | | |
| rs141295666 | T | C | 0.341 | -1.001 | 0.013 | 0.006 | 3.99E-01 | 1.186 | 0.066 | 2.48E-07 | 11337 | 2.34E-03 | 2.66E+01 | | |
| rs148560093 | A | G | -0.298 | 1.065 | 0.982 | 0.996 | 4.30E-01 | 1.351 | 0.065 | 3.97E-06 | 9984 | 2.13E-03 | 2.13E+01 | | |
| rs201738671 | CT | C | -0.349 | -0.095 | 0.015 | 0.009 | 9.21E-01 | 0.965 | 0.075 | 3.64E-06 | 7943 | 2.69E-03 | 2.14E+01 | | |
| rs3338335 | A | T | 0.129 | 0.033 | 0.916 | 0.934 | 9.27E-01 | 0.362 | 0.025 | 3.88E-07 | 1133 | | | | |

| SNPs information of inflammatory proteins with Phantom limb syndrome with pain | | | | | | | | | | | | | | | |
|--|-----------------|---------------|--------------|---------------|--------------|--------------|-------------|--------------|------------|-------------|---------------|---------------------|----------|----------|----------|
| Exposure | SNP | effect_allele | other_allele | beta.exposure | beta.outcome | eaf.exposure | eaf.outcome | pval.outcome | se.outcome | se.exposure | pval.exposure | samplesize.exposure | R2 | f | |
| FGF21 | rs10495032 | T | G | -0.156 | -0.236 | 0.038 | 0.072 | 4.00E-01 | 0.280 | 0.032 | 9.77E-07 | 14720 | 1.63E-03 | 2.40E+01 | |
| | rs12290350 | T | C | -0.069 | -0.113 | 0.217 | 0.180 | 5.65E-01 | 0.196 | 0.015 | 2.02E-06 | 14295 | 1.58E-03 | 2.26E+01 | |
| | rs1260326 | T | C | 0.132 | -0.085 | 0.398 | 0.349 | 5.82E-01 | 0.155 | 0.012 | 1.03E-28 | 14730 | 8.32E-03 | 1.24E+02 | |
| | rs13229619 | A | G | -0.161 | -0.337 | 0.130 | 0.129 | 1.28E-01 | 0.222 | 0.018 | 6.13E-20 | 14743 | 5.64E-03 | 8.36E+01 | |
| | rs142603673 | T | C | -0.235 | 1.037 | 0.023 | 0.009 | 2.12E-01 | 0.832 | 0.046 | 2.35E-07 | 14252 | 1.87E-03 | 2.67E+01 | |
| | rs145127946 | A | G | -0.175 | -0.030 | 0.037 | 0.008 | 9.76E-01 | 0.969 | 0.036 | 1.22E-06 | 14295 | 1.64E-03 | 2.35E+01 | |
| | rs145319387 | G | GT | -0.076 | -0.110 | 0.769 | 0.685 | 4.89E-01 | 0.159 | 0.016 | 3.06E-06 | 10893 | 2.00E-03 | 2.18E+01 | |
| | rs188758663 | A | G | -0.380 | -1.032 | 0.010 | 0.005 | 3.51E-01 | 1.105 | 0.075 | 3.88E-07 | 11668 | 2.20E-03 | 2.57E+01 | |
| | rs2429473 | A | C | -0.081 | -0.185 | 0.795 | 0.808 | 3.24E-01 | 0.188 | 0.015 | 3.32E-08 | 14295 | 2.13E-03 | 3.05E+01 | |
| | rs4700382 | T | C | -0.063 | -0.120 | 0.270 | 0.271 | 4.77E-01 | 0.169 | 0.014 | 3.62E-06 | 14295 | 1.50E-03 | 2.15E+01 | |
| | rs60277384 | T | C | -0.088 | -0.064 | 0.892 | 0.850 | 7.55E-01 | 0.205 | 0.019 | 3.54E-06 | 14742 | 1.46E-03 | 2.15E+01 | |
| | rs7012637 | A | G | -0.061 | -0.197 | 0.481 | 0.508 | 1.86E-01 | 0.149 | 0.012 | 4.61E-07 | 14295 | 1.77E-03 | 2.54E+01 | |
| | rs72965996 | A | G | -0.222 | -0.206 | 0.031 | 0.021 | 6.98E-01 | 0.531 | 0.044 | 6.08E-07 | 13808 | 1.80E-03 | 2.49E+01 | |
| | rs7610704 | T | C | -0.057 | 0.112 | 0.464 | 0.526 | 4.55E-01 | 0.149 | 0.012 | 2.21E-06 | 14295 | 1.56E-03 | 2.24E+01 | |
| | rs838131 | A | C | 0.163 | 0.255 | 0.501 | 0.429 | 9.07E-02 | 0.151 | 0.013 | 6.15E-36 | 14294 | 1.08E-02 | 1.57E+02 | |
| | Osteoprotegerin | rs10142466 | A | G | 0.057 | 0.316 | 0.490 | 0.514 | 1.80E-02 | 0.134 | 0.011 | 1.70E-07 | 14721 | 1.85E-03 | 2.73E+01 |
| | | rs112635299 | T | G | 0.262 | 0.100 | 0.021 | 0.020 | 8.51E-01 | 0.531 | 0.043 | 1.48E-09 | 14733 | 2.48E-03 | 3.66E+01 |
| rs114165349 | | C | G | -0.195 | -0.175 | 0.034 | 0.037 | 6.46E-01 | 0.382 | 0.034 | 1.01E-08 | 14727 | 2.22E-03 | 3.28E+01 | |
| rs1230164 | | C | G | -0.067 | 0.092 | 0.729 | 0.752 | 5.95E-01 | 0.174 | 0.013 | 1.98E-07 | 14285 | 1.89E-03 | 2.71E+01 | |
| rs138408096 | | A | G | -0.201 | -0.019 | 0.021 | 0.013 | 9.79E-01 | 0.711 | 0.042 | 2.09E-06 | 14721 | 1.53E-03 | 2.25E+01 | |
| rs138557324 | | T | C | -0.225 | 0.299 | 0.020 | 0.005 | 7.96E-01 | 1.157 | 0.047 | 1.91E-06 | 14285 | 1.59E-03 | 2.27E+01 | |
| rs1437633 | | A | C | -0.079 | -0.147 | 0.207 | 0.151 | 4.86E-01 | 0.212 | 0.015 | 7.69E-08 | 12851 | 2.24E-03 | 2.89E+01 | |
| rs17600346 | | T | C | -0.157 | 0.304 | 0.963 | 0.954 | 3.95E-01 | 0.357 | 0.030 | 1.54E-07 | 14723 | 1.87E-03 | 2.75E+01 | |
| rs2247769 | | T | C | -0.171 | -0.149 | 0.558 | 0.596 | 3.25E-01 | 0.151 | 0.011 | 1.42E-50 | 14285 | 1.54E-02 | 2.24E+02 | |
| rs271665 | | T | C | -0.067 | -0.390 | 0.748 | 0.681 | 1.45E-02 | 0.159 | 0.013 | 1.90E-07 | 14710 | 1.84E-03 | 2.71E+01 | |
| rs28454196 | | C | G | -0.224 | -0.833 | 0.976 | 0.977 | 1.54E-02 | 0.344 | 0.049 | 3.97E-06 | 14285 | 1.49E-03 | 2.13E+01 | |
| rs376613127 | | A | G | -0.269 | 0.264 | 0.013 | 0.012 | 6.96E-01 | 0.676 | 0.057 | 2.14E-06 | 13154 | 1.70E-03 | 2.25E+01 | |
| rs61968587 | | A | G | -0.267 | -0.060 | 0.984 | 0.994 | 9.53E-01 | 1.034 | 0.057 | 2.47E-06 | 13798 | 1.61E-03 | 2.22E+01 | |
| rs6985207 | | A | C | 0.052 | -0.001 | 0.556 | 0.587 | 9.93E-01 | 0.152 | 0.011 | 4.67E-06 | 14285 | 1.47E-03 | 2.10E+01 | |
| rs704 | | A | G | -0.140 | -0.073 | 0.473 | 0.418 | 6.30E-01 | 0.151 | 0.011 | 1.14E-36 | 14733 | 1.07E-02 | 1.60E+02 | |
| rs74111611 | | T | C | -0.147 | -0.784 | 0.960 | 0.978 | 2.70E-02 | 0.354 | 0.030 | 7.96E-07 | 14732 | 1.65E-03 | 2.44E+01 | |
| rs77507094 | | A | G | 0.197 | -0.621 | 0.979 | 0.964 | 1.28E-01 | 0.408 | 0.043 | 4.64E-06 | 14726 | 1.42E-03 | 2.10E+01 | |
| rs946923 | T | C | 0.172 | 1.015 | 0.971 | 0.989 | 2.03E-01 | 0.798 | 0.038 | 4.77E-06 | 13230 | 1.58E-03 | 2.09E+01 | | |

Supplementary Table S4: SNPs used as instrumental variables in Mendelian Randomization analysis of inflammatory proteins Phantom limb syndrome with pain.

