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**MANUSCRIPT**

# Neurodevelopmental Consequences of Alcohol Abuse in Hispanic Adolescents (Ages 13-19) in the Rio Grande Valley: A Review of Cognitive and Structural Brain Impacts

Marisol Acosta<sup>1</sup>, Jessica P. Carvajal<sup>1,3</sup>, Natalia Del Castillo<sup>1,3</sup>, Valeria I. Garcia<sup>1,2</sup>, Kaitlyn R. Lewis<sup>1,2</sup>, Hannah N. Navarro<sup>1,2</sup>, Dominic R. Policarpio<sup>1,2</sup>, Kai R. Saenz<sup>1,2</sup>

<sup>1</sup>DHR Health High School and Community Outreach

<sup>2</sup>South Texas ISD Health Professions

<sup>3</sup>South Texas ISD Medical Professions

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## Introduction

This literature review examines the neurodevelopmental consequences of alcohol abuse in Hispanic adolescents (ages 13-19) residing in the Rio Grande Valley. The review focuses on the cognitive and structural brain impacts of alcohol abuse within this specific demographic and geographic context. While research on alcohol's effects on the developing brain is extensive, studies specifically targeting Hispanic adolescents in the Rio Grande Valley are limited. This review synthesizes existing literature to identify key findings, research gaps, and areas needing further investigation. The unique cultural and socioeconomic factors within the Rio Grande Valley will be considered where relevant data allows.

## Alcohol Use Patterns in Hispanic Adolescents of the Rio Grande Valley

Understanding the prevalence and patterns of alcohol abuse among Hispanic adolescents in the Rio Grande Valley is crucial for contextualizing the neurodevelopmental consequences. While precise data specific to this demographic and region may be scarce, broader studies on Hispanic populations and border communities offer valuable insights. Research indicates that acculturation significantly influences alcohol use patterns among Hispanics.<sup>4</sup> Men identifying as "very Mexican" or "bicultural" exhibit higher risks of alcohol abuse and dependence compared to their "very Anglo" counterparts.<sup>4</sup> Conversely, women show increased heavy episodic drinking as they acculturate, potentially reflecting shifting social norms surrounding alcohol consumption for females in U.S. society.<sup>4</sup> This suggests a non-linear relationship between acculturation and

alcohol-related disorders, varying by gender and potentially influenced by unique social dynamics prevalent in border regions like the Rio Grande Valley.<sup>4</sup> The study by Courtney and Polich highlights the epidemiological rise of binge drinking among college-age men and its cognitive implications, which may extend to adolescent populations, underscoring the need for further research on this specific demographic in the Rio Grande Valley.<sup>5</sup> Furthermore, research on problem drinking among adolescents (ages 15-19) indicates an average of 5 drinks per occasion and consumption on 38% of days during a monitoring week.<sup>6</sup> This pattern of binge drinking is concerning given its association with serious alcohol-related problems, particularly its impact on frontal lobe functions and working memory.<sup>5,7</sup>

### **Cognitive Impacts of Alcohol Abuse**

Alcohol abuse significantly impacts cognitive function, particularly during adolescence, a critical period of brain development.<sup>1-3</sup> The prefrontal cortex, hippocampus, and cerebellum are especially vulnerable to alcohol's damaging effects, impacting executive functions, memory, and motor control.<sup>1,2</sup> Studies using neuropsychological assessments have demonstrated cognitive deficits in adolescents with alcohol use disorders, including impairments in executive functions, which are crucial for planning, decision-making, and inhibitory control.<sup>2</sup> The study by Crespi et al. utilized joint Independent Component Analysis (jICA) on gray and white matter features to reveal widespread structural changes in brain networks involved in executive functioning, salience, and the default-mode network in individuals with Alcohol Use Disorder (AUD).<sup>2</sup> These neurostructural changes correlated significantly with executive performance, highlighting the link between brain damage and cognitive impairment [2].<sup>2</sup> Furthermore, research indicates that binge drinking is associated with anomalies in brain functioning, particularly affecting brain oscillations linked to motor inhibition and execution.<sup>7</sup> Binge drinkers demonstrated lower oscillatory responses in delta and theta frequency ranges during motor tasks, suggesting underlying neural dysfunction without significant differences in event-related potentials.<sup>7</sup> The bidirectional relationship between inhibitory control and alcohol use is also noteworthy.<sup>8</sup> Impaired inhibitory control can predispose individuals to alcohol misuse, while alcohol consumption further impairs inhibitory control, creating a vicious cycle.<sup>8</sup> Deficits in inhibitory control during adolescence may even predict future alcohol use disorders, highlighting the importance of early intervention.<sup>8</sup> A population-based study linked alcohol and cannabis misuse to impaired cognitive abilities in adolescents, including deficits in recall memory, perceptual reasoning, inhibition, and working memory.<sup>9</sup> The study suggested that these effects could be due to common underlying vulnerabilities, neuroplasticity, neurotoxicity, or developmental sensitivity, underscoring the complexity of the relationship between substance use and cognitive impairment.<sup>9</sup>

