WEBVTT

- NOTE duration:"01:13:00"
- NOTE recognizability:0.926
- NOTE language:en-us
- NOTE Confidence: 0.940253525
- 00:00:00.000 --> 00:00:00.640 Thank you,
- NOTE Confidence: 0.6033236
- 00:00:04.760 --> 00:00:05.280 Okay.
- NOTE Confidence: 0.920264657142857
- $00:00:06.600 \longrightarrow 00:00:08.724$ So thanks for waiting.
- NOTE Confidence: 0.920264657142857
- $00:00:08.724 \dashrightarrow 00:00:12.775$ And so I'll you know shorten this a
- NOTE Confidence: 0.920264657142857
- $00:00:12.775 \rightarrow 00:00:15.560$ bit and focus on the more interesting,
- NOTE Confidence: 0.920264657142857
- 00:00:15.560 --> 00:00:16.745 probably more interesting
- NOTE Confidence: 0.920264657142857
- 00:00:16.745 --> 00:00:19.115 newer material that I have and
- NOTE Confidence: 0.920264657142857
- 00:00:19.115 00:00:21.030 shorten the introductory parts.
- NOTE Confidence: 0.920264657142857
- 00:00:21.030 --> 00:00:23.850 But we'll try to get through
- NOTE Confidence: 0.920264657142857
- $00:00:23.850 \longrightarrow 00:00:25.685$ the get to the to the meat of
- NOTE Confidence: 0.920264657142857
- 00:00:25.685 --> 00:00:26.998 it pretty quickly here and have
- NOTE Confidence: 0.920264657142857
- $00{:}00{:}26.998 \dashrightarrow 00{:}00{:}28.360$ a little time for questions so.
- NOTE Confidence: 0.9402536
- $00{:}00{:}35{.}160 \dashrightarrow 00{:}00{:}36{.}880$ Let's see. No,
- NOTE Confidence: 0.9301902

 $00{:}00{:}36.880 \dashrightarrow 00{:}00{:}38.175$ you have to do it on the

NOTE Confidence: 0.9301902

 $00{:}00{:}38.175 \dashrightarrow 00{:}00{:}39.040$ laptop. Does it not work?

NOTE Confidence: 0.88005245

00:00:45.040 --> 00:00:46.520 That's right. It goes

NOTE Confidence: 0.919390781818182

00:00:47.160 --> 00:00:49.252 okay, okay. So yeah,

NOTE Confidence: 0.919390781818182

 $00:00:49.252 \rightarrow 00:00:53.580$ so we're gonna blaze through the historical.

NOTE Confidence: 0.919390781818182

 $00:00:53.580 \dashrightarrow 00:00:55.956$ Context and the old efficacy data

NOTE Confidence: 0.919390781818182

 $00{:}00{:}55{.}956$ --> $00{:}00{:}58{.}895$ and then I will really spend the

NOTE Confidence: 0.919390781818182

 $00:00:58.895 \rightarrow 00:01:02.030$ majority of the time then presenting

NOTE Confidence: 0.919390781818182

 $00{:}01{:}02{.}030 \dashrightarrow 00{:}01{:}05{.}480$ some hopefully interesting secondary

NOTE Confidence: 0.919390781818182

 $00:01:05.480 \dashrightarrow 00:01:08.172$ analyses we've done with the data

NOTE Confidence: 0.919390781818182

 $00:01:08.172 \dashrightarrow 00:01:12.060$ from the completed randomized trial.

NOTE Confidence: 0.919390781818182

 $00:01:12.060 \longrightarrow 00:01:17.338$ So just this is the sort of kube

NOTE Confidence: 0.919390781818182

 $00{:}01{:}17.338 \dashrightarrow 00{:}01{:}19.820$ bolkoff model of alcohol use disorder

NOTE Confidence: 0.919390781818182

 $00:01:19.820 \longrightarrow 00:01:22.970$ and I think it's a useful model.

NOTE Confidence: 0.919390781818182

00:01:22.970 --> 00:01:25.652 It's, you know,

NOTE Confidence: 0.919390781818182

 $00:01:25.652 \rightarrow 00:01:27.698$ we can argue about whether these

- NOTE Confidence: 0.919390781818182
- $00:01:27.698 \rightarrow 00:01:29.455$ are really separate domains,
- NOTE Confidence: 0.919390781818182
- $00:01:29.455 \rightarrow 00:01:32.365$ but they really capture the functional
- NOTE Confidence: 0.919390781818182
- 00:01:32.370 --> 00:01:37.050 aspects of addiction in incentive salience,
- NOTE Confidence: 0.919390781818182
- $00:01:37.050 \rightarrow 00:01:39.963$ negative negative emotionality,
- NOTE Confidence: 0.919390781818182
- $00:01:39.963 \dashrightarrow 00:01:43.847$ and executive function problems.
- NOTE Confidence: 0.919390781818182
- $00:01:43.850 \dashrightarrow 00:01:48.794$ And we'll come back to this later looking
- NOTE Confidence: 0.919390781818182
- $00:01:48.794 \rightarrow 00:01:52.046$ at the data I was going to for for.
- NOTE Confidence: 0.919390781818182
- $00:01:52.050 \rightarrow 00:01:54.570$ Based on a conversation I had recently
- NOTE Confidence: 0.919390781818182
- 00:01:54.570 -> 00:01:56.832 with with the Jerry Sanacora here,
- NOTE Confidence: 0.919390781818182
- $00:01:56.832 \rightarrow 00:01:59.310$ I was going to say a little
- NOTE Confidence: 0.919390781818182
- $00:01:59.310 \longrightarrow 00:02:00.610$ bit about the how the
- NOTE Confidence: 0.950317
- $00{:}02{:}02{.}970 \dashrightarrow 00{:}02{:}04{.}470$ ideal paradigm of translational science
- NOTE Confidence: 0.950317
- $00:02:04.470 \longrightarrow 00:02:07.530$ is not really what we've been doing here.
- NOTE Confidence: 0.950317
- $00{:}02{:}07{.}530 \dashrightarrow 00{:}02{:}10{.}122$ We've been working with things that
- NOTE Confidence: 0.950317
- $00:02:10.122 \dashrightarrow 00:02:12.203$ were serendipitously discovered and kind
- NOTE Confidence: 0.950317

 $00:02:12.203 \rightarrow 00:02:14.523$ of trying to figure out how they work,

NOTE Confidence: 0.950317

00:02:14.530 --> 00:02:15.980 but I don't think that's

NOTE Confidence: 0.950317

 $00:02:15.980 \longrightarrow 00:02:17.430$ worth spending much time on.

NOTE Confidence: 0.950317

 $00{:}02{:}17{.}430 \dashrightarrow 00{:}02{:}19{.}662$ And I think every body probably also

NOTE Confidence: 0.950317

 $00:02:19.662 \rightarrow 00:02:21.430$ knows what classic psychedelics are.

NOTE Confidence: 0.950317

00:02:21.430 --> 00:02:23.215 And you know that there are other

NOTE Confidence: 0.950317

 $00{:}02{:}23.215 \dashrightarrow 00{:}02{:}24.630$ kinds of psyched elics and so forth.

NOTE Confidence: 0.950317

 $00{:}02{:}24.630 \dashrightarrow 00{:}02{:}26.790$ But we're really talking about

NOTE Confidence: 0.950317

00:02:26.790 --> 00:02:28.390 mostly psilocybin,

NOTE Confidence: 0.950317

 $00{:}02{:}28.390 \dashrightarrow 00{:}02{:}32.790$ but other relatively similar,

NOTE Confidence: 0.950317

 $00:02:32.790 \longrightarrow 00:02:35.750$ mostly 5HT2A agonist or partial

NOTE Confidence: 0.950317

 $00:02:35.750 \longrightarrow 00:02:38.790$ agonist type drugs in the in this talk

NOTE Confidence: 0.91525645

 $00:02:41.870 \longrightarrow 00:02:44.216$ the yeah, this is the history

NOTE Confidence: 0.91525645

00:02:44.216 --> 00:02:47.029 that will kind of blaze through.

NOTE Confidence: 0.91525645

 $00{:}02{:}47.030 \dashrightarrow 00{:}02{:}49.976$ You know, there was a lot of research and

NOTE Confidence: 0.91525645

 $00:02:49.976 \rightarrow 00:02:52.720$ a lot of interest in classic psychedelics,

 $00:02:52.720 \longrightarrow 00:02:55.555$ both as a model of psychosis and

NOTE Confidence: 0.91525645

 $00{:}02{:}55{.}555 \dashrightarrow 00{:}02{:}57{.}722$ then quickly people who became

NOTE Confidence: 0.91525645

 $00:02:57.722 \longrightarrow 00:03:01.694$ interested in the potential therapeutic

NOTE Confidence: 0.91525645

 $00:03:01.694 \rightarrow 00:03:04.830$ uses of classic psychedelics.

NOTE Confidence: 0.91525645

 $00:03:04.830 \longrightarrow 00:03:07.915$ And then that really shot

NOTE Confidence: 0.91525645

 $00:03:07.915 \longrightarrow 00:03:10.973$ shut down around 1970.

NOTE Confidence: 0.91525645

 $00{:}03{:}10{.}973 \dashrightarrow 00{:}03{:}14.677$ For a variety of reasons but there was

NOTE Confidence: 0.91525645

00:03:14.677 --> 00:03:18.904 a fair amount of work done on alcohol

NOTE Confidence: 0.91525645

 $00:03:18.904 \dashrightarrow 00:03:22.618$ use disorder and six randomized trials NOTE Confidence: 0.91525645

 $00{:}03{:}22.618 \dashrightarrow 00{:}03{:}25.764$ which was which I'll I'll get back to

NOTE Confidence: 0.91525645

 $00:03:25.764 \rightarrow 00:03:28.400$ it in a moment showing you some the

NOTE Confidence: 0.91525645

 $00{:}03{:}28{.}400 \dashrightarrow 00{:}03{:}30{.}140$ meta analysis that was done of those.

NOTE Confidence: 0.91525645

 $00{:}03{:}30{.}140 \dashrightarrow 00{:}03{:}33{.}900$ But so now we're in the second wave NOTE Confidence: 0.91525645

 $00:03:33.900 \rightarrow 00:03:37.260$ and this is this goes up to 2020,

NOTE Confidence: 0.91525645

 $00:03:37.260 \rightarrow 00:03:40.106$ we'd probably be up to the ceiling.

00:03:40.106 --> 00:03:44.036 Of this room by now if we included 2021 NOTE Confidence: 0.91525645

 $00{:}03{:}44.036 \dashrightarrow 00{:}03{:}47.240$ and 22 but in any case there's a big

NOTE Confidence: 0.91525645

 $00{:}03{:}47{.}240$ --> $00{:}03{:}51{.}216$ renewal of interest and relatively a NOTE Confidence: 0.91525645

 $00:03:51.216 \rightarrow 00:03:54.840$ relative paucity of real useful data

NOTE Confidence: 0.91525645

 $00{:}03{:}54{.}840 \dashrightarrow 00{:}03{:}57{.}128$ their clinical or mechanistic although

NOTE Confidence: 0.91525645

 $00{:}03{:}57{.}128 \dashrightarrow 00{:}03{:}58{.}836$ that is that is starting to change.

NOTE Confidence: 0.91525645

 $00:03:58.840 \longrightarrow 00:04:02.320$ So so quickly reviewing the

NOTE Confidence: 0.91525645

00:04:02.320 --> 00:04:04.920 efficacy data of for psychedelics,

NOTE Confidence: 0.91525645

00:04:04.920 --> 00:04:07.305 classic psychedelics for alcohol use

NOTE Confidence: 0.91525645

 $00{:}04{:}07{.}305 \dashrightarrow 00{:}04{:}10{.}590$ disorder and other addictions to some extent.

NOTE Confidence: 0.91525645

 $00{:}04{:}10.590 \dashrightarrow 00{:}04{:}12.879$ This is the meta analysis that was

NOTE Confidence: 0.91525645

00:04:12.879 --> 00:04:15.372 done by Krebs and Johansen in published

NOTE Confidence: 0.91525645

 $00:04:15.372 \longrightarrow 00:04:18.326$ in 2012 looking at the six randomized

NOTE Confidence: 0.91525645

00:04:18.326 --> 00:04:20.845 control trials that had been done with

NOTE Confidence: 0.91525645

 $00:04:20.845 \rightarrow 00:04:23.949$ LSD in the treatment of alcohol use disorder.

NOTE Confidence: 0.91525645

 $00:04:23.950 \longrightarrow 00:04:28.120$ And these were they were all

 $00:04:28.120 \rightarrow 00:04:30.070$ double-blind up to Drug Administration.

NOTE Confidence: 0.91525645

 $00{:}04{:}30{.}070 \dashrightarrow 00{:}04{:}34{.}798$ And then sometimes people were knew that

NOTE Confidence: 0.91525645

 $00:04:34.798 \dashrightarrow 00:04:39.810$ they were getting no drugs for example, but.

NOTE Confidence: 0.91525645

 $00:04:39.810 \dashrightarrow 00:04:41.690$ There were some similarities are

NOTE Confidence: 0.91525645

00:04:41.690 --> 00:04:43.570 among all of these studies,

NOTE Confidence: 0.91525645

 $00{:}04{:}43.570$ --> $00{:}04{:}46.450$ almost all males, almost all inpatients,

NOTE Confidence: 0.91525645

 $00:04:46.450 \longrightarrow 00:04:47.870$ high doses of LSD,

NOTE Confidence: 0.91525645

 $00:04:47.870 \longrightarrow 00:04:49.290$ up to 800 micrograms,

NOTE Confidence: 0.91525645

 $00:04:49.290 \dashrightarrow 00:04:52.769$ which is really quite an enormous dose,

NOTE Confidence: 0.91525645

00:04:52.770 --> 00:04:55.495 just some different active and

NOTE Confidence: 0.91525645

 $00{:}04{:}55{.}495 \dashrightarrow 00{:}04{:}57{.}130$ inactive control conditions.

NOTE Confidence: 0.91525645

 $00{:}04{:}57{.}130 \dashrightarrow 00{:}05{:}00{.}460$ And the psychotherapeutic models range

NOTE Confidence: 0.91525645

 $00:05:00.460 \longrightarrow 00:05:03.278$ from really pretty much nonexistent

NOTE Confidence: 0.91525645

 $00{:}05{:}03.278$ --> $00{:}05{:}06.660$ to extensive preparation and and and

NOTE Confidence: 0.91525645

 $00:05:06.660 \rightarrow 00:05:09.420$ psychotherapy afterwards as well.

 $00:05:09.420 \longrightarrow 00:05:12.185$ So the the main finding of this

NOTE Confidence: 0.91525645

 $00:05:12.185 \longrightarrow 00:05:15.448$ was that there was a,

NOTE Confidence: 0.91525645

 $00:05:15.448 \longrightarrow 00:05:18.780$ you know a robust and very consistent

NOTE Confidence: 0.91525645

 $00{:}05{:}18.780 \dashrightarrow 00{:}05{:}20.600$ effect across these trials.

NOTE Confidence: 0.91525645

 $00:05:20.600 \rightarrow 00:05:23.330$ It was really very little evidence

NOTE Confidence: 0.91525645

 $00{:}05{:}23{.}409 \dashrightarrow 00{:}05{:}25{.}877$ of heterogeneity and it was a you

NOTE Confidence: 0.91525645

00:05:25.877 --> 00:05:28.236 know a good moderate sized effect in

NOTE Confidence: 0.91525645

 $00:05:28.236 \rightarrow 00:05:31.047$ spite of all the variability and the

NOTE Confidence: 0.91525645

 $00{:}05{:}31{.}047 \dashrightarrow 00{:}05{:}35{.}240$ over all odds ratio of close to two for.

NOTE Confidence: 0.91525645

 $00{:}05{:}35{.}240$ --> $00{:}05{:}38{.}216$ Abstinence or near abstinence at the NOTE Confidence: 0.91525645

 $00:05:38.216 \rightarrow 00:05:41.160$ first available follow up time point.

NOTE Confidence: 0.91525645

 $00{:}05{:}41{.}160 \dashrightarrow 00{:}05{:}43{.}190$ And those effects appeared to

NOTE Confidence: 0.91525645

 $00{:}05{:}43.190 \dashrightarrow 00{:}05{:}47.200$ persist for six months at least.

NOTE Confidence: 0.91525645

00:05:47.200 --> 00:05:47.800 It's interesting,

NOTE Confidence: 0.947441742857143

00:05:47.800 --> 00:05:48.755 I had noticed this before

NOTE Confidence: 0.947441742857143

 $00:05:48.755 \rightarrow 00:05:50.063$ but every single one of those

- NOTE Confidence: 0.947441742857143
- $00{:}05{:}50{.}063 \dashrightarrow 00{:}05{:}50{.}999$ trials technically failed.
- NOTE Confidence: 0.947441742857143
- $00:05:51.000 \longrightarrow 00:05:52.918$ Yes one of them was like on
- NOTE Confidence: 0.94629164
- $00:05:52.920 \longrightarrow 00:05:54.800$ the border, but that's right.
- NOTE Confidence: 0.94629164
- $00:05:54.800 \rightarrow 00:05:57.604$ And they were all under powered and they all,
- NOTE Confidence: 0.94629164
- $00:05:57.604 \dashrightarrow 00:05:58.878$ but they all look exactly the same.
- NOTE Confidence: 0.94629164
- $00:05:58.880 \rightarrow 00:06:00.399$ But they all look exactly the same.
- NOTE Confidence: 0.94629164
- 00:06:00.400 --> 00:06:02.956 So. So it's yeah, it's interesting.
- NOTE Confidence: 0.94629164
- $00:06:02.960 \longrightarrow 00:06:06.595$ It's just. This is a good lesson
- NOTE Confidence: 0.94629164
- $00:06:06.595 \rightarrow 00:06:08.500$ in not doing underpowered trials
- NOTE Confidence: 0.94629164
- $00:06:08.572 \rightarrow 00:06:11.026$ unless you want to convince people
- NOTE Confidence: 0.94629164
- $00:06:11.026 \longrightarrow 00:06:13.090$ of something that isn't true.
- NOTE Confidence: 0.94629164
- $00{:}06{:}13.090 \dashrightarrow 00{:}06{:}15.334$ So there was one controlled trial
- NOTE Confidence: 0.94629164
- $00:06:15.334 \rightarrow 00:06:18.005$ that maybe is less well known of
- NOTE Confidence: 0.94629164
- $00{:}06{:}18{.}005 \dashrightarrow 00{:}06{:}20{.}834$ LSD for opioid use disorder and this NOTE Confidence: 0.94629164
- 00:06:20.834 --> 00:06:24.278 one was it was randomized but it
- NOTE Confidence: 0.94629164

 $00:06:24.278 \longrightarrow 00:06:26.548$ was not double-blind 74 patients.

NOTE Confidence: 0.94629164

00:06:26.548 --> 00:06:29.914 So you know, not tiny but not

NOTE Confidence: 0.94629164

 $00:06:29.914 \rightarrow 00:06:32.460$ very large and these people,

NOTE Confidence: 0.94629164

 $00:06:32.460 \longrightarrow 00:06:33.990$ these were all.

NOTE Confidence: 0.94629164

 $00:06:33.990 \longrightarrow 00:06:35.350$ People who were on,

NOTE Confidence: 0.773826372857143

 $00{:}06{:}37.510 \dashrightarrow 00{:}06{:}39.505$ they were, they were getting out of

NOTE Confidence: 0.773826372857143

00:06:39.510 --> 00:06:42.126 of jail and they were on parole and

NOTE Confidence: 0.773826372857143

 $00:06:42.126 \dashrightarrow 00:06:44.649$ they would either go out and be on

NOTE Confidence: 0.773826372857143

 $00{:}06{:}44.649 \dashrightarrow 00{:}06{:}47.368$ parole and or they would stay in the

NOTE Confidence: 0.773826372857143

 $00{:}06{:}47.368 \dashrightarrow 00{:}06{:}49.485$ hospital and they'd get this preparation

NOTE Confidence: 0.773826372857143

 $00{:}06{:}49{.}485 \dashrightarrow 00{:}06{:}51{.}550$ and then a high dose LSD session

NOTE Confidence: 0.773826372857143

 $00{:}06{:}51{.}550 \dashrightarrow 00{:}06{:}53{.}708$ and then go to the after care clinic.

