



ACUTE EFFECTS OF Δ -9-
TETRAHYDROCANNABINOL & CANNABIDIOL ON
ATTENTIONAL BIAS IN HEALTHY VOLUNTEERS: A
RANDOMISED, DOUBLE-BLIND, CROSSOVER STUDY

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ATTENTIONAL BIAS

- Unconscious bias towards cannabis stimuli in regular cannabis users (O'Neill et al., 2020) and people with cannabis dependence (Vujanovic et al., 2016)
- Continued cannabis use leads to greater valuing of relevant stimuli
- Stronger valuation with greater frequency of use/dependence
- Therefore, attentional bias can be used as a measure of a person's association with cannabis
- Proxy for addiction-like behaviour

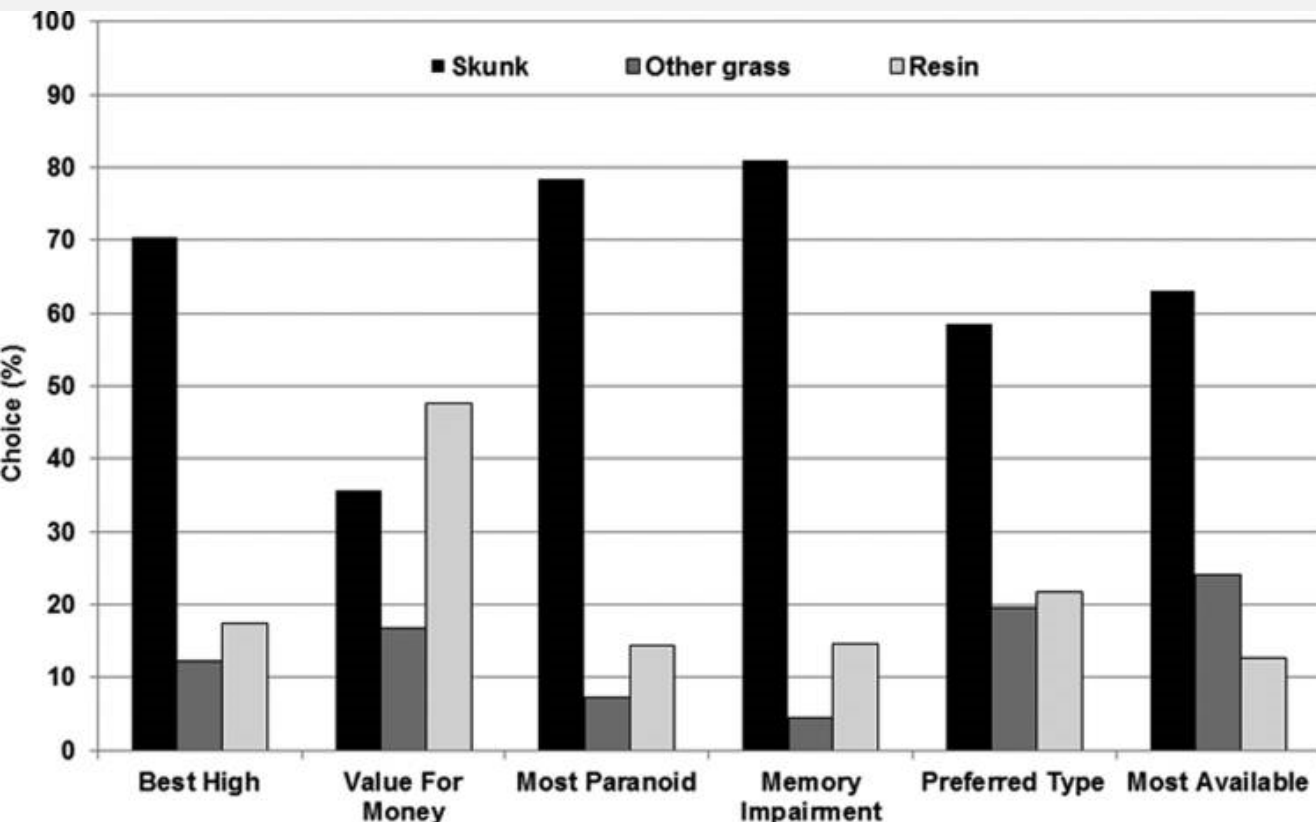


CANNABIS DEPENDENCE

- 9% who try cannabis become dependent (Lopez-Quintero et al., 2011)
- 17% if cannabis use is started in adolescence (Anthony 2006)
- 25-50% among daily users (Hall 2009)

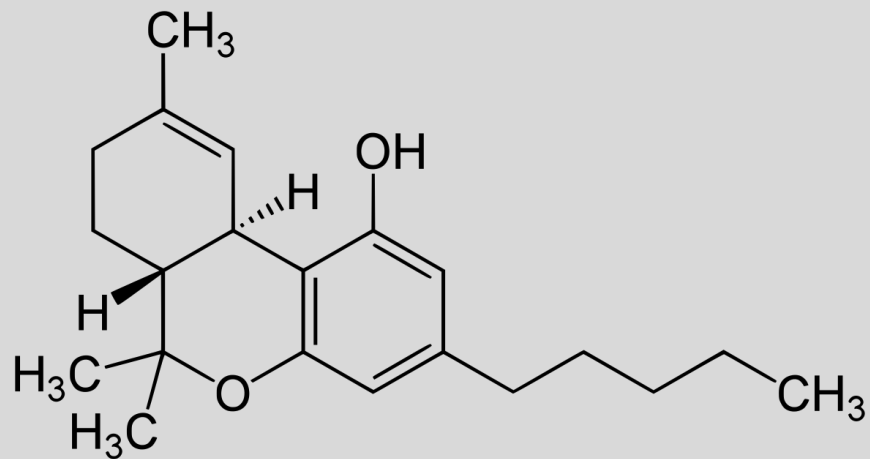


CANNABIS TYPE AND DEPENDENCE

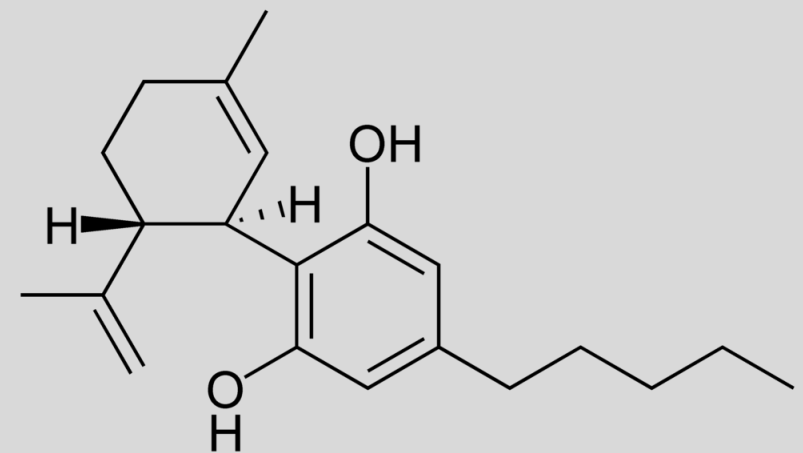


- Frequent use of high potency cannabis predicts dependence severity
- But high potency rated as preferred and most available
- Use of hash and herbal not associated with dependence

CANNABIS SATIVA L.



