

Psychedelic Medicine: A Journey into the History, Cultural Roots, Science, Therapeutics and Potentials

INTERNATIONAL RESEARCH CONGRESS FOR INTEGRATIVE MEDICINE & HEALTH

CLEVELAND, OH

APRIL 12, 2024

10:45 am-12:00 pm



LIVE Q&A! Please Scan QR before we start or Go to <u>https://sli.do</u> and enter #1260685



• Justin Laube, MD: UCLA Center for East-West Medicine & Consultant

• Thais Salles Araujo, MD: UCLA Palliative and Hospice Care

 Leslie Mendoza Temple, MD: Endeavor Health, University of Chicago Pritzker School of Medicine

 Mikhail Kogan, MD: George Washington University, GW Center for Integrative Medicine

 Erika Steinbrenner, MD: Imagine Healthcare, Epiphany Wellness, Symetria Recovery







#### **OVERVIEW**

- Arriving and nesting psychedelics within integrative medicine (Justin) 10 min
- Historical and cultural context of psychedelic substances (Thais) 10 min
- Broad overview of psychedelics from conventional medicine viewpoint (Leslie) 10 min
- Psilocybin case presentation (Misha) 10 min
- Ketamine case presentation (Erika) 10 min
- Ethics, Safety 5 min (Leslie)
- Live Questions & Answers 15-20 min





### **Disclosures**

#### Leslie Mendoza Temple, MD

Scientific Advisory Board Member Ashford International, an independent lab testing and consumer platform for CBD hemp products.

#### Mikhail Kogan, MD

Royalties, Medical Marijuana Book, 2021 and 2023, Penguin Random House





School of Medicine & Health Sciences





# Part 1: Psychedelics within integrative health; their definitions, ancient history & cultural roots

Justin Laube, MD Thais Salles Araujo, MD





Endeavor Health.

| An official website of the United States government  | Here's how you know ~           |                             |                |                         |               |                        |  |  |  |
|--|---------------------------------|-----------------------------|----------------|-------------------------|---------------|------------------------|--|--|--|
| 4 U.S. Department of Health and Human Services   | National Institutes of Health   | tional Institutes of Health |                |                         |               | Información en Español |  |  |  |
| NAtional Center for<br>Complementary and<br>Integrative Health   |                                 |                             | Se             | arch NCCIH              | I             | Q                      |  |  |  |
|  | Health Info F                   | Research G                  | ants & Funding | Training                | News & Events | About NCCIH            |  |  |  |
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| psychedelics   | ychedelics                      |                             |                | Refine Your Results     |               |                        |  |  |  |
| Showing results 1-7 of 7   |                                 |                             |                | Director's Messages (1) |               |                        |  |  |  |
| NIH Psilocybin Research Speaker Series—June 10, 2021 Lecture   NCCIH<br>https://www.nccih.nih.gov/news/events/nih-psilocybin-research-speaker-seriesjune-10-                 |                                 |                             |                | Events (4)              |               |                        |  |  |  |
| 2021-lecture<br>Description 12:30–1:00 p.m. ET – <b>Psychedelics</b> in Headache Medicine: Historicalof  |                                 |                             |                | Press Releases (0)      |               |                        |  |  |  |
| Medicine 1:40–2:15 p.m. ET – Psychedelic Medicine and Ethics Paul S. Appelbaum   |                                 |                             |                | Research Blog Posts (0) |               |                        |  |  |  |
| NIH Psilocybin Research Speaker Series (Ma<br>https://www.nccih.nih.gov/news/events/nih-p<br>2021-lecture  |                                 | riesmay-                    |                | Why                     | <i>י</i> ?    |                        |  |  |  |
| Patient's Perspectives on the Uses of <b>Psychedelics</b> in Medicine Erica Rex, M.A., Journalist  |                                 |                             |                | Federal legal statu     |               |                        |  |  |  |
| NIH Psilocybin Research Speaker Series - June 7, 2021 Lecture   NCCIH<br>https://www.nccih.nih.gov/news/events/nih-psilocybin-research-speaker-seriesjune-7-<br>2021-lecture |                                 |                             |                | Fear                    |               |                        |  |  |  |
| and Anti-Inflammatory Effects of Psychede  | lics in Rodent Preclinical Mode | els Charles                 |                | Silo                    |               |                        |  |  |  |

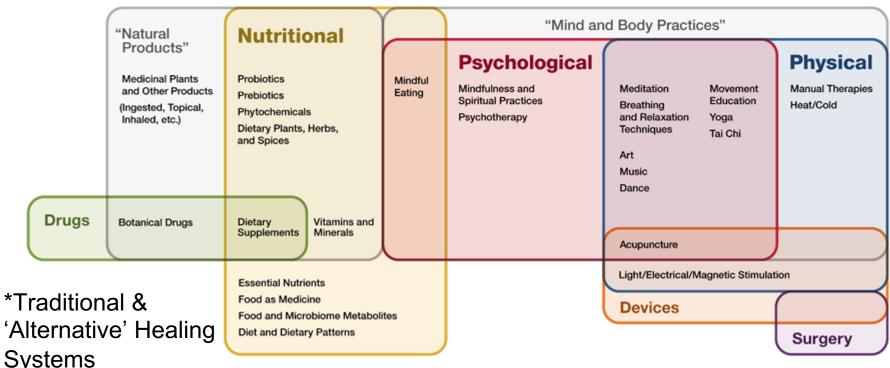
https://www.nccih.nih.gov/about/offices/od/director/past-messages/including-

spirituality-into-a-fuller-picture-of-research-on-whole-person-health

. . .