| Exposure | SNP | effect_allele | other_allele | SNPs information of inflammatory proteins with Small fibre neuropathy | | | | | | | | | | R2 | f |
|-------------|-------------|---------------|--------------|---|--------------|---------------|--------------|---------------|------------|-------------|----------------|----------------------|----------|----------|---|
| | | | | beta.exposure | beta.outcome | ca.f.exposure | ca.f.outcome | p.val.outcome | se.outcome | se.exposure | p.val.exposure | sample.size.exposure | | | |
| CD40LG | rs117191103 | T | C | 0.355 | 0.118 | 0.021 | 0.028 | 4.16E-01 | 0.146 | 0.044 | 4.28E-16 | 14736 | 4.47E-03 | 6.61E+01 | |
| | rs12152184 | A | G | 0.109 | -0.025 | 0.885 | 0.886 | 7.38E-01 | 0.075 | 0.021 | 1.21E-07 | 12402 | 2.25E-03 | 2.80E+01 | |
| | rs12500563 | T | G | 0.314 | 0.551 | 0.988 | 0.998 | 2.51E-01 | 0.480 | 0.069 | 4.76E-06 | 10795 | 1.94E-03 | 2.09E+01 | |
| | rs138897138 | A | G | 0.109 | -0.097 | 0.914 | 0.916 | 2.56E-01 | 0.085 | 0.022 | 4.50E-07 | 14288 | 1.78E-03 | 2.55E+01 | |
| | rs1883832 | T | C | -0.452 | 0.167 | 0.258 | 0.271 | 1.24E-03 | 0.052 | 0.013 | 1.00E-200 | 14736 | 8.03E-02 | 1.22E+03 | |
| | rs3967170 | A | G | 0.079 | -0.028 | 0.135 | 0.159 | 6.72E-01 | 0.065 | 0.017 | 3.46E-06 | 14721 | 1.46E-03 | 2.15E+01 | |
| | rs404842 | A | G | -0.060 | 0.045 | 0.643 | 0.596 | 3.55E-01 | 0.048 | 0.012 | 1.12E-06 | 14287 | 1.66E-03 | 2.37E+01 | |
| | rs4278412 | C | G | 0.198 | 0.201 | 0.020 | 0.039 | 1.04E-01 | 0.123 | 0.041 | 1.67E-06 | 14725 | 1.56E-03 | 2.29E+01 | |
| | rs4664334 | T | C | 0.054 | 0.100 | 0.500 | 0.484 | 3.60E-02 | 0.048 | 0.012 | 4.55E-06 | 14288 | 1.47E-03 | 2.10E+01 | |
| | rs7250371 | T | C | 0.069 | 0.075 | 0.365 | 0.312 | 1.41E-01 | 0.051 | 0.012 | 3.34E-09 | 14722 | 2.37E-03 | 3.50E+01 | |
| | rs73128369 | A | T | -0.179 | 0.132 | 0.941 | 0.944 | 2.07E-01 | 0.104 | 0.027 | 2.53E-11 | 14288 | 3.11E-03 | 4.45E+01 | |
| | rs74794857 | T | C | 0.071 | -0.065 | 0.413 | 0.375 | 1.89E-01 | 0.050 | 0.014 | 3.00E-07 | 11760 | 2.23E-03 | 2.62E+01 | |
| | rs75626299 | C | G | 0.146 | 0.017 | 0.964 | 0.968 | 8.99E-01 | 0.136 | 0.032 | 3.95E-06 | 14728 | 1.44E-03 | 2.13E+01 | |
| | rs8024256 | A | G | -0.056 | 0.117 | 0.309 | 0.293 | 2.44E-02 | 0.052 | 0.012 | 3.41E-06 | 14736 | 1.46E-03 | 2.16E+01 | |
| | rs9526031 | A | C | 0.064 | 0.009 | 0.488 | 0.593 | 8.60E-01 | 0.049 | 0.012 | 9.64E-08 | 14288 | 1.99E-03 | 2.84E+01 | |
| | rs9993874 | A | G | -0.058 | 0.045 | 0.488 | 0.487 | 3.45E-01 | 0.047 | 0.013 | 3.22E-06 | 12403 | 1.74E-03 | 2.17E+01 | |
| | rs113153423 | A | G | 0.191 | 0.266 | 0.972 | 0.981 | 1.19E-01 | 0.171 | 0.041 | 2.60E-06 | 11792 | 1.87E-03 | 2.21E+01 | |
| | rs113563938 | A | T | 0.318 | 0.671 | 0.987 | 0.995 | 5.99E-02 | 0.351 | 0.068 | 2.62E-06 | 11344 | 1.94E-03 | 2.21E+01 | |
| | rs11987533 | A | G | 0.117 | 0.144 | 0.137 | 0.151 | 2.96E-02 | 0.066 | 0.020 | 3.34E-09 | 11344 | 3.07E-03 | 3.50E+01 | |
| | rs12056343 | T | G | -0.076 | 0.040 | 0.749 | 0.721 | 4.54E-01 | 0.053 | 0.017 | 4.55E-06 | 9894 | 2.12E-03 | 2.10E+01 | |
| rs12138611 | C | G | 0.119 | 0.037 | 0.104 | 0.132 | 6.06E-01 | 0.071 | 0.026 | 3.34E-06 | 11344 | 1.90E-03 | 2.16E+01 | | |
| rs143067549 | T | C | -0.417 | -0.161 | 0.011 | 0.002 | 7.21E-01 | 0.450 | 0.091 | 4.69E-06 | 7315 | 2.86E-03 | 2.10E+01 | | |
| rs149111746 | A | G | 0.236 | 0.080 | 0.021 | 0.021 | 6.30E-01 | 0.166 | 0.051 | 3.98E-06 | 11783 | 1.80E-03 | 2.13E+01 | | |
| rs28846828 | T | C | -0.077 | 0.021 | 0.423 | 0.365 | 6.74E-01 | 0.049 | 0.015 | 6.78E-07 | 8817 | 2.79E-03 | 2.47E+01 | | |
| rs59446445 | A | G | 0.174 | 0.272 | 0.032 | 0.022 | 9.63E-02 | 0.163 | 0.038 | 4.61E-06 | 11790 | 1.78E-03 | 2.10E+01 | | |
| rs72785422 | T | C | 0.090 | 0.099 | 0.831 | 0.860 | 1.52E-01 | 0.069 | 0.020 | 4.22E-06 | 11343 | 1.86E-03 | 2.12E+01 | | |
| rs11256610 | A | G | 0.301 | 1.108 | 0.015 | 0.001 | 8.60E-02 | 0.645 | 0.064 | 2.76E-06 | 10850 | 2.02E-03 | 2.20E+01 | | |
| rs116790447 | T | C | 0.173 | 0.106 | 0.052 | 0.039 | 3.95E-01 | 0.124 | 0.037 | 3.16E-06 | 9453 | 2.29E-03 | 2.17E+01 | | |
| rs117613902 | A | G | -0.300 | -0.496 | 0.986 | 0.996 | 1.87E-01 | 0.376 | 0.066 | 4.97E-06 | 10850 | 1.92E-03 | 2.08E+01 | | |
| rs143986178 | A | G | 0.297 | 0.320 | 0.014 | 0.003 | 4.71E-01 | 0.444 | 0.064 | 4.62E-06 | 10850 | 1.96E-03 | 2.13E+01 | | |
| rs145295031 | A | T | 0.306 | 0.030 | 0.014 | 0.013 | 8.89E-01 | 0.213 | 0.065 | 2.20E-06 | 11287 | 1.98E-03 | 2.24E+01 | | |
| rs167018 | A | G | 0.066 | -0.050 | 0.444 | 0.458 | 2.94E-01 | 0.048 | 0.014 | 1.36E-06 | 11336 | 2.05E-03 | 2.33E+01 | | |
| rs2341294 | A | G | -0.132 | -0.245 | 0.072 | 0.083 | 6.95E-03 | 0.091 | 0.028 | 2.51E-06 | 9900 | 2.23E-03 | 2.22E+01 | | |