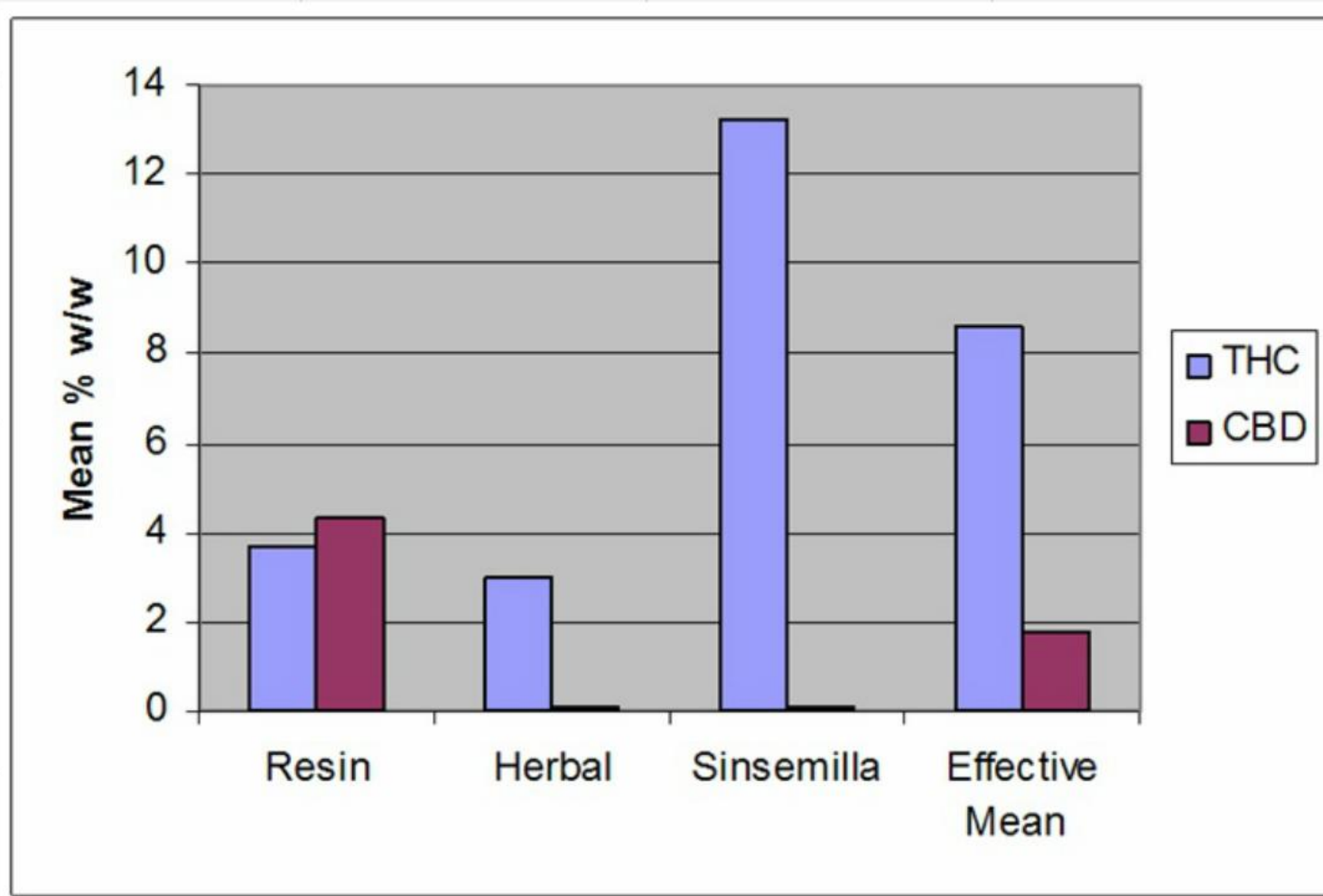
Delta-9-tetrahydrocannabinol (THC)



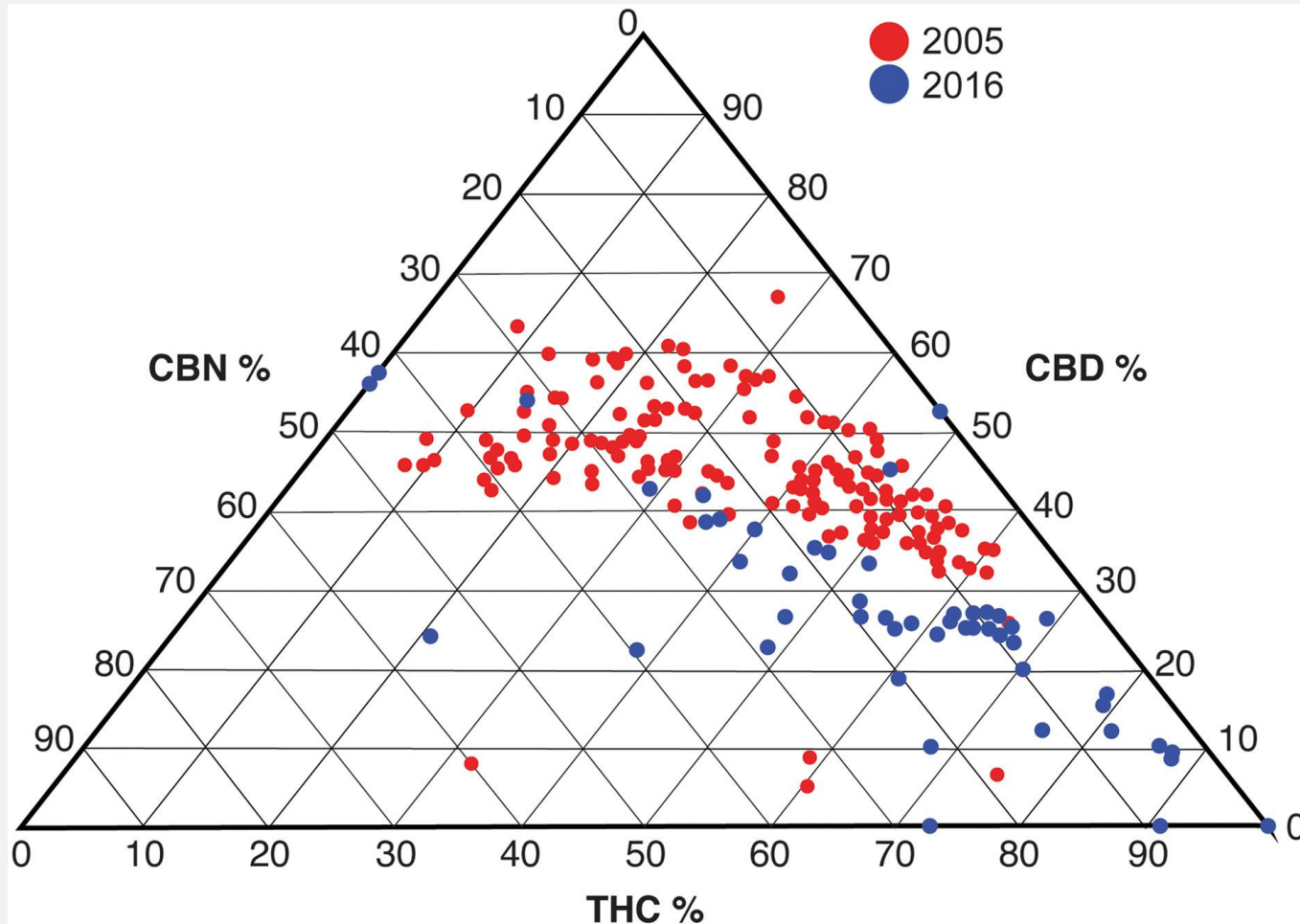
Cannabidiol (CBD)

