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Effects of Sudden Removal of High Social Enrichment upon Monoamine Levels in Cortical and Subcortical Structures of Rat Brain

Spencer Feehan, Rachel Lichtenstein, Hailey Robson, Dr. Deborah S. Kreiss

Background

- Objective: investigate the effects of removal of high social enrichment upon monoamine neurotransmitter levels in cortico-basal ganglia-thalamic circuitry
- Intention: model the sudden loss of social enrichment experienced during the COVID-19 pandemic
- High enrichment: larger cages with multiple toys, human handling, and playdates with 12 same sex non-cagemates
- Neurochemical evaluation: norepinephrine (NE), dopamine (DA), and serotonin (5-HT) were targeted because abnormalities of these neurotransmitters within cortical and subcortical structures underlie psychological disorders

Methods

 Subjects and Enrichment 28 male, 28 female Sprague-Dawley rats Day 34 to 76: all rats experienced high enrichment 4 times a week Day 77 to 106: experimental rats experienced reduction to standard enrichment, whereas control rats experienced continued high social enrichment
 Tissue Extraction Post-mortem bilateral punches obtained on Day 10 from PFC, MC, ACC, OFC, AMY, HYPO, HIPPO, L¹ MT, DS, & VS¹ Punches homogenized² and centrifuged Supernatants extracted and plated
 High Performance Liquid Chromatography (HPLC) Analysis Supernatants analyzed for NE, DA, 5-HT levels^{2,3,4}
 Data Analysis SPSS used for 2-Factor ANOVA analysis of data Two-tailed Student's t-tests used for pairwise comparisons (p<0.05) Data is expressed as mean ± 1 standard error of the mean (SEM) Outliers (± 2.0 standard deviations) eliminated

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Neurotransmitter Levels







5-HT OFC

Male Control







5-HT DS



5-HT VS



5-HT HYPO





Summary of Results

Change in Neurotransmitter Levels							
Monoamines	NE		DA		5-HT		
Sex	М	F	М	F	М	F	
PFC	↓			Ł	♦		
MC	↓	↓	↓		♦	¥	
OFC	↓		↓			¥	
ACC					↓	$\mathbf{\hat{\Lambda}}$	
LT	↓	↓		↓			
МТ			↓			¥	
HYPO	$\mathbf{\hat{\Lambda}}$	↓				¥	
DS	$\mathbf{\Lambda}$	↓			↓	$\mathbf{\hat{T}}$	
VS		↓			↓		
HIPPO			¥				
AMY				¥			

Table 1: change in neurotransmitter levels of experimental rats (underwent removal of enrichment) compared to control rats (underwent continued enrichment).

Conclusions

Removal of enrichment <u>decreased</u> monoamine levels in cortico-basal ganglia-thalamic circuits in both males and females:

Cortical structures: PFC, MC, & OFC
Subcortical structures: LT, MT, & VS

Sex influenced the effects of removal of enrichment:

 In ACC, 5-HT in males decreased, but increased in females

In HYPO, NE in males increased, but decreased in females

In DS, NE in males increased, but decreased in females

In DS, 5-HT in males decreased, but increased in females

Increased understanding of neurochemical changes induced by removal of enrichment will facilitate development of better therapeutic strategies for those experiencing a loss of social enrichment.

References

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2. Bishop C, George JA, Buchta W, Goldenberg AA, Mohamed M, Dickinson SO, Eissa S, and Eskow Jaunarajs KL (2012) Serotonin transporter inhibition attenuates L-DOPA-induced dyskinesia without compromising L-DOPA efficacy in hemi-parkinsonian rats. *European Journal of Neuroscience*. 36:2839-2848

3. HTEC-500, Amuzainc.com

4. EPC-700 Envision software, Amuzainc.com

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