

**UW PACC** Psychiatry and Addictions Case Conference UW Medicine | Psychiatry and Behavioral Sciences

## CANNABIS AND PTSD RECOVERY: PROBLEM OR SOLUTION?

#### MICHELE BEDARD-GILLIGAN, PHD UNIVERSITY OF WASHINGTON







### **GENERAL DISCLOSURES**

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#### **GENERAL DISCLOSURES**

## UW PACC is also supported by Coordinated Care of Washington



#### **SPEAKER DISCLOSURES**

I, Michele Bedard-Gilligan, have the following commercial relationship to disclose:

My husband is Vice President of Supply Chain for 4Front Ventures, a cannabis company. He leads their non cannabis business.



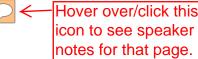
#### **SPEAKER DISCLOSURES**

Michele Bedard-Gilligan, PhD: One financial relationship

#### **PLANNER DISCLOSURES**

The following series planners have no relevant conflicts of<br/>interest to disclose:Mark Duncan MDCameron CaseyBarb McCann PhDBetsy PaynAnna Ratzliff MD PhDDiana RollRick Ries MDCara Towle MSN RNKari Stephens PhDNiambi Kanye





## CASE EXAMPLE: "STELLA"

- Demographics
  - Cis-gender, heterosexual, married, Caucasian/not hispanic 39 year old woman
  - Employed full-time as a tattoo artist
  - Plays in a band
  - Target trauma: domestic violence (physical & sexual abuse) 20+ years prior
- Clinical Presentation
  - Moderate/severe PTSD symptoms (PSS-I = 40)
  - Diagnosis of past MDD, no current comorbidity
  - Daily cannabis use, reported 0.5 grams/per day over last 30 days
  - Cannabis use: "for PMS, sleeplessness"

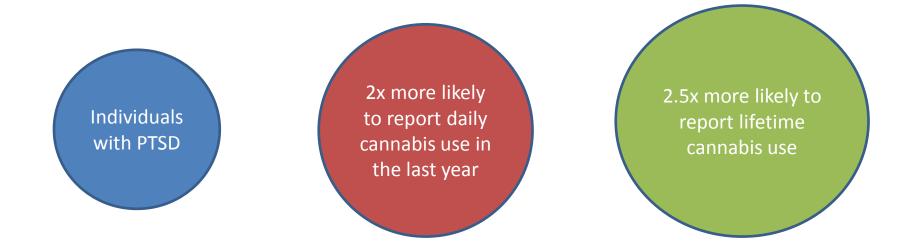


#### **OBJECTIVES**

- 1. Understand the observed relationships between PTSD and cannabis use
- 2. Explain conceptual models of how cannabis might impact PTSD
- 3. Discuss implications of cannabis use for PTSD recovery and treatment

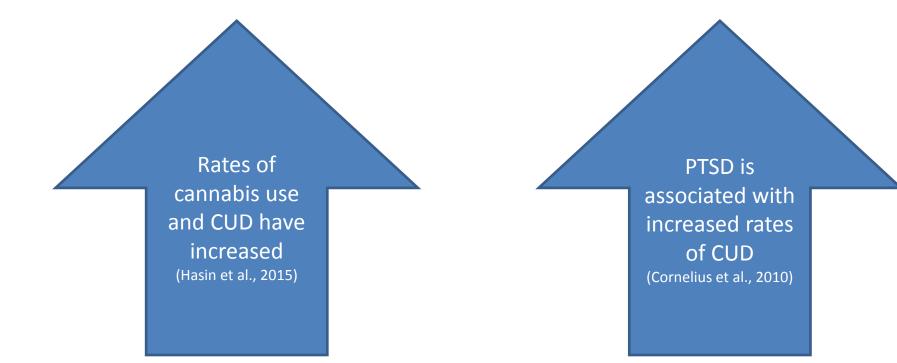


## PTSD IS A PRIMARY REASON THAT INDIVIDUALS SEEK OUT CANNABIS IN STATES THAT HAVE LEGALIZED MEDICAL CANNABIS (BOWLES, 2012)





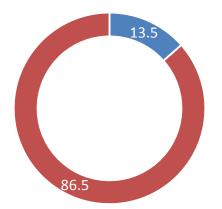
#### CANNABIS USE AND CANNABIS USE DISORDER (CUD) ARE INCREASING IN THE US POPULATION