CANNABIS TYPE AND POTENCY

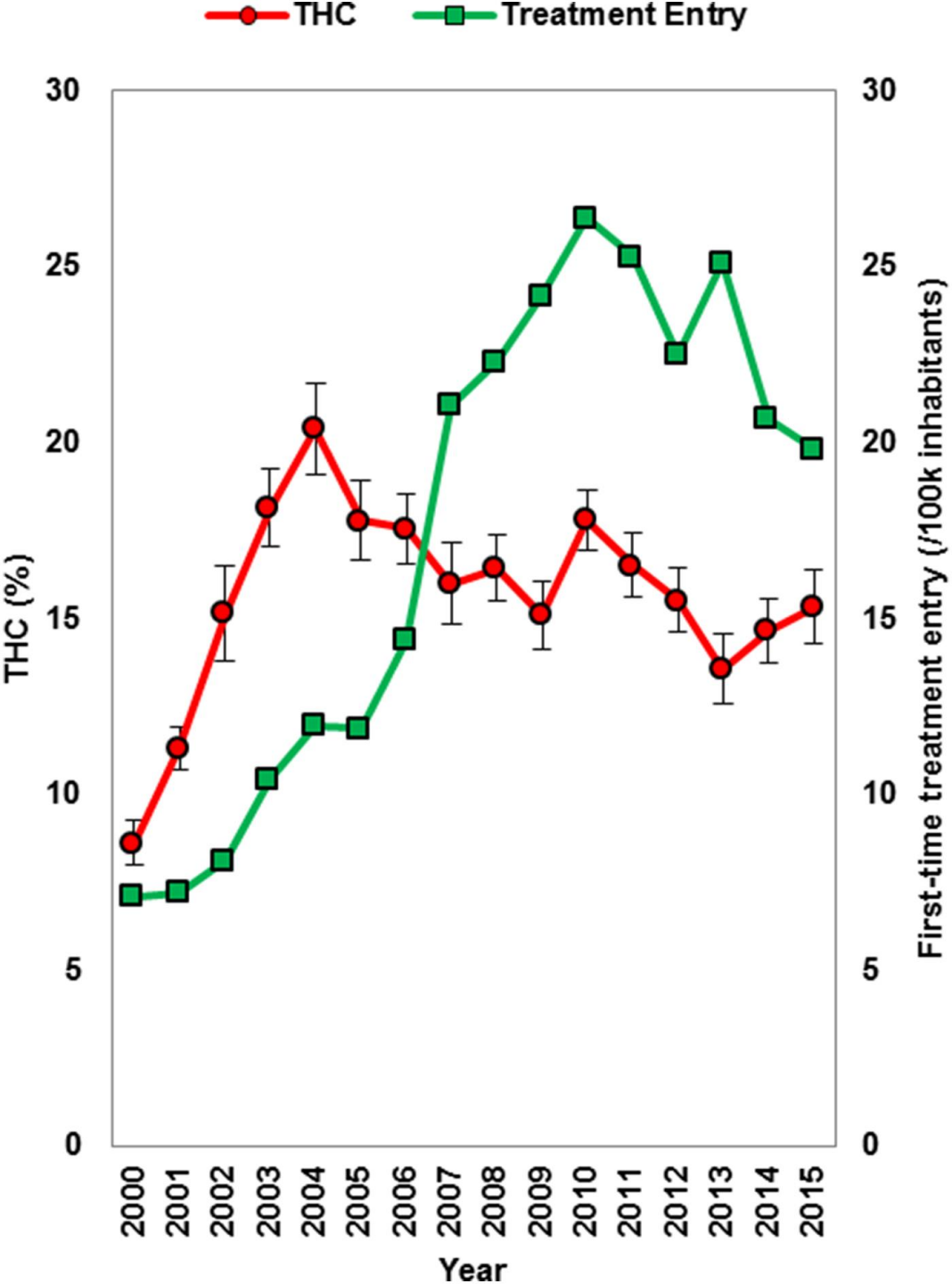
POTTER ET AL 2008



POTENCY CHANGES POTTER ET AL 2018



- Highest **THC** content of resin in 2005 was **10.8%**
- 25% of samples in 2016 exceeded that
- Highest **THC** content of resin in 2016 was **29%**

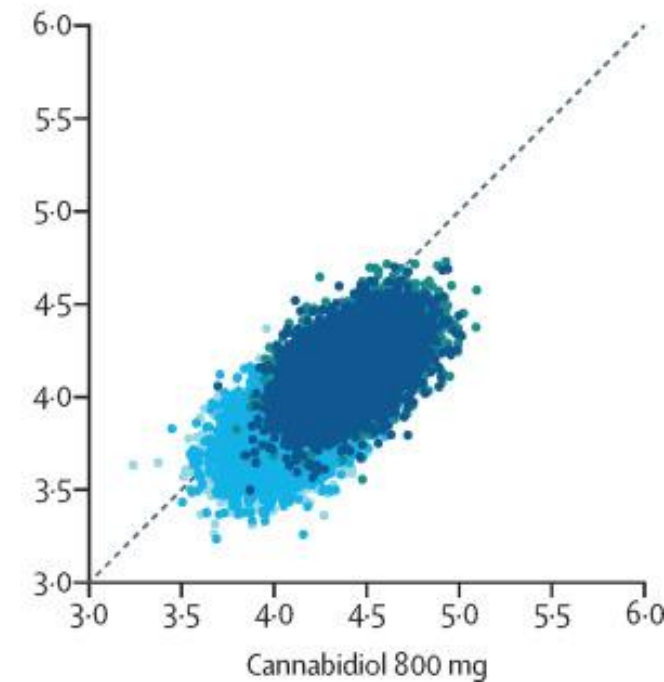
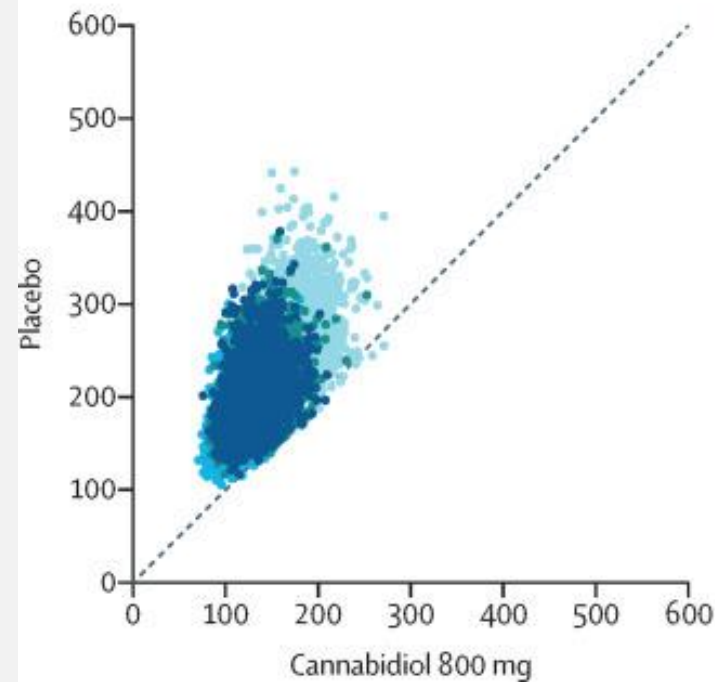
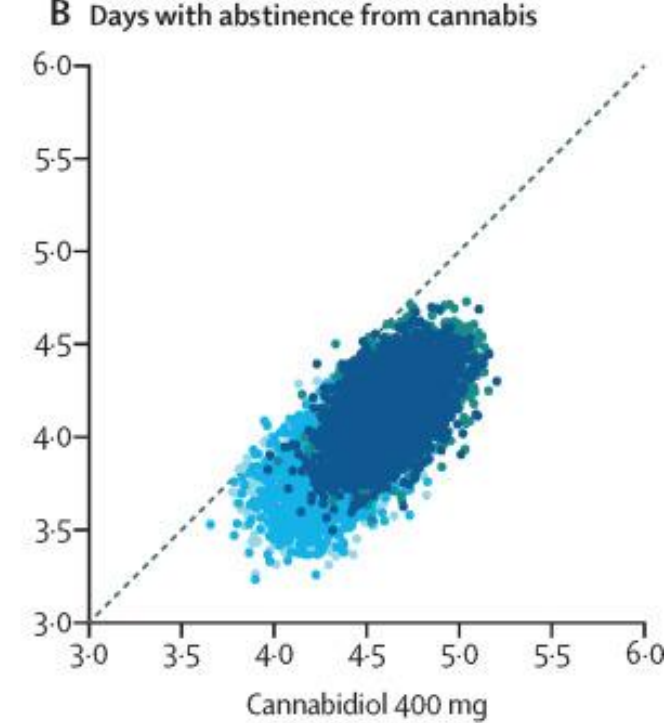
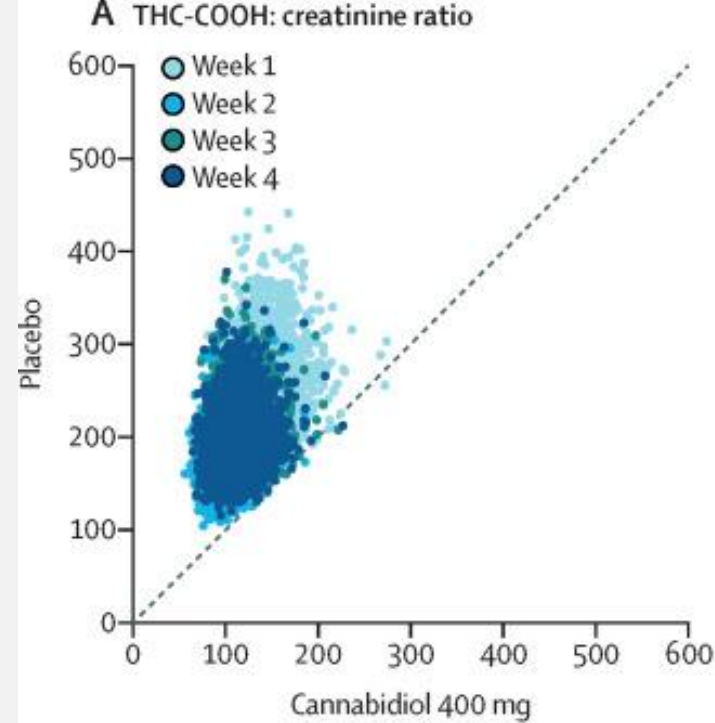


MORE THC, MORE PROBLEMS

- Data from the Netherlands
- More systematic testing compared to UK data
- When cannabis potency (i.e. **THC** content) increases, increase in first time admissions for drug-related issues five years later

CBD TO TREAT DEPENDENCE

- **CBD** trialled in individuals with cannabis use disorder who wanted to quit
- Both 400mg and 800mg **CBD** daily reduced **THC** metabolites in urine and increased number of days not using cannabis over 4 weeks





THC, CBD
AND
ATTENTIONAL
BIAS

- People who use higher **CBD:THC** ratios tend to show reduced attentional bias to cannabis stimuli when sober and when intoxicated with their usual cannabis (Morgan et al., 2010)
- What about in more infrequent users?
- Do we see signs of addiction-like behaviour following acute dosing?



eCBD



- Does **THC** increase attentional bias towards cannabis stimuli?
- Does increasing **CBD:THC** ratio in inhaled cannabis attenuate this effect?