# Where do psychedelics fit?

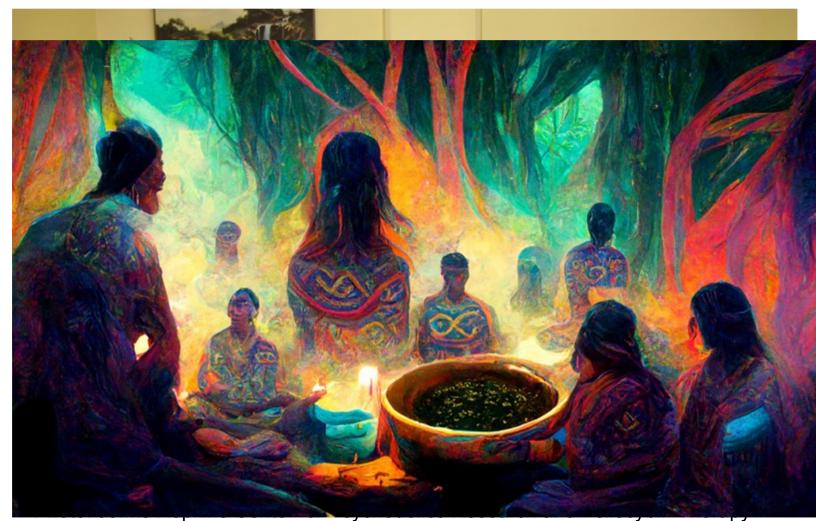


Psychedeli c Assisted Therapy -Medical Model

This is an example of the "Treatment Phase"

Set & Setting

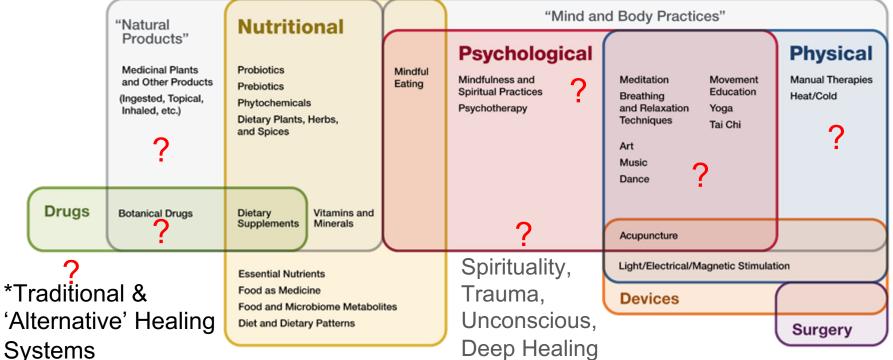
Prep & Integration

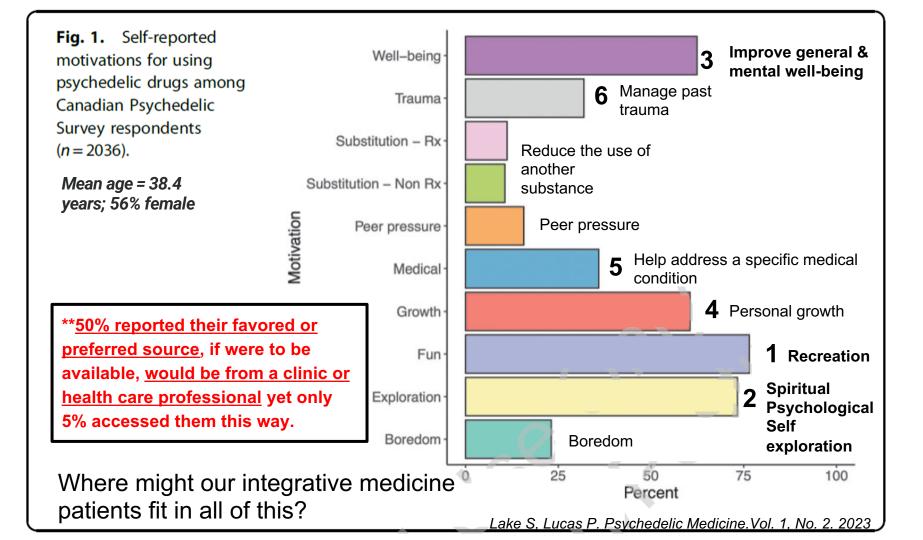




# Where do psychedelics fit?







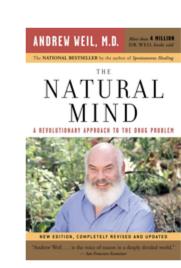
# Remembering our Roots as an IM field - "the Wild West" of Healing and Commitment to our patients

1992













"We realized we had to bring the 'spirit' of healing' back into medicine"

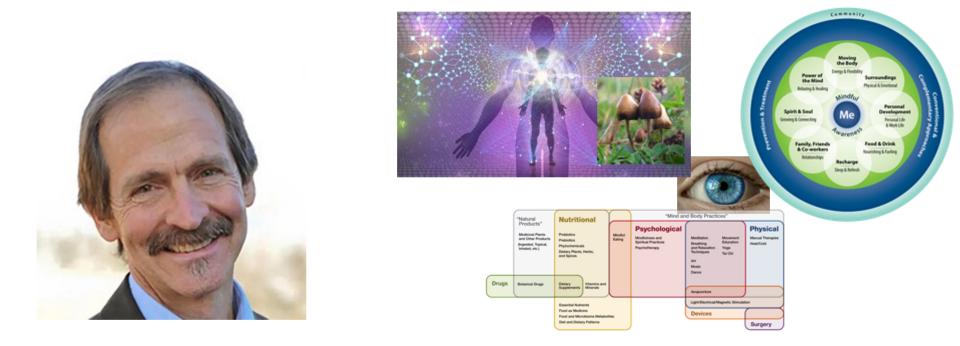
Dr. Gladys McGarey Age 103!

1972 - LSD Dr. Andrew Weil



AND ALTERNATIVE MEDICINE

ACADEMIC CONSORTIUM FOR Integrative 1999 Medicine & Health



"Integrative & Holistic medicine and Psychedelic Medicine come from the same shared roots, share principles core, speak to the same demographic and I think this is a marriage that needs to be supported and encouraged."

