

The Extent of Edgetic Perturbations in the Human Interactome Caused by Population-Specific Mutations

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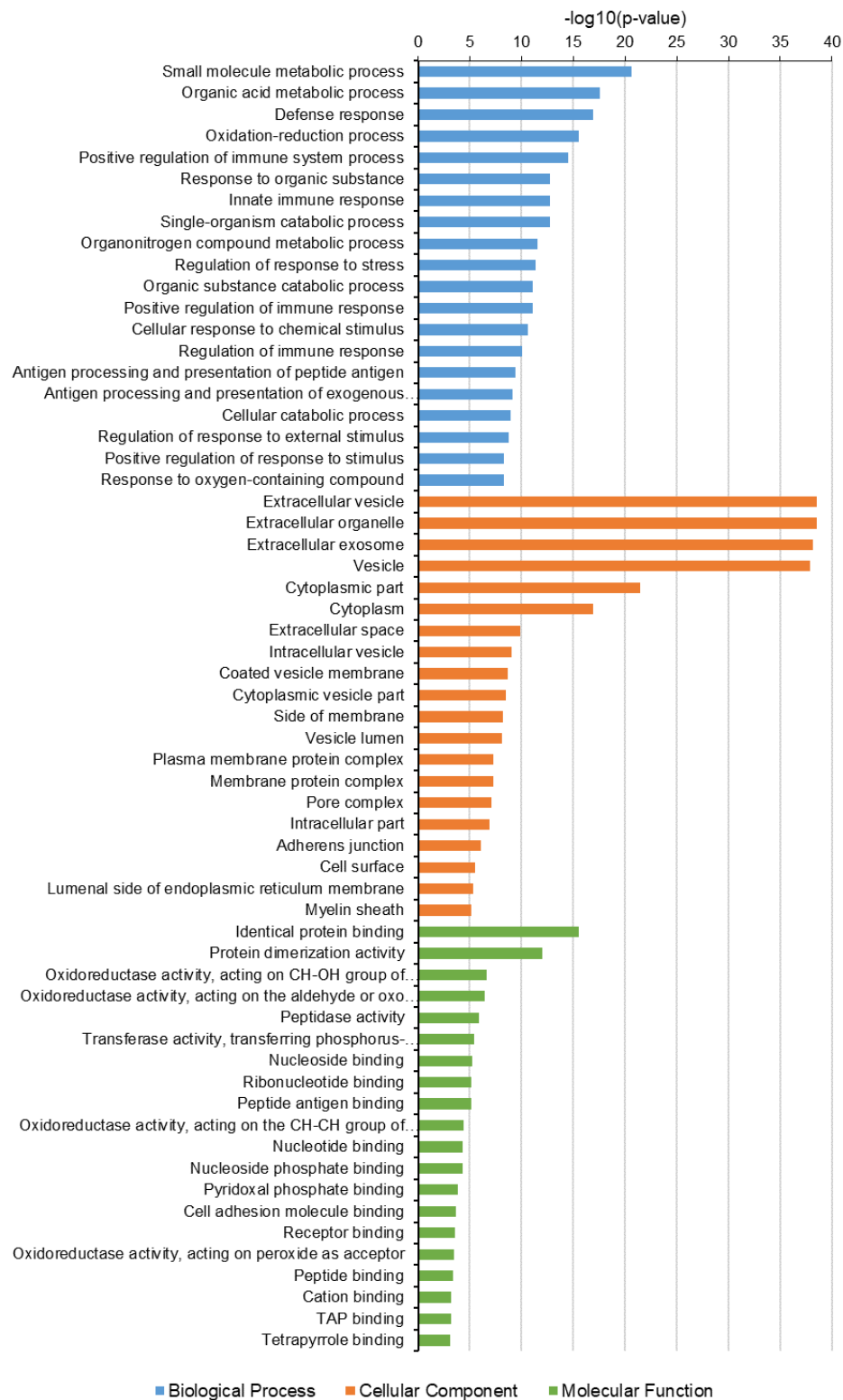
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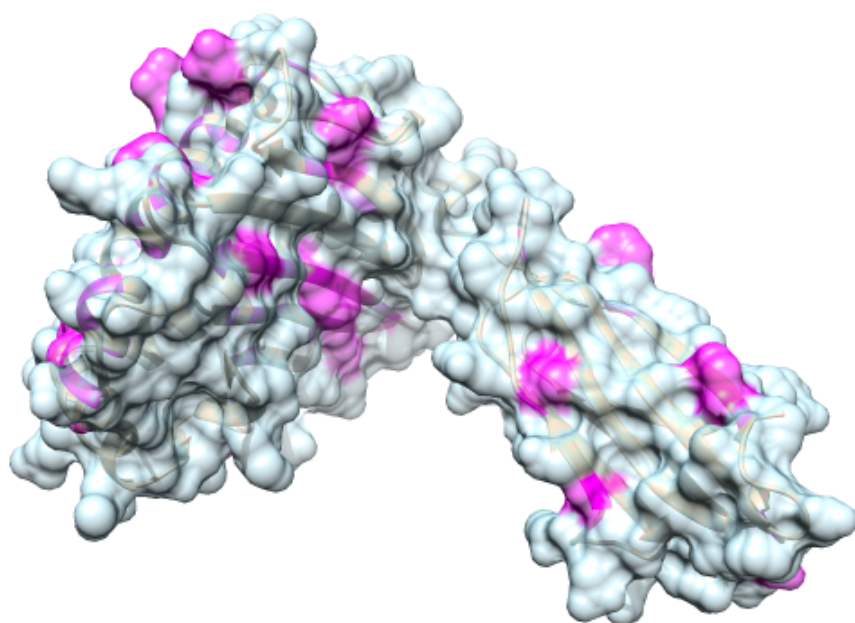