INCLUSION CRITERIA

Infrequent cannabis use (mean cannabis use < 1 /week over the last 12 months)

No past use of synthetic cannabinoids

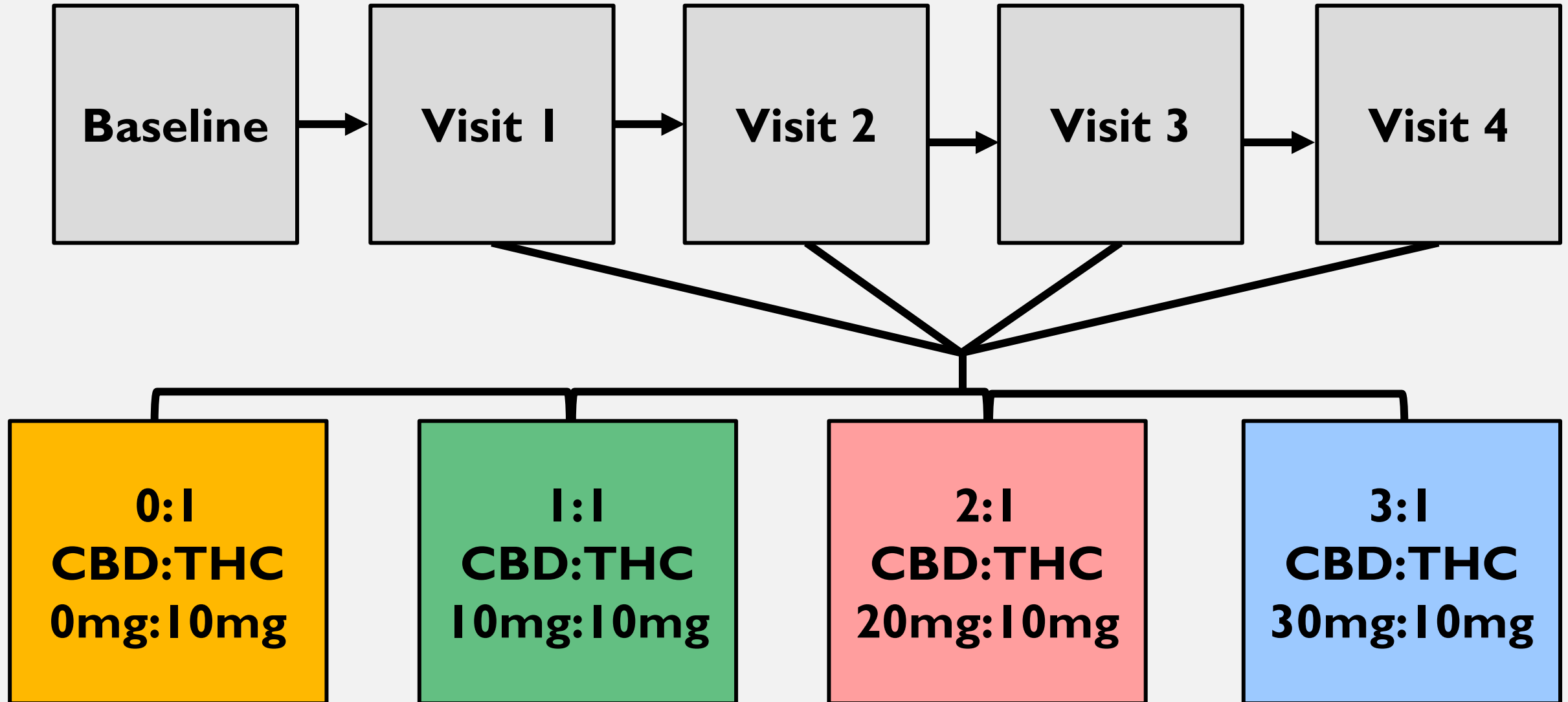
No past or present major mental, physical illness or substance use disorder