| rs60306270 | A | G | -0.089 | -0.141 | 0.140 | 0.087 | 9.46E-02 | 0.084 | 0.019 | 3.51E-06 | 11781 | 1.82E-03 | 2.15E+01 | | |
| rs73169406 | A | G | 0.111 | 0.033 | 0.133 | 0.105 | 6.69E-01 | 0.078 | 0.021 | 8.22E-08 | 11337 | 2.53E-03 | 2.87E+01 | | |
| rs7401175 | A | T | 0.089 | 0.023 | 0.239 | 0.164 | 7.20E-01 | 0.065 | 0.020 | 4.78E-06 | 7943 | 2.63E-03 | 2.09E+01 | | |
| rs112168535 | T | C | -0.155 | 0.357 | 0.956 | 0.961 | 7.92E-03 | 0.134 | 0.032 | 8.90E-07 | 13670 | 1.76E-03 | 2.41E+01 | | |
| rs12075 | A | G | -0.097 | 0.039 | 0.555 | 0.532 | 4.09E-01 | 0.048 | 0.012 | 4.95E-17 | 14733 | 4.75E-03 | 7.03E+01 | | |
| rs12451323 | T | C | 1.025 | -0.147 | 0.884 | 0.895 | 5.69E-02 | 0.077 | 0.017 | 1.00E-200 | 14288 | 1.95E-01 | 3.47E+03 | | |
| rs12523126 | A | T | 0.112 | 0.096 | 0.075 | 0.066 | 3.18E-01 | 0.096 | 0.023 | 1.35E-06 | 14288 | 1.63E-03 | 2.33E+01 | | |
| rs138884871 | T | C | 0.220 | -0.227 | 0.019 | 0.016 | 2.31E-01 | 0.189 | 0.047 | 3.37E-06 | 13672 | 1.58E-03 | 2.16E+01 | | |
| rs144618566 | T | C | 0.517 | -0.144 | 0.017 | 0.004 | 7.10E-01 | 0.388 | 0.054 | 7.15E-22 | 13801 | 6.65E-03 | 9.24E+01 | | |
| rs145516279 | A | G | 0.223 | -0.108 | 0.979 | 0.982 | 5.48E-01 | 0.180 | 0.048 | 3.56E-06 | 14288 | 1.50E-03 | 2.15E+01 | | |
| rs181726805 | A | G | -0.248 | -0.144 | 0.984 | 0.973 | 3.30E-01 | 0.148 | 0.054 | 3.75E-06 | 14736 | 1.45E-03 | 2.14E+01 | | |
| rs35904471 | T | C | 0.076 | 0.014 | 0.266 | 0.329 | 7.91E-01 | 0.051 | 0.017 | 3.99E-06 | 11337 | 1.87E-03 | 2.13E+01 | | |
| rs407595 | A | C | 0.070 | 0.006 | 0.207 | 0.240 | 9.17E-01 | 0.055 | 0.015 | 1.22E-06 | 14287 | 1.57E-03 | 2.25E+01 | | |
| rs41274294 | T | C | 0.882 | 0.097 | 0.968 | 0.970 | 4.84E-01 | 0.139 | 0.036 | 2.19E-135 | 14732 | 4.00E-02 | 6.13E+02 | | |
| rs4722985 | T | C | -0.063 | -0.024 | 0.709 | 0.788 | 6.86E-01 | 0.058 | 0.013 | 2.52E-06 | 14288 | 1.55E-03 | 2.22E+01 | | |
| rs55653410 | A | T | -0.174 | -0.036 | 0.042 | 0.008 | 8.90E-01 | 0.262 | 0.033 | 1.86E-07 | 14288 | 1.90E-03 | 2.72E+01 | | |
| rs62062305 | T | G | 0.082 | 0.095 | 0.855 | 0.854 | 1.58E-01 | 0.067 | 0.017 | 1.16E-06 | 14288 | 1.65E-03 | 2.36E+01 | | |
| rs6685547 | A | G | 0.146 | -0.102 | 0.051 | 0.032 | 4.57E-01 | 0.137 | 0.029 | 4.22E-07 | 13239 | 1.93E-03 | 2.56E+01 | | |
| rs72827004 | C | G | 0.262 | -0.195 | 0.046 | 0.050 | 7.47E-02 | 0.110 | 0.032 | 1.51E-16 | 14728 | 4.61E-03 | 6.81E+01 | | |
| rs74350159 | A | G | 0.280 | -0.363 | 0.012 | 0.009 | 1.34E-01 | 0.242 | 0.061 | 4.19E-06 | 13801 | 1.53E-03 | 2.12E+01 | | |
| rs75573205 | C | G | 0.191 | -0.083 | 0.976 | 0.962 | 5.09E-01 | 0.125 | 0.041 | 2.99E-06 | 14725 | 1.48E-03 | 2.18E+01 | | |
| rs75580960 | T | C | -0.090 | 0.048 | 0.870 | 0.853 | 4.80E-01 | 0.067 | 0.019 | 1.52E-06 | 13224 | 1.75E-03 | 2.31E+01 | | |
| rs78743764 | A | C | -0.223 | 0.400 | 0.019 | 0.017 | 7.77E-02 | 0.182 | 0.048 | 3.53E-06 | 13656 | 1.57E-03 | 2.15E+01 | | |
| rs79567569 | A | C | -0.328 | -0.285 | 0.988 | 0.997 | 4.87E-01 | 0.410 | 0.067 | 9.16E-07 | 13386 | 1.80E-03 | 2.41E+01 | | |
| rs80131163 | A | G | 0.196 | -0.099 | 0.969 | 0.941 | 3.29E-01 | 0.101 | 0.040 | 8.77E-07 | 14730 | 1.64E-03 | 2.42E+01 | | |
| rs948962 | A | C | 0.054 | -0.035 | 0.476 | 0.558 | 4.62E-01 | 0.048 | 0.012 | 4.84E-06 | 14736 | 1.42E-03 | 2.09E+01 | | |
| rs9889517 | A | G | 0.061 | -0.039 | 0.478 | 0.405 | 4.18E-01 | 0.049 | 0.012 | 4.83E-07 | 14288 | 1.77E-03 | 2.53E+01 | | |
| rs116928468 | A | G | -0.257 | 0.129 | 0.019 | 0.062 | 1.96E-01 | 0.100 | 0.054 | 1.69E-06 | 11770 | 1.94E-03 | 2.29E+01 | | |
| rs117867445 | C | G | 0.293 | -0.109 | 0.017 | 0.010 | 6.49E-01 | 0.239 | 0.062 | 2.46E-06 | 10471 | 2.12E-03 | 2.22E+01 | | |
| rs2285179 | A | G | -0.066 | -0.011 | 0.559 | 0.561 | 8.16E-01 | 0.048 | 0.012 | 5.50E-08 | 14288 | 2.06E-03 | 2.95E+01 | | |
| rs2511241 | T | C | 0.177 | -0.221 | 0.916 | 0.923 | 1.32E-02 | 0.089 | 0.024 | 6.38E-14 | 13668 | 4.10E-03 | 5.62E+01 | | |
| rs28444486 | A | G | 0.064 | -0.117 | 0.738 | 0.747 | 3.15E-02 | 0.054 | 0.014 | 2.92E-06 | 14735 | 1.48E-03 | 2.19E+01 | | |
| rs3859189 | A | G | 0.110 | -0.024 | 0.537 | 0.558 | 6.21E-01 | 0.048 | 0.012 | 4.95E-21 | 14729 | 5.98E-03 | 8.85E+01 | | |
| rs4823082 | C | G | -0.159 | 0.011 | 0.036 | 0.034 | 9.30E-01 | 0.130 | 0.035 | 4.32E-06 | 14724 | 1.43E-03 | 2.11E+01 | | |
| rs559651 | T | C | -0.059 | 0.108 | 0.650 | 0.631 | 2.76E-02 | 0.049 | 0.013 | 2.01E-06 | 14715 | 1.53E-03 | 2.26E+01 | | |
| rs61897429 | A | G | -0.085 | -0.009 | 0.139 | 0.140 | 8.97E-01 | 0.068 | 0.017 | 6.28E-07 | 14735 | 1.68E-03 | 2.48E+01 | | |
| rs653178 | T | C | -0.073 | -0.047 | 0.522 | 0.586 | 3.30E-01 | 0.048 | 0.013 | 3.48E-08 | 11785 | 2.57E-03 | 3.04E+01 | | |
| rs7102957 | T | C | -0.080 | -0.035 | 0.863 | 0.805 | 5.64E-01 | 0.060 | 0.017 | 3.46E-06 | 14288 | 1.51E-03 | 2.15E+01 | | |