-Scott Shannon, MD



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#### **Psychedelics** definition

Psychedelics are psychoactive substances able to induce *altered states of consciousness* marked by significant shifts in perception, mood, and thought patterns. Commonly reported experience include an increased sense of *interconnectedness*, *transcendence* of time and space and a felt sense of *sacredness*.

Nichols DE. Psychedelics. Pharmacol Rev. 2016 Apr;68(2):264-355. doi: 10.1124/pr.115.011478. Erratum in: Pharmacol Rev. 2016 Apr;68(2):356. PMID: 26841800; PMCID: PMC4813425.



#### **Psychedelics** definition

These substances have been used for millennia for **ceremonial and therapeutic** purposes, and over the past century, have gained **scientific medical interest**.

Nichols DE. Psychedelics. Pharmacol Rev. 2016 Apr;68(2):264-355. doi: 10.1124/pr.115.011478. Erratum in: Pharmacol Rev. 2016 Apr;68(2):356. PMID: 26841800; PMCID: PMC4813425.



#### **Psychedelics definition**



#### Classic

 Primarily serotonin receptor

#### Non classic •

#### Other mechanisms



lbogaine





# Ayahuasca (DMT) Mescaline

Peyote cactus







#### Evolving terminology



#### Hallucinogens

- 1600's
- Latin root "Mind journeying"



#### **Psychedelics**

- 1956
- Greek origin "Mind manifesting"



#### Entheogens

- 1979
- Greek "entheos", god or divine – "Assessing the divine within"





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#### Ecopsychology: Psychedelics and human evolution







#### **Evolution selection**

Rodríguez Arce JM, Winkelman MJ. Psychedelics, Sociality, and Human Evolution. Front Psychol. 2021 Sep 29;12:729425. doi: 10.3389/fpsyg.2021.729425. PMID: 34659037; PMCID: PMC8514078.

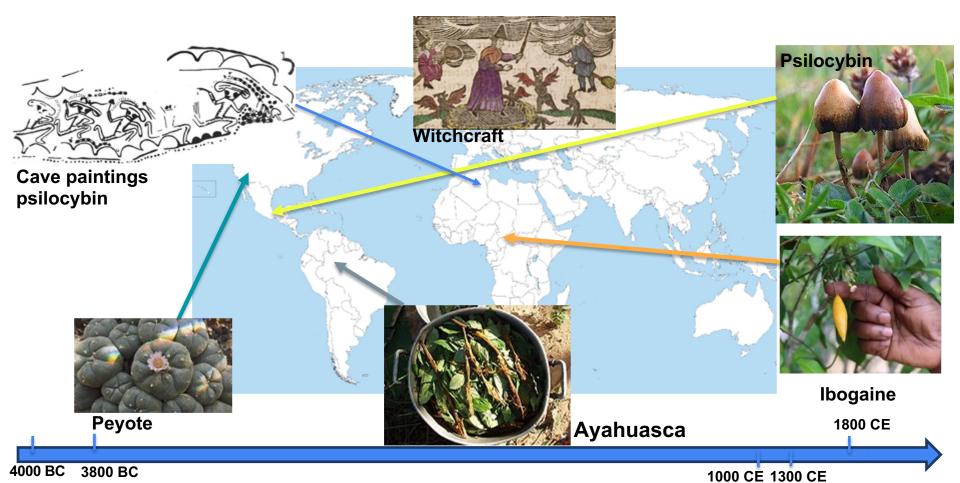
# Shamanism - religion and medicine roots



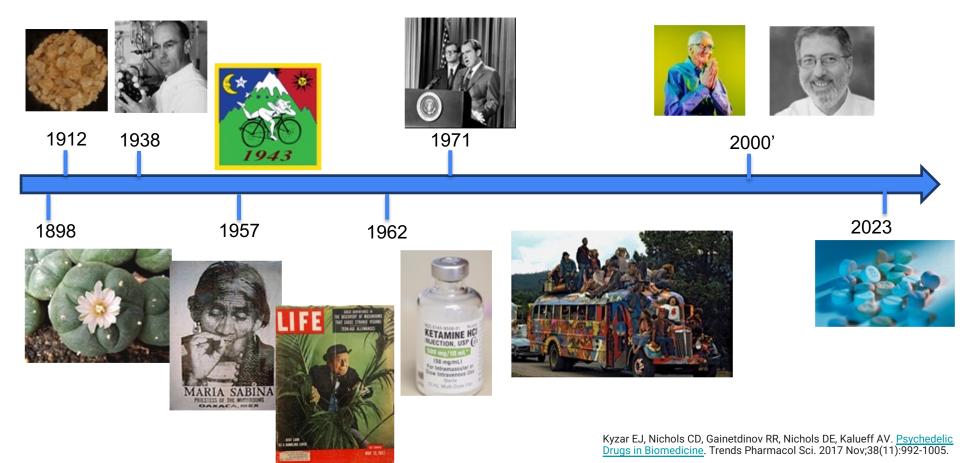
Maria Sabina -Curandera Pajé - "wizard" in tupi-guarani

Breathwork, chanting, dancing, drumming

#### 6000 + years of psychedelics history around the world



#### Psychedelics – Newer biomedical history



### Part 2: Broad overview of psychedelics from conventional medicine viewpoint Psilocybin & Ketamine Case Presentations

Leslie Mendoza Temple, MD Mikhail Kogan, MD

Erika Steinbrenner, MD





Endeavor Health.



LIVE Q&A! Please Scan QR before we start or Go to <u>https://sli.do</u> Enter #1260685

### Broad overview of many, but not all psychedelics

- Mescaline from Peyote, San Pedro and Peruvian Torch cacti
- DMT and Ayahuasca
- 5-Methoxy-DMT or "The Toad"
- Ibogaine
- LSD
- MDMA
- Kratom\*
- Psilocybin (Mikhail Kogan)
- Ketamine (Erika Steinbrenner)

\*not a psychedelic substance







#### Mescaline: Peyote, San Pedro & Peruvian Torch Cacti

**MOA:** Serotonin receptors

**Route:** Oral- fresh or dried cactus; ingest powder or boil as a slurry to drink

**Onset:** 30-60 min for pure mescaline; 2-4 hours for cactus ingestion

Duration: Long duration-12-14 hours

**Effects:** Euphoria, increased tactile sensation; Reduced anxiety, depression, PTSD, and alcoholism; Increased spiritual openness; Used for medicinal purposes as well.