Score < 5 on the Fagerstrom Nicotine Dependence Questionnaire

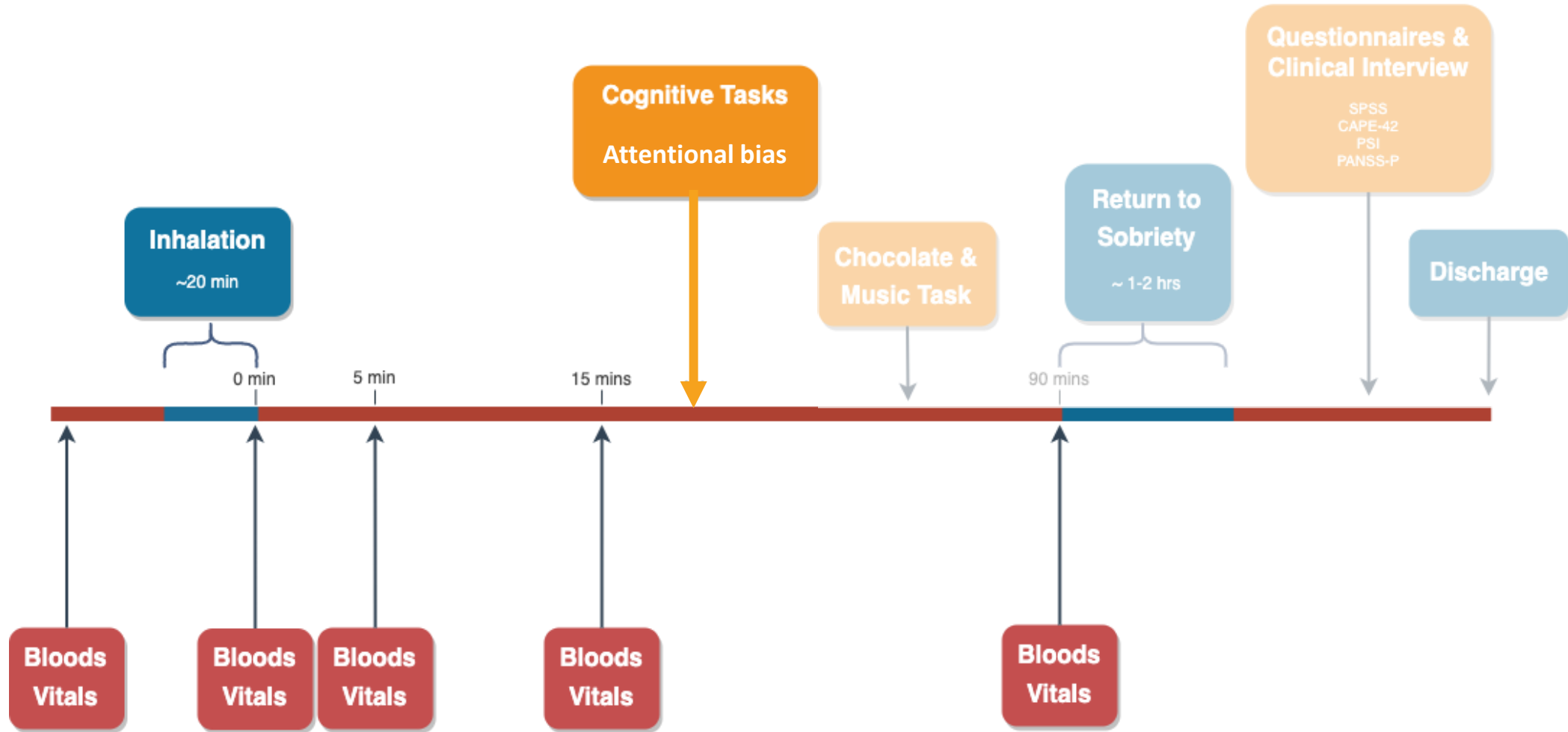
Negative urine drug screen

No past 24-hour use of alcohol or tobacco

DOUBLE BLIND, RANDOMISED CROSSOVER STUDY



n=46



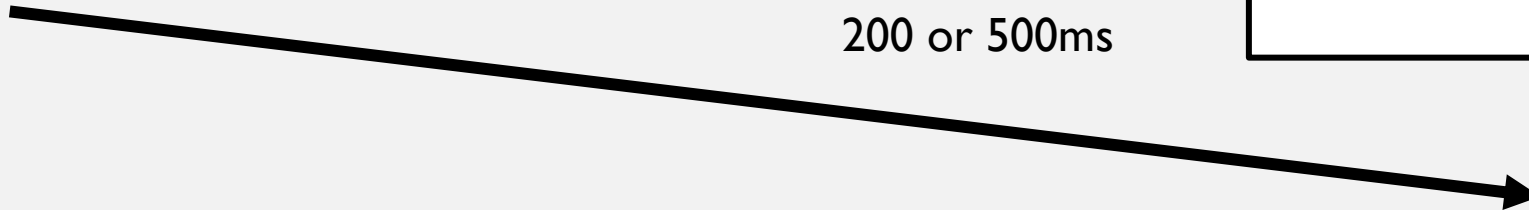
TASK I: IMPLICIT WANTING

+

Fixation
1000ms



Stimulus
200 or 500ms



TASK 2: EXPLICIT LIKING



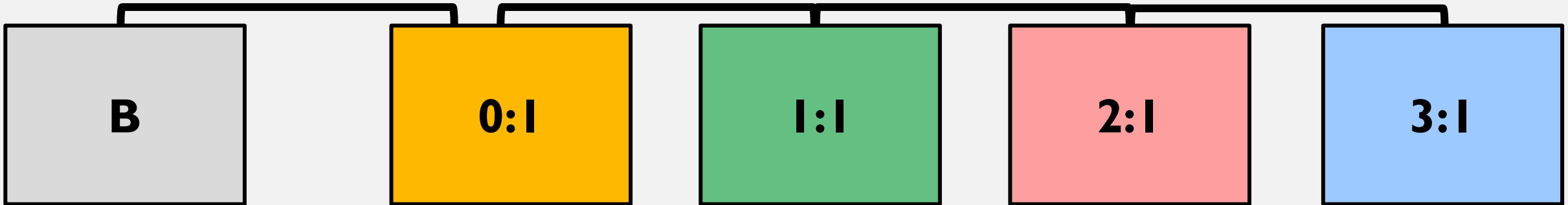
How pleasurable is this image?

-3 -2 -1 0 1 2 3

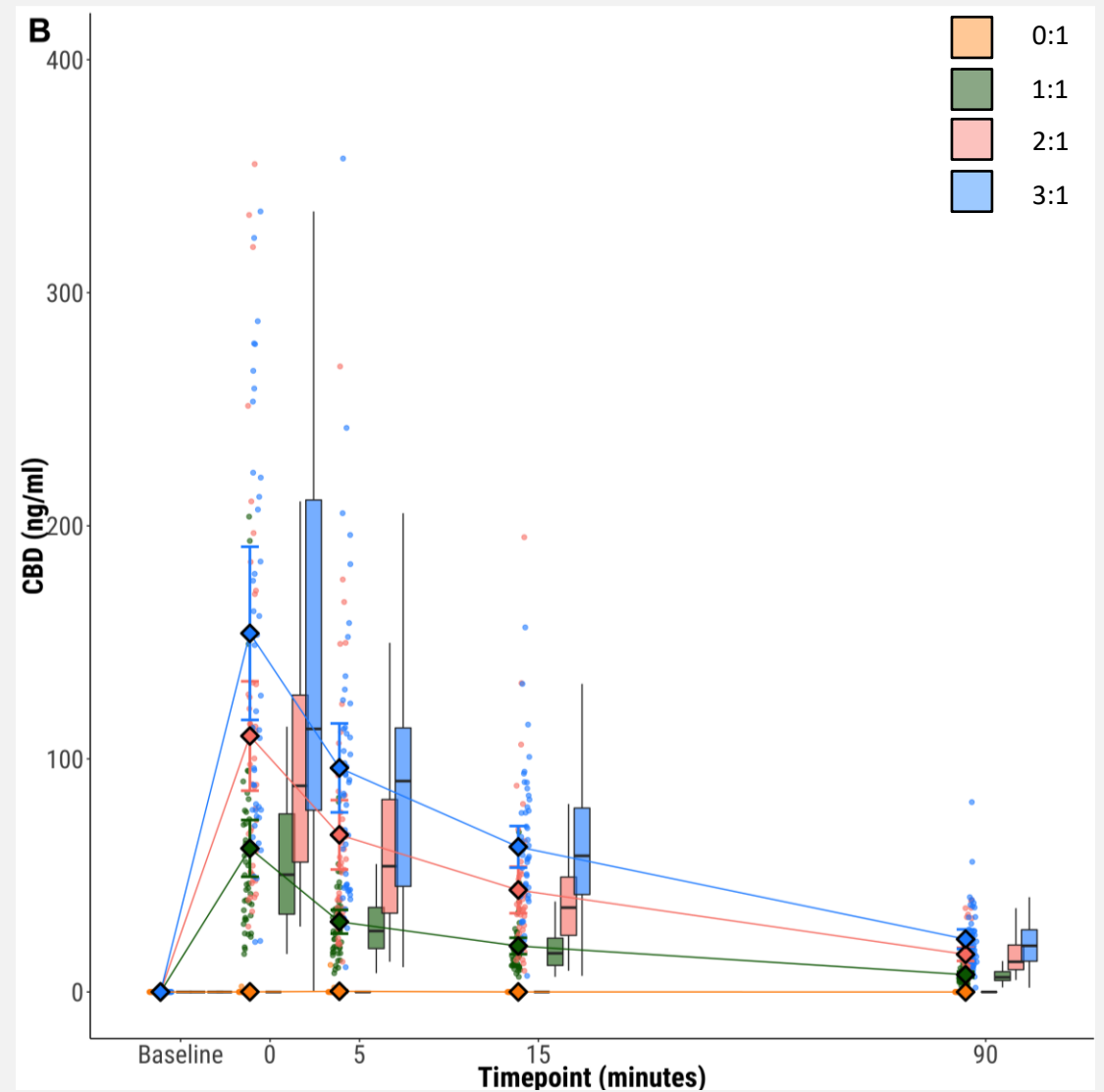
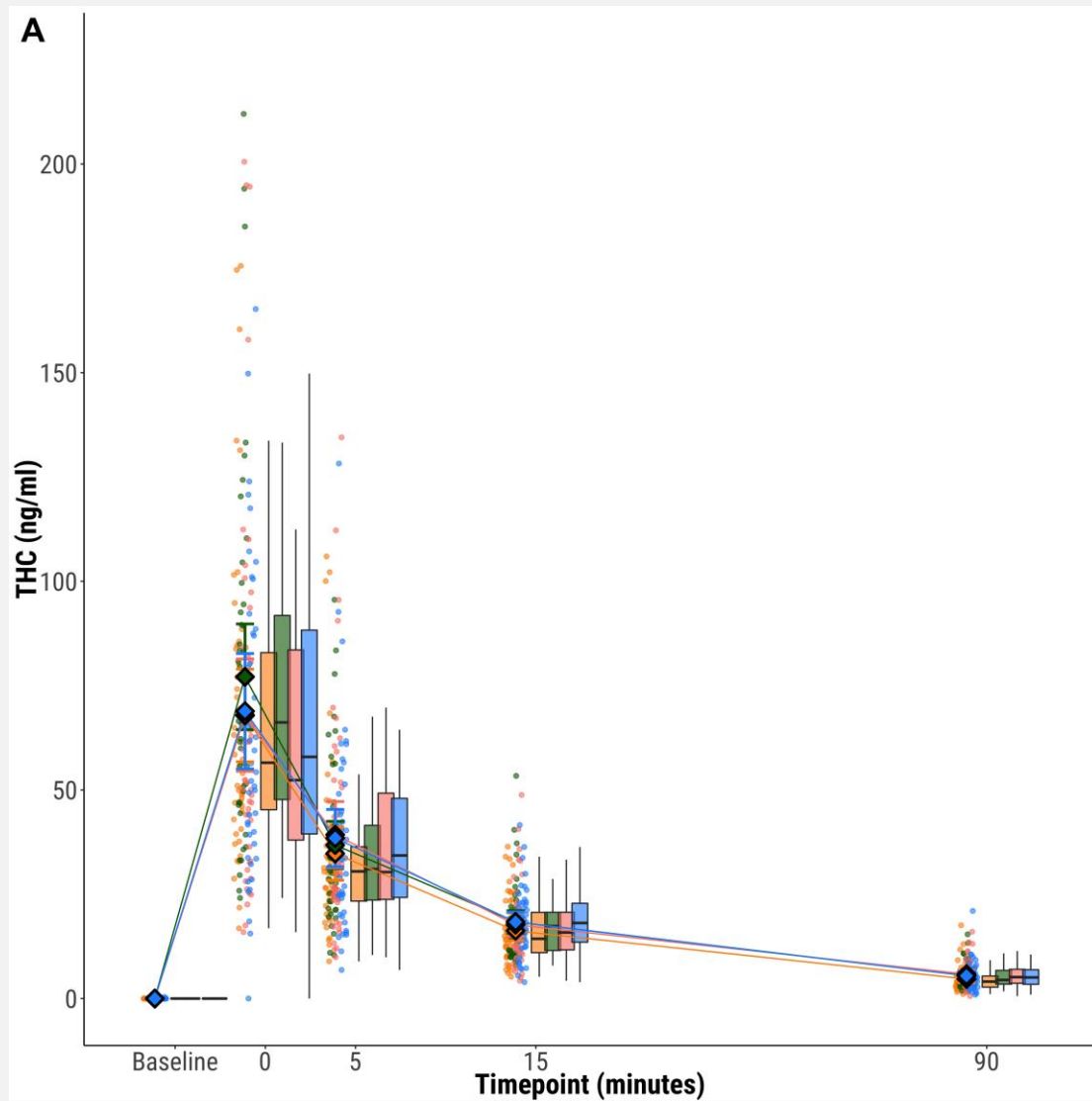
STATISTICAL ANALYSIS