NOTE Confidence: 0.773826372857143

 $00:06:53.710 \longrightarrow 00:06:57.238$ And then both groups were followed for a year

NOTE Confidence: 0.773826372857143

 $00{:}06{:}57{.}238 \dashrightarrow 00{:}07{:}01{.}296$ and part of their parole was daily urine.

NOTE Confidence: 0.773826372857143

 $00{:}07{:}01.296 \dashrightarrow 00{:}07{:}02.620$ Drug monitoring.

NOTE Confidence: 0.773826372857143

 $00{:}07{:}02.620 \dashrightarrow 00{:}07{:}04.720$ So if the outcome data is is

- NOTE Confidence: 0.773826372857143
- $00:07:04.720 \longrightarrow 00:07:06.580$ is certainly of high quality.
- NOTE Confidence: 0.773826372857143
- $00:07:06.580 \longrightarrow 00:07:13.017$ And so the the very kind of low
- NOTE Confidence: 0.773826372857143
- $00:07:13.017 \rightarrow 00:07:15.530$ tech figure on the right there from
- NOTE Confidence: 0.773826372857143
- $00:07:15.608 \dashrightarrow 00:07:18.820$ Archives to General Psychiatry shows
- NOTE Confidence: 0.773826372857143
- 00:07:18.820 --> 00:07:22.500 the cumulative total abstinence
- NOTE Confidence: 0.773826372857143
- $00{:}07{:}22.500 \dashrightarrow 00{:}07{:}26.482$ rates at 369 and 12 months and you
- NOTE Confidence: 0.773826372857143
- $00:07:26.482 \longrightarrow 00:07:28.886$ can see there's a real pretty.
- NOTE Confidence: 0.773826372857143
- $00:07:28.886 \rightarrow 00:07:30.816$ Robust separation starting as early
- NOTE Confidence: 0.773826372857143
- $00{:}07{:}30{.}816 \dashrightarrow 00{:}07{:}33{.}435$ as three months and then for the
- NOTE Confidence: 0.773826372857143
- 00:07:33.435 --> 00:07:35.457 treatment group it really kind of
- NOTE Confidence: 0.773826372857143
- $00:07:35.457 \rightarrow 00:07:37.736$ flattens out around 25% between 9
- NOTE Confidence: 0.773826372857143
- 00:07:37.736 --> 00:07:40.684 and 12 months versus about 5% in the
- NOTE Confidence: 0.773826372857143
- 00:07:40.684 --> 00:07:42.340 in the other group which is you know
- NOTE Confidence: 0.773826372857143
- $00{:}07{:}42.395 \dashrightarrow 00{:}07{:}44.225$ probably what you would have expected.
- NOTE Confidence: 0.773826372857143
- $00{:}07{:}44.230 \dashrightarrow 00{:}07{:}47.006$ The 5% that is in people with you
- NOTE Confidence: 0.773826372857143

 $00:07:47.006 \rightarrow 00:07:49.890$ know this was they weren't getting

NOTE Confidence: 0.773826372857143

 $00{:}07{:}49{.}890 \dashrightarrow 00{:}07{:}52{.}365$ methadone and they were just out in the

NOTE Confidence: 0.773826372857143

 $00:07:52.365 \rightarrow 00:07:56.922$ community and the 30 you know 25% is.

NOTE Confidence: 0.773826372857143

00:07:56.922 --> 00:07:57.928 You know,

NOTE Confidence: 0.773826372857143

00:07:57.930 --> 00:07:59.701 I mean it doesn't sound like the

NOTE Confidence: 0.773826372857143

 $00:07:59.701 \dashrightarrow 00:08:01.066$ greatest outcome, but it it's it,

NOTE Confidence: 0.773826372857143

00:08:01.066 --> 00:08:02.350 it's much better than than what

NOTE Confidence: 0.773826372857143

 $00:08:02.401 \longrightarrow 00:08:03.766$ you otherwise would have seen.

NOTE Confidence: 0.773826372857143

 $00{:}08{:}03.770 \dashrightarrow 00{:}08{:}04.966$ So this was actually,

NOTE Confidence: 0.773826372857143

00:08:04.966 --> 00:08:07.139 I mean I think an interesting study

NOTE Confidence: 0.773826372857143

 $00{:}08{:}07{.}139 \dashrightarrow 00{:}08{:}09{.}047$ that also should be followed up.

NOTE Confidence: 0.773826372857143

00:08:09.050 -> 00:08:12.550 So now jumping forward to the the

NOTE Confidence: 0.773826372857143

 $00:08:12.550 \longrightarrow 00:08:15.335$ first study that was published

NOTE Confidence: 0.773826372857143

 $00:08:15.335 \longrightarrow 00:08:19.760$ of Psychedelic for Addiction in

NOTE Confidence: 0.773826372857143

 $00{:}08{:}19.760 \dashrightarrow 00{:}08{:}21.770$ the 21st century,

NOTE Confidence: 0.773826372857143

00:08:21.770 --> 00:08:25.130 this was Matt Johnson's pilot study,

- NOTE Confidence: 0.773826372857143
- $00:08:25.130 \longrightarrow 00:08:27.790$ open label study, 15 participants.
- NOTE Confidence: 0.773826372857143
- $00{:}08{:}27.790 \dashrightarrow 00{:}08{:}31.108$ And this was two sessions of psilocybin,
- NOTE Confidence: 0.9503169
- 00:08:33.510 --> 00:08:33.670 can't
- NOTE Confidence: 0.939247154
- $00:08:36.710 \longrightarrow 00:08:37.942$ remember exactly the dose,
- NOTE Confidence: 0.939247154
- $00:08:37.942 \longrightarrow 00:08:39.790$ but it was a high dose,
- NOTE Confidence: 0.939247154
- $00:08:39.790 \longrightarrow 00:08:46.270$ 25 milligrams or so. And the people,
- NOTE Confidence: 0.939247154
- $00:08:46.270 \longrightarrow 00:08:48.223$ in fact some of them got a third dose.
- NOTE Confidence: 0.939247154
- $00{:}08{:}48{.}230 \dashrightarrow 00{:}08{:}49{.}838$ And So what you see here
- NOTE Confidence: 0.939247154
- $00:08:49.838 \longrightarrow 00:08:50.910$ is cigarettes per day.
- NOTE Confidence: 0.939247154
- $00:08:50.910 \rightarrow 00:08:54.778$ So this is a continuous outcome, very large.
- NOTE Confidence: 0.939247154
- $00{:}08{:}54.778 \dashrightarrow 00{:}08{:}59.040$ Effect at the six months and the
- NOTE Confidence: 0.939247154
- $00{:}08{:}59{.}040 \dashrightarrow 00{:}09{:}01{.}160$ majority of the participants,
- NOTE Confidence: 0.939247154
- $00{:}09{:}01{.}160 \dashrightarrow 00{:}09{:}03{.}278$ 80% of them in fact were
- NOTE Confidence: 0.939247154
- $00{:}09{:}03.280 \dashrightarrow 00{:}09{:}06.320$ abstinent at that point.
- NOTE Confidence: 0.939247154
- $00:09:06.320 \longrightarrow 00:09:09.071$ They did some longer term follow-ups up
- NOTE Confidence: 0.939247154

 $00:09:09.071 \rightarrow 00:09:12.355$ to 12 months and a follow up paper owning

NOTE Confidence: 0.939247154

00:09:12.355 --> 00:09:15.835 people for anywhere from 16 to 57 months.

NOTE Confidence: 0.939247154

 $00{:}09{:}15.840 \dashrightarrow 00{:}09{:}19.332$ And found that nine out of 15 of them

NOTE Confidence: 0.939247154

 $00:09:19.332 \longrightarrow 00:09:21.515$ still at 60% were still abstinent.

NOTE Confidence: 0.939247154

 $00:09:21.515 \dashrightarrow 00:09:23.055$ So pretty remarkable compared

NOTE Confidence: 0.939247154

00:09:23.055 --> 00:09:24.840 to any available treatment,

NOTE Confidence: 0.939247154

 $00:09:24.840 \longrightarrow 00:09:27.372$ small open label study.

NOTE Confidence: 0.939247154

 $00:09:27.372 \longrightarrow 00:09:31.252$ So there's an ongoing nearly finished

NOTE Confidence: 0.939247154

00:09:31.252 --> 00:09:34.960 open label RCT with I think supposed

NOTE Confidence: 0.939247154

 $00:09:34.960 \rightarrow 00:09:37.752$ to have 80 participants that they're

NOTE Confidence: 0.939247154

00:09:37.752 --> 00:09:40.280 getting pretty close to completing,

NOTE Confidence: 0.939247154

 $00:09:40.280 \rightarrow 00:09:43.230$ which is psilocybin.

NOTE Confidence: 0.939247154

00:09:43.230 --> 00:09:46.320 Versus nicotine replacement,

NOTE Confidence: 0.939247154

 $00:09:46.320 \dashrightarrow 00:09:50.160$ open label RCT and the results look

NOTE Confidence: 0.939247154

 $00:09:50.160 \longrightarrow 00:09:52.380$ not quite as striking as this,

NOTE Confidence: 0.939247154

 $00:09:52.380 \rightarrow 00:09:55.080$ but still highly significant in the

 $00:09:55.080 \rightarrow 00:09:57.300$ interim analyses that they've done.

NOTE Confidence: 0.939247154

 $00{:}09{:}57{.}300 \dashrightarrow 00{:}10{:}01{.}660$ And we're that's that's at Johns Hopkins,

NOTE Confidence: 0.939247154

 $00{:}10{:}01{.}660 \dashrightarrow 00{:}10{:}08{.}818$ yeah at Matt Johnson and we're also with

NOTE Confidence: 0.939247154

00:10:08.820 --> 00:10:12.870 with University of Alabama Peter Hendricks.

NOTE Confidence: 0.939247154

 $00{:}10{:}12{.}870 \dashrightarrow 00{:}10{:}15{.}593$ Matt Johnson and NYU are doing a

NOTE Confidence: 0.939247154

 $00:10:15.593 \rightarrow 00:10:18.429$ three site night of funded study now,

NOTE Confidence: 0.939247154

 $00:10:18.430 \longrightarrow 00:10:22.576$ which is a placebo control for

NOTE Confidence: 0.939247154

 $00:10:22.576 \longrightarrow 00:10:24.906$ cigarettes for for cigarette addiction.

NOTE Confidence: 0.939247154

00:10:24.910 --> 00:10:25.450 Yeah,

NOTE Confidence: 0.939247154

 $00:10:25.450 \rightarrow 00:10:28.750$ so that'll be the first double-blind money.

NOTE Confidence: 0.9402536

 $00:10:30.870 \longrightarrow 00:10:33.588$ So that's that. So alcohol now.

NOTE Confidence: 0.9402536

 $00{:}10{:}33{.}590 \dashrightarrow 00{:}10{:}35{.}550$ So this was published a year later.

NOTE Confidence: 0.9402536

 $00:10:35.550 \longrightarrow 00:10:38.076$ This is the pilot study that.

NOTE Confidence: 0.9402536

 $00{:}10{:}38.080 \dashrightarrow 00{:}10{:}40.130$ My group completed University of

NOTE Confidence: 0.9402536

00:10:40.130 --> 00:10:42.760 New Mexico before I came to NYU,

- $00:10:42.760 \longrightarrow 00:10:43.699$ so very similar,
- NOTE Confidence: 0.9402536
- $00:10:43.699 \rightarrow 00:10:46.320$ kind of designed to the smoking study.
- NOTE Confidence: 0.9402536
- $00:10:46.320 \longrightarrow 00:10:50.640$ We just looked at 10 participants and
- NOTE Confidence: 0.9402536
- 00:10:50.640 --> 00:10:52.666 it was really a feasibility study,
- NOTE Confidence: 0.9402536
- $00:10:52.666 \rightarrow 00:10:55.648$ but we did look at the drinking
- NOTE Confidence: 0.9402536
- $00{:}10{:}55{.}648 \dashrightarrow 00{:}10{:}58{.}388$ outcomes and this is percent.
- NOTE Confidence: 0.9402536
- 00:10:58.388 --> 00:11:00.119 Heavy drinking days.
- NOTE Confidence: 0.9402536
- $00:11:00.120 \longrightarrow 00:11:01.560$ So for anybody that's not
- NOTE Confidence: 0.9402536
- $00{:}11{:}01{.}560 \dashrightarrow 00{:}11{:}02{.}712$ in the alcohol field,
- NOTE Confidence: 0.9402536
- $00:11:02.720 \rightarrow 00:11:04.628$ that's heavy drinking day is for
- NOTE Confidence: 0.9402536
- 00:11:04.628 --> 00:11:06.959 a man it's five or more drinks,
- NOTE Confidence: 0.9402536
- $00:11:06.960 \rightarrow 00:11:08.634$ and for a woman it's four or more drinks.
- NOTE Confidence: 0.9402536
- 00:11:08.640 --> 00:11:11.440 Standard drinks that is,
- NOTE Confidence: 0.9402536
- 00:11:11.440 --> 00:11:12.788 which is, you know,
- NOTE Confidence: 0.9402536
- $00{:}11{:}12.788 \dashrightarrow 00{:}11{:}15.215$ roughly it's .6 of an ounce of
- NOTE Confidence: 0.9402536
- $00:11:15.215 \rightarrow 00:11:16.865$ absolute ethanol and it's about

- NOTE Confidence: 0.9402536
- $00:11:16.865 \rightarrow 00:11:19.728$ a beer or a glass of wine or a
- NOTE Confidence: 0.9402536
- $00:11:19.728 \longrightarrow 00:11:23.453$ shot of hard liquor and so.
- NOTE Confidence: 0.9402536
- 00:11:23.453 --> 00:11:25.691 This is heavy drinking days and
- NOTE Confidence: 0.9402536
- 00:11:25.691 --> 00:11:28.613 you can see the two psilocybin
- NOTE Confidence: 0.9402536
- $00{:}11{:}28.613 \dashrightarrow 00{:}11{:}31.272$ sessions happened at weeks four
- NOTE Confidence: 0.9402536
- $00{:}11{:}31{.}272 \dashrightarrow 00{:}11{:}34{.}296$ and eight after the second and
- NOTE Confidence: 0.9402536
- $00:11:34.296 \rightarrow 00:11:36.760$ third time point on the graph.
- NOTE Confidence: 0.9402536
- $00:11:36.760 \longrightarrow 00:11:39.364$ So you can see there's some
- NOTE Confidence: 0.9402536
- $00{:}11{:}39{.}364 \dashrightarrow 00{:}11{:}41{.}168$ improvement between baseline with
- NOTE Confidence: 0.9402536
- $00:11:41.168 \rightarrow 00:11:44.360$ the first time point in week four.
- NOTE Confidence: 0.9402536
- $00:11:44.360 \longrightarrow 00:11:45.932$ That's while people were
- NOTE Confidence: 0.9402536
- $00{:}11{:}45{.}932 \dashrightarrow 00{:}11{:}47{.}277$ receiving some psychotherapy,
- NOTE Confidence: 0.9402536
- $00:11:47.277 \longrightarrow 00:11:50.619$ which was, part of which was.
- NOTE Confidence: 0.9402536
- $00{:}11{:}50.620 \dashrightarrow 00{:}11{:}53.204$ Motivational interviewing and and
- NOTE Confidence: 0.9402536
- $00:11:53.204 \rightarrow 00:11:55.148$ cognitive behavioral skills training.
- NOTE Confidence: 0.9402536

- $00:11:55.148 \rightarrow 00:11:56.768$ And so they were getting
- NOTE Confidence: 0.9402536
- $00:11:56.768 \longrightarrow 00:11:57.740$ some active treatment.
- NOTE Confidence: 0.9402536
- $00:11:57.740 \longrightarrow 00:11:59.472$ There was some improvement,
- NOTE Confidence: 0.9402536
- $00:11:59.472 \rightarrow 00:12:00.338$ not surprisingly,
- NOTE Confidence: 0.9402536
- $00:12:00.340 \longrightarrow 00:12:03.380$ but after week four,
- NOTE Confidence: 0.9402536
- $00{:}12{:}03{.}380 \dashrightarrow 00{:}12{:}05{.}180$ the second month after the
- NOTE Confidence: 0.9402536
- $00:12:05.180 \longrightarrow 00:12:06.260$ first psilocybin session,
- NOTE Confidence: 0.9402536
- $00:12:06.260 \longrightarrow 00:12:08.836$ there was a marked reduction in
- NOTE Confidence: 0.9402536
- 00:12:08.836 $\operatorname{-->}$ 00:12:11.688 heavy drinking days and the second
- NOTE Confidence: 0.9402536
- $00:12:11.688 \rightarrow 00:12:13.458$ psilocybin session at week eight.
- NOTE Confidence: 0.9402536
- $00{:}12{:}13.460 \dashrightarrow 00{:}12{:}15.860$ And those games were basically
- NOTE Confidence: 0.9402536
- $00:12:15.860 \longrightarrow 00:12:20.369$ maintained out to week 36, which was.
- NOTE Confidence: 0.9402536
- $00:12:20.370 \longrightarrow 00:12:24.409$ 28 weeks after the second psilocybin session.
- NOTE Confidence: 0.9402536
- $00:12:24.410 \longrightarrow 00:12:26.942$ And so we computed the effect
- NOTE Confidence: 0.9402536
- 00:12:26.942 --> 00:12:29.128 sizes versus both the the
- NOTE Confidence: 0.9402536
- $00:12:29.128 \rightarrow 00:12:31.495$ baseline and the week four,

- NOTE Confidence: 0.9402536
- $00:12:31.495 \rightarrow 00:12:34.689$ which is sort of the real baseline because

 $00{:}12{:}34.689 \dashrightarrow 00{:}12{:}37.410$ that's the immediate pre psilocybin

NOTE Confidence: 0.8869315666666667

 $00{:}12{:}39{.}490 \dashrightarrow 00{:}12{:}43{.}570$ baseline and the effect sizes are

NOTE Confidence: 0.8869315666666667

 $00:12:43.570 \longrightarrow 00:12:46.218$ you know on the range of .8 to.

NOTE Confidence: 0.8869315666666667

 $00:12:46.220 \longrightarrow 00:12:48.215$ To 1.0 or a little bit more.

NOTE Confidence: 0.8869315666666667

00:12:48.220 --> 00:12:49.660 So they're, they're good size effect

NOTE Confidence: 0.8869315666666667

00:12:49.660 --> 00:12:51.140 sizes, large effect. The second

NOTE Confidence: 0.94312879

 $00:12:51.140 \longrightarrow 00:12:53.100$ dose was after that third data point.

NOTE Confidence: 0.94312879

 $00:12:53.100 \longrightarrow 00:12:55.252$ So there's no further, there is no further

NOTE Confidence: 0.94312879

 $00:12:55.252 \rightarrow 00:12:56.620$ improvement after the second one. Yeah.

NOTE Confidence: 0.715308533333333

00:12:58.940 --> 00:13:00.179 Intent patients labeled

NOTE Confidence: 0.93522184625

 $00{:}13{:}00{.}460 \dashrightarrow 00{:}13{:}02{.}260$ yeah. And you know and it might have,

NOTE Confidence: 0.93522184625

 $00{:}13{:}02{.}260 \dashrightarrow 00{:}13{:}03{.}835$ you know the effect might have not

NOTE Confidence: 0.93522184625

 $00{:}13{:}03{.}835 \dashrightarrow 00{:}13{:}05{.}298$ persisted as long or it might have.

NOTE Confidence: 0.93522184625

00:13:05.300 --> 00:13:07.659 So we. Yeah. So we don't know,

 $00:13:07.660 \longrightarrow 00:13:09.080$ but that's a big question

NOTE Confidence: 0.93522184625

 $00{:}13{:}09{.}080 \dashrightarrow 00{:}13{:}10{.}500$ too is how many sessions.

NOTE Confidence: 0.93522184625

 $00:13:10.500 \rightarrow 00:13:12.116$ So this is the design of the completed

NOTE Confidence: 0.93522184625

 $00:13:12.116 \rightarrow 00:13:13.579$ trial and this has been published.

NOTE Confidence: 0.93522184625

 $00{:}13{:}13{.}580 \dashrightarrow 00{:}13{:}15{.}362$ I don't want to spend a lot of time.

