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Pick Your Poison: How Aspartame Causes Anxiety

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Pick Your Poison: How Aspartame Causes Anxiety

Abby J. Kersey, Baylee R. Kram, Blessing O. Okosun, and Diane C. Darland Primary Research Project: BIOL491/503 Nutrition and Neuroscience Department of Biology, College of Arts and Sciences

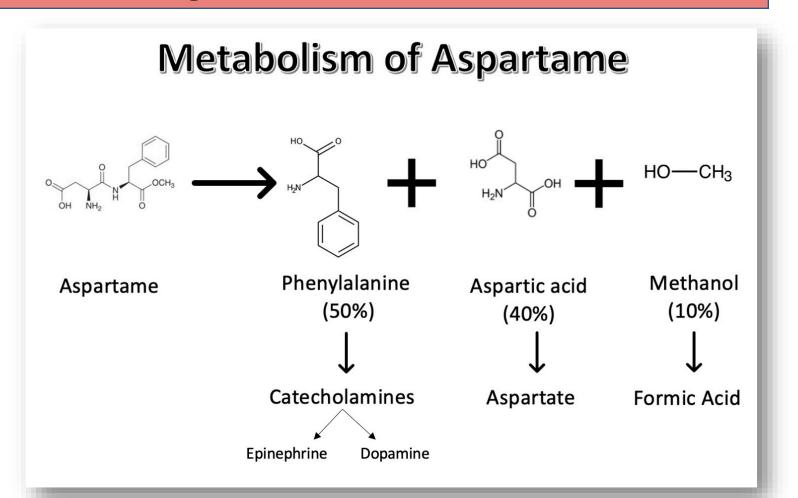
NORTH DAKOTA College of Arts & Sciences

Primary Research Article

Transgenerational transmission of aspartame-induced anxiety and changes in glutamate-GABA signaling and gene expression in the amygdala. Jones, S.K., McCarthy, D.M., Vied, C., Stanwood, G.D., Schatschneider, C., and Bhide, P.G. *PNAS* 2022 119 (49):e2213120119

What is Aspartame?

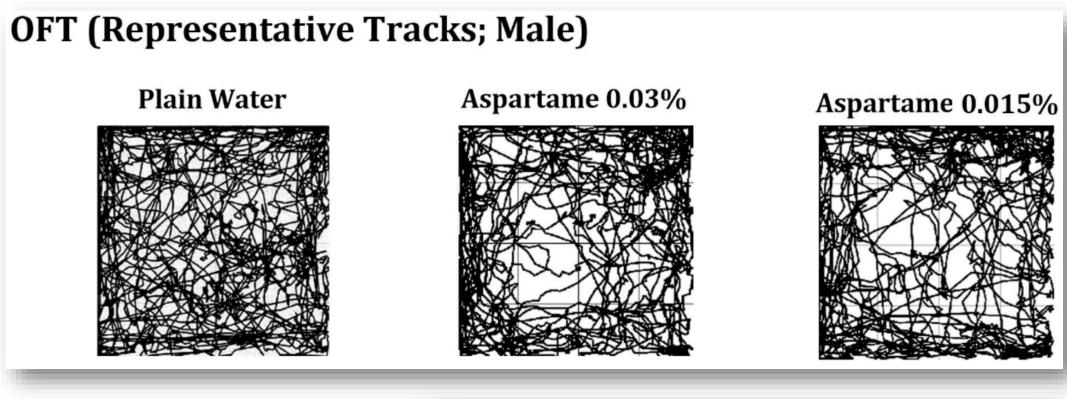
- Artificial sweetener found in ~5,000 diet foods and drinks
- ❖ A methyl ester dipeptide consisting of aspartic acid and phenylalanine



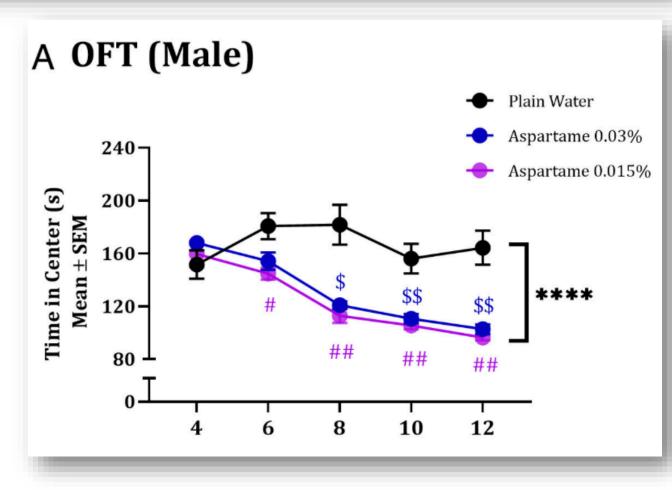
- Breaks down into phenylalanine, aspartic acid, and methanol in the digestive system
- Phenylalanine is a precursor and is converted to tyrosine which is a precursor of catecholamines
- Aspartic acid is an excitatory neurotransmitter

How can anxiety be measured in mice?

