Music, Brains, Drugs & The Beatles

David Hattner, Portland Youth Philharmonic Bill Griesar, PSU/OHSU/Northwest Noggin Aaron Eisen, OHSU/PSU/NW Noggin Jeff Leake, PSU/NW Noggin

Charles Burchfield, Autumnal Fantasy

Northwest Noggin

Community outreach

nwnoggin.org

Art & Brains!

NEURON Myelin Nodes of Ranvier

Axon Axon Axon Soma hillock Terminals Dendrites

nwnoggin.org

nwnoggin.org

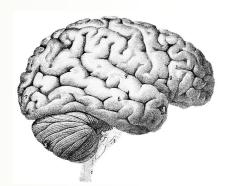
Sleep Anxiety Depression ADHD Autism Development Perception Drugs Memory Language **Racial Bias**

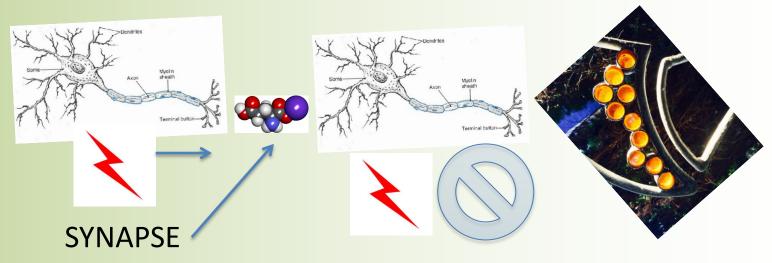
> What you do has relevance GO PLACES

Gilbert & Sullivan

Your brain: made of cells

- Neurons
- Neurons carry *electrical* messages
- Neurons connect chemically across synapses
- Neurotransmitters





All cells have membranes

Neurotransmitter

or drug

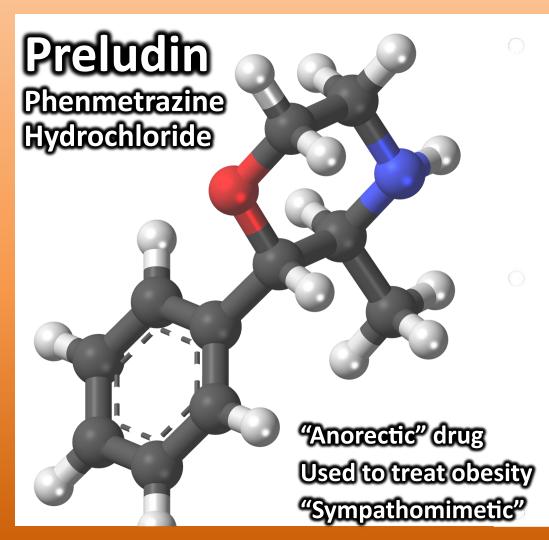
Outside cells, including neurons

Many drugs, including psychedelics, cannot get through, but instead act at RECEPTORS to affect neuron function...

Drugs (chemicals) attach (or "bind") to receptors, changing the activity of affected neurons...

RECEPTOR: "Protein machine"

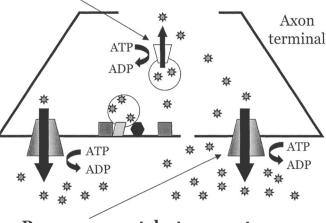
Inside cells



Amphetamine/Meth *reverse* transporters

Especially at DA synapses...

Reverses vesicular transporter



Reverses re-uptake transporter

Amphetamine reversal of both vesicular and re-uptake transporters cause massive DA increase in synaptic cleft...

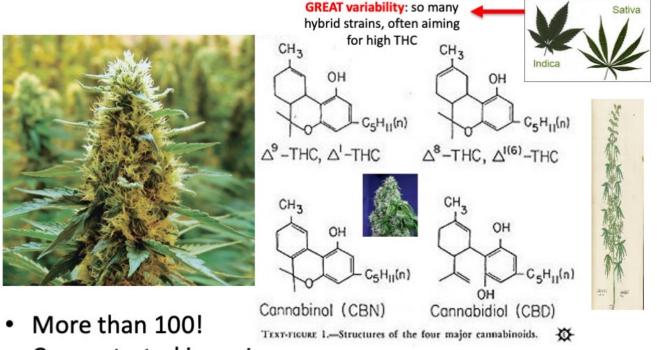
She Loves You I Want to Hold Your Hand

The Beatles 1963

CANNABIS

Cannabis contains cannabinoids

SOURCE: Cannabis, a complex plant: different compounds and different effects on individuals, Atakan (2012)



- Concentrated in resin
- Lots of variability, depending on strain, other factors...

