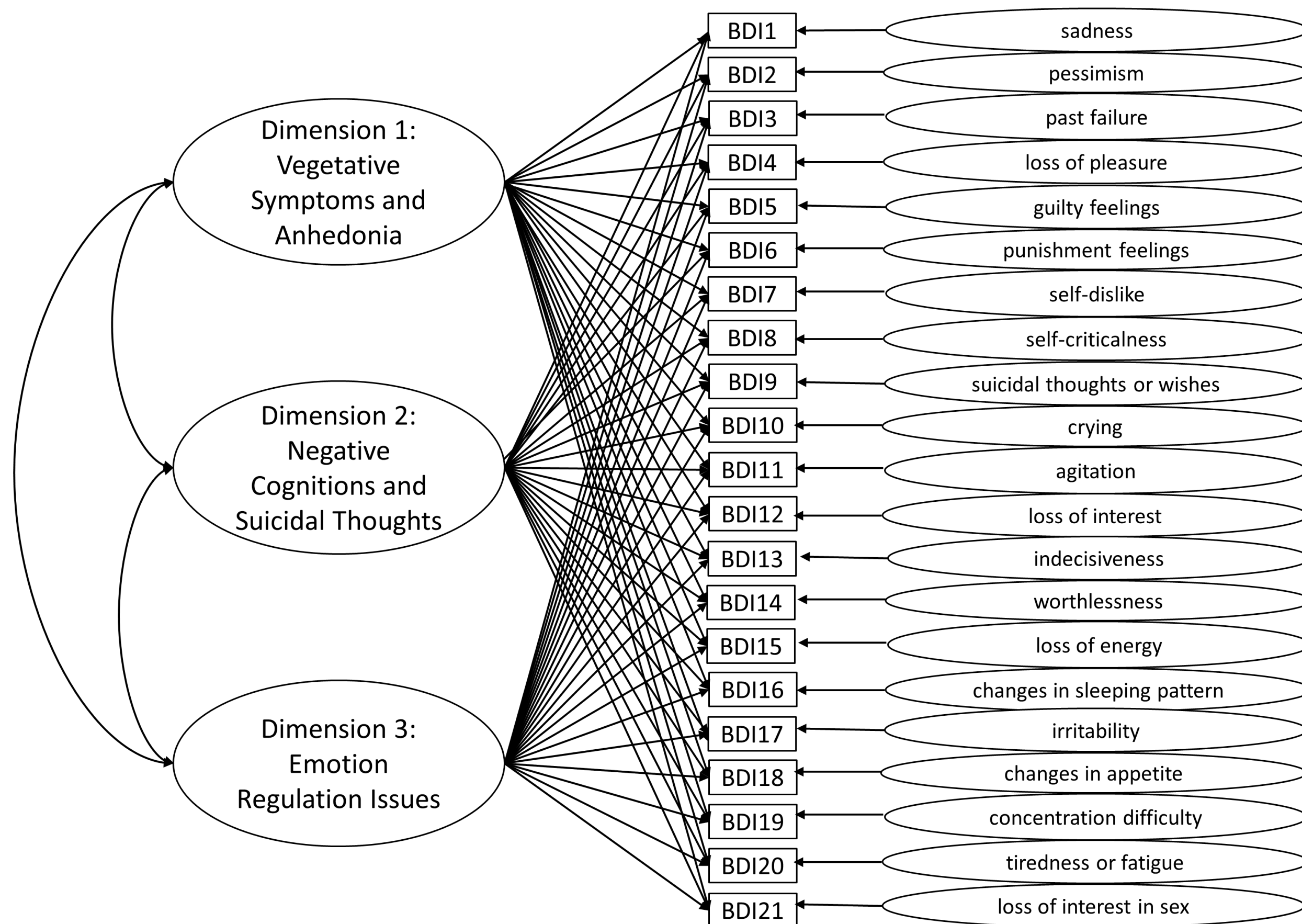


## Introduction

Multiple sclerosis (MS) is an autoimmune neurodegenerative disease that causes inflammation and demyelination in the brain, and it is associated with **increased risk of depression and suicide**; 20-40% of individuals with MS experience depression, a rate 2–4 times greater than the general population.<sup>1,2</sup> As many as **15% of MS-related deaths may be due to suicide**.<sup>1,2</sup>

Preliminary literature suggests that sadness and anhedonic (i.e., loss of pleasure) **aspects of depression may dissociate in MS**.<sup>3</sup> Based on this, **we hypothesized that depression would dissociate into two dimensions**, one characterized by sadness and the other characterized by anhedonia. We predicted that the **anhedonic phenotype would be more strongly related to suicidal ideation**.<sup>4</sup>



**Figure 1.** Rotated Principal Components Analysis of the Beck Depression Inventory II in a Sample of Adults with MS.

## Method

This study used an existing dataset from several studies of individuals with MS, recruited from area MS clinics and the general community. Participants included **117 adults with MS** [Mean (Standard Deviation): age = 47.48 (10.21) years; 78.9% female; 97.3% White non-Hispanic; disease duration = 7.86 (8.30) years; Expanded Disability Status Scale: Median = 3, Range = 0 to 8.0]. Depression was measured using the Beck Depression Inventory-II (BDI-II).<sup>4</sup> Patients' level of **mood symptoms ranged from non-depressed to severely depressed** [10.26 (9.35)].

## Results

The principal components analysis (PCA) was determined appropriate for the BDI-II data; Kaiser-Meyer-Olkin index = .91, Bartlett's test  $p < .001$ . Then, the analysis was run with PCA extraction and a Varimax with Kaiser Normalization rotation, which revealed **three dimensions** (eigenvalue  $> 1$ ). One dominant dimension (eigenvalue 4.80) accounted for 22.88% of variation, with two additional dimensions (eigenvalues 4.72 and 2.88 respectively) that accounted for 22.46% and 13.72% of variation, respectively. Overall, the PCA accounted for 59% of variance in total. Dimension 1 consisted of eight items characterized by **vegetative symptoms and anhedonia**. Dimension 2 included eight items related to **negative cognitions, including pessimism, loss of hope, and suicidal thoughts**. Dimension 3 was comprised of five items pertaining to **sadness and emotion regulation issues**.

## Discussion

This study builds on work previously suggesting a dissociation of MS-related depression symptoms, strengthening and expanding that finding to suggest that **MS-related depression dissociates into three dimensions**. The three dimensions we found consisted of vegetative symptoms and anhedonia (dimension 1), negative cognitions and suicidal thoughts (dimension 2), and sadness and emotion regulation issues (dimension 3).

**Identifying and treating depression may be the most important factor in preventing suicide in patients with MS**,<sup>2</sup> and this study suggests that those who are most affected by negative cognitions such as pessimism and hopelessness, worthlessness, and self-dislike are also at the most risk of suicide, so these cognitions should be screened for in MS patients. The etiology of depression in MS is complex and poorly understood,<sup>1,2</sup> perhaps because it has traditionally been conceptualized as homogenous.<sup>1</sup> This study suggests that similar to recent conceptualizations of depression without MS, depression in MS can be better understood also as a **disorder characterized by multiple distinct phenotypes that can present differently from individual to individual**. These findings also suggest the importance of **negative cognitions in relation to suicidal thoughts**, at least in this sample of MS patients.

Future studies should include non-MS depressed individuals as a comparison. Additionally, this sample had relatively low depression levels, so a replication of this study in individuals with more severe MS-related depression would be beneficial to assess if these dimensions exist across the spectrum of MS-related depression.

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