Side effects: nausea, vomiting, diarrhea

**Considerations:** Peyote is only legal for use in the Native American Church for religious ceremonial purposes, revered like the Eucharist is for Christians. Overharvested; Peyote can take 15-20 years to mature.



`∩

#### DMT (DiMethylTryptamine) & Ayahuasca

Ayahuasca is a combination of two plants from the Amazon basin- a leaf containing DMT and a vine containing an MAO inhibitor.

**MOA:** Serotonin and Sigma-1 receptors; DMT occurs naturally in animals & humans. MAOI component prevents DMT deactivation in gut.

Route: Oral drink (Ayahuasca); smoked leaf (DMT)

**Onset:** Oral 30-60 min; peak at 2-3 hours **Duration:** 4-6+ hours

**Effects:** Visual patterns, contact with 'beings', euphoria, heightened perception; provide access to normally invisible and immaterial worlds

Side effects: "The purge": vomiting, diarrhea; increase BP, HR

**Considerations:** Ayahuasca is used ceremonially; Has most potential drug interactions due to MAO activity.



A member of Brazil's Huni Kui tribe prepares ayahuasca to use in a healing ritual. The psychoactive plant preparation is indigenous to the peoples of the Amazon basin. PHOTOGRAPH BY LUNAE PARRACHO, REUTERS



### 5-MethOxy-DMT: "The Toad" or 5-MeO-DMT

Parotid gland secretions of the Sonoran Desert Toad and the *Anadenanthera* bean of the Amazonian basin

MOA: Serotonin receptor activation

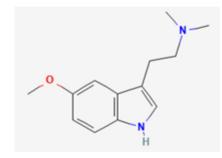
Route: Smoked, vaporized or nasal insufflation

**Onset:** Rapid onset & peaks 2-6 minutes (smoked) **Duration:** 30-60 min

**Effects:** Ego-dissolving, intense mystical experience; potential long lasting positive effects on life satisfaction, depression and anxiety

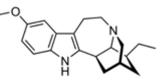
Side effects: Flashbacks, panic attacks

**Considerations:** 5-Methoxy-DMT is about 10 times as potent as DMT. Environmental impact on toad population is significant.









Tryptamine derived from the African shrub, *Tabernanthe iboga;* origin: Bwiti spiritual tradition in Gabon, Africa.

MOA: Serotonin, opioid, NMDA receptors

Route: Oral

**Onset:** Slow: 1-3 hours **Duration:** 24 hours, with prolonged 'afterglow' for weeks

**Effects:** Potential anti-addiction properties for alcohol, cocaine, methamphetamines, opioids, nicotine; profound spiritual experiences/revisit repressed memories

**Side effects:** Cardio: prolonged QT interval; Neurologic/GI: ataxia; nausea, vomiting

**Considerations:** Multiple Ibogaine-related deaths have occurred in non-medical settings likely pre-existing cardiac.

Supply is threatened.





#### LSD: LySergic Acid Diethylamide or "Acid"

**MOA**: Serotonin activity, maybe some dopamine influence

**Dosing:** Almost always orally as LSD-saturated blotter paper

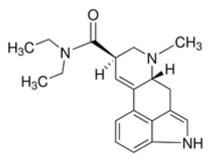
**Onset:** 15-45 minutes; **Peak:** 3-6 hours **Duration:** 12 or more hours

**Effects:** sensory enhancement (taste/smell), visuals, profound life-changing spiritual experiences or personal revelations; connectedness to universe

Side effects: acute anxiety

**Considerations:** long duration makes application of LSD challenging to apply in clinic setting

Notes: No known lethal dose





### MDMA (Ecstasy, Molly): 3,4-

Methylenedioxymethamphetamine

Psychedelic and stimulant effect. FDA just approved a new drug application for PTSD in association with specific therapy.

**MOA**: Increases serotonin, norepinephrine & dopamine by increasing release and decreasing reuptake.

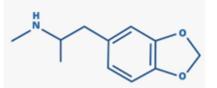
**Onset**: Oral- 45 minutes onset; peaks 15-30 min later.

Duration: 3 hours acutely; can have effects for days afterwards.

**Effects**: Energizing effect, distortions in time and perception, enhanced enjoyment from sensory experiences. Enhances pro-social skills; can increase self-awareness and empathy.

**Side effects**: high blood pressure, jaw-clenching, faintness, panic, hyperthermia, water overload, high-risk sexual behavior, reduced driving safety/motion perception

**Considerations**: Contamination of MDMA may be found with stimulants, bath salts, caffeine, ephedrine, cocaine, heroine, methamphetamine, and others.





#### Kratom (not psychedelic)

Southeast Asia (Thailand); A.k.a. thang, kakuam, thom, ketum, and biak.

MOA: multiple opioid (mu, delta, K) receptors affected;

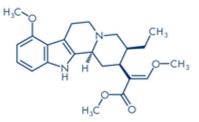
**Effects:** Potential help with opioid dependence, pain management; Low dose: Stimulating. High dose: Sedating. Perceived as a cheaper, more "natural" alternative to opioids.

Side effects: Addiction potential; contamination concerns;

**Considerations:** Overdose looks like an opioid toxidrome and is treated similarly.



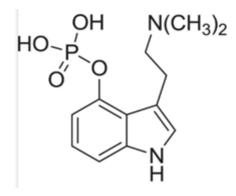
On DEA's "Drug of Concern" List



 Eastlack SC, et al. Kratom-Pharmacology, Clinical Implications, and Outlook: A Comprehensive Review. Pain Ther. 2020 Jun;9(1):55-69.
 Gorelick DA. Kratom: Substance of Abuse or Therapeutic Plant? Psychiatr Clin North Am. 2022 Sep;45(3):415-430.

