

# Polygenic risk scores for bipolar disorders associated with worse global functioning

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## SUBMISSION DETAILS

**What is the Methodological Question Being Addressed?** To investigate the association between genetic risk for bipolar disorders (BD) with measures of global functioning.

**Introduction** Polygenic risk scores for bipolar disorders (PRS-BD) have been related to clinical variables and individual characteristics.<sup>1-3</sup> The aim of this preliminary study was to investigate the association between genetic risk for bipolar disorders (BD) with measures of global functioning.

**Methods** PRS-BD were calculated for adult participants (18-61 years old) with BD and healthy controls using PRSice-2 based on summary statistics from the Psychiatric Genomic Consortium. The psychiatric diagnosis was established according to the Diagnostic and Statistical Manual of Mental Disorders Fourth Edition - Text Revision (DSM-IV-TR). Linear regression analyzes were performed to investigate the association between PRS-BD z scores with functioning, measured by the Global Assessment of Functioning (GAF), which data were log-transformed for the analyzes. Demographic and clinical variables that presented a significant association or a trend for association ( $p < 0.20$ ) with the outcome in the simple linear regression were included in a multiple linear regression model to examine the independent association between PRS-BD and GAF.

**Results** 203 participants (145 females [71.4%] and 58 males [28.6%]) were evaluated: 133 (65.5%) with BD type I diagnosis, 18 (8.9%) with BD type II and 52 (25.6%) healthy controls (Table 1). PRS-BD presented a weak and significant correlation with lower GAF (Spearman correlation  $r = -0.17$ ,  $p = 0.01$ ) (Figure 1). Table 2 shows the simple linear regression analyzes between the independent variables and GAF (Table 2). The association between PRS-BD and GAF remained statistically significant when controlling for ethnicity, depressive and manic/hypomanic symptoms in a multiple linear regression analysis (B coefficient  $-0.02$  [ $-0.03 - -0.003$  95% CI]  $p = 0.01$ ; adjusted  $R^2 = 0.57$ ) (Table 3). Figure 2 shows PRS-BD z scores distribution between the three groups.

**Conclusion** Genetic risk for BD could confer a risk for worse global functioning regardless of psychiatric symptoms. Among the limitations, the lack of controlling for genetic background of the sample should be highlighted.

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## Keywords

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**Guidelines** I have read and understand the Poster Guidelines

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