Cannabinoids act at cannabinoid receptors: CB1 and CB2

dorsal vagal

complex

emesis

CB1 Receptors Neocortex Basal ganglia Amygdala Hypothalamus Hippocampus Cerebellum

Susan R. B. Weiss, Katia D. Howlett, Ruben D. Baler (2017)

cerebral cortex decision making, cognition, & emotinal behavior

caudate nucleus

regulate movements & influence various types of learning

globus pallidus regulate voluntary movements

amygdala responsible for anxiety & stress, emotion & fear, pain

> hypothalamus body temperature, feeding, neuroendocrine function

> > hippocampus ' memory & learning

substantia nigra / important role in reward, addiction, & movement

motor control & coordination

Cannabinoids reduce pain

A large body of literature indicates that cannabinoids suppress behavioral responses to acute and persistent noxious stimulation...(Walker JM, Hohmann AG, 2005)

Co-administration of cannabinoids and opioids allows for pain relief with a lower opioid dose!

Before Tail Flick

(e.g., Wilson AR, Maher L, Morgan MM, 2008)



STILL MORE therapeutic effects



- Multiple sclerosis
 - E.g., "Current status of cannabis treatment of multiple sclerosis," Deutsch et al (2008)
- Epilepsy
 - E.g., "The case for medical marijuana in epilepsy," Maa (2014)
- Cancer
 - E.g., "The combination of cannabidiol and Δ 9-THC enhances the anticancer effects of radiation in an orthotopic murine glioma model," Scott et al (2014)

More therapeutic effects



Appetite stimulation

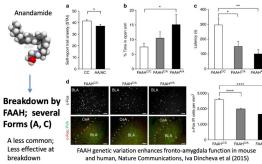
(e.g., Foltin, 1988; Grotenhermen, 2012) Why is this therapeutic?

Nausea relief

(e.g., Parker et al (2011); "The anti-emetic effect of cannabinoids has been shown across a wide variety of animals that are capable of vomiting in response to a toxic challenge." Also studies referenced by the National Cancer Institute at cancer.gov; though chronic use linked to hyperemesis syndrome: Soriano-Co M. 2010)

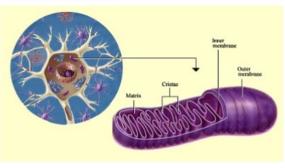
Anxiety: Genetic protection?

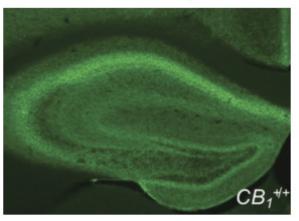
Decreased anxiety in humans and mice with FAAH C385A



A cannabinoid link between mitochondria and memory

Etienne Hebert-Chatelain, et al Nature (2016)





Cellular activity depends on mitochondria Mitochondria site of cellular respiration (ATP) Mitochondria have many CB1 receptors (mtCB1) Cannabinoid action at mtCB1 inhibits respiration Hippocampus starved of energy (less ATP) A mechanism for amnesia..?



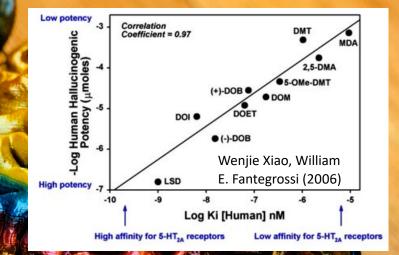
I'm a Loser I'll Follow the Sun Nowhere Man I'm Looking Through You If I Needed Someone 1965