| rs76275710 | A | G | 0.267 | -0.020 | 0.018 | 0.028 | 8.89E-01 | 0.144 | 0.053 | 3.85E-07 | 14735 | 1.75E-03 | 2.58E+01 | | |
| rs80303768 | A | G | -0.152 | -0.008 | 0.037 | 0.012 | 9.70E-01 | 0.214 | 0.033 | 3.35E-06 | 14722 | | | | |

| Exposure | SNP | effect_allele | other_allele | SNPs information of inflammatory proteins with Unspecified neuralgia and neuritis | | | | | | | | | | samplesize/exposure | R2 | f |
|-------------|-------------|---------------|--------------|---|--------------|-------------|------------|--------------|------------|-------------|---------------|----------|----------|---------------------|----|---|
| | | | | beta_exposure | beta_outcome | es_exposure | es_outcome | pval_outcome | se_outcome | se_exposure | pval_exposure | | | | | |
| CCL28 | rs112641086 | T | C | -0.106 | -0.138 | 0.090 | 0.113 | 7.82E-03 | 0.052 | 0.020 | 9.64E-08 | 14731 | 1.93E-03 | 2.84E+01 | | |
| | rs116782490 | A | C | 0.230 | 0.175 | 0.982 | 0.996 | 5.24E-01 | 0.275 | 0.049 | 2.49E-06 | 12633 | 1.73E-03 | 2.22E+01 | | |
| | rs119902 | A | C | 0.068 | -0.068 | 0.366 | 0.366 | 2.61E-01 | 0.054 | 0.112 | 1.53E-06 | 14285 | 1.42E-03 | 2.30E+01 | | |
| | rs143400203 | T | C | 0.181 | -0.061 | 0.024 | 0.032 | 5.00E-01 | 0.090 | 0.039 | 3.79E-06 | 14722 | 1.45E-03 | 2.14E+01 | | |
| | rs1514964 | T | C | -0.126 | -0.044 | 0.949 | 0.959 | 5.82E-01 | 0.080 | 0.026 | 1.72E-06 | 14734 | 1.55E-03 | 2.29E+01 | | |
| | rs199655385 | CT | C | 0.188 | 0.230 | 0.028 | 0.012 | 1.27E-01 | 0.151 | 0.040 | 1.94E-06 | 11336 | 1.99E-03 | 2.26E+01 | | |
| | rs200181753 | T | TG | -0.184 | -0.060 | 0.971 | 0.943 | 3.82E-01 | 0.069 | 0.039 | 2.32E-06 | 11339 | 1.96E-03 | 2.23E+01 | | |
| | rs2479754 | T | C | 0.059 | 0.025 | 0.621 | 0.603 | 2.87E-01 | 0.032 | 0.112 | 1.34E-11 | 14286 | 1.63E-03 | 2.34E+01 | | |
| | rs26231360 | A | C | 0.060 | 0.052 | 0.548 | 0.655 | 1.21E-01 | 0.034 | 0.103 | 3.10E-06 | 11780 | 1.84E-03 | 2.17E+01 | | |
| | rs2866626 | C | G | -0.060 | -0.046 | 0.608 | 0.604 | 1.61E-01 | 0.033 | 0.112 | 8.06E-07 | 14285 | 1.70E-03 | 2.43E+01 | | |
| | rs4866742 | T | C | 0.070 | -0.023 | 0.413 | 0.412 | 4.72E-01 | 0.033 | 0.112 | 8.71E-09 | 12829 | 2.57E-03 | 3.31E+01 | | |
| | rs503990 | A | G | -0.069 | -0.001 | 0.402 | 0.322 | 9.69E-01 | 0.034 | 0.112 | 5.53E-09 | 14285 | 2.37E-03 | 3.40E+01 | | |
| | rs661668 | T | C | 0.053 | 0.051 | 0.541 | 0.546 | 3.03E-01 | 0.032 | 0.112 | 4.14E-06 | 14286 | 1.46E-03 | 2.12E+01 | | |
| | rs7133034 | C | G | 0.218 | 0.003 | 0.980 | 0.991 | 9.83E-01 | 0.172 | 0.047 | 2.87E-06 | 11952 | 1.82E-03 | 2.19E+01 | | |
| | rs7300029 | A | G | -0.194 | -0.071 | 0.035 | 0.030 | 4.63E-01 | 0.046 | 0.036 | 5.02E-08 | 14734 | 2.01E-03 | 2.97E+01 | | |
| | rs76291845 | A | G | 0.270 | 0.243 | 0.986 | 0.985 | 7.13E-02 | 0.135 | 0.054 | 6.53E-07 | 14244 | 1.73E-03 | 2.47E+01 | | |
| | rs7661151 | T | C | 0.053 | 0.055 | 0.632 | 0.627 | 9.93E-02 | 0.033 | 0.112 | 4.70E-06 | 14730 | 1.42E-03 | 2.10E+01 | | |
| | rs7666972 | A | C | -0.285 | 0.132 | 0.018 | 0.002 | 7.29E-01 | 0.381 | 0.051 | 2.63E-08 | 12953 | 2.39E-03 | 3.09E+01 | | |
| | rs7758965 | T | G | 0.139 | 0.066 | 0.956 | 0.969 | 4.78E-01 | 0.082 | 0.028 | 4.77E-06 | 14734 | 1.42E-03 | 2.09E+01 | | |
| | rs78532613 | T | C | -0.184 | -0.103 | 0.027 | 0.021 | 3.66E-01 | 0.113 | 0.040 | 3.04E-06 | 14734 | 1.48E-03 | 2.18E+01 | | |
| | rs7884308 | A | G | -0.168 | -0.081 | 0.038 | 0.023 | 4.62E-01 | 0.111 | 0.032 | 1.83E-07 | 14725 | 1.84E-03 | 2.72E+01 | | |
| | rs80328354 | A | T | -0.284 | 0.132 | 0.988 | 0.991 | 4.42E-01 | 0.171 | 0.060 | 2.11E-06 | 14734 | 1.52E-03 | 2.25E+01 | | |
| | rs1030315 | A | G | -0.056 | -0.023 | 0.443 | 0.446 | 4.71E-01 | 0.032 | 0.112 | 4.61E-06 | 14286 | 1.47E-03 | 2.10E+01 | | |
| | rs1092354 | T | C | -0.045 | -0.017 | 0.997 | 0.997 | 8.24E-01 | 0.035 | 0.076 | 1.23E-04 | 12304 | 1.91E-03 | 2.34E+01 | | |
| | rs1102061 | A | G | 0.077 | 0.038 | 0.817 | 0.777 | 3.33E-01 | 0.029 | 0.116 | 1.84E-06 | 14287 | 1.59E-03 | 2.28E+01 | | |
| | rs11220506 | A | G | -0.070 | 0.024 | 0.188 | 0.260 | 5.12E-01 | 0.027 | 0.115 | 1.79E-06 | 14729 | 1.55E-03 | 2.28E+01 | | |
| | rs11265488 | A | G | -0.062 | 0.018 | 0.468 | 0.442 | 5.83E-01 | 0.032 | 0.112 | 1.24E-07 | 14287 | 1.95E-03 | 2.80E+01 | | |
| | rs11265493 | A | G | 0.212 | 0.040 | 0.407 | 0.396 | 5.43E-03 | 0.032 | 0.112 | 3.99E-71 | 14287 | 2.18E-02 | 3.18E+02 | | |
| | rs114738226 | A | G | -0.180 | 0.043 | 0.039 | 0.045 | 6.23E-01 | 0.087 | 0.034 | 8.65E-08 | 14728 | 1.94E-03 | 2.87E+01 | | |
| | rs139924614 | A | G | 0.162 | 0.230 | 0.967 | 0.992 | 2.15E-01 | 0.185 | 0.035 | 3.33E-06 | 14725 | 1.47E-03 | 2.16E+01 | | |
| rs1434219 | T | C | -0.059 | -0.024 | 0.439 | 0.432 | 4.63E-01 | 0.033 | 0.112 | 1.68E-06 | 14287 | 1.60E-03 | 2.29E+01 | | | |