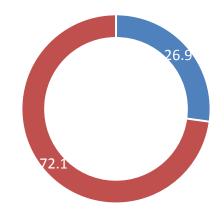
#### RATES OF CANNABIS USE ARE INCREASING IN THOSE SEEKING TREATMENT FOR PTSD

#### 2004 - 2010



Cannabis UseNo Cannabis Use

#### 2011 - 2016

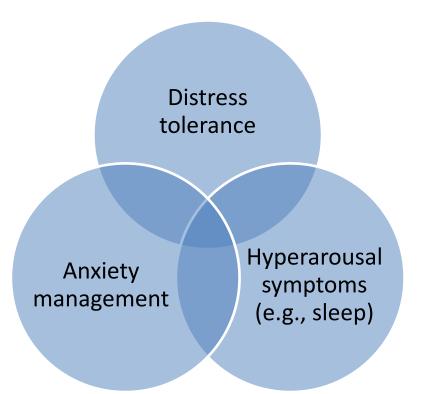


## Cannabis UseNo Cannabis Use



R01MH066347, R01MH066348, R01MH066347

#### WHY DO PATIENTS WITH PTSD USE CANNABIS? SELF-MEDICATION THEORY

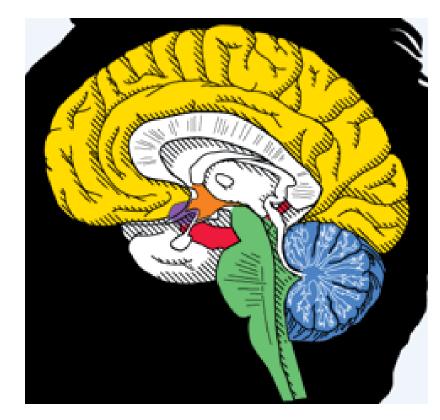




### KEY BRAIN AREAS IMPLICATED IN CANNABIS USE

Targeting the endocannabinoid system as treatment for PTSD (Berardi et al., 2016; Korem et al., 2015; Trezza & Campoglona, 2013) Cannabinoid

receptors  $(CB_1)$  are implicated in processing of fear, stress, emotion, and reward in the brain



#### Cerebral cortex

thinking, perceptual awareness and	men			
	thin	ong		
awareness and	perc	epti	ıal	
	awa	rene	55.8	nd

Altered consciousness; perceptual distortions; memory impairment; occasional delusions and hallucinations

Increased

appetite

#### Hypothalamu

Governs metabolic processes such as appetite

#### Brain stem

Controls many Nausea relief; basic functions rapid heart rate; including reduced blood arousal, the pressure; vomiting reflex, drowsiness blood pressure and heart rate

Also plays a Pain reduction; role in pain reduced sensation, spasticity; muscle tone and reduced tremor movement

#### lippocamp

Is key to Impairment in memory storage memory and recall

#### Cerebellum

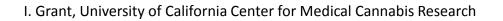
Governs coordination and muscle control Reduced spasticity; impaired coordination

#### Amygdala

Plays a role in emotions reduced anxiety and blocking of traumatic memories in other cases; reduced hostility



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### CANNABIS AS A POTENTIAL TREATMENT FOR PTSD?

- Potential therapeutic benefits
  - Manage the emotional symptoms (e.g., capitalize on anxiolytic effects of cannabis)
  - Increase the process of new learning associated with recovery (i.e., extinction enhancer)





### PTSD IS THE PERSISTENCE OF MALADAPTIVE REACTIONS, NOT THE MERE PRESENCE OF REACTIONS

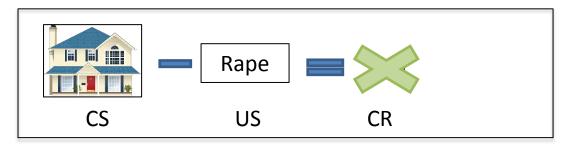
Failure in natural fear extinction thought to be key mechanism in the persistence of PTSD

Mahan & Ressler, 2012



# HOW MIGHT CANNABIS RELATE TO FEAR EXTINCTION?

#### Fear extinction



- Facilitation effect:
  - CB<sub>1</sub> receptors in the ventromedial prefrontal cortex and hippocampus
  - These areas are heavily implicated in extinction learning (e.g., Bouton et al., 2006; Milad et al., 2008)