The Beatles

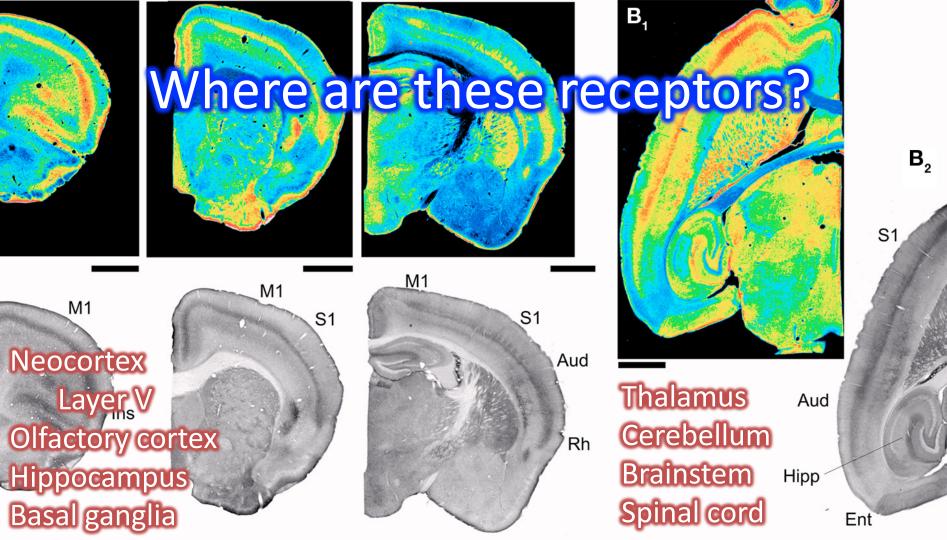
Psychedelics

Chemicals that produce unusual perceptual and cognitive distortions. Many derived from plants, though some are synthetic. Mescaline, psilocin, DMT, LSD

Most psychedelics are agonists at ONE type of SEROTONIN receptor (5-HT2A)



Potency linked directly to hallucinogenic effects



Psychedelics affect the "gating" of sensory input

Changes in perception

"If the doors of perception were cleansed, everything would appear to man as it is, infinite" - William Blake

"The legs, for example, of that chair - how miraculous their tubularity, how supernatural their polished smoothness" - Aldous Huxley, "The Doors of Perception" (1954)

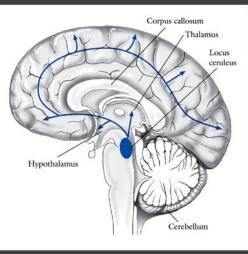
"I looked around me and noticed details of physiognomy that had never struck me before. Each pore in my companion's skin was now visible..." - Solomon Snyder, "Drugs and the Brain"

"I clapped my hands and saw sound waves passing before my eyes" - Solomon Snyder, "Drugs and the Brain" Image courtesy of the Allen Institute for Brain Science

Thalamus

Cortex

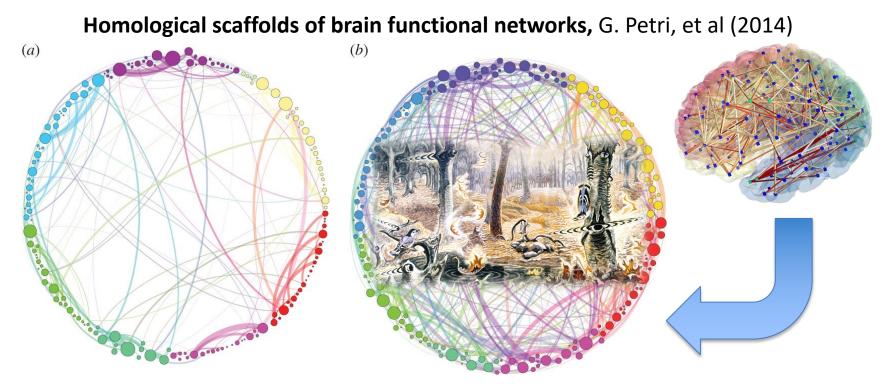
Psychedelics Enhance Sensory Responses in the Locus Coeruleus (LC) via 5-HT_{2A} Receptors



"...it is of interest that the systemic administration of LSD, mescaline, or other psychedelic hallucinogens in rats, although decreasing spontaneous activity, produces a paradoxical facilitation of the activation of LC neurons by sensory stimuli..." (Aghajanian 1980; Rasmussen & Aghajanian 1986)

Ordinary stimuli become extraordinary

Greater functional connectivity



"there is an increased integration between cortical regions in the psilocybin state...One possible by-product of this greater communication across the whole brain is the phenomenon of synesthesia which is often reported in conjunction with the psychedelic state..."

