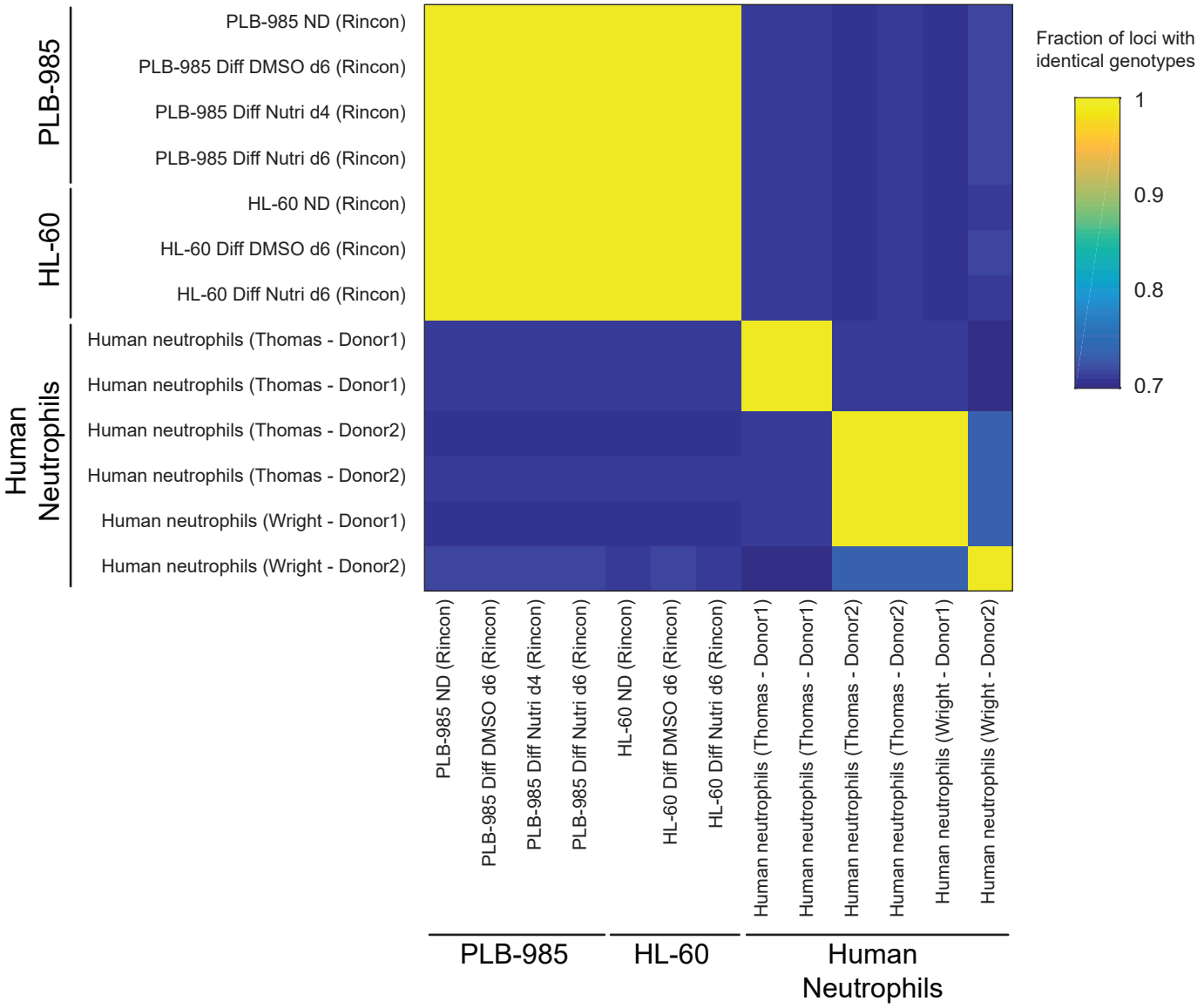Additional File 6