#### Detrimental effect:

- Documented amotivational and anxiolytic effects
- Biphasic effects anxiolytic vs anxiogenic (D'Souza et al., 2004; Viveros et al., 2005)
- Acute impairments on learning, memory, attention, and working memory (Volkow et al., 2016)



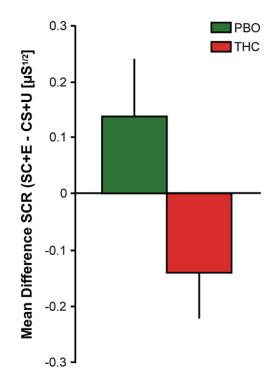
## PSYCHOTHERAPIES BASED ON EXTINCTION LEARNING HAVE LARGE EFFECTS IN DECREASING PTSD SYMPTOMS

		Effect Size
Intervention	N	<i>(g)</i>
All Psychotherapies	76	1.14
Cognitive Therapies	10	1.63
Exposure Therapies	27	1.08
Mixed Cognitive and Exposure	14	1.38
EMDR	11	1.01
All Medications	56	0.42
SSRIs	20	0.48

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## HUMAN STUDIES OF THC AND EXTINCTION

- Administration of THC facilitated short-term extinction as evidenced by decreased skin conductance response when tested 24 hours later (Rabinak et al., 2013)
   THC prevented the recovery of fear
- Facilitation may be limited to acute effects (Klumpers et al., 2012)
- Studies were done in healthy, cannabis naïve volunteers

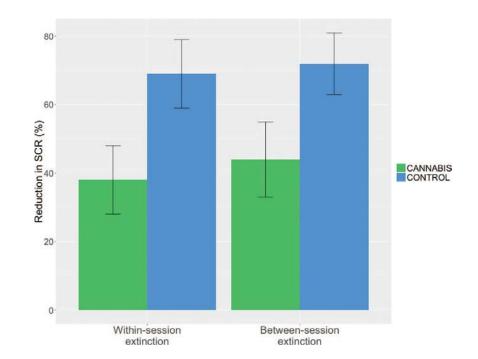




### CHRONIC CANNABIS USERS SHOW DEFICITS IN EXTINCTION LEARNING

- Reduced within-session

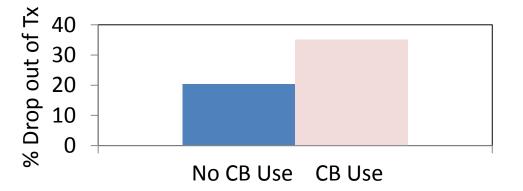
   (d = 0.78) and between
   session (d = 0.76)
   extinction, as measured
   by skin conductance
   response
- Suggests chronic cannabis use impairs, not facilitates, extinction





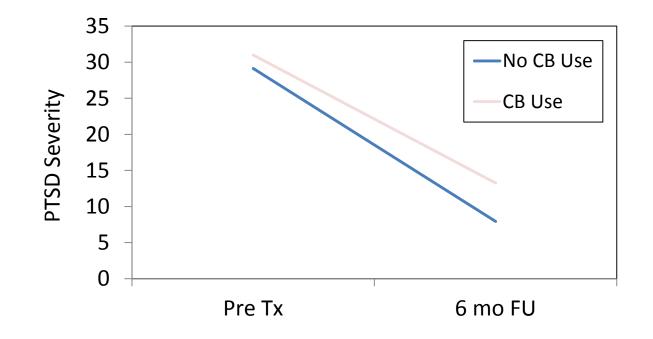
## CANNABIS IS ASSOCIATED WITH HIGHER DROPOUT IN A CLINICAL TRIAL OF AN EXTINCTION-BASED PTSD THERAPY

- In clinical practice, cannabis use may impair PTSD treatment response to an extinction based treatment
- In our clinical trial (N = 200; R01MH066347/R01MH066348) cannabis use predicted
  - Higher likelihood of dropout