Paired t-test
Change from sober to intoxicated

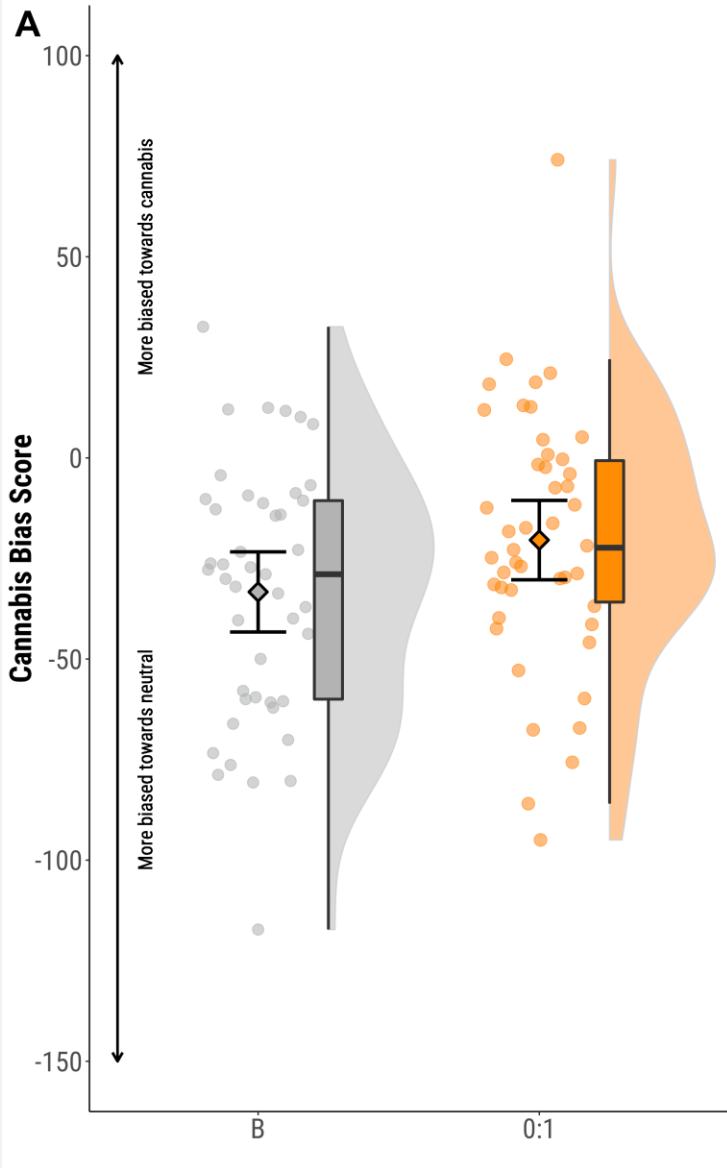
Linear mixed models
CBD effect



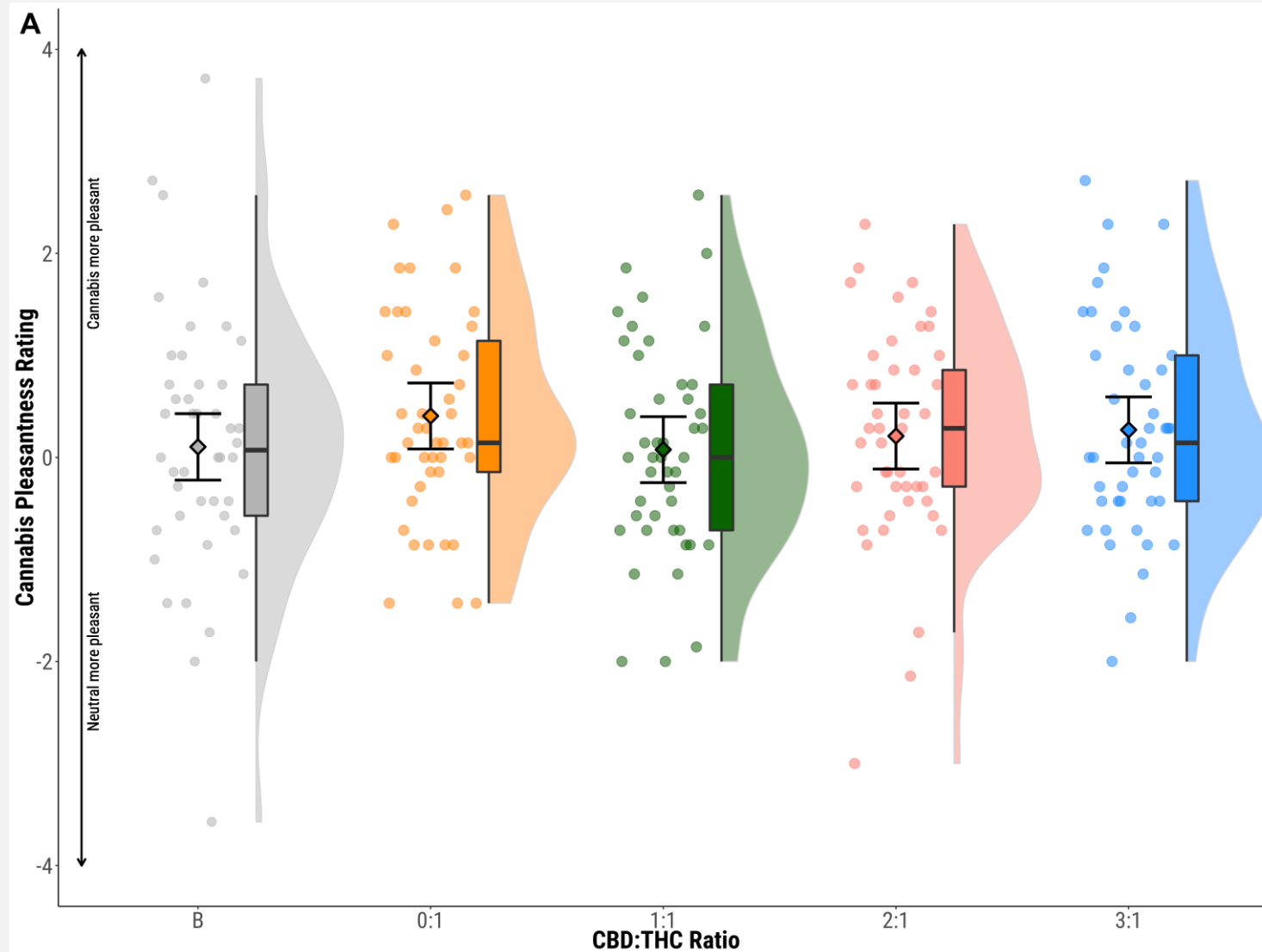