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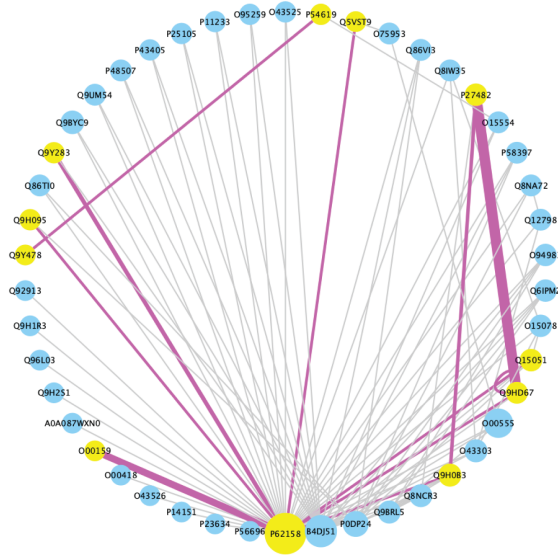
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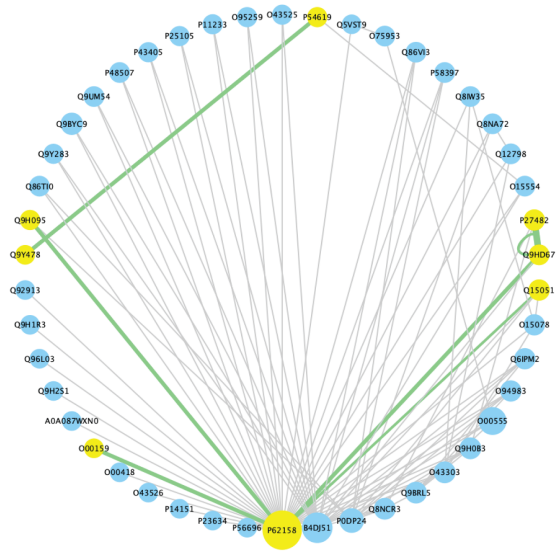
Supplementary Figure S1. Top 20 significant GO terms for genes enriched with disruptive mutations among three basic GO categories: biological process, molecular function and cellular component.



