Adverse Experiences, Hippocampal Connectivity, and Epigenetic Modifications in Adolescent Depression



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Hypothesis

Adverse experiences during adolescence (as environmental factors) may influence brain activity and epigenetic modifications, potentially contributing to the onset and progression of depression.



Study Design

This study employed a case-control design, recruiting 139 adolescents diagnosed with depression (BDI scores: Median = 29.0 (IQR: 21.0–35.0)) and 59 healthy controls (BDI scores: Median = 3.0 (IQR: 1.0–5.0)). Participants underwent structured assessments to evaluate adverse experiences during adolescence. DNA methylation was assessed from peripheral blood samples. Resting-state functional connectivity (rs-FC) analyses were performed using a seed-based approach, with the bilateral hippocampus as the region of interest (voxel-level p < 0.001, cluster-level p < 0.05, GRF corrected).

Results 0.008 0.008 AHR DNAm Lev 13 DNAm 0.006 14 DNAm 0.00 0.000 0.004 0.004 0.004 Ĕ Ħ 0.002 0.002 Ď Fig. 1 DNA Methylation Analysis Fig. 2 rs-FC with Bilateral Hippocampus as Seed Region С В n A r = -0.689 14 DNAm HR

Fig. 3A Relationship Between Adverse Experiences and rs-FC in Adolescents with Depression; 3B-D Healthy Group

Conclusion

Our findings suggest that adverse experiences during adolescence may lead to alterations in hippocampal connectivity and epigenetic modifications, potentially contributing to the pathophysiology of depression.

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