## Structural Brain Impacts of Alcohol Abuse

Neuroimaging studies reveal significant structural brain changes associated with alcohol abuse, particularly in adolescents.<sup>2,10-11</sup> Magnetic resonance imaging (MRI) studies have documented alterations in both gray and white matter volumes indicating significant structural damage during critical developmental periods.<sup>2,10</sup> Specific brain regions, such as the prefrontal cortex, hippocampus, and cerebellum, show greater vulnerability to alcohol's neurotoxic effects.<sup>1,10</sup> Moreover, studies using MRI have demonstrated disrupted cortical volume changes in individuals with prenatal alcohol exposure, suggesting that alcohol's effects on brain structure can begin even before birth.<sup>12</sup> In adolescents, alcohol exposure leads to decreased plasticity and increased loss of cortical structure, highlighting the long-term consequences of alcohol abuse on brain development.<sup>12</sup> Research using Mendelian randomization suggests a causal relationship between heavy alcohol use and reductions in subcortical brain volumes, specifically impacting the amygdala and hippocampus.<sup>11</sup> These subcortical structures play vital roles in emotional regulation and memory, suggesting that alcohol-induced damage in these areas can have significant long-term consequences on mental health and cognitive function.<sup>11</sup> The study by Durazzo et al. found that individuals with alcohol use disorder (AUD) exhibited significant morphological abnormalities in brain regions, particularly the frontal lobe, before treatment.<sup>13</sup> Relapsers after treatment demonstrated greater bilateral frontal gray matter volume deficits compared to those who maintained abstinence, suggesting that persistent frontal gray matter volume deficits may serve as an endophenotype for relapse susceptibility.<sup>13</sup> The study by De Santis et al., comparing men with AUD and rats with excessive alcohol consumption, found comparable DTI alterations, particularly in the corpus callosum and fornix/fimbria, suggesting an underlying process that evolves soon after cessation of alcohol use.<sup>14</sup> These findings highlight the widespread and persistent effects of alcohol abuse on brain structure.

## Neurochemical Changes and Mechanisms

Alcohol's impact on the developing brain involves complex neurochemical changes.<sup>1</sup> Alcohol exposure disrupts normal brain development processes, including alterations in gene expression regulation and molecules involved in cell-cell interactions.<sup>1</sup> It interferes with the mitogenic and growth factor response, enhances free radical formation, and deranges glial cell functions.<sup>1</sup> In both adult and adolescent brains, alcohol damages specific brain areas through mechanisms involving excitotoxicity, free radical formation, and neuroinflammatory damage resulting from the activation of the innate immune system mediated by TLR4 receptors.<sup>1</sup> Alcohol also acts on specific membrane proteins, such as neurotransmitter receptors (e.g., NMDA, GABA-A), ion channels (e.g., L-type Ca<sup>2+</sup> channels, GIRKs), and signaling pathways (e.g., PKA and PKC signaling).<sup>1</sup> These effects contribute to the wide variety of behavioral effects induced by ethanol drinking.<sup>1</sup> Neuroadaptive changes affecting neurotransmission systems, particularly those sensitive to alcohol's acute effects, occur after long-term alcohol consumption.<sup>1</sup> Alcohol-induced maladaptations in the dopaminergic mesolimbic system, abnormal plastic changes in reward-

related brain areas, and genetic and epigenetic factors may contribute to alcohol reinforcement and addiction.<sup>1</sup> Research on adolescent mice has shown that adolescents exhibit resilience to alcohol withdrawal-induced changes in glutamate function in the nucleus accumbens, unlike adults, suggesting age-related differences in neurobiobehavioral responses to alcohol.<sup>15</sup> The study by Sinha highlights the role of chronic stress in addiction and substance abuse, emphasizing the alteration in the HPA axis and autonomic arousal linked to early life stress and its impact on drug use.<sup>16</sup> These neurochemical changes underscore the complexity of alcohol's effects on the brain and the need for further research to understand the specific mechanisms involved.