"Default mode" network

Critical for... Self-reflection Self awareness Rumination

1. VENTROMEDIAL PREFRONTAL CORTEX 2. DORSOMEDIAL PREFRONTAL CORTEX 3. ANTERIOR, HIDDLE, CINCILLATE, CORTEX BUT: increased plasticity

Decreased coordination

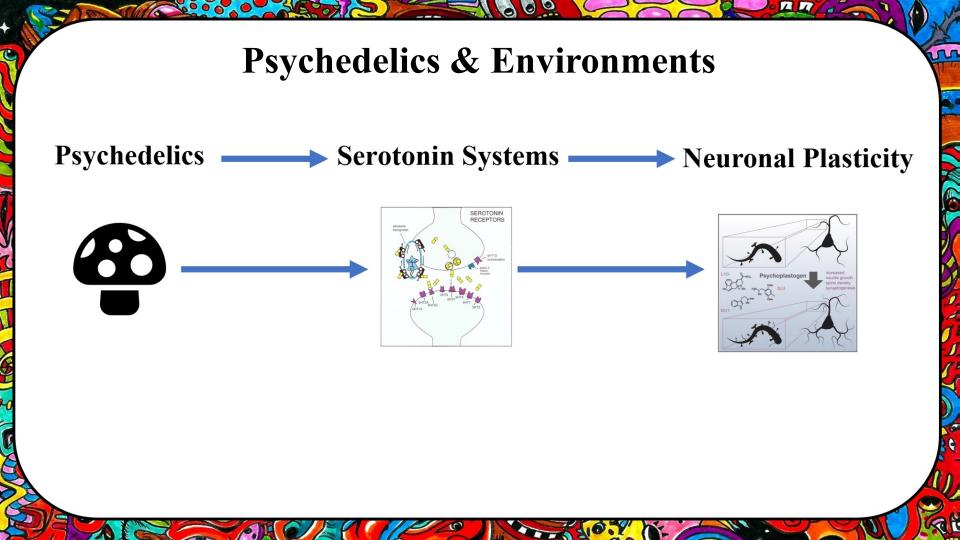
while on psychedelics

"We all live in a yellow submarine..."



Got To Get You Into my Life Yellow Submarine I'm Only Sleeping Love You Too



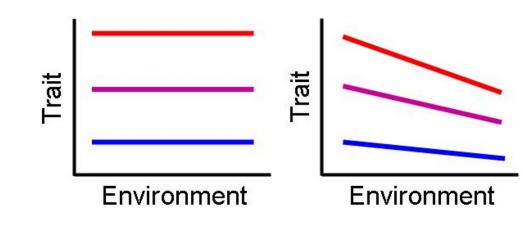


Psychedelics & Environments

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P.H.

No Plasticity Plasticity





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Set & Setting



P.F



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Psychedelics & Environments



The Efficacy of Music Therapy Judith H. Wakim, EdD, RN, CNE, Stephanie Smith, MSN, CRNA, Cherry Guinn, EdD, RN

Undergoing a procedure that requires anesthesia can be anxiety provoking. Anxiety is associated with increases in beart rate and blood pressure and other changes that can have a negative impact properatively; during the induction, maintenance, and emergence phases of anesthesia; and postoperatively. Music therapy is a nonpharmalogical intervention that has the ability to reduce anxiety levels in some patients. This review presents research studies that have been conducted on the effects of music therapy for patients in different clinical settings. In general, the majority of the published articles reviewed revealed that listening to music was beneficial to the patient no matter the setting. Offering a music selection to patients before anesthesia could enhance its positive effect. Perianesthesia nurses could easily develop a protocol for different situations where patients will be exposed to interventions where the use of general or local anesthesia is expected.

Keywords: Perianesthesia anxiety, music therapy, nonpharmacological anxiolytic.