PHARMACOKINETICS



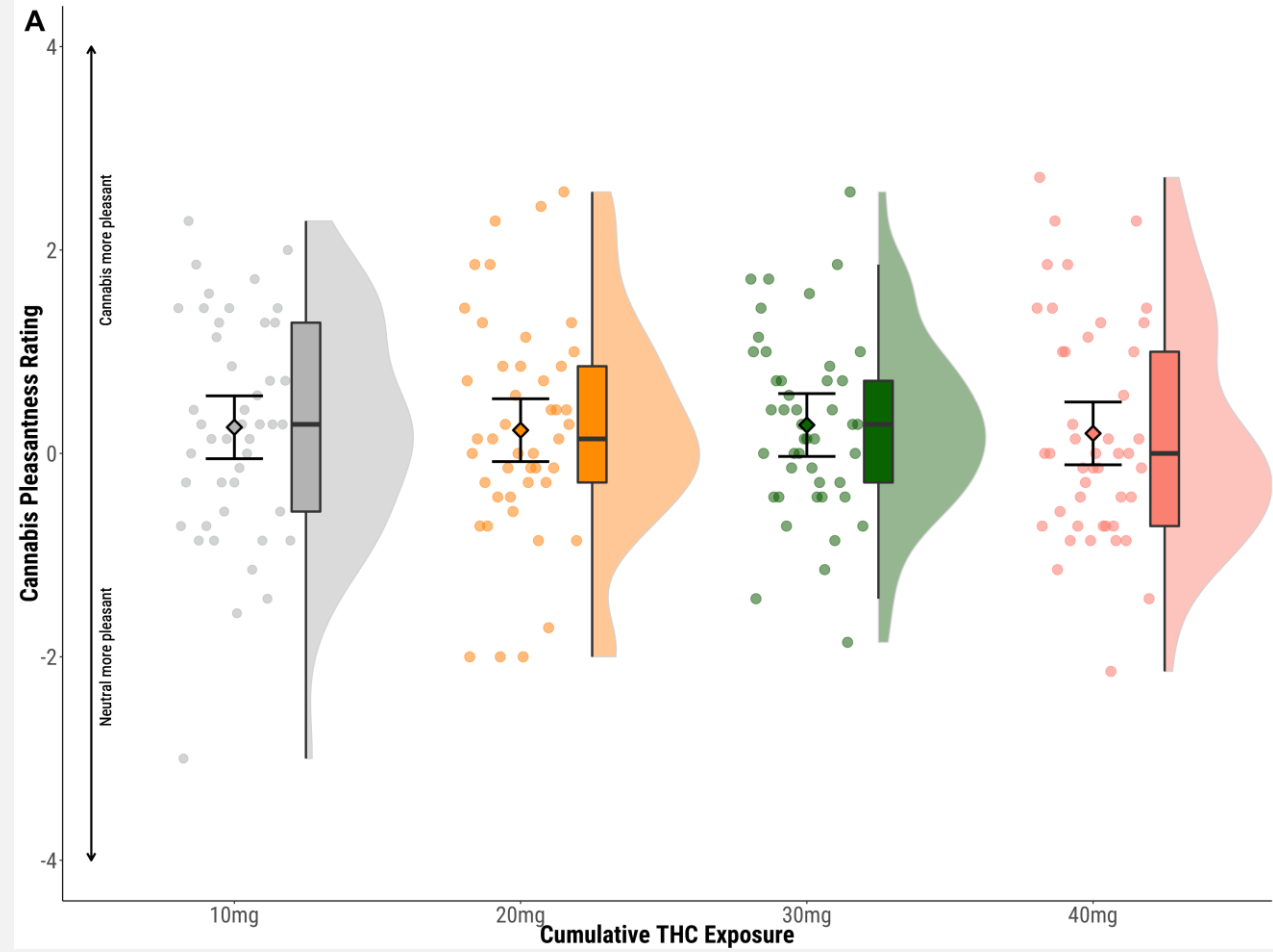
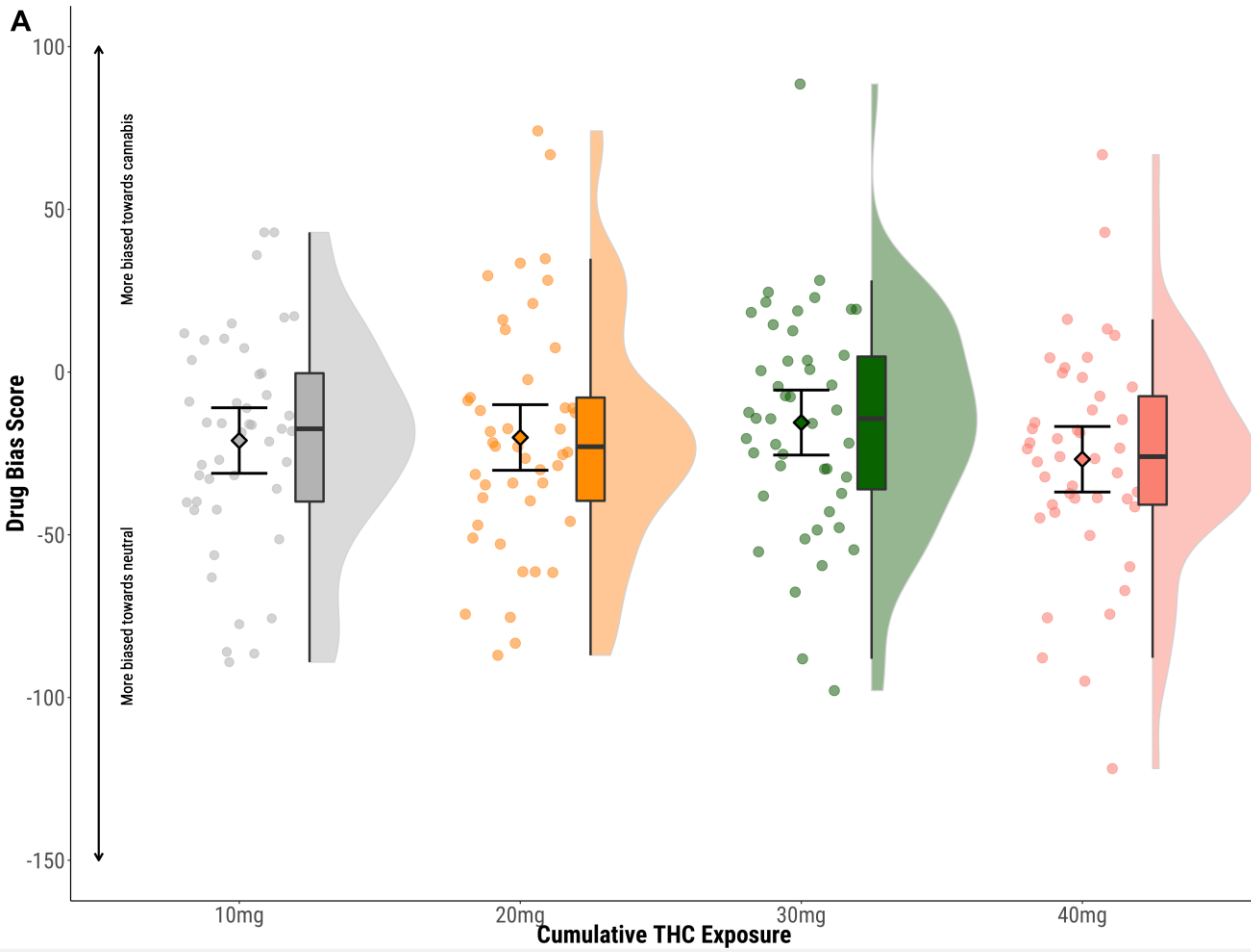
NO EFFECT OF CBD ON IMPLICIT WANTING



