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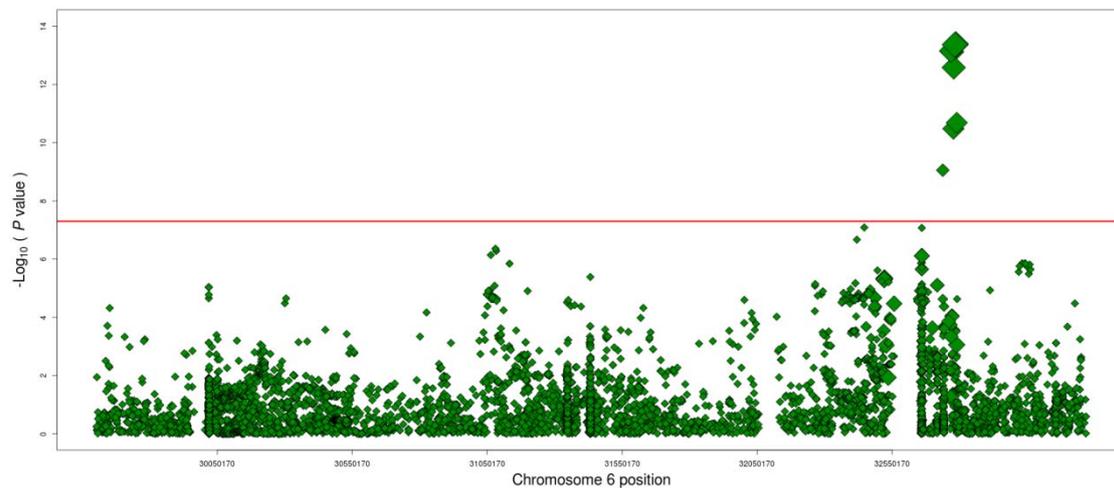
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Study population

The study population comprised 2,465 cases from four vasculitides (GCA, TAK, AAV, and IgAV) and 4,632 unaffected controls, organised as follows: the GCA dataset comprised two independent populations, one from Spain (763 cases and 850 controls) and another from UK (251 cases and 1000 controls); the TAK dataset included two populations, one of European ancestry from North America (USA/Canada; 112 cases and 599 controls) and one from Turkey (339 cases and 516 controls); the AAV dataset included 685 cases and 1,000 controls from UK; and the IgAV dataset comprised a population from Spain (315 cases and 667 controls). GCA and TAK patients were diagnosed following their respective 1990 American College of Rheumatology classification criteria [1, 2]. Additionally, GCA patients were only enrolled if they had a positive temporal artery biopsy, as described [3]. The EMEA clinically-based algorithm was used to classify AAV patients [4]. The IgA vasculitis condition was diagnosed accordingly with both the guidelines included in Michel *et al.* [5] and the American College of Rheumatology classification criteria for this disease [6]. The most relevant clinical data of each case cohort were described elsewhere [7-10]. All participants signed a written informed consent before being enrolled in the study. The procedures were followed according to the ethical standards of the Ethics Committees on human experimentation of the responsible committee on human experimentation (institutional and national) of all participant centres.

Figure S1. Manhattan plot representation of the overall cross-disease meta-analysis of the imputed HLA region. The $-\log_{10}$ of the inverse variance-weighted meta-analysis p-values are plotted against their physical chromosomal position. The diamond size depends on the linkage disequilibrium (r^2) with the lead signal (rs6457617).



References for 'Study population' and Table S4.