NOTE Confidence: 0.93522184625

 $00:13:15.370 \longrightarrow 00:13:17.638$ On this, but just as an example

NOTE Confidence: 0.93522184625

00:13:17.638 --> 00:13:20.009 of sort of a typical design,

NOTE Confidence: 0.93522184625

 $00:13:20.010 \longrightarrow 00:13:21.186$ the way a lot of these

NOTE Confidence: 0.93522184625

 $00:13:21.186 \longrightarrow 00:13:21.970$ studies have been done,

NOTE Confidence: 0.93522184625

 $00{:}13{:}21{.}970 \dashrightarrow 00{:}13{:}24{.}345$ there are two hydro psilocybin

NOTE Confidence: 0.93522184625

00:13:24.345 --> 00:13:25.770 sessions and there's

NOTE Confidence: 0.951994233333333

 $00:13:27.930 \rightarrow 00:13:29.930$ manualized psychotherapy before the first

NOTE Confidence: 0.951994233333333

 $00{:}13{:}29{.}930 \dashrightarrow 00{:}13{:}32{.}929$ one in between the two and afterwards.

NOTE Confidence: 0.951994233333333

 $00{:}13{:}32{.}930 \dashrightarrow 00{:}13{:}37{.}670$ And these are a combination of content

NOTE Confidence: 0.951994233333333

 $00:13:37.670 \longrightarrow 00:13:41.004$ that's intended to help people

NOTE Confidence: 0.951994233333333

 $00{:}13{:}41.004 \dashrightarrow 00{:}13{:}44.706$ get prepared for the sessions and.

 $00:13:44.710 \longrightarrow 00:13:48.278$ Be able to manage the intense subjective

NOTE Confidence: 0.951994233333333

 $00:13:48.278 \rightarrow 00:13:52.550$ effects and to hopefully figure out

NOTE Confidence: 0.951994233333333

 $00:13:52.550 \rightarrow 00:13:56.110$ how to make use of the experience.

NOTE Confidence: 0.951994233333333

 $00:13:56.110 \rightarrow 00:13:58.952$ And so that's the sort of psychedelic

NOTE Confidence: 0.951994233333333

 $00{:}13{:}58{.}952 \dashrightarrow 00{:}14{:}01{.}970$ part of the treatment and the other

NOTE Confidence: 0.951994233333333

 $00:14:01.970 \longrightarrow 00:14:04.850$ part is evidence based treatment for

NOTE Confidence: 0.951994233333333

 $00:14:04.850 \longrightarrow 00:14:07.880$ alcohol use disorder to help them.

NOTE Confidence: 0.951994233333333

 $00:14:07.880 \longrightarrow 00:14:09.680$ Actually make some behavior changes.

NOTE Confidence: 0.951994233333333

00:14:09.680 --> 00:14:12.784 So in this case again it was combination

NOTE Confidence: 0.951994233333333

00:14:12.784 --> 00:14:15.855 of of motivational enhancement the
rapy

NOTE Confidence: 0.951994233333333

 $00:14:15.855 \rightarrow 00:14:20.700$ and some very brief cognitive behavioral

NOTE Confidence: 0.951994233333333

 $00{:}14{:}20.700 \dashrightarrow 00{:}14{:}24.908$ skills based the rapy and so the.

NOTE Confidence: 0.951994233333333

 $00{:}14{:}24{.}908 \dashrightarrow 00{:}14{:}28{.}550$ We followed them in the double-blind

NOTE Confidence: 0.951994233333333

 $00:14:28.663 \longrightarrow 00:14:30.287$ out to week 36.

NOTE Confidence: 0.951994233333333

 $00{:}14{:}30{.}290 \dashrightarrow 00{:}14{:}32{.}222$ There was an open label extension phase

00:14:32.222 --> 00:14:34.167 that I'm not going to talk about,

NOTE Confidence: 0.951994233333333

 $00{:}14{:}34{.}170 \dashrightarrow 00{:}14{:}36{.}200$ but here's the primary outcome from the

NOTE Confidence: 0.951994233333333

 $00{:}14{:}36{.}200 \dashrightarrow 00{:}14{:}38{.}529$ papers which was percent heavy drinking day.

NOTE Confidence: 0.951994233333333

 $00:14:38.530 \longrightarrow 00:14:41.122$ So same thing we were looking at in the

NOTE Confidence: 0.951994233333333

 $00{:}14{:}41{.}122 \dashrightarrow 00{:}14{:}45{.}169$ pilot and you can see very similar curves.

NOTE Confidence: 0.951994233333333

00:14:45.170 --> 00:14:47.735 The psilocy
bin is going to be in red and

NOTE Confidence: 0.951994233333333

 $00{:}14{:}47.735 \dashrightarrow 00{:}14{:}50.678$ the control in blue and all of these slides.

NOTE Confidence: 0.951994233333333

 $00:14:50.678 \longrightarrow 00:14:53.726$ So the again there's a big.

NOTE Confidence: 0.951994233333333

 $00:14:53.730 \longrightarrow 00:14:55.030$ Decrease during those first

NOTE Confidence: 0.951994233333333

 $00:14:55.030 \rightarrow 00:14:56.330$ four weeks of therapy,

NOTE Confidence: 0.951994233333333

 $00:14:56.330 \longrightarrow 00:15:00.160$ but then another large increase in the well,

NOTE Confidence: 0.951994233333333

 $00{:}15{:}00{.}160 \dashrightarrow 00{:}15{:}02{.}740$ decrease in drinking in the psilocy bin

NOTE Confidence: 0.951994233333333

 $00{:}15{:}02.740 \dashrightarrow 00{:}15{:}05.890$ group after the first medication session,

NOTE Confidence: 0.951994233333333

 $00{:}15{:}05{.}890 \dashrightarrow 00{:}15{:}08{.}930$ which then is for the most part maintained.

NOTE Confidence: 0.951994233333333

 $00:15:08.930 \rightarrow 00:15:12.636$ I mean we can you know it looks just that,

NOTE Confidence: 0.951994233333333

00:15:12.636 --> 00:15:13.488 you know,

 $00:15:13.488 \rightarrow 00:15:16.210$ there may be some attenuation of the effect.

NOTE Confidence: 0.951994233333333

 $00:15:16.210 \rightarrow 00:15:20.718$ There really wasn't a time effect.

NOTE Confidence: 0.951994233333333

 $00{:}15{:}20.720 \dashrightarrow 00{:}15{:}22.309$ You know once you take the baseline

NOTE Confidence: 0.951994233333333

 $00{:}15{:}22{.}309 \dashrightarrow 00{:}15{:}25{.}220$ out of this that that was significant

NOTE Confidence: 0.951994233333333

 $00:15:25.220 \rightarrow 00:15:27.596$ but or a group by time interaction.

NOTE Confidence: 0.951994233333333

 $00{:}15{:}27.600 \dashrightarrow 00{:}15{:}31.660$ So they weren't the the separation wasn't

NOTE Confidence: 0.951994233333333

 $00:15:31.660 \rightarrow 00:15:35.400$ diminishing to a significant extent.

NOTE Confidence: 0.951994233333333

00:15:35.400 --> 00:15:37.375 But if you look at it, it looks like,

NOTE Confidence: 0.951994233333333

 $00{:}15{:}37{.}375 \dashrightarrow 00{:}15{:}39{.}085$ you know may be the treatment effects

NOTE Confidence: 0.951994233333333

 $00:15:39.085 \rightarrow 00:15:40.919$ wearing off a little bit but maybe not.

NOTE Confidence: 0.951994233333333

 $00:15:40.920 \dashrightarrow 00:15:44.580$ Yeah, Hedges G for this outcome was .52.

NOTE Confidence: 0.951994233333333

 $00{:}15{:}44{.}580 \dashrightarrow 00{:}15{:}46{.}960$ So it's a solid medium sized effect

NOTE Confidence: 0.951994233333333

 $00{:}15{:}46{.}960 \dashrightarrow 00{:}15{:}49{.}519$ which is much better than you would

NOTE Confidence: 0.951994233333333

 $00{:}15{:}49{.}520 \dashrightarrow 00{:}15{:}52{.}599$ ever see with you know something

NOTE Confidence: 0.951994233333333

 $00{:}15{:}52{.}599 \dashrightarrow 00{:}15{:}55{.}008$ like naltrex one or a camper say so.

 $00:15:55.008 \rightarrow 00:15:56.640$ So that was encouraging.

NOTE Confidence: 0.951994233333333

00:15:56.640 --> 00:15:59.000 Drinks per day is another.

NOTE Confidence: 0.951994233333333

00:15:59.000 --> 00:16:02.200 This is just total quantity of

NOTE Confidence: 0.951994233333333

 $00:16:02.200 \longrightarrow 00:16:05.320$ alcohol divided by the number of

NOTE Confidence: 0.951994233333333

 $00:16:05.320 \dashrightarrow 00:16:08.686$ days and so it looks very similar.

NOTE Confidence: 0.951994233333333

 $00{:}16{:}08{.}690 \dashrightarrow 00{:}16{:}11{.}514$ And you know and that was also significant

NOTE Confidence: 0.951994233333333

 $00:16:11.514 \rightarrow 00:16:13.887$ over the entire follow up period

NOTE Confidence: 0.9335446666666667

 $00{:}16{:}16{.}170 \dashrightarrow 00{:}16{:}18{.}360$ and these are some of the dichotomous

NOTE Confidence: 0.9335446666666667

 $00{:}16{:}18{.}360 \dashrightarrow 00{:}16{:}19{.}970$ outcomes we looked at and I don't

NOTE Confidence: 0.9335446666666667

 $00:16:19.970 \longrightarrow 00:16:21.686$ want to spend too much time on this,

NOTE Confidence: 0.9335446666666667

 $00{:}16{:}21.690 \dashrightarrow 00{:}16{:}23.657$ but again psilocybins in red and the

NOTE Confidence: 0.9335446666666667

00:16:23.657 --> 00:16:26.089 ones that I put in bold there are the,

NOTE Confidence: 0.9335446666666667

 $00:16:26.090 \rightarrow 00:16:27.970$ this is the last week of follow up.

NOTE Confidence: 0.9335446666666667

 $00:16:27.970 \longrightarrow 00:16:28.810$ So that would be

NOTE Confidence: 0.9402536

 $00{:}16{:}31{.}170 \dashrightarrow 00{:}16{:}35{.}660$ 7 to 8. Well, seven months after

NOTE Confidence: 0.9402536

 $00:16:35.660 \rightarrow 00:16:37.640$ the second psilocybin session.

- NOTE Confidence: 0.9402536
- 00:16:37.640 --> 00:16:41.865 And so at that .48 of the psilocybin percent

 $00{:}16{:}41.865 \dashrightarrow 00{:}16{:}45.080$ of the psilocybin patients versus 24% of

NOTE Confidence: 0.9402536

 $00:16:45.080 \rightarrow 00:16:47.595$ the control participants were abstinent,

NOTE Confidence: 0.9402536

 $00:16:47.595 \rightarrow 00:16:50.120$ completely abstinent for that month.

NOTE Confidence: 0.9402536

 $00:16:50.120 \longrightarrow 00:16:53.680$ And $62 \ 1/2$ versus 40% were

NOTE Confidence: 0.9402536

00:16:53.680 --> 00:16:55.280 having no heavy drinking days,

NOTE Confidence: 0.9402536

 $00:16:55.280 \longrightarrow 00:16:58.650$ which is you know, it doesn't doesn't doesn't

NOTE Confidence: 0.9402536

 $00:16:58.650 \rightarrow 00:16:59.800$ mean they're in full remission,

NOTE Confidence: 0.9402536

 $00:16:59.800 \rightarrow 00:17:03.328$ but it's you know they're they're not having.

NOTE Confidence: 0.9402536

 $00:17:03.330 \rightarrow 00:17:05.290$ They're not doing binge drinking at least.

NOTE Confidence: 0.9402536

 $00{:}17{:}05{.}290 \dashrightarrow 00{:}17{:}08{.}778$ So so those effects were all pretty

NOTE Confidence: 0.9402536

00:17:08.778 --> 00:17:12.010 consistent and fairly compelling

NOTE Confidence: 0.9402536

 $00:17:12.010 \rightarrow 00:17:13.570$ and problems related to alcohol.

NOTE Confidence: 0.9402536

 $00{:}17{:}13.570 \dashrightarrow 00{:}17{:}15.058$ This is the SIP short inventory

NOTE Confidence: 0.9402536

 $00{:}17{:}15{.}058 \dashrightarrow 00{:}17{:}16{.}644$ of problems that you know covers

 $00:17:16.644 \longrightarrow 00:17:18.004$ a number of different areas.

NOTE Confidence: 0.9402536

 $00{:}17{:}18.010 \dashrightarrow 00{:}17{:}20.215$ So that was just showing that there

NOTE Confidence: 0.9402536

 $00{:}17{:}20{.}215 \dashrightarrow 00{:}17{:}21{.}476$ was functional significance to

NOTE Confidence: 0.9402536

 $00{:}17{:}21{.}476 \dashrightarrow 00{:}17{:}24{.}178$ these changes as well much larger

NOTE Confidence: 0.9402536

 $00{:}17{:}24.178 \dashrightarrow 00{:}17{:}28.076$ decrease in the psilocybin group so.

NOTE Confidence: 0.9402536

 $00{:}17{:}28.076 \dashrightarrow 00{:}17{:}30.204$ Here's one thing that is not in the

NOTE Confidence: 0.9402536

 $00{:}17{:}30{.}204 \dashrightarrow 00{:}17{:}32{.}396$ paper that I wanted to show you before

NOTE Confidence: 0.9402536

00:17:32.396 --> 00:17:34.679 I talk about mechanisms a little bit.

NOTE Confidence: 0.9402536

 $00{:}17{:}34.680 \dashrightarrow 00{:}17{:}36.558$ It's just in terms of outcomes.

NOTE Confidence: 0.9402536

 $00{:}17{:}36{.}560 \dashrightarrow 00{:}17{:}40{.}520$ This is a little bit complicated to look at,

NOTE Confidence: 0.9402536

00:17:40.520 - 00:17:45.120 but we were interested in seeing,

NOTE Confidence: 0.9402536

 $00:17:45.120 \rightarrow 00:17:47.297$ did it matter whether people had already

NOTE Confidence: 0.9402536

 $00{:}17{:}47{.}297 \dashrightarrow 00{:}17{:}49{.}741$ stopped or cut down on their drinking

NOTE Confidence: 0.9402536

 $00:17:49.741 \rightarrow 00:17:51.913$ before they received the sill assignment?

NOTE Confidence: 0.9402536

 $00:17:51.920 \longrightarrow 00:17:53.390$ Did you know in terms of what

NOTE Confidence: 0.9402536

 $00:17:53.390 \longrightarrow 00:17:54.800$ the size of the effect is?

- NOTE Confidence: 0.9402536
- 00:17:54.800 --> 00:17:56.920 And so?
- NOTE Confidence: 0.9402536
- $00:17:56.920 \rightarrow 00:18:01.480$ I divided the sample into people who were
- NOTE Confidence: 0.9402536
- 00:18:01.480 --> 00:18:03.640 still drinking at a problematic level,
- NOTE Confidence: 0.9402536
- $00{:}18{:}03.640 \dashrightarrow 00{:}18{:}06.935$ so that would be WHO risk level
- NOTE Confidence: 0.9402536
- $00{:}18{:}06{.}935 \dashrightarrow 00{:}18{:}08{.}715$ of two or greater.
- NOTE Confidence: 0.9402536
- 00:18:08.720 --> 00:18:13.104 So that's at least moderate risk,
- NOTE Confidence: 0.9402536
- 00:18:13.104 --> 00:18:16.160 moderate, severe or very severe
- NOTE Confidence: 0.9402536
- $00:18:16.160 \longrightarrow 00:18:17.875$ or less than that which would be,
- NOTE Confidence: 0.9402536
- 00:18:17.880 --> 00:18:18.768 you know,
- NOTE Confidence: 0.9402536
- $00:18:18.768 \longrightarrow 00:18:20.544$ basically within relatively safe
- NOTE Confidence: 0.9402536
- $00{:}18{:}20{.}544 \dashrightarrow 00{:}18{:}22{.}320$ guidelines which are comparable,
- NOTE Confidence: 0.9402536
- $00:18:22.320 \longrightarrow 00:18:23.712$ not exactly the same,
- NOTE Confidence: 0.9402536
- $00{:}18{:}23.712 \dashrightarrow 00{:}18{:}26.130$ but they're comparable to the NI AAA.
- NOTE Confidence: 0.9402536
- 00:18:26.130 --> 00:18:28.290 Guidelines for for safe drinking.
- NOTE Confidence: 0.9402536
- $00:18:28.290 \longrightarrow 00:18:31.682$ And so the ones in this with the
- NOTE Confidence: 0.9402536

 $00:18:31.682 \rightarrow 00:18:34.646$ solid lines are the ones who have

NOTE Confidence: 0.9402536

 $00{:}18{:}34.650 \dashrightarrow 00{:}18{:}38.526$ who are still drinking at a,

NOTE Confidence: 0.9402536

 $00{:}18{:}38{.}530 \dashrightarrow 00{:}18{:}41{.}374$ you know at a clinically significant NOTE Confidence: 0.9402536

 $00{:}18{:}41{.}374 \dashrightarrow 00{:}18{:}44{.}662$ level and the ones with the dotted

NOTE Confidence: 0.9402536

00:18:44.662 --> 00:18:48.106 lines were abstinent or just low low

NOTE Confidence: 0.9402536

 $00{:}18{:}48{.}106 \dashrightarrow 00{:}18{:}50{.}452$ intensity drinking and so obviously the. NOTE Confidence: 0.9402536

00:18:50.452 --> 00:18:53.259 People who are not drinking as much

NOTE Confidence: 0.9402536

 $00{:}18{:}53.259 \dashrightarrow 00{:}18{:}55.776$ have fewer heavy drinking days less

NOTE Confidence: 0.9402536

00:18:55.776 --> 00:19:00.590 than 10% at baseline and they do

NOTE Confidence: 0.9402536

 $00{:}19{:}00{.}590 \dashrightarrow 00{:}19{:}04{.}164$ both groups maintain that almost it's NOTE Confidence: 0.9402536

 $00{:}19{:}04{.}164 \dashrightarrow 00{:}19{:}06{.}782$ it's just a flat line for both groups

NOTE Confidence: 0.9402536

 $00{:}19{:}06{.}782 \dashrightarrow 00{:}19{:}09{.}096$ there's it's really no evidence that NOTE Confidence: 0.9402536

 $00:19:09.096 \longrightarrow 00:19:11.490$ the psilocybin or the treatment is

NOTE Confidence: 0.9402536

00:19:11.490 --> 00:19:13.510 doing anything further after that

NOTE Confidence: 0.9402536

 $00{:}19{:}13{.}510$ --> $00{:}19{:}15{.}638$ that that where the therapy I mean NOTE Confidence: 0.9402536

 $00:19:15.638 \rightarrow 00:19:18.173$ but the it's a different story with

- NOTE Confidence: 0.9402536
- $00:19:18.173 \rightarrow 00:19:20.501$ the people who are still symptomatic.

00:19:20.510 --> 00:19:21.802 There's a, you know,

NOTE Confidence: 0.9402536

 $00{:}19{:}21.802 \dashrightarrow 00{:}19{:}24.664$ a much bigger drop in the psilocybin group

NOTE Confidence: 0.9402536

00:19:24.664 --> 00:19:26.866 and it, you know, even looks in this,

NOTE Confidence: 0.9402536

 $00:19:26.870 \longrightarrow 00:19:27.626$ this is, you know,

NOTE Confidence: 0.9402536

 $00:19:27.626 \longrightarrow 00:19:29.041$ not a huge sample now if you're

NOTE Confidence: 0.9402536

00:19:29.041 --> 00:19:30.290 dividing it in half, right?

NOTE Confidence: 0.9402536

 $00:19:30.290 \longrightarrow 00:19:30.590$ But.