| rs147033439 | A | G | 0.205 | -0.040 | 0.025 | 0.015 | 7.68E-01 | 0.135 | 0.041 | 5.88E-07 | 14731 | 1.69E-03 | 2.49E+01 | | | |
| rs14948216 | A | C | -0.215 | -0.143 | 0.018 | 0.015 | 2.89E-01 | 0.134 | 0.047 | 4.97E-06 | 14725 | 1.41E-03 | 2.08E+01 | | | |
| rs1578467 | C | G | -0.064 | -0.023 | 0.376 | 0.357 | 3.23E-01 | 0.035 | 0.114 | 4.53E-06 | 14287 | 1.47E-03 | 2.10E+01 | | | |
| rs2554697 | A | G | 0.063 | 0.017 | 0.635 | 0.665 | 6.17E-01 | 0.034 | 0.113 | 2.88E-06 | 11782 | 1.85E-03 | 2.19E+01 | | | |
| rs3184504 | T | C | 0.116 | 0.077 | 0.476 | 0.406 | 1.86E-02 | 0.033 | 0.113 | 1.27E-19 | 11784 | 6.92E-03 | 8.21E+01 | | | |
| rs3426779 | T | G | 0.063 | -0.001 | 0.248 | 0.218 | 9.78E-01 | 0.039 | 0.114 | 4.99E-06 | 14287 | 1.46E-03 | 2.08E+01 | | | |
| rs3828139 | T | C | 0.074 | 0.033 | 0.468 | 0.518 | 2.98E-01 | 0.032 | 0.112 | 4.05E-10 | 14286 | 2.73E-03 | 3.91E+01 | | | |
| rs4672486 | A | G | 0.066 | -0.015 | 0.453 | 0.450 | 6.30E-01 | 0.032 | 0.112 | 1.61E-08 | 14720 | 2.16E-03 | 3.19E+01 | | | |
| rs5272834 | T | C | 0.070 | 0.060 | 0.699 | 0.709 | 3.69E-01 | 0.035 | 0.113 | 3.81E-08 | 14729 | 2.12E-03 | 3.17E+01 | | | |
| rs6928352 | A | G | -0.107 | 0.082 | 0.079 | 0.081 | 1.33E-01 | 0.059 | 0.022 | 5.87E-07 | 14731 | 1.69E-03 | 2.50E+01 | | | |
| rs72837690 | T | C | -0.078 | -0.056 | 0.641 | 0.577 | 8.79E-02 | 0.033 | 0.113 | 5.37E-10 | 14287 | 2.69E-03 | 3.85E+01 | | | |
| rs78142752 | T | C | 0.229 | 0.102 | 0.018 | 0.024 | 3.29E-01 | 0.104 | 0.030 | 4.15E-06 | 14734 | 1.44E-03 | 2.12E+01 | | | |
| rs11120539 | T | C | -0.056 | -0.008 | 0.494 | 0.308 | 8.06E-01 | 0.032 | 0.112 | 2.53E-06 | 14287 | 1.53E-03 | 2.21E+01 | | | |
| rs118117380 | A | G | 0.236 | 0.091 | 0.019 | 0.009 | 5.97E-01 | 0.172 | 0.051 | 3.18E-07 | 14287 | 1.53E-03 | 2.17E+01 | | | |
| rs11930290 | A | T | -0.185 | 0.067 | 0.975 | 0.986 | 6.30E-01 | 0.139 | 0.039 | 2.11E-06 | 12848 | 1.75E-03 | 2.25E+01 | | | |
| rs12195292 | A | G | -0.077 | -0.057 | 0.863 | 0.866 | 4.30E-01 | 0.047 | 0.117 | 3.72E-06 | 14710 | 1.45E-03 | 2.14E+01 | | | |
| rs181409232 | A | C | -0.222 | -0.052 | 0.019 | 0.015 | 6.95E-01 | 0.132 | 0.048 | 3.43E-06 | 12846 | 1.68E-03 | 2.16E+01 | | | |
| rs20050353 | T | TA | 0.226 | -0.105 | 0.979 | 0.995 | 6.49E-01 | 0.242 | 0.047 | 1.27E-06 | 10852 | 2.16E-03 | 2.35E+01 | | | |
| rs2391462 | A | G | -0.053 | -0.023 | 0.237 | 0.215 | 3.21E-01 | 0.037 | 0.114 | 1.63E-06 | 14286 | 1.47E-03 | 2.10E+01 | | | |
| rs24515123 | A | G | 0.170 | 0.068 | 0.030 | 0.030 | 4.63E-01 | 0.093 | 0.035 | 9.75E-07 | 14730 | 1.63E-03 | 2.40E+01 | | | |
| rs368215473 | A | G | 0.123 | 0.055 | 0.094 | 0.073 | 3.74E-01 | 0.062 | 0.026 | 1.77E-06 | 9397 | 2.42E-03 | 2.88E+01 | | | |
| rs370203222 | C | G | 0.106 | -0.009 | 0.100 | 0.139 | 8.43E-01 | 0.047 | 0.022 | 1.84E-06 | 11759 | 1.93E-03 | 2.28E+01 | | | |
| rs3811621 | C | G | 0.116 | 0.071 | 0.290 | 0.295 | 4.17E-02 | 0.035 | 0.113 | 6.63E-19 | 14286 | 5.49E-03 | 7.88E+01 | | | |
| rs3120599 | A | T | 0.061 | 0.023 | 0.706 | 0.731 | 5.20E-01 | 0.036 | 0.113 | 1.30E-11 | 14287 | 3.20E-03 | 4.25E+01 | | | |
| rs711205 | T | C | 0.071 | 0.006 | 0.750 | 0.718 | 8.65E-01 | 0.036 | 0.114 | 1.95E-07 | 14287 | 1.89E-03 | 2.71E+01 | | | |
| rs6723743 | A | T | 0.094 | 0.024 | 0.873 | 0.903 | 6.00E-01 | 0.055 | 0.019 | 3.65E-07 | 14287 | 1.81E-03 | 2.59E+01 | | | |
| rs75271339 | A | T | -0.415 | -0.004 | 0.014 | 0.035 | 9.65E-01 | 0.086 | 0.089 | 3.12E-06 | 9947 | 2.18E-03 | 2.17E+01 | | | |
| rs794007 | T | C | 0.069 | 0.031 | 0.241 | 0.216 | 4.28E-01 | 0.039 | 0.114 | 1.97E-06 | 12844 | 1.76E-03 | 2.26E+01 | | | |
| rs11120866 | T | C | -0.060 | -0.014 | 0.604 | 0.615 | 6.65E-01 | 0.033 | 0.113 | 2.40E-06 | 11786 | 1.83E-03 | 2.22E+01 | | | |
| rs112271527 | A | ATGG | 0.444 | 0.340 | 0.844 | 0.837 | 3.19E-01 | 0.049 | 0.013 | 2.58E-07 | 8389 | 3.08E-03 | 3.75E+01 | | | |
| rs112790107 | A | G | 0.509 | -0.026 | 0.066 | 0.027 | 7.98E-01 | 0.106 | 0.106 | 1.64E-06 | 9645 | 2.33E-03 | 2.30E+01 | | | |
| rs11674185 | A | G | -0.059 | -0.013 | 0.364 | 0.352 | 6.90E-01 | 0.034 | 0.113 | 4.69E-06 | 11787 | 1.78E-03 | 2.10E+01 | | | |
| rs117238330 | T | C | 0.227 | -0.197 | 0.022 | 0.012 | 1.93E-01 | 0.152 | 0.045 | 5.67E-07 | 11781 | 2.12E-03 | 2.50E+01 | | | |
| rs1205489 | T | C | 0.151 | -0.041 | 0.046 | 0.047 | 6.53E-01 | 0.035 | 0.032 | 2.76E-06 | 10725 | 2.04E-03 | 2.20E+01 | | | |
| rs1412253 | A | G | -0.091 | -0.047 | 0.838 | 0.836 | 3.72E-01 | 0.051 | 0.019 | 2.20E-06 | 11778 | 1.90E-03 | 2.24E+01 | | | |
| rs1438038 | T | C | 0.516 | -0.072 | 0.317 | 0.347 | 3.42E-02 | 0.034 | 0.113 | 1.00E-200 | 11788 | 1.25E-01 | 1.68E+03 | | | |