Supplementary Figure S2. HLA-A is one of the top genes carrying high number of normal disruptive mutation in the HLA gene family.

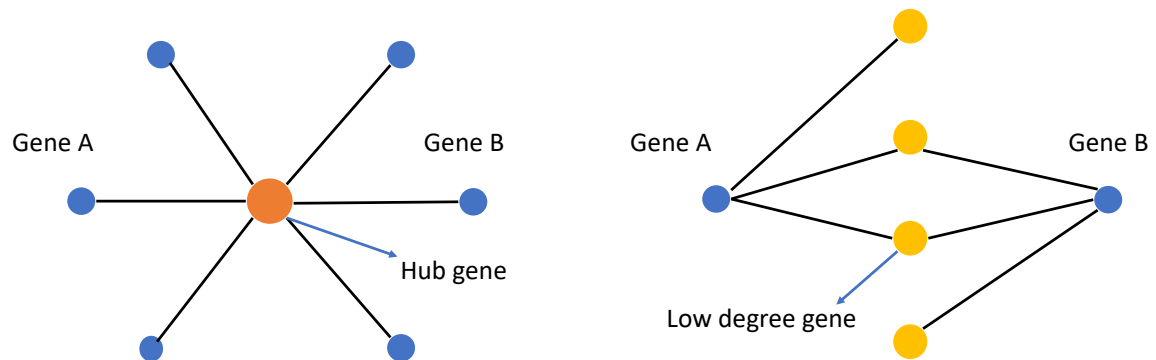


East Asian

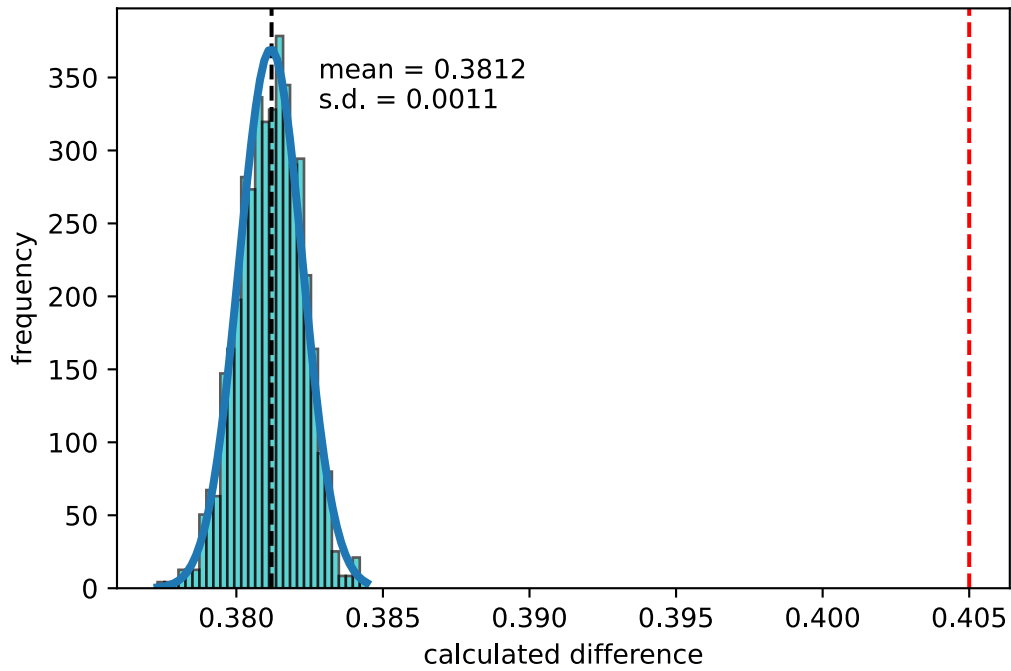


American

Supplementary Figure S3. Distinct rewiring patterns in interactome modules associated with arrhythmias in East Asians and Americans. The East Asian population has 11 proteins that carry mutations (yellow nodes) resulting in 11 rewired interactions (magenta edges), whereas the American population has 6 mutant proteins (yellow nodes) with 6 rewired interactions (green edges). Additionally, the higher allele frequency of a mutation and the corresponding rewired interaction is represented by the increasing thickness of edges, where one can see that the East Asian population has higher frequency of these mutations.



Supplementary Figure S4. Illustration of the concept of Diffusion State Distance (DSD). DSD downweights the influence of the hub gene and favors low degree gene.



Supplementary Figure S5. The reported normalized difference of the total detrimental mutations occurring in one protein between any two major populations is statistically significantly larger compared to the population of randomly shuffled edgetic profiles. The reported Z-score is ~21 and P-Value < 0.001. We also note that the randomized difference from 1,000 experiments falls within a very tight range.