NOTE Confidence: 0.936899133333333

 $00:19:32.900 \longrightarrow 00:19:35.820$ 42 total in the in the higher risk

NOTE Confidence: 0.936899133333333

 $00{:}19{:}35{.}820 \dashrightarrow 00{:}19{:}38{.}115$ group but still the you know it's

NOTE Confidence: 0.936899133333333

00:19:38.115 -> 00:19:39.830 a bigger effect and if anything it

NOTE Confidence: 0.936899133333333

00:19:39.887 $\operatorname{-->}$ 00:19:41.543 looks like the groups are diverging

NOTE Confidence: 0.936899133333333

00:19:41.543 --> 00:19:43.014 over time that the psilocybin

NOTE Confidence: 0.936899133333333

00:19:43.014 --> 00:19:44.898 people continue to do better.

NOTE Confidence: 0.936899133333333

 $00{:}19{:}44{.}900 \dashrightarrow 00{:}19{:}46{.}764$ So it didn't look exactly like that for

 $00:19:46.764 \rightarrow 00:19:48.780$ all the outcomes for total abstinence,

NOTE Confidence: 0.936899133333333

 $00:19:48.780 \longrightarrow 00:19:51.300$ there was some of the people who were

NOTE Confidence: 0.936899133333333

00:19:51.300 --> 00:19:53.456 abstinent, you know or close to abstinent,

NOTE Confidence: 0.936899133333333

 $00:19:53.460 \longrightarrow 00:19:56.300$ they still they benefited some

NOTE Confidence: 0.936899133333333

 $00{:}19{:}56{.}300 \dashrightarrow 00{:}19{:}59{.}180$ but for these continuous measures

NOTE Confidence: 0.936899133333333

 $00:19:59.180 \longrightarrow 00:20:00.950$ it it looks like they're it.

NOTE Confidence: 0.936899133333333

 $00{:}20{:}00{.}950 \dashrightarrow 00{:}20{:}03{.}438$ You really get a larger effect for the

NOTE Confidence: 0.936899133333333

 $00:20:03.438 \rightarrow 00:20:06.241$ people who are still symptomatic and so non.

NOTE Confidence: 0.936899133333333

 $00{:}20{:}06{.}241 \dashrightarrow 00{:}20{:}08{.}227$ So that has implications both for

NOTE Confidence: 0.936899133333333

 $00:20:08.227 \rightarrow 00:20:10.349$ treatment and for study design in terms

NOTE Confidence: 0.936899133333333

 $00:20:10.349 \rightarrow 00:20:12.748$ of who you want to have in the trial.

NOTE Confidence: 0.88056946

 $00:20:15.310 \longrightarrow 00:20:17.908$ So okay, how does this work?

NOTE Confidence: 0.88056946

00:20:17.910 --> 00:20:22.412 If it does work, so they're just in

NOTE Confidence: 0.88056946

 $00{:}20{:}22{.}412 \dashrightarrow 00{:}20{:}24{.}629$ real most of this in the last few years.

NOTE Confidence: 0.88056946

 $00:20:24.630 \rightarrow 00:20:26.758$ There are quite a few studies now looking

NOTE Confidence: 0.88056946

 $00:20:26.758 \longrightarrow 00:20:29.393$ at the effects of classic psychedelics in.

- NOTE Confidence: 0.88056946
- 00:20:29.393 --> 00:20:31.331 Animal models of of alcohol use
- NOTE Confidence: 0.88056946
- $00{:}20{:}31{.}331 \dashrightarrow 00{:}20{:}33{.}501$ disorder and it's really only three
- NOTE Confidence: 0.88056946
- $00:20:33.501 \rightarrow 00:20:35.775$ of those actually are with psilocybin.
- NOTE Confidence: 0.88056946
- $00{:}20{:}35.780 \dashrightarrow 00{:}20{:}39.938$ There's a number of other compounds
- NOTE Confidence: 0.88056946
- $00:20:39.940 \longrightarrow 00:20:44.512$ including psilocybin and variety of
- NOTE Confidence: 0.88056946
- $00{:}20{:}44.512 \dashrightarrow 00{:}20{:}47.660$ different animal models including
- NOTE Confidence: 0.88056946
- 00:20:47.660 --> 00:20:49.940 just alcohol consumption,
- NOTE Confidence: 0.88056946
- $00:20:49.940 \rightarrow 00:20:52.855$ voluntary consumption preference,
- NOTE Confidence: 0.88056946
- $00:20:52.855 \longrightarrow 00:20:53.530$ selfadministration,
- NOTE Confidence: 0.88056946
- $00{:}20{:}53.530 \dashrightarrow 00{:}20{:}55.555$ behavioral sensitization and
- NOTE Confidence: 0.88056946
- $00:20:55.555 \rightarrow 00:20:59.070$ conditions place preference, so.
- NOTE Confidence: 0.88056946
- 00:20:59.070 --> 00:21:01.470 Different variety of different models
- NOTE Confidence: 0.88056946
- $00{:}21{:}01{.}470 \dashrightarrow 00{:}21{:}04{.}776$ and 15 out of 18 of these that I I
- NOTE Confidence: 0.88056946
- 00:21:04.776 --> 00:21:08.590 was able to find had at least some
- NOTE Confidence: 0.88056946
- $00:21:08.590 \rightarrow 00:21:11.710$ one of the psychedelic conditions
- NOTE Confidence: 0.88056946

00:21:11.710 --> 00:21:15.910 had had a larger effect than control

NOTE Confidence: 0.88056946

 $00{:}21{:}15{.}910 \dashrightarrow 00{:}21{:}19{.}666$ and there are a few negative studies.

NOTE Confidence: 0.88056946

 $00{:}21{:}19.670 \dashrightarrow 00{:}21{:}22.094$ One of them was by the same group NOTE Confidence: 0.88056946

 $00:21:22.094 \rightarrow 00:21:23.984$ that published this the second study

NOTE Confidence: 0.88056946

 $00{:}21{:}23{.}984 \dashrightarrow 00{:}21{:}26{.}180$ that that I that's in the

NOTE Confidence: 0.88056946

 $00{:}21{:}26.180 \dashrightarrow 00{:}21{:}27.710$ bottom of the slide here.

NOTE Confidence: 0.88056946

 $00{:}21{:}27{.}710 \dashrightarrow 00{:}21{:}31{.}620$ They using a different model in this study,

NOTE Confidence: 0.88056946

 $00:21:31.620 \rightarrow 00:21:34.341$ they were able to demonstrate that

NOTE Confidence: 0.88056946

 $00:21:34.341 \longrightarrow 00:21:37.825$ psilocybin did significantly reduce

NOTE Confidence: 0.88056946

00:21:37.825 --> 00:21:41.309 alcohol self self administration

NOTE Confidence: 0.88056946

 $00{:}21{:}41{.}310 \dashrightarrow 00{:}21{:}47{.}201$ and it also restored the M blue

NOTE Confidence: 0.88056946

 $00{:}21{:}47{.}201 \dashrightarrow 00{:}21{:}50{.}024$ 2 receptor gene expression in the

NOTE Confidence: 0.88056946

 $00{:}21{:}50{.}024 \dashrightarrow 00{:}21{:}51{.}709$ rats that were alcohol dependent.

NOTE Confidence: 0.88056946

 $00{:}21{:}51{.}710$ --> $00{:}21{:}55{.}427$ And so what's the significance of that?

NOTE Confidence: 0.88056946

 $00{:}21{:}55{.}430 \dashrightarrow 00{:}21{:}57{.}932$ They also demonstrated in a different

NOTE Confidence: 0.88056946

 $00{:}21{:}57{.}932 \dashrightarrow 00{:}22{:}00{.}373$ cohort of animals that knocked down

- NOTE Confidence: 0.88056946
- $00{:}22{:}00{.}373 \dashrightarrow 00{:}22{:}03{.}710$ of this of the M blue tube receptor

 $00{:}22{:}03.710 \dashrightarrow 00{:}22{:}05.790$ was associated with executive

NOTE Confidence: 0.88056946

00:22:05.790 --> 00:22:07.870 function impairments and increased

NOTE Confidence: 0.88056946

 $00{:}22{:}07{.}870 \dashrightarrow 00{:}22{:}11{.}110$ Q induced reinstatement and so.

NOTE Confidence: 0.88056946

 $00:22:11.110 \longrightarrow 00:22:13.598$ So the story is that this

NOTE Confidence: 0.88056946

 $00:22:13.598 \longrightarrow 00:22:15.550$ is a plausible model.

NOTE Confidence: 0.88056946

00:22:15.550 -> 00:22:17.910 They haven't connected the dots,

NOTE Confidence: 0.88056946

00:22:17.910 --> 00:22:19.085 assume they're working on that

NOTE Confidence: 0.88056946

 $00:22:19.085 \longrightarrow 00:22:19.790$ right now probably.

NOTE Confidence: 0.88056946

 $00{:}22{:}19.790 \dashrightarrow 00{:}22{:}22.454$ But they haven't you know shown that this

NOTE Confidence: 0.88056946

 $00:22:22.454 \rightarrow 00:22:25.368$ is what happens when psilocybin treated.

NOTE Confidence: 0.88056946

 $00{:}22{:}25{.}370 \dashrightarrow 00{:}22{:}25{.}668$ Rats.

NOTE Confidence: 0.88056946

00:22:25.668 --> 00:22:26.562 But it's a,

NOTE Confidence: 0.88056946

 $00:22:26.562 \rightarrow 00:22:28.530$ it's a hypothesis and a testable 1.

NOTE Confidence: 0.88056946

 $00:22:28.530 \longrightarrow 00:22:31.127$ So we'll see how that pans out.

- 00:22:31.130 --> 00:22:31.490 But
- NOTE Confidence: 0.94377579
- $00{:}22{:}34{.}370 \dashrightarrow 00{:}22{:}36{.}827$ that's, you know, one of the more
- NOTE Confidence: 0.94377579
- $00:22:36.827 \rightarrow 00:22:38.955$ sophisticated efforts to really come up
- NOTE Confidence: 0.94377579
- $00:22:38.955 \rightarrow 00:22:41.321$ with an explanation on a molecular level.
- NOTE Confidence: 0.943128857142857
- $00:22:41.330 \longrightarrow 00:22:42.646$ Do you know where that knockdown was?
- NOTE Confidence: 0.943128857142857
- $00{:}22{:}42.650 \dashrightarrow 00{:}22{:}45.330$ Was that medial Pfc or was it global or was I
- NOTE Confidence: 0.86817964
- $00:22:48.530 \longrightarrow 00:22:49.350$ think it was global?
- NOTE Confidence: 0.86817964
- $00{:}22{:}49{.}350 \dashrightarrow 00{:}22{:}50{.}930$ I think it was. I think it were.
- NOTE Confidence: 0.9402536
- $00{:}22{:}53.010 \dashrightarrow 00{:}22{:}54.610$ Yeah, I think I think so.
- NOTE Confidence: 0.9191548025
- $00:23:01.330 \longrightarrow 00:23:04.132$ Okay. There's been more done in
- NOTE Confidence: 0.9191548025
- $00{:}23{:}04{.}132 \dashrightarrow 00{:}23{:}06{.}610$ animal models of depression and
- NOTE Confidence: 0.9191548025
- $00:23:06.610 \rightarrow 00:23:09.250$ anxiety and stress related disorders
- NOTE Confidence: 0.9191548025
- $00:23:09.250 \longrightarrow 00:23:11.806$ and there have been, you know,
- NOTE Confidence: 0.9191548025
- $00:23:11.810 \rightarrow 00:23:14.090$ persisting effects demonstrated
- NOTE Confidence: 0.9191548025
- $00:23:14.090 \rightarrow 00:23:15.855$ with classic psychedelics and
- NOTE Confidence: 0.9191548025
- 00:23:15.855 --> 00:23:18.346 in a number of different models,

- NOTE Confidence: 0.9191548025
- $00:23:18.346 \rightarrow 00:23:21.610$ including for swim test,
- NOTE Confidence: 0.9191548025
- $00:23:21.610 \longrightarrow 00:23:22.888$ aversive foot shock
- NOTE Confidence: 0.950317
- $00:23:25.290 \longrightarrow 00:23:28.258$ another study looked at.
- NOTE Confidence: 0.950317
- 00:23:28.260 --> 00:23:31.298 Male preference for sucrose and female urine,
- NOTE Confidence: 0.950317
- 00:23:31.300 00:23:33.980 which is a sort of an hedonic well,
- NOTE Confidence: 0.950317
- $00{:}23{:}33{.}980 \dashrightarrow 00{:}23{:}35{.}898$ it's the, it's it's a hedonic response.
- NOTE Confidence: 0.950317
- $00:23:35.900 \rightarrow 00:23:38.616$ So it's they recover their interest in
- NOTE Confidence: 0.950317
- $00:23:38.620 \rightarrow 00:23:41.497$ things that they should be interested in.
- NOTE Confidence: 0.950317
- $00{:}23{:}41{.}500$ --> $00{:}23{:}46{.}540$ There's studies with DMT demonstrating NOTE Confidence: $0{.}950317$
- $00{:}23{:}46{.}540 \dashrightarrow 00{:}23{:}50{.}580$ anxiolytic and antidepressant effects and
- NOTE Confidence: 0.950317
- $00{:}23{:}50{.}580 \dashrightarrow 00{:}23{:}56{.}220$ also the facilitation of fear extinction.
- NOTE Confidence: 0.950317
- $00{:}23{:}56{.}220 \dashrightarrow 00{:}23{:}59{.}804$ So, so this all looks promising and
- NOTE Confidence: 0.950317
- $00{:}23{:}59{.}804 \dashrightarrow 00{:}24{:}02{.}252$ and fairly consistent across you know NOTE Confidence: 0.950317
- $00{:}24{:}02{.}252 \dashrightarrow 00{:}24{:}04{.}380$ a number of of different systems
- NOTE Confidence: 0.943175857142857
- $00{:}24{:}06{.}460 \dashrightarrow 00{:}24{:}09{.}248$ neuroplastogenic effects you know
- NOTE Confidence: 0.943175857142857

 $00:24:09.248 \rightarrow 00:24:11.850$ been clearly demonstrated including

NOTE Confidence: 0.943175857142857

 $00{:}24{:}11.850 \dashrightarrow 00{:}24{:}13.890$ spinogenesis and eptogenesis and

NOTE Confidence: 0.943175857142857

 $00{:}24{:}13.890 \dashrightarrow 00{:}24{:}16.440$ and some extent neurogenesis that's

NOTE Confidence: 0.943175857142857

 $00:24:16.513 \rightarrow 00:24:18.739$ you know the significance of that.

NOTE Confidence: 0.943175857142857

 $00:24:18.740 \longrightarrow 00:24:20.228$ Well the significance of all of

NOTE Confidence: 0.943175857142857

 $00:24:20.228 \longrightarrow 00:24:22.098$ this is you know less than clear,

NOTE Confidence: 0.943175857142857

00:24:22.100 --> 00:24:24.480 I mean cocaine will.

NOTE Confidence: 0.943175857142857

00:24:24.480 --> 00:24:26.782 Cause neuroplastic changes, I mean,

NOTE Confidence: 0.943175857142857

 $00:24:26.782 \longrightarrow 00:24:28.830$ so it doesn't mean that this is all

NOTE Confidence: 0.943175857142857

 $00{:}24{:}28.830 \dashrightarrow 00{:}24{:}31.504$ great and causing anything but it you

NOTE Confidence: 0.943175857142857

 $00{:}24{:}31{.}504 \dashrightarrow 00{:}24{:}34{.}632$ know this does this is something that

NOTE Confidence: 0.943175857142857

 $00:24:34.632 \rightarrow 00:24:36.815$ reliably happens and it you know that's

NOTE Confidence: 0.943175857142857

 $00{:}24{:}36.815 \dashrightarrow 00{:}24{:}38.510$ the case with ketamine as well so.

NOTE Confidence: 0.922931007692308

 $00{:}24{:}41.000 \dashrightarrow 00{:}24{:}43.664$ But we'll see and there's also

NOTE Confidence: 0.922931007692308

 $00:24:43.664 \rightarrow 00:24:44.996$ these antiinflammatory effects

NOTE Confidence: 0.922931007692308

 $00:24:44.996 \rightarrow 00:24:47.609$ of classic psychedelics which you
- NOTE Confidence: 0.922931007692308
- $00:24:47.609 \rightarrow 00:24:49.343$ know really it's not clear what
- NOTE Confidence: 0.922931007692308
- $00:24:49.343 \longrightarrow 00:24:50.999$ the significance is at this point,
- NOTE Confidence: 0.922931007692308
- $00:24:51.000 \rightarrow 00:24:51.732$ but they're pretty,
- NOTE Confidence: 0.922931007692308
- $00:24:51.732 \rightarrow 00:24:52.952$ they're quite pronounced and some
- NOTE Confidence: 0.922931007692308
- 00:24:52.952 --> 00:24:55.975 of them occur at very, very low doses,
- NOTE Confidence: 0.922931007692308
- $00:24:55.975 \rightarrow 00:24:59.200$ sub sub experiential doses that.
- NOTE Confidence: 0.922931007692308
- $00:24:59.200 \longrightarrow 00:25:03.536$ So it's whole other possible mechanism
- NOTE Confidence: 0.922931007692308
- $00:25:03.536 \rightarrow 00:25:06.440$ of action at least for some,
- NOTE Confidence: 0.922931007692308
- $00{:}25{:}06{.}440 \dashrightarrow 00{:}25{:}07{.}880$ you know some conditions perhaps.
- NOTE Confidence: 0.941259872
- 00:25:10.470 --> 00:25:12.458 And then, you know, we always talk
- NOTE Confidence: 0.941259872
- $00:25:12.458 \rightarrow 00:25:14.669$ about these drugs AS5HD2A agonists or
- NOTE Confidence: 0.941259872
- $00{:}25{:}14.669 \dashrightarrow 00{:}25{:}17.387$ partial agonists that that is true.
- NOTE Confidence: 0.941259872
- $00:25:17.390 \rightarrow 00:25:19.308$ And a lot of the effects definitely
- NOTE Confidence: 0.941259872
- $00{:}25{:}19{.}308 \dashrightarrow 00{:}25{:}23{.}229$ seem to be mediated by that receptor
- NOTE Confidence: 0.941259872
- $00{:}25{:}23.230 \dashrightarrow 00{:}25{:}25.522$ and you know, we can attenuate
- NOTE Confidence: 0.941259872

 $00:25:25.522 \rightarrow 00:25:27.549$ those effectors by blocking it

NOTE Confidence: 0.9377376925

 $00{:}25{:}29{.}630 \dashrightarrow 00{:}25{:}32{.}178$ or knocking it out.

NOTE Confidence: 0.9377376925

 $00{:}25{:}32{.}180 \dashrightarrow 00{:}25{:}34{.}882$ But there are a couple of studies

NOTE Confidence: 0.9377376925

 $00:25:34.882 \rightarrow 00:25:36.947$ demonstrating that in animal models

NOTE Confidence: 0.9377376925

 $00{:}25{:}36{.}947 \dashrightarrow 00{:}25{:}39{.}521$ that you can get the neurotrophic

NOTE Confidence: 0.9377376925

 $00{:}25{:}39{.}521 \dashrightarrow 00{:}25{:}41{.}779$ and antidepressant effects

NOTE Confidence: 0.932202801

00:25:44.220 --> 00:25:49.700 at a while blocking the 5H T to A receptor,

NOTE Confidence: 0.932202801

 $00:25:49.700 \rightarrow 00:25:52.044$ at least to the extent that you don't

NOTE Confidence: 0.932202801

 $00{:}25{:}52.044 \dashrightarrow 00{:}25{:}55.046$ get the head twitch which is considered

NOTE Confidence: 0.932202801

 $00:25:55.046 \rightarrow 00:25:58.858$ to be a that's a I don't know if it's,

NOTE Confidence: 0.932202801

 $00:25:58.860 \rightarrow 00:25:59.945$ I wouldn't say it's the

NOTE Confidence: 0.932202801

 $00{:}25{:}59{.}945 \dashrightarrow 00{:}26{:}01{.}354$ animal equivalent, but it's.