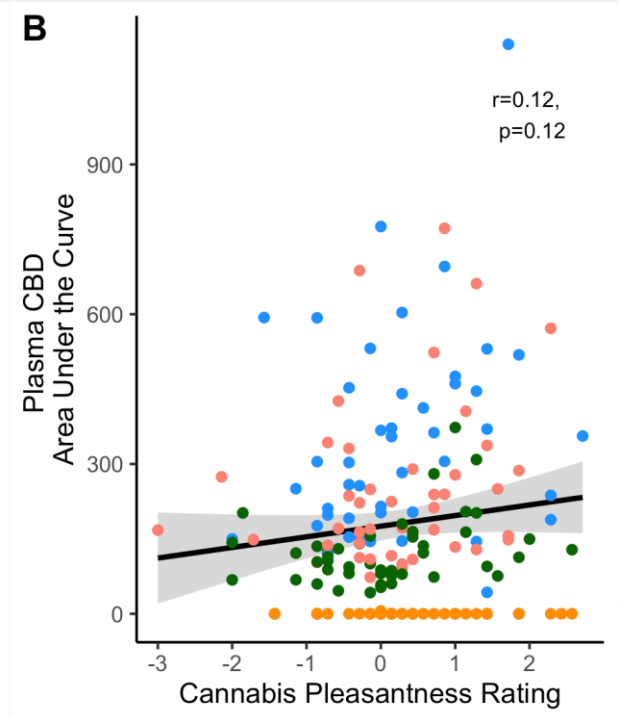
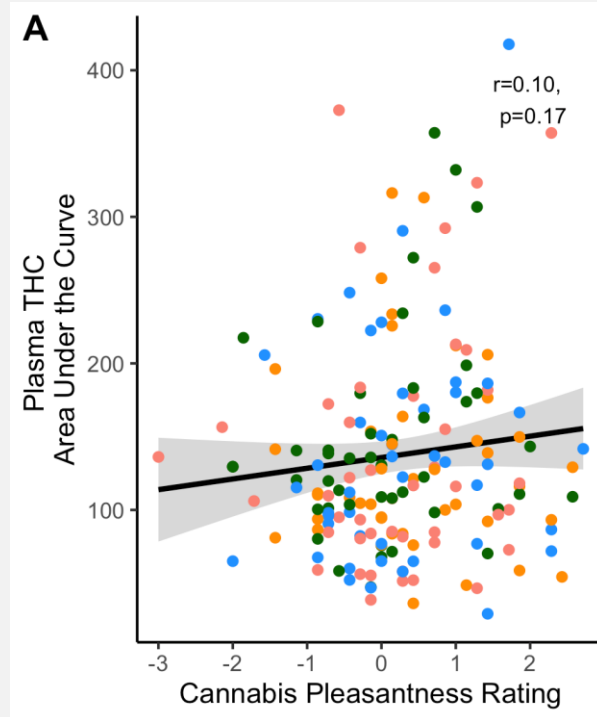
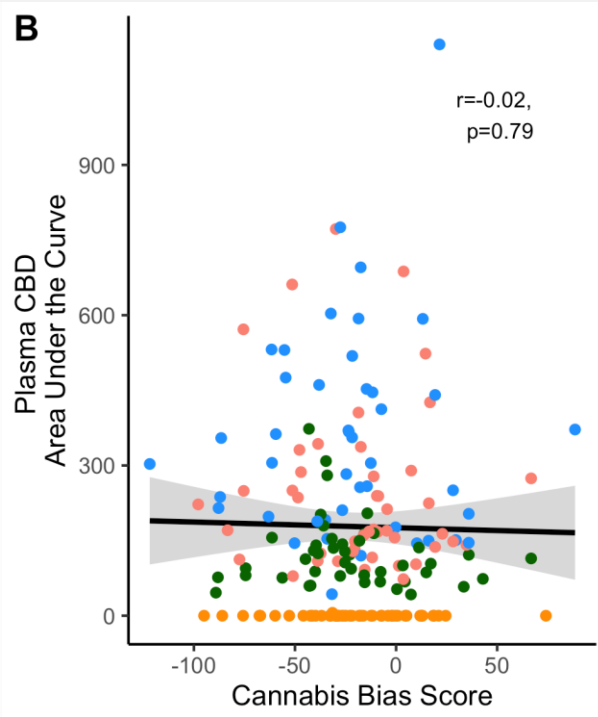
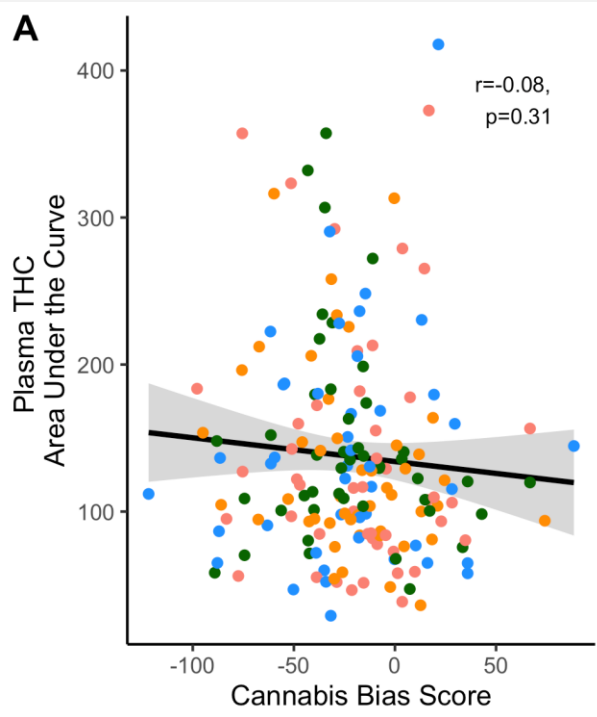
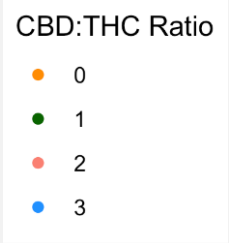
NO EFFECT OF THC OR CBD ON EXPLICIT LIKING



NO EFFECT OF CUMULATIVE THC EXPOSURE



NO RELATIONSHIP BETWEEN PLASMA THC OR CBD ON IMPLICIT WANTING OR EXPLICIT LIKING





DISCUSSION

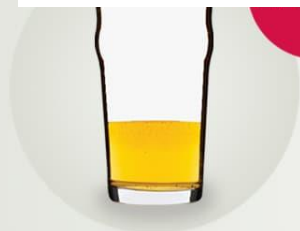
- **THC** acutely increases implicit wanting of cannabis in infrequent users without noticeable changes in explicit liking
- At most common **CBD:THC** ratios, no evidence that **CBD** protects against this
- No evidence that cumulative **THC** exposure increases bias
- Acute effects versus chronic effects
- Reducing **THC** dose



‘Standard THC units’: a proposal to standardize dose across all cannabis products and methods of administration

Tom P. Freeman^{1,2,3}  & Valentina Lorenzetti⁴ 

Addiction and Mental Health Group (AIM), Department of Psychology, University of Bath, Bath, UK,¹ National Addiction Centre, King’s College London, London, UK,² Clinical Psychopharmacology Unit, University College London, London, UK³ and School of Behavioural and Health Sciences, Australian Catholic University, Fitzroy, VIC, Australia⁴



Standard
4.5% cider



Standard
13% wine



Standard
40% whiskey



Standard
4% beer



Standard 4%
alcopop (275ml)

250ml

SUMMARY

- Attentional bias is associated with cannabis dependence
- **THC** acutely increases implicit bias towards cannabis stimuli in infrequent users
- Co-administered **CBD** may not be preventive of addiction
- Infrequent use alone may not increase bias further
- One way to better inform use is through a standard **THC** unit



MAIN STUDY FINDINGS

Associations of alcohol and substance use with mental health

KNOWLEDGE MARKET 2 (K2)

IN PROGRAMME

Friday, 25 November, 2022 - 10:50 to 12:20

[View presentations](#)

Chair



Goodwin Laura

Senior Lecturer In Mental Health, Lancaster University,
United Kingdom

Oral presentation

4. Can we make cannabis safer? An experimental study of four CBD: THC ratios in healthy volunteers

10:50 to 12:20

Knowledge market 2 (K2)

Amir Englund

ARTICLE **OPEN**

[Check for updates](#)

Does cannabidiol make cannabis safer? A randomised, double-blind, cross-over trial of cannabis with four different CBD:THC ratios

Amir Englund^{1,2}, Dominic Oliver², Edward Chesney², Lucy Chester², Jack Wilson³, Simina Sovi², Andrea De Micheli², John Hodsoll⁴, Paolo Fusar-Poli^{2,5}, John Strang^{1,6}, Robin M. Murray², Tom P. Freeman⁷ and Philip McGuire²



ACKNOWLEDGEMENTS

eCBD

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Jack Wilson

Lucy Chester

Simina Sovi

Stina Wigroth

John Hodson

John Strang

Robin Murray

Tom Freeman

Paolo Fusar-Poli

Philip McGuire



June 2019 – Data collection complete!



QUESTIONS?