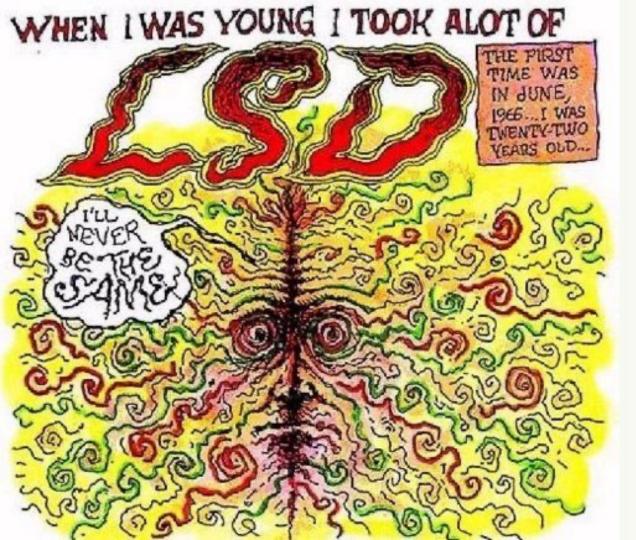
© 2010 by American Society of PeriAnesthesia Nurses



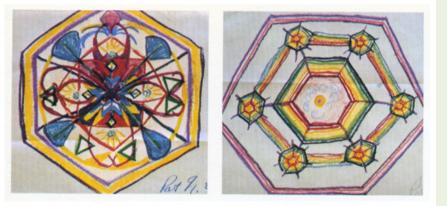


Hey Bulldog Only a Northern Song Your Mother Should Know





Robert Crumb "LSD"



Is there a link between creativity and hallucinogens?

Distortion of body extremities is illustrated in a sophisticated way by a professional painter. This drawing was executed by a well-known Czech artist after the recovery from LSD intoxication.

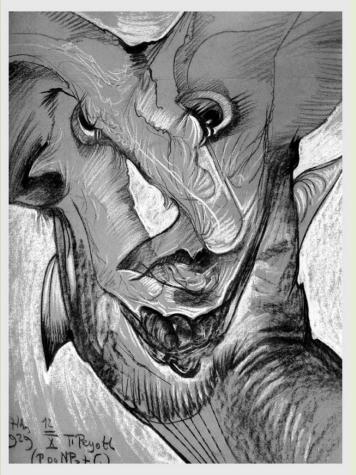
Mandalas created before (left) and during (right) an LSD session conducted in 1972 at the Maryland Psychiatric Center

In 1952 Lászlo Mátéfi described how an experimental subject under the influence of a hallucinogen experienced a discrepancy between his intention and performance while making a portrait:

I see the object correctly but draw it falsely; my hands won't follow it.... This desire to paint is harder and harder for me to perform since the expanse of my experience pulls me more and more into it. Myself, the drawing, and the surroundings create a unity--and that hinders me because I cannot concentrate on the model. I have the need to bring everything including the painted picture into the surface of the image. Had the painting process been more of a technical success, I would have been able to produce a fantastically good work

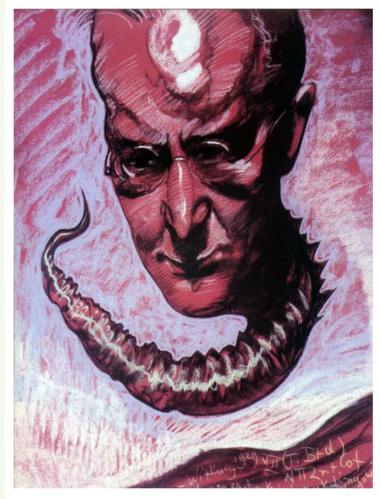


Stanislaw Witkacy "Teodora Bialynickiego-Birul" 1929



Witkacy made this portrait under the influence of mescaline.

> Witkacy made this portrait under the influence of peyote.



Stanislaw Witkacy "Neny Stachurskiej" 1929



Marijke Dunham, outfits for the Beatles

"Drugs were a big part of it, the aesthetic has a bad name due to that. I mean Van Gogh painted his strange paintings and there's been no mention of drugs, but drugs definitely had an influence as far as the cause was concerned, with the strange moving lines, definitely ... and since I'm not ashamed about it I don't have to hide it. That's where it comes from." -Marijke Dunham "I didn't take LSD, I didn't want to be that far from my senses"-Peter Blake





PORTLAND YOUTH PHILHARMONIC

DAVID HATTNER, MUSICAL DIRECTOR