NOTE Confidence: 0.932202801

 $00:26:01.354 \rightarrow 00:26:05.250$ If it's the kind of dose that would,

NOTE Confidence: 0.932202801

 $00{:}26{:}05{.}250 \dashrightarrow 00{:}26{:}07{.}357$ it's the animal counterpart to the to

NOTE Confidence: 0.932202801

 $00{:}26{:}07{.}357 \dashrightarrow 00{:}26{:}09{.}140$ the subjective effects and and so it's

NOTE Confidence: 0.932202801

00:26:09.140 --> 00:26:10.919 a good you know an animal models if

 $00:26:10.919 \longrightarrow 00:26:12.641$ there's a head twitch then that drug

NOTE Confidence: 0.932202801

00:26:12.641 --> 00:26:14.290 if you give a comparable dose to a

NOTE Confidence: 0.932202801

 $00:26:14.290 \longrightarrow 00:26:15.767$ human it's going to be a psychedelic.

NOTE Confidence: 0.932202801

 $00:26:15.770 \longrightarrow 00:26:19.370$ So that's so the point of this is

NOTE Confidence: 0.932202801

 $00{:}26{:}19.370 \dashrightarrow 00{:}26{:}21.770$ you know not that to a agonism isn't

NOTE Confidence: 0.932202801

 $00{:}26{:}21.770 \dashrightarrow 00{:}26{:}23.760$ important but that it may be possible

NOTE Confidence: 0.932202801

 $00{:}26{:}23.760 \dashrightarrow 00{:}26{:}25.810$ to get some of these the rapeutic

NOTE Confidence: 0.932202801

 $00:26:25.810 \longrightarrow 00:26:29.540$ benefits without.

NOTE Confidence: 0.932202801

 $00{:}26{:}29{.}540 \dashrightarrow 00{:}26{:}31{.}969$ The mind altering effects or at least

NOTE Confidence: 0.932202801

 $00:26:31.969 \longrightarrow 00:26:34.617$ without you know the same extent of them.

NOTE Confidence: 0.932202801

 $00{:}26{:}34{.}620 \dashrightarrow 00{:}26{:}37{.}460$ So that remains to be demonstrated in people,

NOTE Confidence: 0.932202801

 $00{:}26{:}37{.}460 \dashrightarrow 00{:}26{:}41{.}196$ but that's that's a possibility Okay.

NOTE Confidence: 0.932202801

00:26:41.196 --> 00:26:44.524 So now I'm going to show you what

NOTE Confidence: 0.932202801

 $00{:}26{:}44{.}524 \dashrightarrow 00{:}26{:}46{.}613$ we can say about it again that you

NOTE Confidence: 0.932202801

 $00{:}26{:}46.613 \dashrightarrow 00{:}26{:}48.404$ know the trial that we completed

00:26:48.404 --> 00:26:50.094 wasn't wasn't a mechanistic study,

NOTE Confidence: 0.932202801

 $00{:}26{:}50{.}100 \dashrightarrow 00{:}26{:}53{.}316$ but we had a lot of self report

NOTE Confidence: 0.932202801

 $00{:}26{:}53.316 \dashrightarrow 00{:}26{:}57.710$ measures and we did a small pilot.

NOTE Confidence: 0.932202801

00:26:57.710 --> 00:26:59.182 FM, RI study that,

NOTE Confidence: 0.932202801

 $00{:}26{:}59{.}182 \dashrightarrow 00{:}27{:}00{.}924$ I'll show you some of the results

NOTE Confidence: 0.932202801

 $00:27:00.924 \longrightarrow 00:27:02.150$ of which are you know.

NOTE Confidence: 0.932202801

00:27:02.150 --> 00:27:02.686 Again,

NOTE Confidence: 0.932202801

 $00:27:02.686 \rightarrow 00:27:07.189$ it's very small but it's you know somewhat

NOTE Confidence: 0.94276945

 $00{:}27{:}09{.}430 \dashrightarrow 00{:}27{:}12{.}370$ interesting I think.

NOTE Confidence: 0.94276945

 $00{:}27{:}12.370 \dashrightarrow 00{:}27{:}15.770$ So starting with some of the

NOTE Confidence: 0.94276945

 $00:27:15.770 \longrightarrow 00:27:17.706$ obvious self report measures,

NOTE Confidence: 0.94276945

 $00:27:17.710 \rightarrow 00:27:23.630$ craving was significantly attenuated with.

NOTE Confidence: 0.94276945

00:27:23.630 --> 00:27:25.930 Psilocybin and it happened

NOTE Confidence: 0.94276945

00:27:25.930 --> 00:27:28.490 right away at week four,

NOTE Confidence: 0.94276945

 $00{:}27{:}28{.}490 \dashrightarrow 00{:}27{:}29{.}790$ I mean following week four.

NOTE Confidence: 0.94276945

00:27:29.790 --> 00:27:32.641 So it'd be week five and was maintained

 $00:27:32.641 \rightarrow 00:27:35.486$ pretty much over the the entire period.

NOTE Confidence: 0.94276945

 $00{:}27{:}35{.}486 \dashrightarrow 00{:}27{:}39{.}014$ So these just quick or quickly Orient you to

NOTE Confidence: 0.94276945

 $00:27:39.014 \rightarrow 00:27:41.870$ these slides because I'll show four of them.

NOTE Confidence: 0.94276945

 $00:27:41.870 \rightarrow 00:27:46.350$ I think the where it says week four,

NOTE Confidence: 0.94276945

 $00:27:46.350 \rightarrow 00:27:48.943$ that's really the baseline in this MMRM.

NOTE Confidence: 0.94276945

 $00:27:48.943 \longrightarrow 00:27:49.882$ So that's not.

NOTE Confidence: 0.94276945

 $00:27:49.882 \rightarrow 00:27:52.319$ Part of the repeated measure that's that's a,

NOTE Confidence: 0.94276945

 $00{:}27{:}52{.}320 \dashrightarrow 00{:}27{:}53{.}740$ that's the baseline covariate

NOTE Confidence: 0.94276945

 $00:27:53.740 \longrightarrow 00:27:55.515$ and where it says baseline,

NOTE Confidence: 0.94276945

 $00:27:55.520 \longrightarrow 00:27:56.815$ that's the beginning of the

NOTE Confidence: 0.94276945

 $00{:}27{:}56.815 \dashrightarrow 00{:}27{:}58.597$ study and that's in there just to

NOTE Confidence: 0.94276945

 $00{:}27{:}58.597 \dashrightarrow 00{:}28{:}00{.}116$ kind of show you what they were,

NOTE Confidence: 0.94276945

 $00{:}28{:}00{.}120 \dashrightarrow 00{:}28{:}01{.}116$ where they were at the beginning.

NOTE Confidence: 0.94276945

 $00{:}28{:}01{.}120 \dashrightarrow 00{:}28{:}02{.}640$ But that's not actually part of the model.

NOTE Confidence: 0.94276945

00:28:02.640 --> 00:28:04.980 So maybe you shouldn't have

 $00:28:04.980 \longrightarrow 00:28:05.800$ connected the dots, but

NOTE Confidence: 0.931427584615385

 $00{:}28{:}08{.}240 \dashrightarrow 00{:}28{:}10{.}438$ so then all of the points after

NOTE Confidence: 0.931427584615385

 $00:28:10.438 \longrightarrow 00:28:12.320$ that five weeks 5 through 36,

NOTE Confidence: 0.931427584615385

 $00:28:12.320 \longrightarrow 00:28:14.580$ those are the repeated measure.

NOTE Confidence: 0.931427584615385

 $00{:}28{:}14.580 \dashrightarrow 00{:}28{:}16.645$ And So what you're interested in here

NOTE Confidence: 0.931427584615385

00:28:16.645 --> 00:28:18.988 in terms of the treatment effect is

NOTE Confidence: 0.931427584615385

 $00{:}28{:}18{.}988 \dashrightarrow 00{:}28{:}21{.}082$ called the treatment effect and it

NOTE Confidence: 0.931427584615385

 $00:28:21.145 \rightarrow 00:28:23.419$ was highly significant and the time

NOTE Confidence: 0.931427584615385

 $00{:}28{:}23.419 \dashrightarrow 00{:}28{:}25.774$ effect is called assessment and that's

NOTE Confidence: 0.931427584615385

 $00:28:25.774 \rightarrow 00:28:27.859$ also highly significant is they're,

NOTE Confidence: 0.931427584615385

 $00:28:27.860 \longrightarrow 00:28:29.690$ they're both groups are going

NOTE Confidence: 0.931427584615385

 $00{:}28{:}29{.}690 \dashrightarrow 00{:}28{:}31{.}990$ downhill and the time by treatment

NOTE Confidence: 0.931427584615385

 $00{:}28{:}31{.}990 \dashrightarrow 00{:}28{:}34{.}310$ assessment is you know whether the

NOTE Confidence: 0.931427584615385

 $00{:}28{:}34{.}310 \dashrightarrow 00{:}28{:}36{.}686$ slopes are the same after people

NOTE Confidence: 0.931427584615385

 $00{:}28{:}36.686 \dashrightarrow 00{:}28{:}38.898$ after receiving the psilocybin.

NOTE Confidence: 0.931427584615385

 $00:28:38.900 \longrightarrow 00:28:40.292$ And so they weren't

00:28:40.292 --> 00:28:41.336 significantly different here,

NOTE Confidence: 0.931427584615385

 $00:28:41.340 \longrightarrow 00:28:43.220$ but you know maybe close.

NOTE Confidence: 0.916508510333333

 $00:28:44.140 \longrightarrow 00:28:45.700$ It is interesting that the cravings

NOTE Confidence: 0.916508510333333

 $00{:}28{:}45.700 \dashrightarrow 00{:}28{:}47.276$ continues to angle down throughout most

NOTE Confidence: 0.916508510333333

 $00{:}28{:}47{.}276 \dashrightarrow 00{:}28{:}49{.}244$ of the followup as opposed to your other

NOTE Confidence: 0.916508510333333

 $00{:}28{:}49{.}296 \dashrightarrow 00{:}28{:}50{.}880$ outcome measures where you got your

NOTE Confidence: 0.916508510333333

 $00{:}28{:}50{.}880 \dashrightarrow 00{:}28{:}52{.}536$ your strongest effect at week five,

NOTE Confidence: 0.916508510333333

 $00:28:52.536 \longrightarrow 00:28:54.920$ the five to eight time point and then

NOTE Confidence: 0.916508510333333

 $00{:}28{:}54{.}984 \dashrightarrow 00{:}28{:}56{.}937$ if anything flat or a ramp adrift

NOTE Confidence: 0.916508510333333

00:28:56.937 -> 00:28:59.499 up well may not be significant.

NOTE Confidence: 0.9335446666666667

00:28:59.620 --> 00:29:01.897 Well, I mean I think it's a real effect,

NOTE Confidence: 0.9335446666666667

 $00{:}29{:}01{.}900 \dashrightarrow 00{:}29{:}04{.}388$ it's the, the best way to decrease craving

NOTE Confidence: 0.9335446666666667

 $00{:}29{:}04{.}388 \dashrightarrow 00{:}29{:}06{.}897$ is to get people to stop drinking.

NOTE Confidence: 0.9335446666666667

00:29:06.900 --> 00:29:08.988 So you know it always goes

NOTE Confidence: 0.9335446666666667

 $00:29:08.988 \rightarrow 00:29:10.620$ down if people were asked.

00:29:10.620 --> 00:29:13.168 So you know downstream this, you know this.

NOTE Confidence: 0.9335446666666667

00:29:13.168 --> 00:29:15.718 You know, like we will maybe try to spend,

NOTE Confidence: 0.9335446666666667

 $00:29:15.720 \longrightarrow 00:29:17.380$ those people are not drinking

NOTE Confidence: 0.9335446666666667

 $00:29:17.380 \rightarrow 00:29:19.040$ because we decreased their craving.

NOTE Confidence: 0.9335446666666667

 $00:29:19.040 \longrightarrow 00:29:20.140$ But after week five,

NOTE Confidence: 0.9335446666666667

 $00:29:20.140 \longrightarrow 00:29:21.240$ if they stop drinking,

NOTE Confidence: 0.9335446666666667

 $00:29:21.240 \longrightarrow 00:29:22.758$ then their craving is also going

NOTE Confidence: 0.9335446666666667

00:29:22.758 --> 00:29:24.240 down because they stop drinking.

NOTE Confidence: 0.9335446666666667

 $00:29:24.240 \longrightarrow 00:29:28.720$ So it it's it's not a very strong causal

NOTE Confidence: 0.9335446666666667

 $00:29:28.720 \rightarrow 00:29:30.239$ even suggestion that we can make here,

NOTE Confidence: 0.9335446666666667

00:29:30.240 --> 00:29:31.200 but it's, you know,

NOTE Confidence: 0.9335446666666667

00:29:31.200 --> 00:29:32.188 it's a possibility, right?

NOTE Confidence: 0.9335446666666667

00:29:32.188 --> 00:29:33.528 If we don't know the

NOTE Confidence: 0.9335446666666667

 $00:29:33.528 \rightarrow 00:29:34.600$ direction of the causality,

NOTE Confidence: 0.941930833333333

 $00{:}29{:}37{.}200 \dashrightarrow 00{:}29{:}41{.}196$ this is something kind of like.

NOTE Confidence: 0.941930833333333

 $00:29:41.200 \rightarrow 00:29:44.920$ Craving it's temptation from the Alcohol

00:29:44.920 --> 00:29:46.360 Abstinent Selfefficacy questionnaire.

NOTE Confidence: 0.941930833333333

 $00:29:46.360 \rightarrow 00:29:49.240$ So this is hypothetical craving really.

NOTE Confidence: 0.941930833333333

00:29:49.240 --> 00:29:51.416 If you were you know in a walking

NOTE Confidence: 0.941930833333333

 $00:29:51.416 \longrightarrow 00:29:53.380$ down the street and you passed

NOTE Confidence: 0.941930833333333

 $00:29:53.380 \rightarrow 00:29:55.480$ your favorite bar and your buddy

NOTE Confidence: 0.941930833333333

 $00:29:55.480 \longrightarrow 00:29:57.836$ said come on in don't be a square,

NOTE Confidence: 0.941930833333333

 $00:29:57.840 \longrightarrow 00:30:00.836$ how tempted would you be to drink

NOTE Confidence: 0.941930833333333

 $00:30:00.840 \longrightarrow 00:30:03.144$ and and you rate a bunch of 20

NOTE Confidence: 0.941930833333333

 $00{:}30{:}03{.}144 \dashrightarrow 00{:}30{:}07{.}110$ different things like that and so.

NOTE Confidence: 0.941930833333333

 $00{:}30{:}07{.}110 \dashrightarrow 00{:}30{:}09{.}182$ We see in this case the separation

NOTE Confidence: 0.941930833333333

00:30:09.182 --> 00:30:10.881 didn't really happen until after

NOTE Confidence: 0.941930833333333

 $00:30:10.881 \rightarrow 00:30:12.389$ the second psilocybin session,

NOTE Confidence: 0.941930833333333

 $00:30:12.390 \longrightarrow 00:30:13.314$ which is you know,

NOTE Confidence: 0.941930833333333

 $00{:}30{:}13{.}314 \dashrightarrow 00{:}30{:}15{.}934$ so that we saw that in a couple of these

NOTE Confidence: 0.941930833333333

 $00{:}30{:}15{.}934 \dashrightarrow 00{:}30{:}18{.}630$ measures and then it is maintained.

 $00{:}30{:}18.630 \dashrightarrow 00{:}30{:}22.650$ So in this case we we do have a treatment

NOTE Confidence: 0.941930833333333

 $00{:}30{:}22.650 \dashrightarrow 00{:}30{:}24.832$ by assessment interaction and yeah,

NOTE Confidence: 0.941930833333333

 $00{:}30{:}24.832 \dashrightarrow 00{:}30{:}26.837$ I don't know about that.

NOTE Confidence: 0.941930833333333

00:30:26.840 --> 00:30:28.884 Delay, but it it does look different

NOTE Confidence: 0.941930833333333

 $00{:}30{:}28{.}884 \dashrightarrow 00{:}30{:}30{.}839$ from some of the other curves.

NOTE Confidence: 0.956354982

 $00{:}30{:}30{.}960 \dashrightarrow 00{:}30{:}33{.}120$ How much does this measure correlate

NOTE Confidence: 0.956354982

 $00:30:33.120 \longrightarrow 00:30:34.560$ with other clinical outcomes?

NOTE Confidence: 0.956354982

 $00:30:34.560 \rightarrow 00:30:36.560$ Just it seems like a much more cognitive,

NOTE Confidence: 0.956354982

00:30:36.560 --> 00:30:42.760 you know, measure than the behavioral or I

NOTE Confidence: 0.942083229090909

 $00:30:42.760 \longrightarrow 00:30:44.158$ think it's, I mean people often

NOTE Confidence: 0.942083229090909

 $00:30:44.158 \longrightarrow 00:30:45.520$ call it a craving measure.

NOTE Confidence: 0.942083229090909

00:30:45.520 --> 00:30:46.765 I mean I think and you know,

NOTE Confidence: 0.942083229090909

00:30:46.765 --> 00:30:48.705 craving is a reasonable

NOTE Confidence: 0.942083229090909

00:30:48.705 --> 00:30:50.800 predictor of of outcome.

NOTE Confidence: 0.942083229090909

00:30:50.800 - 00:30:53.600 I mean it's fairly consistent,

NOTE Confidence: 0.942083229090909

 $00:30:53.600 \rightarrow 00:30:56.860$ you know, moderately predictive.

- NOTE Confidence: 0.942083229090909
- 00:30:56.860 --> 00:30:58.320 Okay selfefficacy is also
- NOTE Confidence: 0.942083229090909
- $00:30:58.320 \longrightarrow 00:30:59.780$ a pretty good predictor.
- NOTE Confidence: 0.942083229090909
- 00:30:59.780 --> 00:31:01.467 I mean in general if you're doing
- NOTE Confidence: 0.942083229090909
- 00:31:01.467 --> 00:31:02.580 just treatment outcome studies,
- NOTE Confidence: 0.942083229090909
- 00:31:02.580 --> 00:31:04.806 if people say I'm, I am confident
- NOTE Confidence: 0.942083229090909
- $00:31:04.806 \rightarrow 00:31:07.577$ I can stop that it means something.
- NOTE Confidence: 0.942083229090909
- $00:31:07.580 \rightarrow 00:31:14.328$ And so you can see here there was yeah,
- NOTE Confidence: 0.942083229090909
- $00:31:14.328 \longrightarrow 00:31:16.912$ this is 1 where there's a there
- NOTE Confidence: 0.942083229090909
- $00:31:16.912 \longrightarrow 00:31:18.172$ isn't a treatment by assessment
- NOTE Confidence: 0.942083229090909
- $00:31:18.172 \rightarrow 00:31:19.421$ interaction that's going to sound
- NOTE Confidence: 0.942083229090909
- $00:31:19.421 \longrightarrow 00:31:20.615$ surprising if you look at the.
- NOTE Confidence: 0.940253533333333
- 00:31:22.880 --> 00:31:24.266 But you're not, well you're not
- NOTE Confidence: 0.940253533333333
- $00:31:24.266 \rightarrow 00:31:25.314$ including the very baseline,
- NOTE Confidence: 0.940253533333333
- $00{:}31{:}25{.}320 \dashrightarrow 00{:}31{:}26{.}762$ yeah, but it's it still looks like
- NOTE Confidence: 0.940253533333333
- $00:31:26.762 \longrightarrow 00:31:27.961$ the slopes would be different
- NOTE Confidence: 0.940253533333333

 $00:31:27.961 \longrightarrow 00:31:29.037$ but apparently they're not.

NOTE Confidence: 0.940253533333333

 $00{:}31{:}29{.}040 \dashrightarrow 00{:}31{:}31{.}392$ But there is this you know pretty

NOTE Confidence: 0.940253533333333

 $00:31:31.392 \rightarrow 00:31:33.224$ strong treatment effect and it's

NOTE Confidence: 0.940253533333333

 $00:31:33.224 \rightarrow 00:31:36.144$ it's persists overtime and so

NOTE Confidence: 0.940253533333333

 $00:31:36.144 \longrightarrow 00:31:39.335$ that's that's good and that is

NOTE Confidence: 0.940253533333333

00:31:39.335 --> 00:31:41.345 something that tends to predict good NOTE Confidence: 0.940253533333333

 $00{:}31{:}41{.}345 \dashrightarrow 00{:}31{:}43{.}564$ outcomes and then self compassion

NOTE Confidence: 0.9402535333333333

 $00:31:43.564 \rightarrow 00:31:46.262$ is kind of a complicated construct,

NOTE Confidence: 0.940253533333333

 $00{:}31{:}46.262 \dashrightarrow 00{:}31{:}48.656$ but I think it's it's highly relevant.