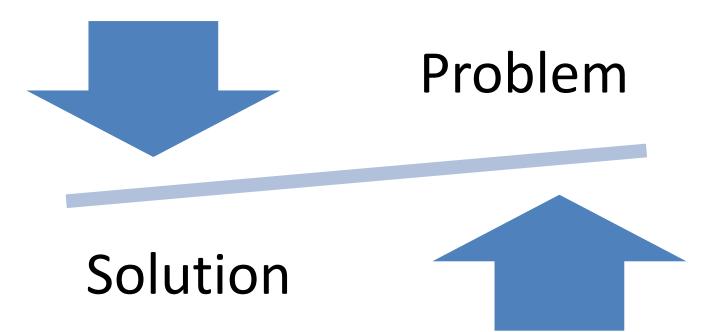


#### CANNABIS USE ALSO ASSOCIATED WITH LESS PTSD RECOVERY IN THAT SAME TRIAL



Bedard-Gilligan et al., 2018

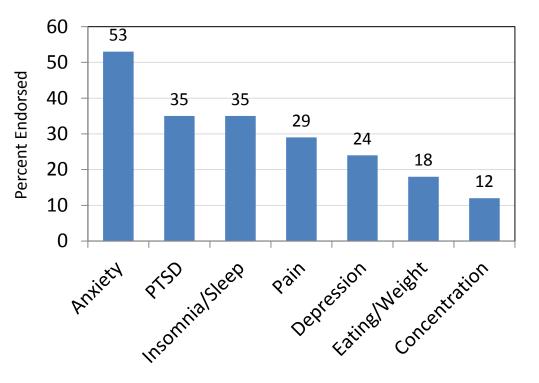
#### EFFECTS OF CANNABIS ON RECOVERY ARE UNCLEAR





# THE FUNCTION OF CANNABIS: PATIENT VIEWS

- Patients perceive clinical benefits to their cannabis use
  - "Marijuana makes it so I do not have to take any heavy narcotic pain medications"
  - "Marijuana replaces an obscene amount of things that I have been prescribed and tried in the past with less successful results"
  - "I use it to calm my anxiety and focus more on self-care"



Diverse reasons for cannabis use (N = 17)

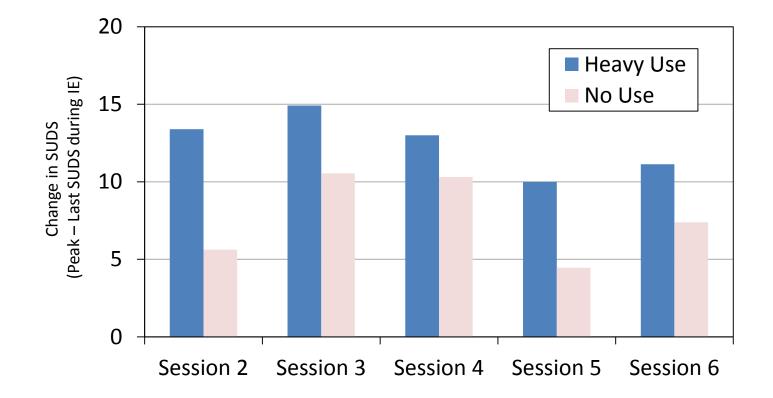


### ONGOING TRIAL OF EXPOSURE THERAPY FOR PTSD AND CANNABIS USE (R34 DA040034)

- Key questions/issues:
  - Does cannabis use get in the way of treatment?
    - How does it affect engagement and fear extinction?
    - How does it affect cognitive ability to process?
  - Does cannabis use change with treatment?