### **Psilocybin (Mushrooms)**

- MOA: Serotonin receptor agonist, particularly targeting 5-HT2A receptors
- **Route:** Typically oral, consumed in dried mushroom form or as an extract (note most trials to date use injection of pure extract)
- **Onset:** 20-40 minutes after ingestion
- **Duration:** Effects last 4-6 hours, with possible after effects lasting up to 24 hours
- **Effects:** Euphoria, altered perception of time and space, visual and auditory hallucinations, introspection, spiritual experiences
- **Side Effects:** Nausea, anxiety, paranoia, increased heart rate, pupil dilation, derealization or depersonalization
- **Contraindications:** History of psychosis or psychotic disorders, severe cardiovascular disease, pregnancy
- **Combining with SSRIs/SNRIs**: May diminish effects due to serotonin receptor competition; potential for increased risk of serotonin syndrome, use caution





#### Case: Psilocybin

44 year old woman with fibromyalgia (headache, neck, back), anxiety, insomnia, severe childhood trauma due to abuse and abandonment.

Anxiety so severe unable to function at least few days/month.

No response to therapy, 2 SSRIs landed her in the ER with severe side effects including dizziness, nausea, and severe restlessness.

- Stellate ganglion block failed for anxiety.
- Integrative approach: acupuncture, B complex, fish oil, CBD helped minimally
- Patient decided to participate in a small group psilocybin retreat with her husband.

1 weekend 1:1 patient:trained therapist ratio. 1 day prep before experience including yoga, meditation, group/individual expectation settings, and half day processing after 6 hour psilocybin experience. Individual 1 hour therapy session 1 week after the retreat. Total individual/group therapy time about 12 hours.









## Reflection written for the retreat sponsor (1 week after the retreat)



"This weekend was intense! Full of gifts, teachings, and insights. I am still deeply processing, and I am sure more will unfold from here. I can already say that I feel incredible gratitude and healing. Words really do fail, but I'd approximate the experience to all three levels of my Reiki trainings/attunements all boosted at once?

I have some guesses about why a single experience can be so longlasting. I have strong memories that I'm sure I'll come back to again and again. I felt intensely loved, supported, grounded, guided -- a true sense that there are answers and signs and signals all around. Because the memories feel emotional and embodied, I believe they will last."



JAMA Psychiatry. 2021 May; 78(5): 1–9. Published online 2020 Nov 4. doi: 10.1001/jamapsychiatry.2020.3285 PMCID: PMC7643046 PMID: <u>33146667</u>

#### Effects of Psilocybin-Assisted Therapy on Major Depressive Disorder

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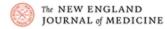
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The effect sizes were large at week 5 (Cohen d = 2.5; 95% CI, 1.4-3.5; P < .001) and week 8 (Cohen d = 2.6; 95% CI, 1.5-3.7; P < .001). The QIDS-SR documented a rapid decrease in mean (SD) depression score from baseline to day 1 after session 1 (16.7 [3.5] vs 6.3 [4.4]; Cohen d = 2.6; 95% CI, 1.8-3.5; P < .001), which remained statistically significantly reduced through the week 4 follow-up (6.0 [5.7]; Cohen d = 2.3; 95% CI, 1.5-3.0; P < .001). In the overall sample, 17 participants (71%) at week 1 and 17 (71%) at week 4 had a clinically significant response to the intervention ( $\geq$ 50% reduction in GRID-HAMD score), and 14 participants (58%) at week 1 and 13 participants (54%) at week 4 were in remission ( $\leq$ 7 GRID-HAMD score).



#### **IGINAL ARTICLE**

≔

subrutinib or Ibrutinib in lapsed or Refractory Chronic nphocytic Leukemia





EDITORIAL Thiazide-like versus Thiazide Diuretics - Finally, an Answer? ORIGINAL ARTICLE Once-Weekly Semaglutide in Adolescents with Obesity



SUBSCRIBE

**OR RENEW** 

Trial of Bere (B-VEC) for Epidermoly

#### **ORIGINAL ARTICLE**

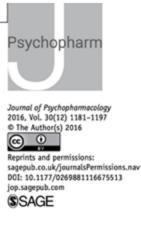
#### Trial of Psilocybin versus Escitalopram for Depression

Robin Carhart-Harris, Ph.D., Bruna Giribaldi, B.Sc., Rosalind Watts, D.Clin.Psy., Michelle Baker-Jones, B.A., Ashleigh Murphy-Beiner, M.Sc., Roberta Murphy, M.D., Jonny Martell, M.D., Allan Blemings, M.Sc., David Erritzoe, M.D., and David J. Nutt, M.D.

2 doses of 25mg 3 weeks apart plus daily placebo) and 29 to the escitalopram group. The mean scores on the QIDS-SR-16 at baseline were 14.5 in the psilocybin group and 16.4 in the escitalopram group. The mean (±SE) changes in the scores from baseline to week 6 were -8.0±1.0 points in the psilocybin group and  $-6.0\pm1.0$  in the escitalopram group, for a between-group difference of 2.0 points (95% confidence interval [CI], -5.0 to 0.9) (P=0.17). A QIDS-SR-16 response occurred in 70% of the patients in the psilocybin group and in 48% of those in the escitalopram group, for a between-group difference of 22 percentage points (95% CI, -3 to 48); QIDS-SR-16 remission occurred in 57% and 28%, respectively, for a between-group difference of 28 percentage points (95% CI, 2 to 54). Other secondary outcomes generally favored psilocybin over escitalopram, but the analyses were not corrected for multiple comparisons. The incidence of adverse events was similar in the trial groups.