| rs148323905 | T | C | 0.288 | 0.264 | 0.017 | 0.014 | 5.99E-02 | 0.138 | 0.061 | 2.12E-06 | 11341 | 1.98E-03 | 2.25E+01 | | | |
| rs148797749 | A | G | -0.171 | 0.017 | 0.036 | 0.047 | 8.21E-01 | 0.077 | 0.036 | 2.32E-06 | 11785 | 1.89E-03 | 2.23E+01 | | | |
| rs15024893 | A | G | 0.255 | 0.047 | 0.985 | 0.983 | 7.11E-01 | 0.128 | 0.055 | 3.71E-06 | 11298 | 1.89E-03 | 2.14E+01 | | | |
| rs18984846 | A | G | 0.025 | 0.004 | 0.025 | 0.015 | 3.98E-01 | 0.066 | 0.066 | 3.91E-06 | 9880 | 2.13E-03 | 2.19E+01 | | | |
| rs192746810 | A | G | -0.184 | -0.050 | 0.046 | 0.039 | 3.11E-01 | 0.084 | 0.040 | 3.11E-06 | 11341 | 1.91E-03 | 2.17E+01 | | | |
| rs20000616 | CA | C | -0.608 | -0.228 | 0.021 | 0.012 | 1.21E-01 | 0.147 | 0.063 | 3.20E-22 | 7943 | | | | | |

| SNPs information of Neuropathic pain with inflammatory proteins | | | | | | | | | | | | | | | |
|---|-----------------------|-------------|---------------|--------------|---------------|--------------|-------------|------------|------------|--------------|-------------|---------------|--------------------|----------|--------|
| Exposure | Inflammatory proteins | SNP | effect_allele | other_allele | beta.exposure | beta.outcome | ef.exposure | ef.outcome | se.outcome | pval.outcome | se.exposure | pval.exposure | sampleize.exposure | R2 | f |
| Glossopharyngeal nerve disease | NT-3 | rs117369515 | T | C | 2.619 | -0.098 | 0.002 | 0.010 | 0.081 | 2.23E-01 | 0.611 | 1.83E-05 | 435435 | 4.22E-05 | 18.357 |
| | | rs144095072 | A | G | 1.229 | -0.097 | 0.034 | 0.019 | 0.049 | 4.75E-02 | 0.283 | 1.42E-05 | 435435 | 4.33E-05 | 18.838 |
| Phantom limb syndrome with pain | IL-10RA | rs114130563 | C | A | 1.093 | -0.024 | 0.043 | 0.057 | 0.030 | 4.14E-01 | 0.236 | 3.62E-06 | 262495 | 8.17E-05 | 21.456 |
| | | rs6113819 | C | T | -0.770 | 0.016 | 0.242 | 0.216 | 0.017 | 3.32E-01 | 0.173 | 9.08E-06 | 262495 | 7.50E-05 | 19.695 |
| | | rs6900341 | G | T | 0.595 | -0.032 | 0.377 | 0.388 | 0.013 | 1.79E-02 | 0.128 | 3.11E-06 | 262495 | 8.28E-05 | 21.748 |
| | | rs114130563 | C | A | 1.093 | -0.018 | 0.043 | 0.057 | 0.030 | 5.39E-01 | 0.236 | 3.62E-06 | 262495 | 8.17E-05 | 21.456 |
| | | rs6113819 | C | T | -0.770 | 0.025 | 0.242 | 0.216 | 0.017 | 1.42E-01 | 0.173 | 9.08E-06 | 262495 | 7.50E-05 | 19.695 |
| | TNFSF12 | rs6900341 | G | T | 0.595 | -0.024 | 0.377 | 0.388 | 0.013 | 7.35E-02 | 0.128 | 3.11E-06 | 262495 | 8.28E-05 | 21.748 |
| | | rs114130563 | C | A | 1.093 | -0.024 | 0.043 | 0.056 | 0.026 | 3.62E-01 | 0.236 | 3.62E-06 | 262495 | 8.17E-05 | 21.456 |
| | | rs6113819 | C | T | -0.770 | 0.008 | 0.242 | 0.214 | 0.015 | 5.93E-01 | 0.173 | 9.08E-06 | 262495 | 7.50E-05 | 19.695 |
| | | rs6900341 | G | T | 0.595 | -0.034 | 0.377 | 0.389 | 0.012 | 4.18E-03 | 0.128 | 3.11E-06 | 262495 | 8.28E-05 | 21.748 |
| | | rs113586666 | A | T | -0.668 | 0.029 | 0.044 | 0.048 | 0.031 | 3.51E-01 | 0.138 | 1.28E-06 | 492478 | 4.76E-05 | 23.449 |
| Small fibre neuropathy | IL-2RB | rs141736886 | T | C | 0.764 | -0.026 | 0.012 | 0.015 | 0.057 | 6.50E-01 | 0.167 | 4.82E-06 | 492478 | 4.25E-05 | 20.907 |
| | | rs150320807 | G | T | 0.481 | -0.047 | 0.039 | 0.0328 | 0.039 | 2.32E-01 | 0.105 | 4.26E-06 | 492478 | 4.29E-05 | 21.144 |
| | | rs1685391 | G | A | -0.259 | 0.006 | 0.252 | 0.293 | 0.014 | 6.70E-01 | 0.056 | 3.73E-06 | 492478 | 4.34E-05 | 21.397 |
| | | rs34795132 | C | G | -0.472 | 0.071 | 0.072 | 0.079 | 0.025 | 4.77E-03 | 0.103 | 4.35E-06 | 492478 | 4.28E-05 | 21.103 |
| | | rs41279051 | T | C | -0.265 | -0.016 | 0.237 | 0.205 | 0.016 | 3.23E-01 | 0.058 | 4.26E-06 | 492478 | 4.29E-05 | 21.145 |
| | | rs6700449 | A | G | 0.227 | 0.004 | 0.317 | 0.269 | 0.015 | 7.89E-01 | 0.049 | 3.49E-06 | 492478 | 4.37E-05 | 21.528 |
| | | rs78722174 | T | C | 0.980 | -0.011 | 0.008 | 0.016 | 0.070 | 8.74E-01 | 0.199 | 8.69E-07 | 492478 | 4.91E-05 | 24.198 |
| | | rs881921 | A | G | 0.225 | -0.010 | 0.604 | 0.541 | 0.013 | 4.65E-01 | 0.049 | 3.73E-06 | 492478 | 4.35E-05 | 21.401 |
| | | rs9539607 | T | C | -0.277 | 0.023 | 0.224 | 0.247 | 0.015 | 1.35E-01 | 0.059 | 2.26E-06 | 492478 | 4.54E-05 | 22.363 |
| | | rs113586666 | A | T | -0.668 | 0.033 | 0.044 | 0.049 | 0.032 | 2.98E-01 | 0.138 | 1.28E-06 | 492478 | 4.76E-05 | 23.449 |
| | IL-5 | rs141736886 | T | C | 0.764 | -0.066 | 0.012 | 0.015 | 0.058 | 2.58E-01 | 0.167 | 4.82E-06 | 492478 | 4.25E-05 | 20.907 |
| | | rs150320807 | G | T | 0.481 | 0.013 | 0.039 | 0.033 | 0.040 | 7.38E-01 | 0.105 | 4.26E-06 | 492478 | 4.29E-05 | 21.144 |
| | | rs1685391 | G | A | -0.259 | 0.011 | 0.252 | 0.293 | 0.015 | 4.61E-01 | 0.056 | 3.73E-06 | 492478 | 4.34E-05 | 21.397 |
| | | rs34795132 | C | G | -0.472 | 0.043 | 0.072 | 0.079 | 0.025 | 8.79E-02 | 0.103 | 4.35E-06 | 492478 | 4.28E-05 | 21.103 |
| | | rs41279051 | T | C | -0.265 | 0.030 | 0.237 | 0.206 | 0.017 | 7.19E-02 | 0.058 | 4.26E-06 | 492478 | 4.29E-05 | 21.145 |