1. Arend WP, Michel BA, Bloch DA, Hunder GG, Calabrese LH, Edworthy SM, et al. The American College of Rheumatology 1990 criteria for the classification of Takayasu arteritis. *Arthritis and rheumatism*. 1990; 33:1129-1134.
2. Hunder GG, Bloch DA, Michel BA, Stevens MB, Arend WP, Calabrese LH, et al. The American College of Rheumatology 1990 criteria for the classification of giant cell arteritis. *Arthritis and rheumatism*. 1990; 33:1122-1128.
3. Gonzalez-Gay MA, Martinez-Dubois C, Agudo M, Pompei O, Blanco R, Llorca J. Giant cell arteritis: epidemiology, diagnosis, and management. *Current rheumatology reports*. 2010; 12:436-442.
4. Watts R, Lane S, Hanslik T, Hauser T, Hellmich B, Koldingsnes W, et al. Development and validation of a consensus methodology for the classification of the ANCA-associated vasculitides and polyarteritis nodosa for epidemiological studies. *Annals of the rheumatic diseases*. 2007; 66:222-227.
5. Michel BA, Hunder GG, Bloch DA, Calabrese LH. Hypersensitivity vasculitis and Henoch-Schonlein purpura: a comparison between the 2 disorders. *The Journal of rheumatology*. 1992; 19:721-728.
6. Mills JA, Michel BA, Bloch DA, Calabrese LH, Hunder GG, Arend WP, et al. The American College of Rheumatology 1990 criteria for the classification of Henoch-Schonlein purpura. *Arthritis and rheumatism*. 1990; 33:1114-1121.
7. Carmona FD, Mackie SL, Martin JE, Taylor JC, Vaglio A, Eyre S, et al. A large-scale genetic analysis reveals a strong contribution of the HLA class II region to giant cell arteritis susceptibility. *American journal of human genetics*. 2015; 96:565-580.
8. Lopez-Mejias R, Carmona FD, Castaneda S, Genre F, Remuzgo-Martinez S, Sevilla-Perez B, et al. A genome-wide association study suggests the HLA Class II region as the major susceptibility locus for IgA vasculitis. *Scientific reports*. 2017; 7:5088.
9. Lyons PA, Rayner TF, Trivedi S, Holle JU, Watts RA, Jayne DR, et al. Genetically distinct subsets within ANCA-associated vasculitis. *The New England journal of medicine*. 2012; 367:214-223.

10. Saruhan-Direskeneli G, Hughes T, Aksu K, Keser G, Coit P, Aydin SZ, et al. Identification of multiple genetic susceptibility loci in Takayasu arteritis. *American journal of human genetics*. 2013; 93:298-305.
11. Carmona FD, Vaglio A, Mackie SL, Hernandez-Rodriguez J, Monach PA, Castaneda S, et al. A Genome-wide Association Study Identifies Risk Alleles in Plasminogen and P4HA2 Associated with Giant Cell Arteritis. *American journal of human genetics*. 2017; 100:64-74.
12. Merkel PA, Xie G, Monach PA, Ji X, Ciavatta DJ, Byun J, et al. Identification of Functional and Expression Polymorphisms Associated With Risk for Antineutrophil Cytoplasmic Autoantibody-Associated Vasculitis. *Arthritis Rheumatol*. 2017; 69:1054-1066.
13. Xie G, Roshandel D, Sherva R, Monach PA, Lu EY, Kung T, et al. Association of granulomatosis with polyangiitis (Wegener's) with HLA-DPB1*04 and SEMA6A gene variants: evidence from genome-wide analysis. *Arthritis and rheumatism*. 2013; 65:2457-2468.
14. Terao C, Yoshifuji H, Kimura A, Matsumura T, Ohmura K, Takahashi M, et al. Two susceptibility loci to Takayasu arteritis reveal a synergistic role of the IL12B and HLA-B regions in a Japanese population. *American journal of human genetics*. 2013; 93:289-297.
15. Renauer PA, Saruhan-Direskeneli G, Coit P, Adler A, Aksu K, Keser G, et al. Identification of Susceptibility Loci in IL6, RPS9/LILRB3, and an Intergenic Locus on Chromosome 21q22 in Takayasu Arteritis in a Genome-Wide Association Study. *Arthritis Rheumatol*. 2015; 67:1361-1368.