Psilocybin produces substantial and sustained decreases in depression and anxiety in patients with life-threatening cancer: A randomized double-blind trial

Roland R Griffiths<sup>1,2</sup>, Matthew W Johnson<sup>1</sup>, Michael A Carducci<sup>3</sup>, Annie Umbricht<sup>1</sup>, William A Richards<sup>1</sup>, Brian D Richards<sup>1</sup>, Mary P Cosimano<sup>1</sup> and Margaret A Klinedinst<sup>1</sup>



"Participants, staff, and community observers rated participant moods, attitudes, and behaviors throughout the study. High-dose psilocybin produced large decreases in clinician- and self-rated measures of depressed mood and anxiety, along with increases in quality of life, life meaning, and optimism, and decreases in death anxiety. At 6-month follow-up, these changes were sustained, with about 80% of participants continuing to show clinically significant decreases in depressed mood and anxiety. Participants attributed improvements in attitudes about life/self, mood, relationships, and spirituality to the high-dose experience, with >80% endorsing moderately or greater increased well-being/life satisfaction."

### What about psilocybin microdosing?



#### Sci Rep. 2021; 11: 22479.

Puk

#### PMCID: PMC8602275

The present study describes microdosing practices, motivations and mental health among a sample of self-selected microdosers (n = 4050) and non-microdosers (n = 4653) via a mobile application. Psilocybin was the most commonly used microdose substances in our sample (85%) and we identified diverse microdose practices with regard to dosage, frequency, and the practice of *stacking* which involves combining psilocybin with non-psychedelic substances such as Lion's Mane mushrooms, chocolate, and niacin. Microdosers were generally similar to nonmicrodosing controls with regard to demographics, but were more likely to report a history of mental health concerns. Among individuals reporting mental health concerns, microdosers exhibited lower levels of depression, anxiety, and stress across gender. Health and wellness-related motives were the most prominent motives across microdosers in general, and were more prominent among females and among individuals who reported mental health concerns. Our results indicate health and wellness motives and perceived mental health benefits among microdosers, and highlight the need for further research into the mental health consequences of microdosing including studies with rigorous longitudinal designs.

## Back to our case



Pt started taking 250 mg of dried mushroom once every 3 days in combination with Niacin and Lion's Mane mushroom in addition to her ongoing integrative care.

Weekly virtual 30 min therapy sessions

8 weeks follow up - anxiety mostly resolved as insomnia. No lost working days, feels more at peace and much more hopeful going forward.

Fibromyalgia with some improvement; Pain around periods still severe, Headache was slightly better.

Concerns with microdosing or frequent use of Psilocybin, LSD, and others?



JOURNA

#### 👌 Open access 🗏 🐵 🛞 🖉 Review article 👘 First published online January 12, 2024

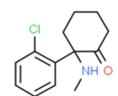
## Microdosing psychedelics and the risk of cardiac fibrosis and valvulopathy: Comparison to known cardiotoxins



#### Abstract

Though microdosing psychedelics has become increasingly popular, its long-term effects on cardiac health remain unknown. Microdosing most commonly involves ingesting sub-threshold doses of lysergic acid diethylamide (LSD), psilocybin, or other psychedelic drugs 2–4 times a week for at least several weeks, but potentially months or years. Concerningly, both LSD and psilocybin share structural similarities with medications which raise the risk of cardiac fibrosis and valvulopathy when taken regularly, including methysergide, pergolide, and fenfluramine. 3,4-Methylenedioxymethamphetamine, which is also reportedly used for microdosing, is likewise associated with heart valve damage when taken chronically. In this review, we evaluate the evidence that microdosing LSD, psilocybin, and other psychedelics for several months or more could raise the risk of cardiac fibrosis. We discuss the relationship between drug-induced cardiac fibrosis and the 5-HT2B receptor, and we make recommendations for evaluating the safety of microdosing psychedelics in future studies.

## Ketamine



MOA: NMDA-receptor antagonist

Route: IV, IN, IM, SL, oral

Onset/Duration: variable depending on route

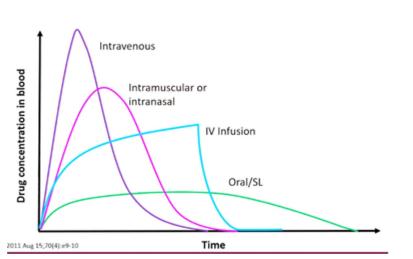
**Effects**: deep sense of relaxation, geometric patterns, dilation of time, dissociation, "birds' eye" view of one's life

#### Side effects:

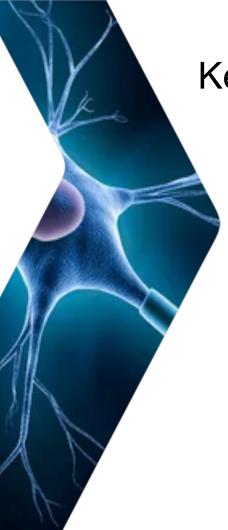
<u>Acute</u>: Mild and transient ↑ in BP/HR, Nausea, Vertigo <u>Chronic</u>: LUTS, liver toxicity

**Contraindications**: <u>Active</u> mania/psychosis, acute alcohol intoxication, uncontrolled HTN

**Medical clearance:** Hx aortic dissection, ICH, aneurysms, arrhythmia, cystitis, pregnancy





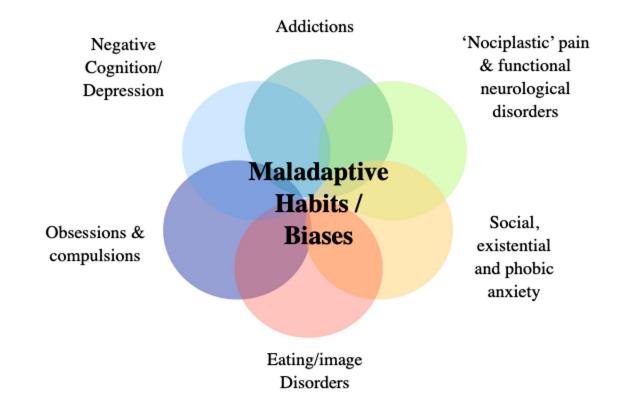


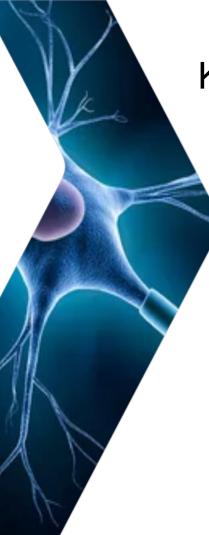
## Ketamine: NMDA-antagonism

### **SIMILARITIES**

- Disruption of Default Mode Network
- Consciousness Expansion
- Neuroplasticity  $\rightarrow$  Catalyst of Change
- Set, Setting, Skillset, Support all influence outcome