* What is the open field test? Mice with anxiety spend relatively shorter time in the center areas than mice without anxiety.

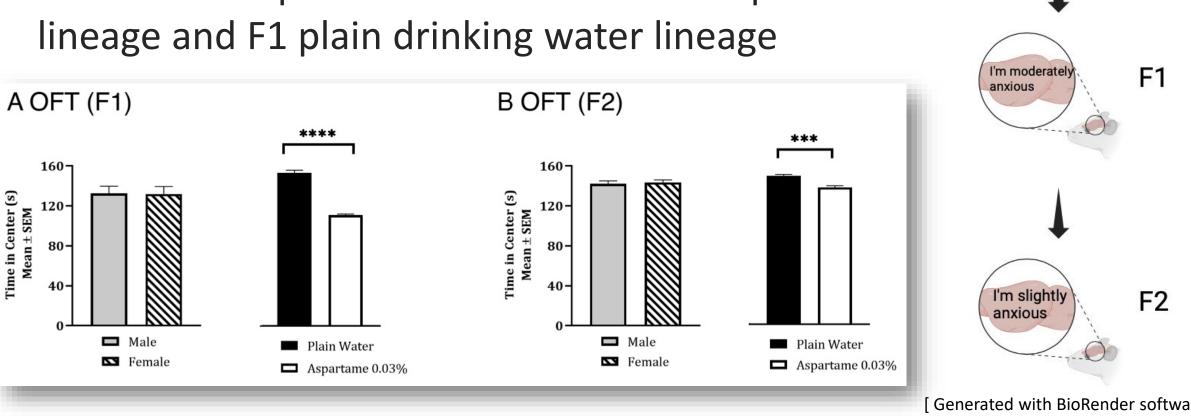


Anxiety phenotypes began at 6 weeks of aspartame exposure and persisted until 12 weeks



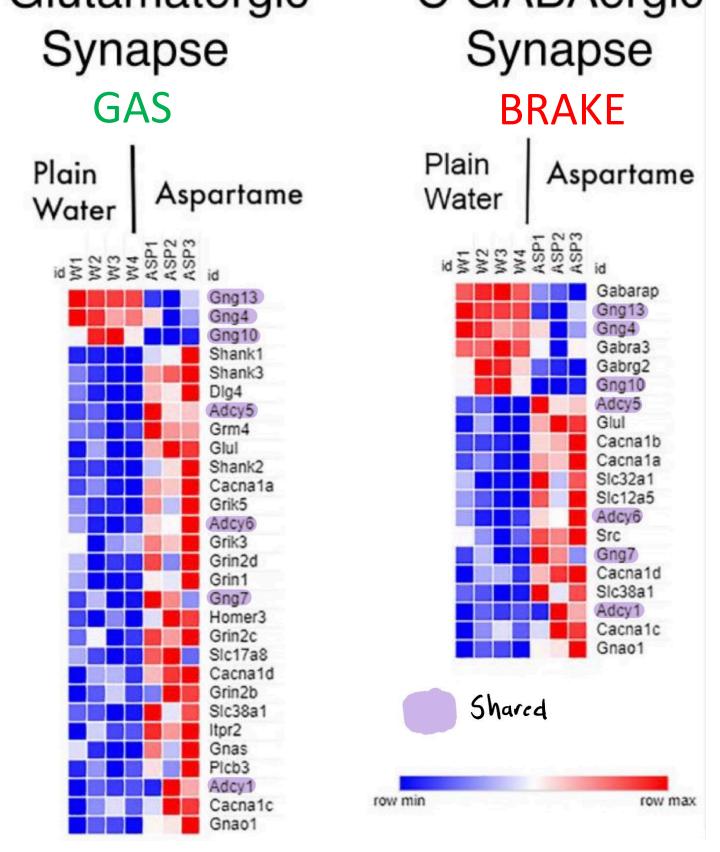
Is aspartame-induced anxiety "inherited" in mice?

- Transgenerational Transmission-phenotypes that are inherited by multiple generations descending from the directly exposed individuals
- ❖ F1 mice were produced from three F0 male mice lineages following 12 wk: 0.03% & 0.015% aspartame exposure, and plain water
- F2 mice were produced from F1 0.03% aspartame lineage and F1 plain drinking water lineage



How does aspartame treatment change gene expression in mice brains?