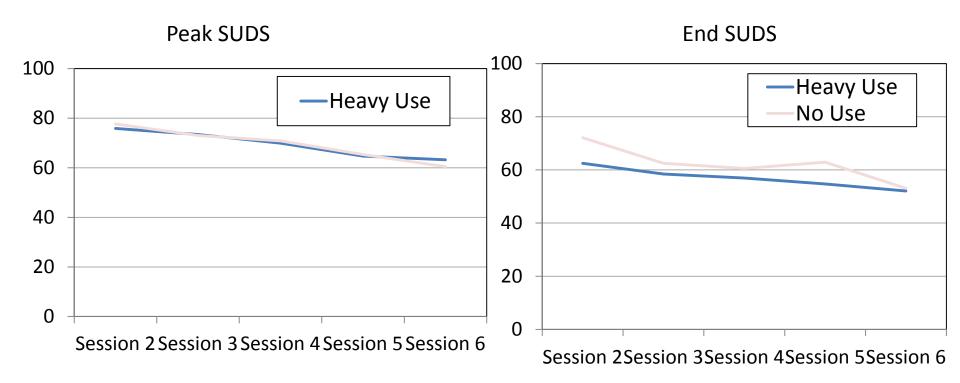


#### WITHIN SESSION EXTINCTION PATTERNS BY CANNABIS USE GROUP (*N* = 31)



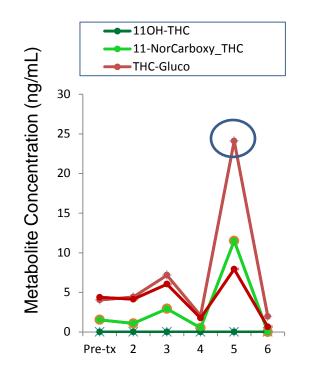


## BETWEEN SESSION EXTINCTION PATTERNS BY CANNABIS USE GROUP (N = 31)





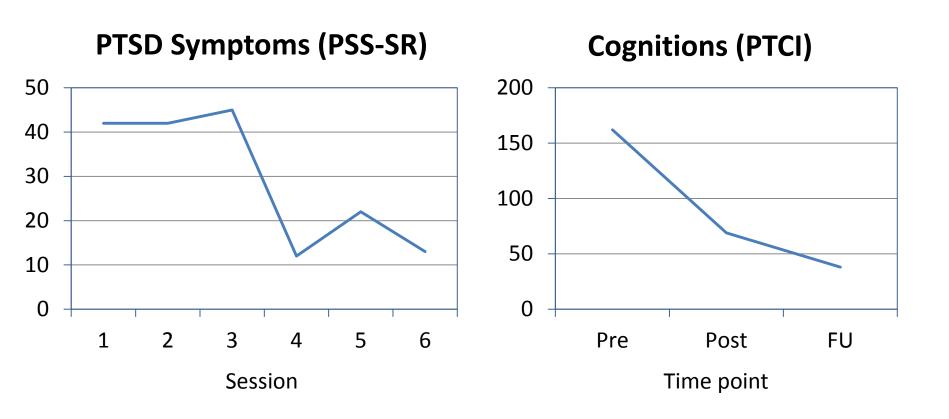
## CASE EXAMPLE "STELLA": HIGHER COGNITIVE FUNCTIONING DURING PROCESSING?



Session 5					
PTSD (PSS-SR)	22				
Depression (QIDs)	14				
Time since last cannabis use	1 hour				
Perceived high	2/10				
Strain	Strawberry Banana Sherbert				
Pre-SUDs	30				
Post-SUDs	15				



#### CHANGES IN PTSD SYMPTOMS AND TRAUMA RELATED COGNITIONS FOR "STELLA"





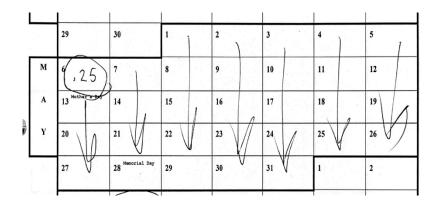
#### "STELLA'S" PRE TO FOLLOW-UP CANNABIS USE

#### Pre-Tx TLFB Cannabis Use

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E	",5	12.5	<sup>13</sup> .5	14 valentine's bay	15,5	16°°, 5	<sup>17</sup> , 5
В	18 . 5	19 President's bay	20 ,5	21.5	22 .S	23.5	24 .5
	<sup>25</sup> ,5	26,5	27.5 (	28.5	1	2	3

#### Follow-up TLFB Cannabis Use





#### **PERSONALIZING TREATMENTS**

- One size fits all approach to treatment means we are missing important subgroups
- Meeting patients "where they are at"
- Many using cannabis may not seek or comply with treatment that requires reductions/abstinence in use
- Doesn't mean we need to abandon treatments we know work
- Simple treatment adjustments may improve adherence and create hope
  - Fewer sessions in compressed time frame with no HW

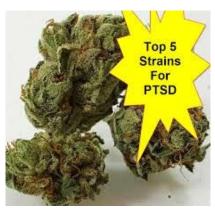


## CANNABIS AND PTSD: PROBLEM OR SOLUTION?

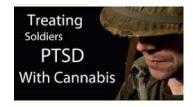




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