

BDNF Val66Met and DRD2 Taq1A polymorphisms interact to influence PTSD symptom severity: A preliminary investigation in a South African population

Sian M.J. Hemmings, Lindi I. Martin, Marisa Klopper, Lize van der Merwe, Lisa Aitken, Erika de Wit, Gillian F. Black, Eileen G. Hoal, Gerhard Walzl and Soraya Seedat

Abstract

Background: We evaluated the role that selected variants in serotonin transporter (*5-HTT*), dopamine receptor 2 (*DRD2*) and brain-derived neurotrophic factor (*BDNF*) genes play in PTSD symptom severity in an at-risk population. We also investigated the interaction between the genetic variants to determine whether these variables and the interactions between the variables influenced the severity of PTSD symptoms.

Methods: PTSD symptoms were quantitatively assessed using the Davidson Trauma Scale (DTS) in 150 participants from an at-risk South African population. All participants were genotyped for the *5-HTTLPR*, *DRD2 Taq1A* and *BDNF Val66Met* polymorphisms. Gene-gene interactions were investigated using various linear models. All analyses were adjusted for age, gender, major depressive disorder diagnosis, level of resilience, level of social support and alcohol dependence.

Results: A significant interaction effect between *DRD2 Taq1A* and *BDNF Val66Met* variants on DTS score was observed. On the background of the *BDNF Val66Val* genotype, DTS score increased significantly with the addition of a *DRD2 Taq1A A1* allele. However, on the *BDNF Met66* allele background, the addition of an *A1* allele was found to reduce total DTS score.

Conclusions: This study provides preliminary evidence for an epistatic interaction between *BDNF Val66Met* and *DRD2 Taq1A* polymorphisms on the severity of PTSD symptoms, where both too little and too much dopamine can result in increased PTSD symptom severity.

1. Introduction

Posttraumatic stress disorder (PTSD) is an anxiety disorder that develops following exposure to a life-threatening event (APA, 1994). However, not all trauma-exposed individuals develop PTSD. The lifetime prevalence of exposure to traumatic events varies between 40% and 90%, depending on the sample investigated, whereas the lifetime prevalence of PTSD in trauma-exposed individuals has been estimated at approximately 9% (Breslau et al., 2012). Numerous risk factors for developing PTSD have been elucidated, including those pertaining to the traumatic event, such as type, severity and duration of trauma, poor social support and childhood adversity (reviewed in Yehuda and LeDoux, 2007). In addition,

substance dependence. *Am J Med Genet B Neuropsychiatr Genet* 2006;141B(4):387-93.

Zimet GD, Dahlem NW, Zimet SG, Farley GK. The multidimensional scale of perceived social support. *J Pers Assess* 1988;52:30-41.