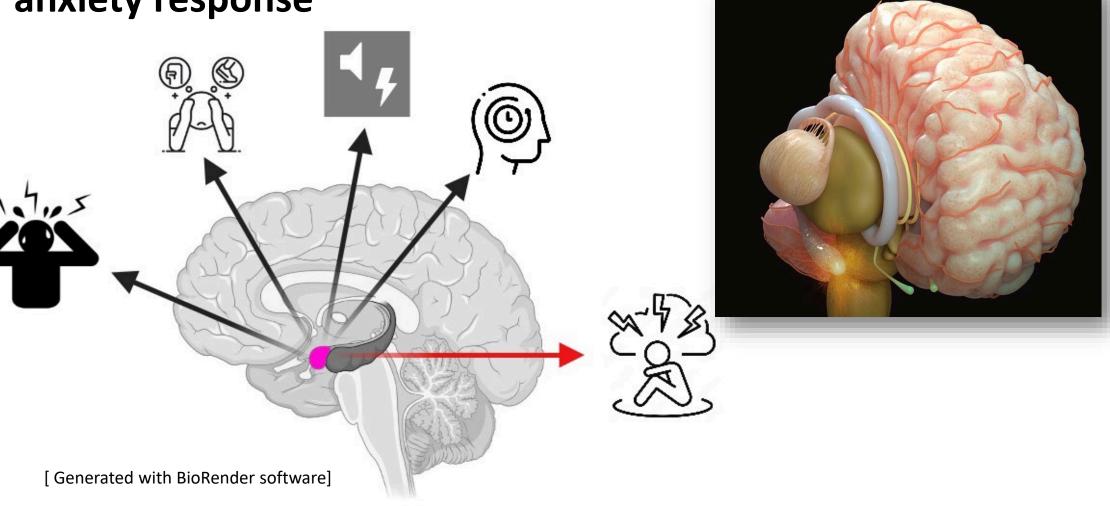
- Glutamatergic Synapse- Glutamate is the major excitatory neurotransmitter at almost all synapses in the vertebrate central nervous system (CNS) *Acts as the gas pedal*
- **❖GABAergic Synapse** maintains the balance between excitatory and inhibitory synaptic transmission, and plays a crucial role in normal function and long-term homeostasis of the neuronal circuits *Acts B Glutamatergic C GABAergic as the brake pedal*
- Aspartame consumption shifts the excitationinhibition equilibrium in the amygdala toward excitation
- Outcome: the amygdala now has double gas pedals, and no brake pedal.



What does the Amygdala do?

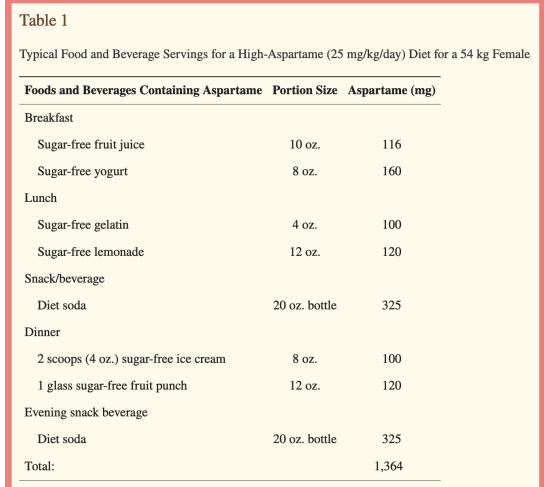
- Part of the Limbic Loop which acts as the center of emotions, behavior, and memory
- ❖ Neural Excitation-inhibition equilibrium in the amygdala plays a central role in the regulation of anxiety

Stimulation of the amygdala causes an increased stress and anxiety response



How does aspartame affect humans?

- Neurobehavioral effects of aspartame consumption. Lindseth, G.N., Coolahan, S.E., Petros, T.V., and Lindseth, P.D. Res Nurs Health 2014 37(3):185-193.
- FDA "safe" recommended *daily* value is 50 mg per 1 kg (RDA)
- ❖ Aspartame exposure in students was 50% of RDA for 8 days
- Students reported impaired working memory and spatial orientation



Variable	M	SD	Paired t-test	p	
Spatial orientation					
High-aspartame	14.1	4.2	2.4	.03*	
Low-aspartame	16.6	4.3			
Working memory					
High-aspartame	730.0	152.7	1.5	N.S.	
Low-aspartame	761.1	201.6			
Mood (irritability)					
High-aspartame	33.4	9.0	3.4	.002**	
Low-aspartame	30.5	7.3			
Depression					
High-aspartame	36.8	7.0	3.8	.001**	
Low-aspartame	34.4	6.2			

- Vulnerable populations are at greater risk with consecutive aspartame consumption
- ❖ 13/40 participants dropped from study after only 7 days
- Aspartame group reported more symptoms than placebo group

Take Home Message: Limit Your **Aspartame Consumption!**