## **Common Factor in Mental Illness?**





## Ketamine: Legality, Access

- Schedule III Controlled Substance (legal)
- Doses for mental health lower than for anesthesia
- **ONLY** intranasal form (Spravato) FDA-approved for mental health:
  - 1. Treatment-Resistant Depression
  - 2. Acute SI
- Still frequently given from a purely biological/medication standpoint



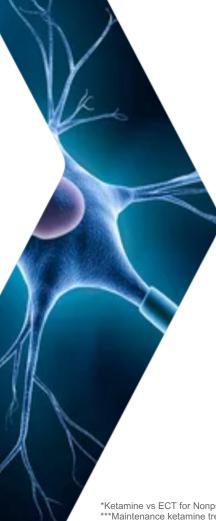
## Ketamine vs Spravato

# Ketamine for the treatment of major depression: a systematic review and meta-analysis

Stevan Nikolin,<sup>a,b,\*</sup> Anthony Rodgers,<sup>d</sup> Andreas Schwaab,<sup>c</sup> Anees Bahji,<sup>e,f</sup> Carlos Zarate, Jr.<sup>g</sup> Gustavo Vazquez,<sup>g</sup> and Colleen Loo<sup>a,b,d</sup>

- "Our findings suggested that <u>effect sizes for depression severity, as</u> well as response and remission rates, were numerically <u>greater for</u> racemic ketamine than esketamine."
- Differences were evident in initial effects, ongoing treatment, and lasting effects after the final dose.

The Lancet. Vol 62. August 2023.



## Ketamine for Depression

- Response rate ~55%\*
- Antidepressant effects are rapid but transitory
- In most patients who respond well to a single dose of ketamine, the benefits disappear within two weeks
- Study found maintenance ketamine treatment to be effective up to 5 years\*\*
- Tachyphylaxis, cognitive impairment, addiction, and serious renal and urinary problems seem uncommon.

\*Ketamine vs ECT for Nonpsychotic Treatment-Resistant Major Depression." Anand A, et al. NEJM (2023). \*\*\*Maintenance ketamine treatment for depression: a systematic review of efficacy, safety, and tolerability." Smith-Apeldoorn, S. Y., Veraart, J. KE., Spijker, J., Kamphuis, J., & Schoevers, R. A. *The Lancet Psychiatry*,(2022).



## Ketamine for SI

#### Ketamine for suicidality: An umbrella review

Ahmad Shamabadi<sup>1,2</sup> | Ali Ahmadzade<sup>1</sup> | Alireza Hasanzadeh<sup>1,2</sup>

- IV Ketamine can significantly reduce suicidal ideation (SI) within 40 minutes
  - $\rightarrow$  Several studies showed efficacy at 2, 4, and 24 hours post-treatment
- **Temporary** Effect may last up to 10 days
- Mixed results regarding esketamine efficacy.

## **Ketamine: Other Applications**

Ketamine for the treatment of mental health and substance use disorders: comprehensive systematic review

Zach Walsh\*, Ozden Merve Mollaahmetoglu\*, Joseph Rootman, Shannon Golsof, Johanna Keeler, Beth Marsh, David J. Nutt and Celia J. A. Morgan

- Less robust evidence but still positive and short-lived benefits in <u>bipolar disorder</u>, <u>social</u> and generalized anxiety, <u>obsessive</u>-compulsive disorders, <u>post-traumatic stress disorders</u>, <u>substance use disorders (alcohol, opiate, cocaine)</u>, and <u>eating disorders</u>
- Research from clinical use of KAP and qualitative studies in the field suggest that <u>sufficient</u> preparation before the experience, a clinical and professional setting, and trusting and <u>supportive relationships with staff is crucial</u>
- Given findings that <u>ketamine's therapeutic benefits can be extended with psychological</u> therapy, it is advisable to provide ketamine treatment alongside a psychological therapy

## Ketamine: Patient Case- Brief Vignette

34 Y F with history of MDD

Current Meds: Desvenlafaxine 50 mg/day

<u>Prior Meds</u>: citalopram, fluoxetine, sertraline, bupropion, lamotrigine, methylphenidate

Prior Admissions: 2016 - SI

Therapist: sees weekly

 $\frac{PHQ9}{21 \rightarrow 17 \rightarrow 10 \rightarrow 4 \rightarrow 3 \rightarrow 3}$ Met with therapist after each infusion