| | | rs6700449 | A | G | 0.227 | -0.011 | 0.317 | 0.271 | 0.015 | 4.81E-01 | 0.049 | 3.49E-06 | 492478 | 4.37E-05 | 21.528 |
| | | rs78722174 | T | C | 0.980 | -0.079 | 0.008 | 0.016 | 0.070 | 2.58E-01 | 0.199 | 8.69E-07 | 492478 | 4.91E-05 | 24.198 |
| | | rs881921 | A | G | 0.225 | -0.026 | 0.604 | 0.541 | 0.013 | 5.51E-02 | 0.049 | 3.73E-06 | 492478 | 4.35E-05 | 21.401 |
| | | rs9539607 | T | C | -0.277 | 0.005 | 0.224 | 0.247 | 0.015 | 7.60E-01 | 0.059 | 2.26E-06 | 492478 | 4.54E-05 | 22.363 |
| | | rs114767765 | T | C | 0.427 | 0.018 | 0.022 | 0.019 | 0.046 | 7.01E-01 | 0.093 | 4.64E-06 | 359521 | 5.84E-05 | 20.981 |
| Neuralgia and neuritis, unspecified | CCL23 | rs116769657 | T | C | 0.584 | -0.029 | 0.012 | 0.018 | 0.054 | 5.97E-01 | 0.120 | 1.20E-06 | 359521 | 6.56E-05 | 23.585 |
| | | rs117432663 | A | G | 0.217 | -0.011 | 0.121 | 0.071 | 0.023 | 6.32E-01 | 0.046 | 2.61E-06 | 359521 | 6.14E-05 | 22.086 |
| | | rs1338460 | G | A | 0.154 | -0.011 | 0.614 | 0.655 | 0.013 | 3.87E-01 | 0.033 | 3.48E-06 | 359521 | 5.99E-05 | 21.530 |
| | | rs142461741 | C | T | -0.644 | 0.018 | 0.020 | 0.022 | 0.043 | 6.70E-01 | 0.138 | 2.92E-06 | 359521 | 6.08E-05 | 21.870 |
| | | rs150102867 | G | A | 0.506 | 0.073 | 0.018 | 0.014 | 0.063 | 2.49E-01 | 0.106 | 1.64E-06 | 359521 | 6.39E-05 | 22.971 |
| | | rs1522153 | A | G | 0.163 | -0.015 | 0.685 | 0.705 | 0.013 | 2.39E-01 | 0.035 | 3.19E-06 | 359521 | 6.03E-05 | 21.696 |
| | | rs1854137 | G | C | -0.149 | 0.010 | 0.623 | 0.601 | 0.012 | 4.29E-01 | 0.032 | 4.03E-06 | 359521 | 5.91E-05 | 21.251 |
| | | rs186696265 | T | C | 0.539 | 0.006 | 0.016 | 0.018 | 0.050 | 9.00E-01 | 0.107 | 4.35E-07 | 359521 | 7.10E-05 | 25.530 |
| | | rs1919935 | C | T | -0.163 | 0.004 | 0.319 | 0.281 | 0.013 | 7.45E-01 | 0.035 | 3.06E-06 | 359521 | 6.06E-05 | 21.779 |
| | | rs2811465 | C | T | -0.185 | 0.013 | 0.215 | 0.230 | 0.015 | 3.88E-01 | 0.040 | 4.03E-06 | 359521 | 5.91E-05 | 21.250 |
| | | rs28537856 | G | C | 0.721 | -0.026 | 0.007 | 0.016 | 0.070 | 7.12E-01 | 0.155 | 3.51E-06 | 359521 | 5.98E-05 | 21.515 |
| | | rs62100052 | T | C | 0.283 | -0.038 | 0.062 | 0.068 | 0.027 | 1.60E-01 | 0.062 | 4.38E-06 | 359521 | 5.87E-05 | 21.091 |
| | | rs7310651 | T | C | 0.158 | -0.006 | 0.289 | 0.323 | 0.013 | 6.62E-01 | 0.035 | 4.51E-06 | 359521 | 5.85E-05 | 21.036 |
| | | rs73207239 | C | A | -0.285 | 0.003 | 0.082 | 0.071 | 0.023 | 8.99E-01 | 0.062 | 4.89E-06 | 359521 | 5.81E-05 | 20.878 |
| | | rs75317505 | C | G | 0.894 | -0.110 | 0.004 | 0.020 | 0.046 | 1.79E-02 | 0.193 | 3.82E-06 | 359521 | 5.94E-05 | 21.355 |
| | CXCL5 | rs114767765 | T | C | 0.427 | 0.011 | 0.022 | 0.019 | 0.046 | 8.15E-01 | 0.093 | 4.64E-06 | 359521 | 5.84E-05 | 20.981 |
| | | rs116769657 | T | C | 0.584 | 0.020 | 0.012 | 0.018 | 0.055 | 7.16E-01 | 0.120 | 1.20E-06 | 359521 | 6.56E-05 | 23.585 |
| | | rs117432663 | A | G | 0.217 | 0.026 | 0.121 | 0.071 | 0.024 | 2.72E-01 | 0.046 | 2.61E-06 | 359521 | 6.14E-05 | 22.086 |
| | | rs1338460 | G | A | 0.154 | 0.006 | 0.614 | 0.655 | 0.013 | 6.17E-01 | 0.033 | 3.48E-06 | 359521 | 5.99E-05 | 21.530 |
| | | rs142461741 | C | T | -0.644 | -0.001 | 0.020 | 0.022 | 0.044 | 9.80E-01 | 0.138 | 2.92E-06 | 359521 | 6.08E-05 | 21.870 |
| | | rs150102867 | G | A | 0.506 | 0.101 | 0.018 | 0.014 | 0.064 | 1.14E-01 | 0.106 | 1.64E-06 | 359521 | 6.39E-05 | 22.971 |
| | | rs1522153 | A | G | 0.163 | 0.004 | 0.685 | 0.705 | 0.013 | 7.41E-01 | 0.035 | 3.19E-06 | 359521 | 6.03E-05 | 21.696 |
| | | rs1854137 | G | C | -0.149 | -0.005 | 0.623 | 0.602 | 0.012 | 7.04E-01 | 0.032 | 4.03E-06 | 359521 | 5.91E-05 | 21.251 |
| | | rs186696265 | T | C | 0.539 | 0.068 | 0.016 | 0.018 | 0.051 | 1.81E-01 | 0.107 | 4.35E-07 | 359521 | 7.10E-05 | 25.530 |
| | | rs1919935 | C | T | -0.163 | -0.009 | 0.319 | 0.281 | 0.013 | 5.12E-01 | 0.035 | 3.06E-06 | 359521 | 6.06E-05 | 21.779 |
| | | rs2811465 | C | T | -0.185 | -0.010 | 0.215 | 0.230 | 0.015 | 5.15E-01 | 0.040 | 4.03E-06 | 359521 | 5.91E-05 | 21.250 |
| | | rs28537856 | G | C | 0.721 | 0.081 | 0.007 | 0.0156 | 0.071 | 2.51E-01 | 0.155 | 3.51E-06 | 359521 | 5.98E-05 | 21.515 |
| | | rs62100052 | T | C | 0.283 | 0.051 | 0.062 | 0.068 | 0.027 | 6.32E-02 | 0.062 | 4.38E-06 | 359521 | 5.87E-05 | 21.091 |
| | | rs7310651 | T | C | 0.158 | 0.006 | 0.289 | 0.323 | 0.013 | 6.59E-01 | 0.035 | 4.51E-06 | 359521 | 5.85E-05 | 21.036 |
| | | rs73207239 | C | A | -0.285 | 1.00E-04 | 0.082 | 0.071 | 0.023 | 9.97E-01 | 0.062 | 4.89E-06 | 359521 | 5.81E-05 | 20.878 |
| rs75317505 | C | G | 0.894 | -0.037 | 0.004 | 0.021 | 0.047 | 4.29E-01 | 0.193 | 3.82E-06 | 359521 | 5.94E-05 | 21.355 | | |

Supplementary Table S7: SNPs used as instrumental variables in Mendelian Randomization analysis of neuropathic pain on inflammatory proteins.