NOTE Confidence: 0.940253533333333

00:31:48.660 --> 00:31:49.260 I mean, for one thing,

NOTE Confidence: 0.940253533333333

 $00{:}31{:}49{.}260 \dashrightarrow 00{:}31{:}50{.}856$ the effects are really this is I

NOTE Confidence: 0.940253533333333

 $00{:}31{:}50.856 \dashrightarrow 00{:}31{:}52.317$ think the largest effect that we

NOTE Confidence: 0.940253533333333

 $00{:}31{:}52{.}317 \dashrightarrow 00{:}31{:}53{.}775$ saw in any of the question naires.

NOTE Confidence: 0.941691338571429

 $00:31:55.900 \rightarrow 00:31:58.616$ And there are two subscales to this,

NOTE Confidence: 0.941691338571429

 $00:31:58.620 \longrightarrow 00:32:02.624$ and one of them is sort of the

NOTE Confidence: 0.941691338571429

 $00:32:02.624 \rightarrow 00:32:03.696$ opposite of self compassion.

 $00{:}32{:}03.700 \dashrightarrow 00{:}32{:}05.140$ It's it's self criticism.

NOTE Confidence: 0.941691338571429

 $00:32:05.140 \longrightarrow 00:32:07.762$ It's the extent to which you, you know,

NOTE Confidence: 0.941691338571429

 $00:32:07.762 \longrightarrow 00:32:09.379$ are always like beating up on yourself.

NOTE Confidence: 0.941691338571429

00:32:09.380 --> 00:32:10.884 Or if you make a mistake you say

NOTE Confidence: 0.941691338571429

00:32:10.884 --> 00:32:12.140 I'm stupid, I always mess up.

NOTE Confidence: 0.941691338571429

00:32:12.140 --> 00:32:13.020 I'll never do right.

NOTE Confidence: 0.915843438571429

 $00:32:15.700 \longrightarrow 00:32:17.488$ You know that kind of negative

NOTE Confidence: 0.915843438571429

 $00:32:17.488 \longrightarrow 00:32:19.739$ self talk is a big part of it.

NOTE Confidence: 0.915843438571429

 $00:32:19.740 \longrightarrow 00:32:22.218$ And the other half is self compassion,

NOTE Confidence: 0.915843438571429

 $00:32:22.220 \longrightarrow 00:32:23.258$ which is kind of the opposite.

NOTE Confidence: 0.915843438571429

00:32:23.260 --> 00:32:24.260 It's like saying, you know,

NOTE Confidence: 0.915843438571429

 $00{:}32{:}24{.}260 \dashrightarrow 00{:}32{:}27{.}180$ well put it in perspective,

NOTE Confidence: 0.915843438571429

 $00:32:27.180 \longrightarrow 00:32:29.148$ you know you're you're not so bad and

NOTE Confidence: 0.915843438571429

00:32:29.148 --> 00:32:30.779 you're doing your best or whatever.

NOTE Confidence: 0.915843438571429

 $00{:}32{:}30{.}780 \dashrightarrow 00{:}32{:}33{.}052$ You know you have good qualities or

00:32:33.052 --> 00:32:34.588 you know it's okay to make a mistake

NOTE Confidence: 0.915843438571429

 $00:32:34.588 \dashrightarrow 00:32:36.335$ or what those those sorts of things.

NOTE Confidence: 0.915843438571429

 $00:32:36.340 \rightarrow 00:32:39.946$ So it's it's cognitive and effective.

NOTE Confidence: 0.915843438571429

00:32:39.950 --> 00:32:41.770 You know I don't know you know

NOTE Confidence: 0.915843438571429

 $00:32:41.770 \longrightarrow 00:32:43.332$ so if we want to you know in

NOTE Confidence: 0.915843438571429

 $00:32:43.332 \longrightarrow 00:32:44.828$ terms of the three domains,

NOTE Confidence: 0.915843438571429

 $00:32:44.830 \dashrightarrow 00:32:47.901$ is this executive control or or you

NOTE Confidence: 0.915843438571429

 $00:32:47.901 \rightarrow 00:32:49.798$ know negative or you know less negative

NOTE Confidence: 0.915843438571429

 $00{:}32{:}49{.}798 \dashrightarrow 00{:}32{:}51{.}548$ affect it's probably both but I it

NOTE Confidence: 0.915843438571429

 $00:32:51.548 \rightarrow 00:32:53.550$ doesn't really map on to those that well.

NOTE Confidence: 0.915843438571429

00:32:53.550 --> 00:32:55.894 But I think it is it is interesting

NOTE Confidence: 0.915843438571429

 $00:32:55.894 \rightarrow 00:32:58.238$ in in capturing something that

NOTE Confidence: 0.915843438571429

 $00:32:58.238 \rightarrow 00:32:59.878$ changes you know both cognitively

NOTE Confidence: 0.915843438571429

 $00:32:59.878 \longrightarrow 00:33:01.190$ and effectively for people.

NOTE Confidence: 0.916288475

 $00:33:03.830 \dashrightarrow 00:33:05.720$ So that's interesting and we you know

NOTE Confidence: 0.916288475

 $00:33:05.720 \rightarrow 00:33:07.965$ we're working on a paper where I mean it

- NOTE Confidence: 0.916288475
- $00:33:07.965 \rightarrow 00:33:09.890$ looks like they're you know there is some.

 $00{:}33{:}09{.}890 \dashrightarrow 00{:}33{:}13{.}122$ It it does predict some amount of the

NOTE Confidence: 0.916288475

 $00:33:13.122 \rightarrow 00:33:15.090$ the the drinking outcome variance

NOTE Confidence: 0.94427896

00:33:16.050 - 00:33:17.210 that is the biggest factor.

NOTE Confidence: 0.93412381111111

 $00:33:17.930 \longrightarrow 00:33:19.040$ The self compassion is the

NOTE Confidence: 0.93412381111111

 $00:33:19.040 \longrightarrow 00:33:19.928$ biggest of the individual.

NOTE Confidence: 0.93412381111111

 $00:33:19.930 \longrightarrow 00:33:20.958$ Well that's the biggest

NOTE Confidence: 0.93412381111111

 $00:33:20.958 \rightarrow 00:33:22.243$ effect just in between group.

NOTE Confidence: 0.93412381111111

00:33:22.250 --> 00:33:23.670 Yeah group effect in terms

NOTE Confidence: 0.93412381111111

 $00:33:23.670 \longrightarrow 00:33:25.090$ of like the self report,

NOTE Confidence: 0.93412381111111

00:33:25.090 - 00:33:27.310 the effects of psilocybin on

NOTE Confidence: 0.93412381111111

 $00{:}33{:}27{.}310 \dashrightarrow 00{:}33{:}29{.}290$ the self report itself. Have

NOTE Confidence: 0.9335446666666667

 $00:33:29.290 \rightarrow 00:33:31.920$ you done any multivariate analysis

NOTE Confidence: 0.9335446666666667

00:33:31.920 --> 00:33:34.125 across these like you could do a?

NOTE Confidence: 0.94226628

 $00:33:38.340 \longrightarrow 00:33:39.260$ I'm blanking on the analysis

00:33:39.260 --> 00:33:40.180 I thought you could do,

NOTE Confidence: 0.94226628

 $00:33:40.180 \longrightarrow 00:33:42.378$ but something that looks at the relationship

NOTE Confidence: 0.94226628

 $00:33:42.378 \rightarrow 00:33:44.339$ between these two measures over time,

NOTE Confidence: 0.94226628

 $00{:}33{:}44{.}340 \dashrightarrow 00{:}33{:}47{.}956$ because the curves as you've pointed as

NOTE Confidence: 0.94226628

 $00{:}33{:}47.956 \dashrightarrow 00{:}33{:}50.058$ we've gone through the curves and the

NOTE Confidence: 0.94226628

 $00{:}33{:}50{.}058 \dashrightarrow 00{:}33{:}52{.}252$ point at which they separate are the tank.

NOTE Confidence: 0.94226628

00:33:52.252 --> 00:33:53.020 They're not identical,

NOTE Confidence: 0.94226628

 $00:33:53.020 \rightarrow 00:33:54.098$ which could be noise in the data,

NOTE Confidence: 0.94226628

 $00{:}33{:}54.100 \dashrightarrow 00{:}33{:}56.704$ but you might wonder if you could

NOTE Confidence: 0.94226628

 $00{:}33{:}56{.}704 \dashrightarrow 00{:}33{:}59{.}059$ make some inferences across about the.

NOTE Confidence: 0.94226628

 $00{:}33{:}59{.}060 \dashrightarrow 00{:}34{:}00{.}370$ You know, the unfolding of

NOTE Confidence: 0.94226628

 $00:34:00.370 \longrightarrow 00:34:01.900$ different phases of the effect of

NOTE Confidence: 0.931966094117647

 $00{:}34{:}01{.}900 \dashrightarrow 00{:}34{:}03{.}826$ no, I would love to and I mean that

NOTE Confidence: 0.931966094117647

 $00:34:03.826 \rightarrow 00:34:05.896$ is something we can do and you know

NOTE Confidence: 0.931966094117647

 $00:34:05.900 \longrightarrow 00:34:07.124$ the drinking outcomes we have in

NOTE Confidence: 0.931966094117647

 $00:34:07.124 \rightarrow 00:34:08.577$ these one month, one month bins.

 $00{:}34{:}08{.}577 \dashrightarrow 00{:}34{:}11{.}687$ So we can we can tease that out and we could

NOTE Confidence: 0.931966094117647

 $00{:}34{:}11.687 \dashrightarrow 00{:}34{:}14.262$ do cross like kind of analysis like that.

NOTE Confidence: 0.931966094117647

 $00:34:14.262 \longrightarrow 00:34:15.588$ I mean the problem is there's

NOTE Confidence: 0.931966094117647

 $00:34:15.588 \longrightarrow 00:34:16.857$ just so many we could do,

NOTE Confidence: 0.931966094117647

 $00{:}34{:}16{.}860 \dashrightarrow 00{:}34{:}19{.}292$ you know so you don't want to take

NOTE Confidence: 0.931966094117647

 $00:34:19.292 \dashrightarrow 00:34:21.957$ too many bites of the apple here but.

NOTE Confidence: 0.931966094117647

 $00:34:21.960 \longrightarrow 00:34:23.358$ And it's not a huge sample,

NOTE Confidence: 0.931966094117647

 $00:34:23.360 \rightarrow 00:34:25.480$ so I don't want to over analyze this,

NOTE Confidence: 0.931966094117647

 $00:34:25.480 \longrightarrow 00:34:26.944$ but I think you know those

NOTE Confidence: 0.931966094117647

 $00:34:26.944 \rightarrow 00:34:27.920$ are really good questions.

NOTE Confidence: 0.931966094117647

 $00:34:27.920 \dashrightarrow 00:34:31.560$ And another thing that we will do

NOTE Confidence: 0.931966094117647

00:34:31.560 - 00:34:36.360 is try to do some machine learning

NOTE Confidence: 0.931966094117647

00:34:36.360 --> 00:34:38.760 approach to try to identify likely

NOTE Confidence: 0.931966094117647

 $00{:}34{:}38{.}760 \dashrightarrow 00{:}34{:}41{.}110$ responders and we can do that

NOTE Confidence: 0.931966094117647

 $00{:}34{:}41{.}110 \dashrightarrow 00{:}34{:}42{.}594$ based on baseline characteristics.

 $00:34:42.600 \longrightarrow 00:34:45.936$ We could also do that based on the

NOTE Confidence: 0.931966094117647

 $00{:}34{:}45{.}936 \dashrightarrow 00{:}34{:}48{.}577$ subjective effects or or you know week 5.

NOTE Confidence: 0.931966094117647

00:34:48.580 --> 00:34:50.748 Outcomes you know week five values of of

NOTE Confidence: 0.931966094117647

 $00:34:50.748 \rightarrow 00:34:53.099$ some of these self-reports or you know,

NOTE Confidence: 0.931966094117647

 $00{:}34{:}53{.}100 \dashrightarrow 00{:}34{:}55{.}164$ so you can include you look at the

NOTE Confidence: 0.931966094117647

 $00:34:55.164 \rightarrow 00:34:56.632$ pretreatment things but then you could NOTE Confidence: 0.931966094117647

 $00:34:56.632 \rightarrow 00:34:58.365$ also see you know if there's something

NOTE Confidence: 0.931966094117647

00:34:58.365 - 00:35:00.172 that something in the immediate

NOTE Confidence: 0.931966094117647

 $00{:}35{:}00{.}172 \dashrightarrow 00{:}35{:}02{.}380$ response to treatment that will predict

NOTE Confidence: 0.931966094117647

00:35:02.441 --> 00:35:04.336 the longer term drinking outcomes.

NOTE Confidence: 0.925158445

 $00{:}35{:}06{.}900 \dashrightarrow 00{:}35{:}12{.}420$ So how about how about the nature of the.

NOTE Confidence: 0.925158445

 $00:35:12.420 \longrightarrow 00:35:13.646$ Intoxication effect. Yeah.

NOTE Confidence: 0.925158445

 $00:35:13.646 \longrightarrow 00:35:14.776$ So we'll get to that.

NOTE Confidence: 0.925158445

 $00{:}35{:}14.780 \dashrightarrow 00{:}35{:}16.327$ That's So that's the other that's the

NOTE Confidence: 0.925158445

 $00:35:16.327 \dashrightarrow 00:35:18.017$ new piece that I want to show you too.

NOTE Confidence: 0.925158445

 $00:35:18.020 \rightarrow 00:35:20.218$ We have taken some look at that.

00:35:20.220 --> 00:35:24.140 So this is, you know it's well

NOTE Confidence: 0.925158445

 $00:35:24.140 \longrightarrow 00:35:27.085$ known that psilocybin tends to

NOTE Confidence: 0.925158445

 $00:35:27.085 \rightarrow 00:35:28.695$ produce some personality changes.

NOTE Confidence: 0.925158445

 $00:35:28.695 \rightarrow 00:35:31.110$ That increase in openness is the one

NOTE Confidence: 0.925158445

 $00{:}35{:}31{.}174$ --> $00{:}35{:}33{.}412$ that's been demonstrated the most times

NOTE Confidence: 0.925158445

 $00{:}35{:}33{.}412 \dashrightarrow 00{:}35{.}35{.}578$ decrease in neuroticism was that it was

NOTE Confidence: 0.925158445

 $00:35:35.580 \dashrightarrow 00:35:37.818$ had been demonstrated before as well.

NOTE Confidence: 0.925158445

 $00:35:37.820 \longrightarrow 00:35:39.938$ So we saw.

NOTE Confidence: 0.925158445

 $00:35:39.940 \longrightarrow 00:35:41.840$ Yeah between group differences in

NOTE Confidence: 0.925158445

 $00:35:41.840 \longrightarrow 00:35:44.329$ and in change in that neuroticism

NOTE Confidence: 0.925158445

00:35:44.329 - > 00:35:46.377 extraversion and openness and

NOTE Confidence: 0.925158445

 $00{:}35{:}46{.}377 \dashrightarrow 00{:}35{:}48{.}937$ and within the psilocybin group,

NOTE Confidence: 0.925158445

 $00{:}35{:}48{.}940 \dashrightarrow 00{:}35{:}50{.}700$ conscient iousness also increased but you

NOTE Confidence: 0.925158445

 $00:35:50.700 \rightarrow 00:35:54.097$ can see it did in the control group as well.

NOTE Confidence: 0.925158445

00:35:54.100 --> 00:35:56.916 And you know I think I,

 $00:35:56.916 \rightarrow 00:36:00.244$ I don't want to over reify these personality

NOTE Confidence: 0.925158445

 $00:36:00.244 \rightarrow 00:36:02.196$ dimensions either because you know,

NOTE Confidence: 0.925158445

 $00{:}36{:}02.196 \dashrightarrow 00{:}36{:}04.020$ I mean we we like to think about

NOTE Confidence: 0.925158445

 $00{:}36{:}04.074 \dashrightarrow 00{:}36{:}05.398$ them as fixed characteristics

NOTE Confidence: 0.925158445

00:36:05.398 --> 00:36:07.053 but if you stop drinking,

NOTE Confidence: 0.925158445

 $00{:}36{:}07{.}060 \dashrightarrow 00{:}36{:}08{.}758$ you know you're going to act

NOTE Confidence: 0.925158445

 $00:36:08.758 \rightarrow 00:36:10.100$ different and you're going to.

NOTE Confidence: 0.925158445

00:36:10.100 --> 00:36:10.700 You know,

NOTE Confidence: 0.925158445

00:36:10.700 --> 00:36:13.100 I was taught in in my clinical training,

NOTE Confidence: 0.925158445

00:36:13.100 --> 00:36:13.846 you know,

NOTE Confidence: 0.925158445

00:36:13.846 --> 00:36:16.315 don't ever try to make a personality

NOTE Confidence: 0.925158445

 $00:36:16.315 \rightarrow 00:36:17.615$ disorder diagnosis somebody who's

NOTE Confidence: 0.925158445

00:36:17.615 --> 00:36:18.907 actively addicted to something

NOTE Confidence: 0.925158445

00:36:18.907 - 00:36:20.485 because they're going to act like

NOTE Confidence: 0.925158445

 $00{:}36{:}20.485 \dashrightarrow 00{:}36{:}22.260$ some body with a personality disorder.

NOTE Confidence: 0.925158445

00:36:22.260 --> 00:36:24.130 And I think, you know, there's some,

- NOTE Confidence: 0.925158445
- $00:36:24.130 \longrightarrow 00:36:25.780$ there is some truth to that.

00:36:25.780 --> 00:36:27.372 But but you know,

NOTE Confidence: 0.925158445

 $00:36:27.372 \rightarrow 00:36:29.362$ these were pretty robust differences

NOTE Confidence: 0.925158445

 $00{:}36{:}29{.}362 \dashrightarrow 00{:}36{:}31{.}819$ and it's particularly interesting

NOTE Confidence: 0.925158445

 $00:36:31.820 \longrightarrow 00:36:34.564$ if you look at the facets that

NOTE Confidence: 0.925158445

 $00:36:34.564 \rightarrow 00:36:36.247$ actually changed because there's

NOTE Confidence: 0.925158445

 $00{:}36{:}36{.}247 \dashrightarrow 00{:}36{:}38{.}387$ a lot inside of neuroticism.

NOTE Confidence: 0.925158445

 $00:36:38.390 \rightarrow 00:36:40.945$ And what actually changed significantly

NOTE Confidence: 0.925158445

 $00:36:40.945 \longrightarrow 00:36:42.989$ within the psilocybin group,

NOTE Confidence: 0.925158445

00:36:42.990 --> 00:36:44.330 less depression,

NOTE Confidence: 0.925158445

 $00:36:44.330 \longrightarrow 00:36:47.010$ less impulsiveness and less

NOTE Confidence: 0.925158445

00:36:47.010 -> 00:36:48.350 emotional vulnerability.

NOTE Confidence: 0.925158445

00:36:48.350 - 00:36:49.370 And the extraversion,

NOTE Confidence: 0.925158445

 $00{:}36{:}49{.}370 \dashrightarrow 00{:}36{:}51{.}410$ it wasn't that they wanted to

NOTE Confidence: 0.925158445

 $00:36:51.410 \longrightarrow 00:36:53.586$ go out and party but not drink.