Maintenance: Q4 weeks Tried extending further but noticed worsening mood

# Part 3: Regulatory, Ethics, Safety

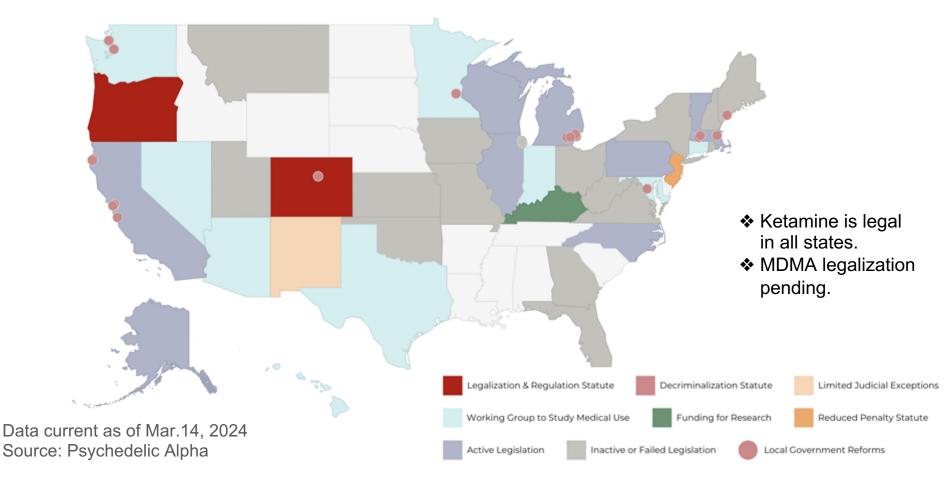




School of Medicine & Health Sciences



## **Psychedelic legislation in the United States**



### **Diversity, Equity & Inclusion in Psychedelic Integration Therapy**

"Psychedelic-assisted therapy and the current psychedelic renaissance is embedded within a White-dominant medical framework.

There is a lack of diversity within the field of psychedelic researchers, with Indigenous people and people of color underrepresented both as researchers, therapists, and participants in studies."



George JR, Michaels TI, Sevelius J, Williams MT. The psychedelic renaissance and the limitations of a White-dominant medical framework: a call for indigenous and ethnic minority inclusion. J Psychedelic Stud. 2020;4:4–15. Michaels TI, Purdon J, Collins A, Williams MT. Inclusion of people of color in psychedelic-assisted psychotherapy: a review of the literature. BMC Psychiatry. 2018;18:1–9.

## Cultural roots, Westernization

"Psychedelics have been part of the spiritual practices and cultures of Indigenous people throughout the world and have historically been frequently condemned by Western cultures.

As a result, the "discovery" of psychedelicassisted therapy by Western medicine has been criticized as another example of colonialism or cultural appropriation that repeats a history of oppression."





George JR, Michaels TI, Sevelius J, Williams MT. The psychedelic renaissance and the limitations of a White-dominant medical framework: a call for indigenous and ethnic minority inclusion. J Psychedelic Stud. 2020;4:4–15.

Michaels TI, Purdon J, Collins A, Williams MT. Inclusion of people of color in psychedelic-assisted psychotherapy: a review of the literature. BMC Psychiatry. 2018;18:1–9.

# Call to action: To strike a balance between seemingly opposing forces.

Engineer safe supply of psychedelic substances when legislation allows.

Keep the cost of integration therapy within reach.

Dialogue with traditional healers from the communities that started these practices while promoting ethical, safe use for patients.



# Viewer submitted and voted Q & A - Follow along on your device





LIVE Q&A! Please Scan QR Go to <u>https://sli.do</u> and enter #1260685

Presenter Q&A Web Browser Reference Link



### Thank you!

- Justin Laube, MD: UCLA
- Thais Salles Araujo, MD: UCLA
- Leslie Mendoza Temple, MD: Endeavor Health/ University of Chicago
- Mikhail Kogan, MD: George Washington University
- Erika Steinbrenner, MD: Imagine Healthcare, Epiphany Wellness, Symetria Recovery

## Extra slides



Banisteriopsis caapi Harmine tetrahydroharmine harmaline

## Supplementation Dietary Preparation

Magnesium L Threonate 144 mg BID NAC 1200 mg BID Alpha lipoic acid 200 mg Vitamin C 1000 mg BID 5HTP 100 mg with EGCG 500 mg Electrolytes **B6** CDP-Choline 500 mg Melatonin 3-10 mg

1 week before/after:
No red/heavy meats,
alcohol, spicy food
2 days before:
No fermented foods
1 day before/after:
Minimize sugar

# No food 5 hrs prior to ceremony

## How to engage but stay safe as a healthcare professional.

- "Avoid facilitating access to psychedelics or prohibited substances in any way and avoid providing a space wherein psychedelics would be used.
- 2. Refrain from coordinating work with underground guides. Referring a client to an underground therapist is a clear and obvious form of knowingly facilitating access to prohibited substances.
- 3. In contrast, receiving referrals <u>from</u> an 'underground' guide entails less legal risk. Members of the public are free to refer to whatever practitioner they wish."



Pilecki, B., Luoma, J.B., Bathje, G.J. *et al.* Ethical and legal issues in psychedelic harm reduction and integration therapy.*Harm Reduct J* 18, 40 (2021).

## Could our integrative health patient also be seeking out and be supported by psychedelic therapies?

 Table 2. Patient Percentages of Reasons for Seeking Care at IM Clinic and Goals (n = 4182)
 BraveNet PBRN 2008-2011

|   | Extremely/  |                         |                       |
|---|-------------|-------------------------|-----------------------|
|   | Quite a Lot | Conditions<br>Addressed | Percent<br>(n = 4182) |
| Reasons for seeking care at IM clinic   |             |                         | · · · · ·             |
| I want to improve my health and wellness now to prevent future problems           | 83.85       | Pain (chronic)          | 33.1                  |
| I want to try new options for my health care                                      | 76.65       | Fatigue                 | 10.2                  |
|   |             | Hyperlipidemia          | 10.0                  |
| To maximize my health regardless of whether or not my illness is curable          | 74.62       | Pain (acute)            | 9.7                   |
| To be in a place that acknowledges the connection between mind, body, spirit, and | 70.33       | Stress                  | 9.3                   |
| community   |             | Wellness visit          | 8.5                   |
| To receive objective, medical advice on nonconventional approaches                | 67.24       | Cancer                  | 8.3                   |
| To receive care in a safe, healing environment                                    | 66.71       | Weight                  | 8.0                   |
|   |             | Anxiety                 | 7.7                   |
| A place where I can receive care from a multidisciplinary team                    | 58.48       | Depression              | 7.2                   |

Ruth Q. Wolever, PhD et. al EXPLORE November/December 2012, Vol. 8, No.