### **Risk And Protective Factors**

Several risk and protective factors influence the development of alcohol abuse and its neurodevelopmental consequences in Hispanic adolescents in the Rio Grande Valley. Socioeconomic factors, including poverty and limited access to resources, may increase vulnerability to alcohol abuse.<sup>4,17,18</sup> Cultural factors, such as acculturation and family traditions surrounding alcohol consumption, also play a significant role.<sup>4</sup> Parental attitudes toward alcohol use, parental education level, and family support systems can impact adolescent drinking behavior.<sup>18-19</sup> Peer influence, school environment, and community-level factors all contribute to the complex interplay of risk factors.<sup>19-20</sup> Conversely, protective factors, such as strong family bonds, positive peer relationships, school connectedness, and engagement in prosocial activities, can buffer against alcohol abuse.<sup>19-20</sup> The study by Zucker et al. highlights the importance of early developmental processes, noting that developmental pathways to underage drinking often begin in early childhood and involve numerous risk and vulnerability factors.<sup>21</sup> This underscores the need for comprehensive prevention strategies that address multiple risk factors across different developmental stages. The study by Assari found that parental educational attainment affects youth inhibitory control, which in turn predicts high-risk behaviors, including substance use.<sup>18</sup> However, there were diminished returns for Non-Hispanic Black youth regarding the positive effects of parental education on behaviors compared to Non-Hispanic White youth, suggesting the need to address societal barriers that hinder the effectiveness of family resources.<sup>18</sup>

### **Research Gaps and Future Directions**

Despite the existing literature, significant research gaps remain concerning the neurodevelopmental consequences of alcohol abuse in Hispanic adolescents in the Rio Grande Valley. More research is needed to:

*Characterize alcohol use patterns:* Detailed epidemiological studies are needed to precisely define the prevalence, patterns, and types of alcohol use within this specific population. This

would involve collecting data on binge drinking frequency, quantity of alcohol consumed, and age of initiation.

*Investigate specific neurocognitive deficits:* Further research should utilize neuropsychological assessments to identify specific cognitive deficits associated with alcohol abuse in Hispanic adolescents in the Rio Grande Valley, focusing on executive functions, memory, attention, and processing speed.

*Assess structural brain changes:* More neuroimaging studies, such as MRI and DTI, are necessary to determine the specific structural brain changes associated with alcohol abuse in this population, comparing them to age-matched controls and considering potential confounding factors such as socioeconomic status, acculturation, and co-occurring mental health disorders.

*Explore neurochemical mechanisms:* Research should examine the neurochemical mechanisms underlying alcohol's effects on the developing brain in Hispanic adolescents. This would involve investigating alterations in neurotransmitter systems, gene expression, and cellular processes.

*Evaluate the impact of cultural and socioeconomic factors:* Studies should examine how cultural factors (e.g., acculturation, family traditions) and socioeconomic factors (e.g., poverty, access to resources) interact with alcohol abuse to influence neurodevelopmental outcomes.

*Develop culturally appropriate interventions:* Research should focus on developing and testing culturally appropriate prevention and intervention programs for Hispanic adolescents in the Rio Grande Valley. These programs should consider the unique cultural contexts and address multiple risk and protective factors.

*Conduct longitudinal studies:* Longitudinal studies are crucial to understanding the long-term neurodevelopmental consequences of alcohol abuse, tracking cognitive and structural brain changes over time, and examining the impact on academic achievement, mental health, and social adjustment.

## **Conclusion**

Alcohol abuse during adolescence poses significant neurodevelopmental risks, impacting both cognitive function and brain structure. While existing research provides valuable insights into the effects of alcohol on the developing brain, more targeted studies are needed to understand the specific consequences for Hispanic adolescents in the Rio Grande Valley. Addressing the identified research gaps will enable the development of effective prevention and intervention strategies tailored to this population, ultimately promoting their long-term health and well-being. The unique social, cultural, and economic factors within this community must be considered to create culturally sensitive and effective interventions. The integration of epidemiological data, neuropsychological assessments, neuroimaging techniques, and a strong focus on community

engagement are essential for advancing our understanding of this critical public health issue. Further research will be vital in understanding the complex interplay between alcohol abuse, cultural factors, and neurodevelopment within this vulnerable population. Only through a comprehensive approach incorporating multiple disciplines can effective strategies be developed to mitigate the long-term consequences of alcohol abuse on the lives of Hispanic adolescents in the Rio Grande Valley.

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