- $00{:}36{:}53{.}590 \dashrightarrow 00{:}36{:}56{.}264$ It was actually increase in positive emotion.
- NOTE Confidence: 0.925158445
- 00:36:56.270 --> 00:36:56.900 So, you know,
- NOTE Confidence: 0.925158445
- 00:36:56.900 00:36:58.961 I don't know why that doesn't end up in
- NOTE Confidence: 0.925158445
- 00:36:58.961 --> 00:37:00.470 neuroticism, but it's it's, you know,
- NOTE Confidence: 0.925158445
- $00:37:00.470 \longrightarrow 00:37:03.949$ it's more of a positive affect thing.
- NOTE Confidence: 0.925158445
- 00:37:03.950 --> 00:37:05.300 And in openness,
- NOTE Confidence: 0.925158445
- 00:37:05.300 00:37:07.550 it was increased openness to,
- NOTE Confidence: 0.925158445
- 00:37:07.550 --> 00:37:09.860 you know, fantasies,
- NOTE Confidence: 0.925158445
- $00:37:09.860 \dashrightarrow 00:37:12.998$ internal thoughts and and feelings.
- NOTE Confidence: 0.925158445
- 00:37:12.998 --> 00:37:14.910 So, you know, I didn't, you know,
- NOTE Confidence: 0.925158445
- $00:37:14.910 \longrightarrow 00:37:16.830$ when we talk about cognitive flexibility,
- NOTE Confidence: 0.925158445
- $00{:}37{:}16.830 \dashrightarrow 00{:}37{:}17.494$ this could be one.
- NOTE Confidence: 0.925158445
- 00:37:17.494 --> 00:37:18.778 I mean that can mean a lot
- NOTE Confidence: 0.925158445
- $00:37:18.778 \longrightarrow 00:37:19.507$ of different things.
- NOTE Confidence: 0.925158445
- $00:37:19.510 \rightarrow 00:37:22.366$ But this is one kind of flexibility perhaps
- NOTE Confidence: 0.925158445
- $00:37:22.366 \rightarrow 00:37:25.470$ that that we're saying that could be useful.

- NOTE Confidence: 0.925158445
- $00:37:25.470 \longrightarrow 00:37:27.038$ And then increased deliberation.
- NOTE Confidence: 0.925158445
- 00:37:27.038 --> 00:37:28.480 So, you know,
- NOTE Confidence: 0.925158445
- $00{:}37{:}28{.}480 \dashrightarrow 00{:}37{:}30{.}055$ decreased impulsiveness and
- NOTE Confidence: 0.925158445
- 00:37:30.055 --> 00:37:31.480 increased deliberation, you know,
- NOTE Confidence: 0.925158445
- $00:37:31.480 \longrightarrow 00:37:32.590$ does this mean they've they had,
- NOTE Confidence: 0.925158445
- $00:37:32.590 \longrightarrow 00:37:35.299$ they have better?
- NOTE Confidence: 0.925158445
- 00:37:35.300 --> 00:37:37.076 Inhibitory control perhaps at
- NOTE Confidence: 0.925158445
- 00:37:37.076 > 00:37:39.296 least that's what they think.
- NOTE Confidence: 0.925158445
- 00:37:39.300 --> 00:37:39.780 So,
- NOTE Confidence: 0.939723947368421
- $00:37:42.060 \rightarrow 00:37:43.845$ so that's suggestive but don't
- NOTE Confidence: 0.939723947368421
- $00:37:43.845 \longrightarrow 00:37:46.758$ want to take it too far but they
- NOTE Confidence: 0.939723947368421
- 00:37:46.758 --> 00:37:48.852 are you know good size changes.
- NOTE Confidence: 0.939723947368421
- $00:37:48.860 \longrightarrow 00:37:50.306$ So, so here's the really the
- NOTE Confidence: 0.939723947368421
- $00{:}37{:}50{.}306 \dashrightarrow 00{:}37{:}51{.}850$ really new part that these next
- NOTE Confidence: 0.939723947368421
- $00{:}37{:}51.850 \dashrightarrow 00{:}37{:}53.416$ two things I have not presented
- NOTE Confidence: 0.939723947368421

 $00{:}37{:}53.416 \dashrightarrow 00{:}37{:}55.088$ before and be interested to see

NOTE Confidence: 0.939723947368421

 $00{:}37{:}55{.}088 \dashrightarrow 00{:}37{:}56{.}458$ what people think about them.

NOTE Confidence: 0.939723947368421

 $00:37:56.460 \longrightarrow 00:37:59.910$ So we had four, how are we doing for time.

NOTE Confidence: 0.939723947368421

00:37:59.910 --> 00:38:00.230 All right,

NOTE Confidence: 0.9285129666666667

 $00{:}38{:}00{.}550 \dashrightarrow 00{:}38{:}04{.}330$ time is at the time is. It's a construct.

NOTE Confidence: 0.9285129666666667

 $00:38:04.330 \longrightarrow 00:38:06.682$ I don't want to over reapply the

NOTE Confidence: 0.9285129666666667

 $00{:}38{:}06{.}682 \dashrightarrow 00{:}38{:}09{.}109$ construct and I shouldn't take it instead

NOTE Confidence: 0.9301902

 $00:38:09.110 \longrightarrow 00:38:09.990$ of the Uber, but

NOTE Confidence: 0.94226628

 $00{:}38{:}12{.}830 \dashrightarrow 00{:}38{:}17{.}430$ so we used four question naires

NOTE Confidence: 0.94226628

 $00:38:17.430 \longrightarrow 00:38:21.096$ after each of the psilocybin or.

NOTE Confidence: 0.94226628

00:38:21.100 --> 00:38:22.018 Diphen Hydramine sessions.

NOTE Confidence: 0.94226628

 $00:38:22.018 \longrightarrow 00:38:23.854$ And I'll tell you more about

NOTE Confidence: 0.94226628

 $00:38:23.854 \rightarrow 00:38:25.180$ what those were shortly.

NOTE Confidence: 0.94226628

 $00{:}38{:}25{.}180 \dashrightarrow 00{:}38{:}26{.}998$ There are other questionnaires that are

NOTE Confidence: 0.94226628

 $00{:}38{:}26{.}998 \dashrightarrow 00{:}38{:}29{.}046$ now available that we didn't have when

NOTE Confidence: 0.94226628

00:38:29.046 --> 00:38:30.859 I wrote this protocol and you know,

- NOTE Confidence: 0.94226628
- $00{:}38{:}30{.}860 \dashrightarrow 00{:}38{:}32{.}216$ I kind of wish we did.
- NOTE Confidence: 0.94226628
- $00:38:32.220 \longrightarrow 00:38:35.864$ Like the, there's one called the
- NOTE Confidence: 0.94226628
- 00:38:35.864 --> 00:38:37.217 Emotional Breakthrough Inventory.
- NOTE Confidence: 0.94226628
- $00:38:37.220 \rightarrow 00:38:39.908$ There's one called the that specifically
- NOTE Confidence: 0.94226628
- $00:38:39.908 \rightarrow 00:38:44.460$ gets at the challenging experiences.
- NOTE Confidence: 0.94226628
- 00:38:44.460 --> 00:38:45.066 And you know,
- NOTE Confidence: 0.94226628
- $00:38:45.066 \rightarrow 00:38:48.960$ I think there's even a couple more now, but.
- NOTE Confidence: 0.94226628
- $00{:}38{:}48{.}960 \dashrightarrow 00{:}38{:}52{.}537$ And I decided that it it's a
- NOTE Confidence: 0.94226628
- $00{:}38{:}52{.}537 \dashrightarrow 00{:}38{:}54{.}490$ little bit complicated because not
- NOTE Confidence: 0.94226628
- $00:38:54.490 \rightarrow 00:38:56.040$ everyone gets the second session.
- NOTE Confidence: 0.94226628
- 00:38:56.040 --> 00:38:56.931 Most people did,
- NOTE Confidence: 0.94226628
- $00{:}38{:}56{.}931 \dashrightarrow 00{:}38{:}58{.}713$ but there are people who didn't
- NOTE Confidence: 0.94226628
- $00:38:58.720 \rightarrow 00:39:03.360$ and because they didn't want to.
- NOTE Confidence: 0.94226628
- 00:39:03.360 --> 00:39:05.633 I mean, a couple people dropped out, man.
- NOTE Confidence: 0.94226628
- $00{:}39{:}05{.}633 \dashrightarrow 00{:}39{:}07{.}198$ Actually nobody really dropped out.
- NOTE Confidence: 0.94226628

- 00:39:07.200 --> 00:39:07.760 They continued
- NOTE Confidence: 0.96641844
- $00{:}39{:}07{.}880 \dashrightarrow 00{:}39{:}09{.}080$ psychotherapy, but they
- NOTE Confidence: 0.96641844
- $00:39:09.080 \longrightarrow 00:39:09.880$ declined psychotherapy
- NOTE Confidence: 0.87031739625
- 00:39:09.880 --> 00:39:12.280 in the follow up, but just said,
- NOTE Confidence: 0.87031739625
- $00{:}39{:}12{.}280 \dashrightarrow 00{:}39{:}13{.}636$ you know, that was too much
- NOTE Confidence: 0.87031739625
- $00:39:13.636 \longrightarrow 00:39:15.079$ for me or boring for me.
- NOTE Confidence: 0.87031739625
- $00{:}39{:}15{.}079 \dashrightarrow 00{:}39{:}17{.}190$ So and we we wanted to be
- NOTE Confidence: 0.87031739625
- $00:39:17.190 \longrightarrow 00:39:19.460$ able to let people do that. So
- NOTE Confidence: 0.9452853
- $00{:}39{:}20{.}340 \dashrightarrow 00{:}39{:}21{.}912$ and the rent of course people
- NOTE Confidence: 0.9452853
- $00{:}39{:}21{.}912 \dashrightarrow 00{:}39{:}23{.}347$ getting the same drug diphenhydromin
- NOTE Confidence: 0.9452853
- $00:39:23.347 \longrightarrow 00:39:24.977$ or 07 in both sessions.
- NOTE Confidence: 0.9452853
- $00:39:24.980 \longrightarrow 00:39:27.780$ So if they are pretty sure they
- NOTE Confidence: 0.9452853
- $00{:}39{:}27.780 \dashrightarrow 00{:}39{:}29.424$ got place bo the first time they're
- NOTE Confidence: 0.9452853
- $00:39:29.424 \rightarrow 00:39:33.540$ going to then they know yeah so
- NOTE Confidence: 0.93514248
- $00:39:33.540 \longrightarrow 00:39:36.012$ so there wasn't a significantly different
- NOTE Confidence: 0.93514248
- $00:39:36.012 \rightarrow 00:39:39.174$ number of of dropouts in the in the but

 $00:39:39.174 \longrightarrow 00:39:41.245$ there were more people who declined

NOTE Confidence: 0.93514248

 $00{:}39{:}41.245 \dashrightarrow 00{:}39{:}43.497$ the second diphenhydromin session.

NOTE Confidence: 0.95031694375

00:39:44.550 -> 00:39:45.870 And you can imagine they declined

NOTE Confidence: 0.95031694375

 $00{:}39{:}45{.}870 \dashrightarrow 00{:}39{:}46{.}682$ for different reasons and

NOTE Confidence: 0.95031694375

 $00:39:46.682 \rightarrow 00:39:48.070$ they pouring versus too much,

NOTE Confidence: 0.938097092857143

00:39:49.270 --> 00:39:51.300 right. So, you know, so you can

NOTE Confidence: 0.938097092857143

 $00:39:51.300 \dashrightarrow 00:39:53.510$ either look just at the first session,

NOTE Confidence: 0.938097092857143

00:39:53.510 - 00:39:55.594 but then I think you know, you might

NOTE Confidence: 0.938097092857143

 $00:39:55.594 \dashrightarrow 00:39:58.142$ be missing the boat because it just,

NOTE Confidence: 0.938097092857143

 $00:39:58.142 \longrightarrow 00:39:59.638$ you know, clinically people

NOTE Confidence: 0.938097092857143

 $00{:}39{:}59{.}638 \dashrightarrow 00{:}40{:}01{.}202$ could have a terrible, you know,

NOTE Confidence: 0.938097092857143

 $00{:}40{:}01{.}202 \dashrightarrow 00{:}40{:}03{.}204$ a miserable experience one time in a.

NOTE Confidence: 0.938097092857143

 $00{:}40{:}03.210 \dashrightarrow 00{:}40{:}05.262$ Be a tivic experience the next time and and

NOTE Confidence: 0.938097092857143

 $00{:}40{:}05{.}262 \dashrightarrow 00{:}40{:}07{.}247$ and we you know we definitely saw that.

NOTE Confidence: 0.938097092857143

 $00:40:07.250 \longrightarrow 00:40:08.789$ So I, you know I think we'd lose a

 $00:40:08.789 \longrightarrow 00:40:10.369$ lot of information if we did that.

NOTE Confidence: 0.938097092857143

00:40:10.370 --> 00:40:12.962 So what I ended up doing was just

NOTE Confidence: 0.938097092857143

 $00:40:12.962 \rightarrow 00:40:15.888$ to add consider it sort of a dose,

NOTE Confidence: 0.938097092857143

 $00:40:15.890 \longrightarrow 00:40:17.370$ you know dose effect,

NOTE Confidence: 0.938097092857143

 $00{:}40{:}17.370 \dashrightarrow 00{:}40{:}20.506$ how much so how much of A particular

NOTE Confidence: 0.938097092857143

00:40:20.506 --> 00:40:22.197 kind of subjective experience did

NOTE Confidence: 0.938097092857143

 $00{:}40{:}22.197 \dashrightarrow 00{:}40{:}24.572$ you get and I mean I don't know

NOTE Confidence: 0.938097092857143

 $00:40:24.572 \longrightarrow 00:40:26.306$ if it's really additive or what

NOTE Confidence: 0.938097092857143

00:40:26.306 --> 00:40:27.570 but that's what I did.

NOTE Confidence: 0.938097092857143

 $00:40:27.570 \longrightarrow 00:40:29.978$ So if they didn't have a second

NOTE Confidence: 0.938097092857143

 $00{:}40{:}29{.}978 \dashrightarrow 00{:}40{:}32{.}419$ session they just get 0 for that one.

NOTE Confidence: 0.938097092857143

 $00:40:32.420 \longrightarrow 00:40:33.484$ And if they and and if they

NOTE Confidence: 0.938097092857143

 $00:40:33.484 \longrightarrow 00:40:34.580$ did have a second session,

NOTE Confidence: 0.938097092857143

 $00:40:34.580 \longrightarrow 00:40:36.380$ then the two are added together

NOTE Confidence: 0.938097092857143

 $00{:}40{:}36{.}380 \dashrightarrow 00{:}40{:}38{.}515$ and then there's a lot of different

NOTE Confidence: 0.938097092857143

00:40:38.515 --> 00:40:39.704 drinking outcomes you could

- NOTE Confidence: 0.938097092857143
- $00:40:39.704 \rightarrow 00:40:41.416$ look at and a lot of you know,
- NOTE Confidence: 0.938097092857143
- $00{:}40{:}41{.}420 \dashrightarrow 00{:}40{.}44{.}096$ other outcomes you could look at.
- NOTE Confidence: 0.938097092857143
- $00:40:44.100 \longrightarrow 00:40:46.540$ So to keep it simple,
- NOTE Confidence: 0.938097092857143
- $00:40:46.540 \longrightarrow 00:40:49.124$ what we're going to look at here is
- NOTE Confidence: 0.938097092857143
- $00:40:49.124 \rightarrow 00:40:50.900$ the correlations between you know,
- NOTE Confidence: 0.938097092857143
- 00:40:50.900 --> 00:40:54.300 all these different Christina,
- NOTE Confidence: 0.938097092857143
- $00:40:54.300 \rightarrow 00:40:54.460$ sorry.
- NOTE Confidence: 0.948639664166667
- $00:40:57.340 \longrightarrow 00:40:59.195$ So we're going to look at the
- NOTE Confidence: 0.948639664166667
- $00{:}40{:}59{.}195 \dashrightarrow 00{:}41{:}00{.}433$ correlation between these these
- NOTE Confidence: 0.948639664166667
- $00:41:00.433 \rightarrow 00:41:02.294$ subjective effects scores.
- NOTE Confidence: 0.948639664166667
- 00:41:02.294 --> 00:41:05.242 And they're drinking outcomes
- NOTE Confidence: 0.948639664166667
- $00{:}41{:}05{.}242 \dashrightarrow 00{:}41{:}10{.}100$ for the whole subsequent seven
- NOTE Confidence: 0.948639664166667
- $00{:}41{:}10.100 \dashrightarrow 00{:}41{:}13.940$ months to two months to month.
- NOTE Confidence: 0.948639664166667
- 00:41:13.940 --> 00:41:16.012 Yeah, well, second month of follow up
- NOTE Confidence: 0.948639664166667
- $00{:}41{:}16.012 \dashrightarrow 00{:}41{:}18.303$ after it's the first month after the
- NOTE Confidence: 0.948639664166667

 $00:41:18.303 \rightarrow 00:41:20.319$ second psilocybin session all the way

NOTE Confidence: 0.948639664166667

 $00{:}41{:}20.376 \dashrightarrow 00{:}41{:}22.536$ out to to the end of the double-blind.

NOTE Confidence: 0.948639664166667

00:41:22.540 --> 00:41:24.502 And we're going to look at

NOTE Confidence: 0.948639664166667

 $00:41:24.502 \rightarrow 00:41:26.400$ this for the whole sample.

NOTE Confidence: 0.948639664166667

 $00:41:26.400 \longrightarrow 00:41:28.220$ I don't think the whole sample is

NOTE Confidence: 0.948639664166667

 $00{:}41{:}28{.}220 \dashrightarrow 00{:}41{:}29{.}334$ necessarily all that informative

NOTE Confidence: 0.948639664166667

 $00{:}41{:}29{.}334 \dashrightarrow 00{:}41{:}31{.}273$ because I don't know what it mean.

NOTE Confidence: 0.948639664166667

00:41:31.280 --> 00:41:32.132 If you know,

NOTE Confidence: 0.948639664166667

 $00:41:32.132 \longrightarrow 00:41:33.836$ if we looked at blood pressure

NOTE Confidence: 0.948639664166667

 $00:41:33.836 \longrightarrow 00:41:35.916$ as a mediator or you know this,

NOTE Confidence: 0.948639664166667

 $00:41:35.920 \longrightarrow 00:41:37.120$ we're not a mediation yet.

NOTE Confidence: 0.948639664166667

00:41:37.120 --> 00:41:39.196 But let's just say you know,

NOTE Confidence: 0.948639664166667

00:41:39.200 --> 00:41:40.000 well, what do you know,

NOTE Confidence: 0.948639664166667

 $00:41:40.000 \rightarrow 00:41:42.296$ the psilocybin people have much higher blood

NOTE Confidence: 0.948639664166667

 $00:41:42.296 \rightarrow 00:41:44.398$ pressure increase in and what do you know,

NOTE Confidence: 0.948639664166667

 $00:41:44.400 \longrightarrow 00:41:46.000$ they they drank less.

- NOTE Confidence: 0.948639664166667
- 00:41:46.000 --> 00:41:48.306 Let's see if there's a mediation,
- NOTE Confidence: 0.948639664166667
- $00:41:48.306 \longrightarrow 00:41:49.836$ if there's a relationship there.
- NOTE Confidence: 0.948639664166667
- $00{:}41{:}49{.}840 \dashrightarrow 00{:}41{:}50{.}820$ And we could demonstrate that
- NOTE Confidence: 0.948639664166667
- $00:41:50.820 \rightarrow 00:41:53.100$ and that would be dumb, right?
- NOTE Confidence: 0.948639664166667
- $00{:}41{:}53.100 \dashrightarrow 00{:}41{:}54.400$ So.
- NOTE Confidence: 0.948639664166667
- 00:41:54.400 --> 00:41:55.835 So I don't know what you know
- NOTE Confidence: 0.948639664166667
- 00:41:55.835 00:41:57.079 the whole sample really means.
- NOTE Confidence: 0.948639664166667
- $00:41:57.080 \rightarrow 00:41:59.744$ It's a lot more convincing if we can
- NOTE Confidence: 0.948639664166667
- $00{:}41{:}59{.}744 \dashrightarrow 00{:}42{:}01{.}070$ demonstrate a relationship within
- NOTE Confidence: 0.948639664166667
- $00:42:01.070 \longrightarrow 00:42:03.050$ the psilocybin group that like when
- NOTE Confidence: 0.948639664166667
- $00:42:03.050 \rightarrow 00:42:05.516$ they got this particular kind of experience,
- NOTE Confidence: 0.948639664166667
- $00:42:05.520 \longrightarrow 00:42:07.239$ they did better.
- NOTE Confidence: 0.948639664166667
- 00:42:07.240 --> 00:42:09.438 And based on what I showed you,
- NOTE Confidence: 0.948639664166667
- $00{:}42{:}09{.}440 \dashrightarrow 00{:}42{:}11.060$ with the effect being larger and
- NOTE Confidence: 0.948639664166667
- $00{:}42{:}11.060 \dashrightarrow 00{:}42{:}12.959$ the people here were still drinking,
- NOTE Confidence: 0.94654315875

00:42:15.760 --> 00:42:19.300 I then removed sequentially

NOTE Confidence: 0.94654315875

 $00{:}42{:}19{.}300 \dashrightarrow 00{:}42{:}22{.}840$ the people who were.