STROBE-MR checklist of recommended items to address in reports of Mendelian randomization studies

| Item No. | Section | Checklist item | Page No. | Relevant text from manuscript |
|----------|---|--|----------|---|
| 1 | TITLE and ABSTRACT | Indicate Mendelian randomization (MR) as the study's design in the title and/or the abstract if that is a main purpose of the study | | Title: Association of circulating inflammatory proteins with neuropathic pain: A Mendelian randomization |
| | INTRODUCTION | | | |
| 2 | Background | Explain the scientific background and rationale for the reported study. What is the exposure? Is a potential causal relationship between exposure and outcome plausible? Justify why MR is a helpful method to address the study question | | Exposure involved 91 inflammatory protein GWAS data with an outcome of SP. The use of MR Methods can not only study the causal effects of inflammatory proteins on SP but also show potential in identifying prospective biomarkers associated with SP and assessing risk and protective factors. |
| 3 | Objectives | State specific objectives clearly, including pre-specified causal hypotheses (if any). State that MR is a method that, under specific assumptions, intends to estimate causal effects | | MR is a statistical method that utilizes genetic variations as a tool to evaluate causal relationships between inflammatory proteins and SP. |
| | METHODS | | | |
| 4 | Study design and data sources | Present key elements of the study design early in the article. Consider including a table listing sources of data for all phases of the study. For each data source contributing to the analysis, describe the following: Setting: Describe the study design and the underlying population, if possible. Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection, when available. Participants: Give the eligibility criteria, and the sources and methods of selection of participants. Report the sample size, and whether any power or sample size calculations were carried out prior to the main analysis. Describe measurement, quality control and selection of genetic variants. For each exposure, outcome, and other relevant variables, describe methods of assessment and diagnostic criteria for diseases. Provide details of ethics committee approval and participant informed consent, if relevant. | | |
| | a) | Setting: Describe the study design and the underlying population, if possible. Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection, when available. | | This study describes the study design and underlying population, and specific populations and numbers have been reported. |
| | b) | Participants: Give the eligibility criteria, and the sources and methods of selection of participants. Report the sample size, and whether any power or sample size calculations were carried out prior to the main analysis. | | This paper reports the specific sources of GWAS data, as well as details on sample size and population demographics. |
| | c) | Describe measurement, quality control and selection of genetic variants. | | This paper describes the sources of instrumental variables and details the selection process and criteria for instrumental variables. Inflammatory proteins as exposed, NP as outcome in Finland in the database are described. |
| | d) | For each exposure, outcome, and other relevant variables, describe methods of assessment and diagnostic criteria for diseases. | | |
| | e) | Provide details of ethics committee approval and participant informed consent, if relevant. | | \ |
| 5 | Assumptions | Explicitly state the three core IV assumptions for the main analysis (relevance, independence and exclusion restriction) as well | | Figure 1 illustrates the three core assumptions. |
| 6 | Statistical methods: main analysis | Describe statistical methods and statistics used | | |
| | a) | Describe how quantitative variables were handled in the analyses (i.e., scale, units, model) | | For MR analysis, we use Wald the wire method to analyze only a IV of the protein, fixed effects inverse variance weighting method (IVW) applies to have two or three IVs protein, random effects IVW method suitable for protein with four or more IVs. |
| | b) | Describe how genetic variants were handled in the analyses and, if applicable, how their weights were selected | | (1) SNPs associated with exposure at the genome-wide significance level (GWAS, $P \leq 5 \times 10^{-6}$), (2) The clustering of LD threshold is set to $r^2 = 0.001$, clustering the window size is set to 10000KB (3) IVs with F-statistic < 10 were considered as weak IVs and excluded from subsequent analysis. The rest of the SNP for subsequent analysis of MR. |
| | c) | Describe the MR estimator (e.g. two-stage least squares, Wald ratio) and related statistics. Detail the included covariates and, in case of two-sample MR, whether the same covariate set was used for adjustment in the two samples | | For MR analysis, we use Wald the wire method to analyze only a IV of the protein, fixed effects inverse variance weighting method (IVW) applies to have two or three IVs protein, random effects IVW method suitable for protein with four or more IVs. |
| | d) | Explain how missing data were addressed | | This paper does not have any missing values. |
| | e) | If applicable, indicate how multiple testing was addressed | | This paper did not employ multivariable MR. |
| 7 | Assessment of assumptions | Describe any methods or prior knowledge used to assess the assumptions or justify their validity | | This paper reports the use of F-statistics to estimate statistical power. |
| 8 | Sensitivity analyses and additional analyses | Describe any sensitivity analyses or additional analyses performed (e.g. comparison of effect estimates from different approaches, independent replication, bias analytic techniques, validation of instruments, simulations) | | This paper employs The Cochran's Q to assess the heterogeneity of individual causal effects. MR-Egger's intercept term and MR-PRESSO method are also used to evaluate horizontal pleiotropy. Furthermore, F-statistics are used to assess statistical power. |
| 9 | Software and pre-registration | | | |
| | a) | Name statistical software and package(s), including version and settings used | | This paper utilized R software, with R packages including TwoSample MR and MR-PRESSO, all of which are available for free on the official R software website. |
| | b) | State whether the study protocol and details were pre-registered (as well as when and where) | | This paper does not provide registration information such as time and address. |
| | RESULTS | | | |
| 10 | Descriptive data | | | |
| | a) | Report the numbers of individuals at each stage of included studies and reasons for exclusion. Consider use of a flow diagram | | In the methodology section, the paper provides information about the sample population size, while the results section does not include demographic information about the population. Additionally, the paper does not present details about excluded individuals or the reasons for their exclusion. |
| | b) | Report summary statistics for phenotypic exposure(s), outcome(s), and other relevant variables (e.g. means, SDs, proportions) | | This paper provides a detailed description of the exposure and outcome data. |
| | c) | If the data sources include meta-analyses of previous studies, provide the assessments of heterogeneity across these studies | | The original data does not include meta-analysis. |
| | d) | For two-sample MR: i. Provide justification of the similarity of the genetic variant-exposure associations between the exposure and outcome samples ii. Provide information on the number of individuals who overlap between the exposure and outcome studies | | This paper falls under the category of two-sample MR with all sample populations originating from Europe, resulting in significant population heterogeneity. There is no information about overlapping populations, and there are no common subjects. |
| 11 | Main results | | | |
| | a) | Report the associations between genetic variant and exposure, and between genetic variant and outcome, preferably on an interpretable scale | | This paper in the MR results section describes the relationship between instrumental variables and the exposure and outcome. |
| | b) | Report MR estimates of the relationship between exposure and outcome, and the measures of uncertainty from the MR analysis, on an interpretable scale, such as odds ratio or relative risk per SD difference | | This paper reports the odds ratio (OR) and 95% confidence interval (CI) for the association between exposure and outcome in the MR results. |
| | c) | If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period | | This paper reports which inflammatory proteins are risk for PN in the MR results. |
| | d) | Consider plots to visualize results (e.g. forest plot, scatterplot of associations between genetic variants and outcome versus between genetic variants and exposure) | | Figure 2 and Figure 3. |
| 12 | Assessment of assumptions | | | |
| | a) | Report the assessment of the validity of the assumptions | | This paper reports the use of F-statistics to estimate statistical power. |
| | b) | Report any additional statistics (e.g., assessments of heterogeneity across genetic variants, such as I^2 , Q statistic or E-value) | | This paper employs The Cochran's Q to assess the heterogeneity of individual causal effects. MR-Egger's intercept term is also used to evaluate horizontal pleiotropy. Furthermore, F-statistics are used to assess statistical power. |
| 13 | Sensitivity analyses and additional analyses | | | |
| | a) | Report any sensitivity analyses to assess the robustness of the main results to violations of the assumptions | | The Cochran's Q to assess the heterogeneity of individual causal effects. MR-Egger's intercept term is also used to evaluate horizontal pleiotropy. |
| | b) | Report results from other sensitivity analyses or additional analyses | | The results of sensitivity analyses are reported. |
| | c) | Report any assessment of direction of causal relationship (e.g., bidirectional MR) | | Two-way MR Method is adopted in this paper. |
| | d) | When relevant, report and compare with estimates from non-MR analyses | | Results with other non-MR analyses are not reported in this report |
| | e) | Consider additional plots to visualize results (e.g., leave-one-out analyses) | | Supplementary figure |

| DISCUSSION | | | |
|-------------------|------------------------------|--|---|
| 14 | Key results | Summarize key results with reference to study objectives | This paper summarizes key MR results in Supplementary tables |
| 15 | Limitations | Discuss limitations of the study, taking into account the validity of the IV assumptions, other sources of potential bias, and imprecision. Discuss both direction and magnitude of any potential bias and any efforts to address them. | This paper discusses the limitations of the study, including data sources, the three major assumptions, and the analytical process, providing detailed content. |
| 16 | Interpretation | | |
| | a) | Meaning: Give a cautious overall interpretation of results in the context of their limitations and in comparison with other studies | This paper dedicates a substantial portion to discussing |
| | b) | Mechanism: Discuss underlying biological mechanisms that could drive a potential causal relationship between the investigated exposure and the outcome, and whether the gene-environment equivalence assumption is reasonable. Use causal language carefully, clarifying that IV estimates may provide causal effects only under certain assumptions | Various positive proteins provide a reasonable interpretation of MR results by comparing them with multiple previously published studies. |
| | c) | Clinical relevance: Discuss whether the results have clinical or public policy relevance, and to what extent they inform effect sizes of possible interventions | The various biological mechanisms by which inflammatory proteins are involved in neuropathic pain are discussed. |
| 17 | Generalizability | Discuss the generalizability of the study results (a) to other populations, (b) across other exposure periods/timings, and (c) across other levels of exposure | Limitations of the study and the population used are reported. |
| OTHER INFORMATION | | | |
| 18 | Funding | Describe sources of funding and the role of funders in the present study and, if applicable, sources of funding for the databases and original study or studies on which the present study is based | No funding was received for the current study. |
| 19 | Data and data sharing | Provide the data used to perform all analyses or report where and how the data can be accessed, and reference these sources in the article. Provide the statistical code needed to reproduce the results in the article, or report whether the code is publicly accessible and if so, where | The data in this manuscript are all available in publicly available databases and are referenced by link in the article. |
| 20 | Conflicts of Interest | All authors should declare all potential conflicts of interest | The authors have no relevant financial or non-financial interests to disclose. |

Supplementary Table S8: STROBE-MR checklist of recommended items to address in reports of Mendelian randomization studies