NOTE Confidence: 0.94654315875

 $00:42:22.840 \longrightarrow 00:42:24.478$ At a level of base of 0,

NOTE Confidence: 0.94654315875

 $00:42:24.480 \rightarrow 00:42:26.285$ meaning abstinent people who were

NOTE Confidence: 0.94654315875

 $00:42:26.285 \longrightarrow 00:42:28.090$ abstinent before they got the

NOTE Confidence: 0.94654315875

00:42:28.147 --> 00:42:31.040 psilocybin or or diagonidro mean,

NOTE Confidence: 0.94654315875

 $00:42:31.040 \longrightarrow 00:42:35.155$ and then people who were at the

NOTE Confidence: 0.94654315875

 $00:42:35.155 \longrightarrow 00:42:37.920$ low level And then so I remember

NOTE Confidence: 0.94654315875

 $00{:}42{:}37{.}920 \dashrightarrow 00{:}42{:}38{.}880$ first the abstinent people,

NOTE Confidence: 0.94654315875

 $00:42:38.880 \longrightarrow 00:42:40.470$ then the low level drinkers

NOTE Confidence: 0.94654315875

 $00{:}42{:}40{.}470 \dashrightarrow 00{:}42{:}42{.}040$ and then the moderate drinkers.

NOTE Confidence: 0.94654315875

 $00:42:42.040 \longrightarrow 00:42:45.330$ So you get smaller and smaller groups

NOTE Confidence: 0.94654315875

 $00:42:45.330 \rightarrow 00:42:46.916$ so you know less and less power,

NOTE Confidence: 0.94654315875

 $00:42:46.920 \longrightarrow 00:42:49.720$ but you can still look at what the

NOTE Confidence: 0.94654315875

 $00:42:49.720 \longrightarrow 00:42:51.660$ correlations are and if they're.

NOTE Confidence: 0.94654315875

 $00:42:51.660 \rightarrow 00:42:56.636$ The same or different so so these people

- NOTE Confidence: 0.94654315875
- 00:42:56.636 --> 00:42:59.900 may unless you're doing this work,
- NOTE Confidence: 0.94654315875
- $00:42:59.900 \longrightarrow 00:43:01.898$ you may not know these scales
- NOTE Confidence: 0.94654315875
- $00:43:01.898 \longrightarrow 00:43:03.820$ so I'll describe them briefly.
- NOTE Confidence: 0.94654315875
- $00{:}43{:}03{.}820 \dashrightarrow 00{:}43{:}07{.}380$ This is the the five D5 dimensional
- NOTE Confidence: 0.94654315875
- $00{:}43{:}07{.}380 \dashrightarrow 00{:}43{:}09{.}580$ altered states of consciousness scale.
- NOTE Confidence: 0.94654315875
- $00{:}43{:}09{.}580 \dashrightarrow 00{:}43{:}12{.}143$ It's the one from the the Zurich
- NOTE Confidence: 0.94654315875
- 00:43:12.143 --> 00:43:14.558 group front full inviters group.
- NOTE Confidence: 0.94654315875
- 00:43:14.560 --> 00:43:20.973 And it is you know it's a well you
- NOTE Confidence: 0.94654315875
- $00{:}43{:}20.973 \dashrightarrow 00{:}43{:}24.065$ know it's constructed psychometrically
- NOTE Confidence: 0.94654315875
- 00:43:24.065 --> 00:43:27.960 and and well validated on some
- NOTE Confidence: 0.94654315875
- $00{:}43{:}27{.}960 \dashrightarrow 00{:}43{:}29{.}960$ you know medium size samples.
- NOTE Confidence: 0.94654315875
- $00{:}43{:}29{.}960 \dashrightarrow 00{:}43{:}31{.}836$ And so it has these factors as
- NOTE Confidence: 0.94654315875
- $00:43:31.836 \longrightarrow 00:43:33.801$ five but these are the first
- NOTE Confidence: 0.94654315875
- $00{:}43{:}33{.}801 \dashrightarrow 00{:}43{:}35{.}904$ three here are the main ones that
- NOTE Confidence: 0.94654315875
- $00:43:35.904 \rightarrow 00:43:37.480$ people generally care about.
- NOTE Confidence: 0.94654315875

 $00:43:37.480 \longrightarrow 00:43:40.273$ First one is is what they call

NOTE Confidence: 0.94654315875

 $00:43:40.273 \rightarrow 00:43:42.006$ oceanic boundlessness because they

NOTE Confidence: 0.94654315875

 $00:43:42.006 \rightarrow 00:43:43.870$ think mystical experiences is.

NOTE Confidence: 0.94654315875

 $00:43:43.870 \longrightarrow 00:43:44.454$ Hocus pocus.

NOTE Confidence: 0.94654315875

 $00:43:44.454 \rightarrow 00:43:46.790$ So, so this is much more scientific sounding,

NOTE Confidence: 0.94654315875

00:43:46.790 --> 00:43:47.190 right?

NOTE Confidence: 0.950317

 $00:43:49.230 \rightarrow 00:43:50.990$ It probably sounds better in German, but

NOTE Confidence: 0.944027375

 $00:43:53.510 \rightarrow 00:43:56.750$ it's anyway it's really this this you know,

NOTE Confidence: 0.944027375

 $00{:}43{:}56.750 \dashrightarrow 00{:}43{:}59.710$ on eness, bliss, connectedness.

NOTE Confidence: 0.944027375

00:43:59.710 --> 00:44:02.590 Meaning it's it's basically it maps

NOTE Confidence: 0.944027375

 $00{:}44{:}02.590 \dashrightarrow 00{:}44{:}05.624$ pretty tightly on to what the other

NOTE Confidence: 0.944027375

 $00:44:05.624 \rightarrow 00:44:07.624$ groups have called mystical experience.

NOTE Confidence: 0.944027375

 $00{:}44{:}07.630 \dashrightarrow 00{:}44{:}11.270$ So I heard George Goldsmith also,

NOTE Confidence: 0.944027375

00:44:11.270 --> 00:44:12.770 you know, talking down the

NOTE Confidence: 0.944027375

00:44:12.770 --> 00:44:13.670 mystical experience questionnaires,

NOTE Confidence: 0.944027375

 $00:44:13.670 \longrightarrow 00:44:15.014$ and we use the scientific one it's

- NOTE Confidence: 0.944027375
- $00:44:15.014 \rightarrow 00:44:16.149$ and they're measuring the same thing,
- NOTE Confidence: 0.944027375
- $00:44:16.150 \longrightarrow 00:44:17.910$ but they've got a a,
- NOTE Confidence: 0.944027375
- $00{:}44{:}17{.}910 \dashrightarrow 00{:}44{:}21{.}422$ a better name for it I think so And
- NOTE Confidence: 0.944027375
- $00:44:21.422 \rightarrow 00:44:23.630$ then visionary restructuralization
- NOTE Confidence: 0.944027375
- $00:44:23.630 \longrightarrow 00:44:28.172$ that means really the the perceptual
- NOTE Confidence: 0.944027375
- $00{:}44{:}28.172 \dashrightarrow 00{:}44{:}32.482$ effects you know alters altered
- NOTE Confidence: 0.944027375
- $00:44:32.482 \rightarrow 00:44:36.070$ perception and then dread of ego.
- NOTE Confidence: 0.944027375
- $00:44:36.070 \rightarrow 00:44:40.228$ The solution is sort of the bad trick factor,
- NOTE Confidence: 0.944027375
- 00:44:40.230 --> 00:44:43.985 anxiety, sense of impending doom
- NOTE Confidence: 0.944027375
- $00:44:43.985 \longrightarrow 00:44:46.989$ or annihilation or whatever.
- NOTE Confidence: 0.944027375
- $00:44:46.990 \longrightarrow 00:44:48.625$ And then the general scores
- NOTE Confidence: 0.944027375
- $00{:}44{:}48.625 \dashrightarrow 00{:}44{:}50.750$ is the sum of those three.
- NOTE Confidence: 0.944027375
- $00:44:50.750 \rightarrow 00:44:54.030$ So and then looking across the the columns,
- NOTE Confidence: 0.944027375
- $00{:}44{:}54{.}030 \dashrightarrow 00{:}44{:}56{.}670$ the 1st is the whole sample.
- NOTE Confidence: 0.944027375
- $00:44:56.670 \rightarrow 00:44:59.225$ Then we have the whole psilocybin group,
- NOTE Confidence: 0.944027375

 $00:44:59.230 \longrightarrow 00:45:02.366$ 44 people who were not abstinent before

NOTE Confidence: 0.944027375

 $00:45:02.366 \longrightarrow 00:45:05.838$ they got the the psilocybin now.

NOTE Confidence: 0.944027375

 $00{:}45{:}05{.}840 \dashrightarrow 00{:}45{:}10{.}000$ 21 people who were drinking at least at the

NOTE Confidence: 0.8928256632

 $00:45:12.480 \longrightarrow 00:45:14.034$ medium risk level and then 14

NOTE Confidence: 0.8928256632

 $00{:}45{:}14.034 \dashrightarrow 00{:}45{:}16.015$ who are at least were who were

NOTE Confidence: 0.8928256632

00:45:16.015 --> 00:45:17.773 high or very high after they've

NOTE Confidence: 0.8928256632

 $00{:}45{:}17.773 \dashrightarrow 00{:}45{:}19.596$ gotten the four weeks of the rapy.

NOTE Confidence: 0.8928256632

 $00:45:19.600 \longrightarrow 00:45:21.850$ So because people had improved quite

NOTE Confidence: 0.8928256632

 $00{:}45{:}21.850 \dashrightarrow 00{:}45{:}25.472$ a bit and So what you see is that the

NOTE Confidence: 0.8928256632

00:45:25.472 --> 00:45:27.239 oceanic boundlessness does seem to

NOTE Confidence: 0.8928256632

 $00{:}45{:}27{.}239 \dashrightarrow 00{:}45{:}29{.}514$ be pulling most of the weight here.

NOTE Confidence: 0.8928256632

 $00:45:29.520 \longrightarrow 00:45:31.164$ So that the the.

NOTE Confidence: 0.8928256632

 $00:45:31.164 \longrightarrow 00:45:33.630$ The the correlate you know the

NOTE Confidence: 0.8928256632

 $00:45:33.720 \rightarrow 00:45:35.504$ the significant, you know,

NOTE Confidence: 0.8928256632

 $00:45:35.504 \rightarrow 00:45:36.464$ I don't know what significance

NOTE Confidence: 0.8928256632

 $00:45:36.464 \rightarrow 00:45:37.360$ means in this context,
$00:45:37.360 \longrightarrow 00:45:40.665$ but the P values that are you

NOTE Confidence: 0.8928256632

00:45:40.665 --> 00:45:43.360 know under point O five are are

NOTE Confidence: 0.8928256632

 $00:45:43.360 \longrightarrow 00:45:45.110$ bold and red and the ones that

NOTE Confidence: 0.8928256632

 $00{:}45{:}45{.}110 \dashrightarrow 00{:}45{:}46{.}946$ are trend level or or not bold and

NOTE Confidence: 0.8928256632

00:45:46.946 --> 00:45:48.720 red just so you can kind of see

NOTE Confidence: 0.8928256632

 $00{:}45{:}48.720 \dashrightarrow 00{:}45{:}50.869$ them but you could look at the the

NOTE Confidence: 0.8928256632

 $00{:}45{:}50{.}869 \dashrightarrow 00{:}45{:}54{.}163$ correlations just as well and they're

NOTE Confidence: 0.8928256632

 $00:45:54.163 \rightarrow 00:45:56.640$ probably more interesting actually,

NOTE Confidence: 0.8928256632

 $00:45:56.640 \longrightarrow 00:45:59.170$ but what you see is.

NOTE Confidence: 0.8928256632

00:45:59.170 --> 00:45:59.794 Yeah,

NOTE Confidence: 0.8928256632

 $00{:}45{:}59{.}794 \dashrightarrow 00{:}46{:}02{.}290$ the oceanic boundlessness factor

NOTE Confidence: 0.8928256632

 $00{:}46{:}02.290 \dashrightarrow 00{:}46{:}06.010$ is correlated with drinks per day

NOTE Confidence: 0.9301902

 $00:46:08.250 \longrightarrow 00:46:10.976$ at each at each level,

NOTE Confidence: 0.9301902

 $00{:}46{:}10.976 \dashrightarrow 00{:}46{:}12.386$ and the correlation gets higher

NOTE Confidence: 0.9301902

 $00:46:12.386 \longrightarrow 00:46:14.303$ as you get to the people who

 $00:46:14.303 \rightarrow 00:46:15.603$ are drinking more and more.

NOTE Confidence: 0.9301902

 $00{:}46{:}15.610 \dashrightarrow 00{:}46{:}19.342$ So whether you experience this

NOTE Confidence: 0.9301902

00:46:19.342 --> 00:46:21.634 oceanic boundlessness had a

NOTE Confidence: 0.9301902

 $00{:}46{:}21.634 \dashrightarrow 00{:}46{:}24.081$ stronger effect on whether you

NOTE Confidence: 0.9301902

00:46:24.081 --> 00:46:26.290 drank less if you were drinking

NOTE Confidence: 0.9301902

 $00:46:26.290 \rightarrow 00:46:28.390$ more before you got the psilocybin.

NOTE Confidence: 0.9301902

 $00{:}46{:}28{.}390 \dashrightarrow 00{:}46{:}29{.}308$ So that's so

NOTE Confidence: 0.92616476

 $00:46:29.310 \longrightarrow 00:46:31.478$ there is just a I mean there's

NOTE Confidence: 0.92616476

00:46:31.478 --> 00:46:33.542 a floor effect thing there that

NOTE Confidence: 0.92616476

 $00{:}46{:}33{.}542 \dashrightarrow 00{:}46{:}35{.}263$ people who aren't drinking much NOTE Confidence: 0.92616476

 $00:46:35.263 \rightarrow 00:46:37.510$ don't have far to go that's true.

NOTE Confidence: 0.92616476

 $00{:}46{:}37{.}510 \dashrightarrow 00{:}46{:}39{.}110$ So you would expect increased

NOTE Confidence: 0.938705325384616

 $00:46:39.750 \longrightarrow 00:46:40.818$ but they could but they couldn't

NOTE Confidence: 0.938705325384616

 $00:46:40.818 \longrightarrow 00:46:42.148$ go the I mean they could they

NOTE Confidence: 0.943607995

 $00:46:42.150 \longrightarrow 00:46:43.470$ could have gone the other right.

NOTE Confidence: 0.943607995

 $00{:}46{:}43.470 \dashrightarrow 00{:}46{:}45.442$ But I would expect larger I would

- NOTE Confidence: 0.943607995
- $00{:}46{:}45{.}442 \dashrightarrow 00{:}46{:}47{.}822$ expect larger correlations with a
- NOTE Confidence: 0.943607995
- $00:46:47.822 \longrightarrow 00:46:49.730$ higher baseline for this analysis
- NOTE Confidence: 0.943607995
- $00{:}46{:}49{.}730 \dashrightarrow 00{:}46{:}51{.}388$ more room to more room to improve.
- NOTE Confidence: 0.943607995
- $00{:}46{:}51{.}390 \dashrightarrow 00{:}46{:}52{.}655$ Yeah no that that you have to
- NOTE Confidence: 0.943607995
- 00:46:52.655 --> 00:46:53.390 do is I don't know how you.
- NOTE Confidence: 0.938995675
- $00{:}46{:}54{.}990 \dashrightarrow 00{:}46{:}57{.}470$ Well, if it's working, you would if it's
- NOTE Confidence: 0.938995675
- $00{:}46{:}57{.}470 \dashrightarrow 00{:}46{:}59{.}350$ something you would expect that. But
- NOTE Confidence: 0.94427896
- $00:46:59.630 \longrightarrow 00:47:00.750$ yeah, no, this supports that.
- NOTE Confidence: 0.94427896
- 00:47:00.750 --> 00:47:01.202 It's working.
- NOTE Confidence: 0.94427896
- 00:47:01.202 --> 00:47:02.510 I'm not sure it supports.
- NOTE Confidence: 0.94427896
- $00:47:02.510 \longrightarrow 00:47:04.494$ It's working better in the people who
- NOTE Confidence: 0.94427896
- $00:47:04.494 \rightarrow 00:47:05.910$ are drinking more as opposed to just.
- NOTE Confidence: 0.94427896
- $00:47:05.910 \longrightarrow 00:47:06.855$ It's working right.
- NOTE Confidence: 0.94427896
- $00{:}47{:}06.855 \dashrightarrow 00{:}47{:}08.430$ Pretty well across the board.
- NOTE Confidence: 0.94427896
- $00:47:08.430 \longrightarrow 00:47:09.718$ And you see it more in the people
- NOTE Confidence: 0.94427896

- $00:47:09.718 \longrightarrow 00:47:10.603$ that are drinking, right.
- NOTE Confidence: 0.94427896
- $00:47:10.603 \longrightarrow 00:47:10.949$ That's all.
- NOTE Confidence: 0.9452853
- $00:47:11.070 \rightarrow 00:47:13.662$ Well, it's, yeah, they're just more to do.
- NOTE Confidence: 0.9452853
- $00:47:13.662 \longrightarrow 00:47:14.842$ I mean, there's more benefit
- NOTE Confidence: 0.9452853
- $00:47:14.842 \rightarrow 00:47:18.430$ to be had, I guess. So, yeah.
- NOTE Confidence: 0.9452853
- $00:47:18.430 \longrightarrow 00:47:20.390$ And then, you know, the other.
- NOTE Confidence: 0.9486397066666667
- $00{:}47{:}20.880 \dashrightarrow 00{:}47{:}22.056$ It is interesting that the oceanic
- NOTE Confidence: 0.9486397066666667
- 00:47:22.056 --> 00:47:23.280 boundless just jumps out as carrying.
- NOTE Confidence: 0.938240864
- $00{:}47{:}24.120 \dashrightarrow 00{:}47{:}26.720$ Yeah. And the other ones,
- NOTE Confidence: 0.938240864
- $00{:}47{:}26.720 \dashrightarrow 00{:}47{:}28.335$ the correlations don't really even
- NOTE Confidence: 0.938240864
- $00{:}47{:}28{.}335 \dashrightarrow 00{:}47{:}30{.}793$ go up except for the general which
- NOTE Confidence: 0.938240864
- $00:47:30.793 \longrightarrow 00:47:32.570$ includes and that's it's oceanic
- NOTE Confidence: 0.938240864
- $00:47:32.570 \longrightarrow 00:47:34.400$ boundless as it's pulling that one.
- NOTE Confidence: 0.938240864
- $00:47:34.400 \rightarrow 00:47:38.630$ So, so that's so that's interesting.
- NOTE Confidence: 0.938240864
- $00:47:38.630 \longrightarrow 00:47:40.390$ This is the mystical
- NOTE Confidence: 0.938240864
- $00:47:40.390 \longrightarrow 00:47:41.270$ experience questionnaire.

- NOTE Confidence: 0.938240864
- $00:47:41.270 \longrightarrow 00:47:43.496$ So this one there are you know
- NOTE Confidence: 0.938240864
- $00:47:43.496 \longrightarrow 00:47:45.068$ there's there's been factor
- NOTE Confidence: 0.938240864
- $00:47:45.068 \longrightarrow 00:47:46.709$ analyses demonstrating this.
- NOTE Confidence: 0.938240864
- $00{:}47{:}46.710 \dashrightarrow 00{:}47{:}47.667$ This four factor,
- NOTE Confidence: 0.938240864
- $00{:}47{:}47{.}667 \dashrightarrow 00{:}47{:}49{.}900$ it's been done a couple of different
- NOTE Confidence: 0.938240864
- $00{:}47{:}49.960 \dashrightarrow 00{:}47{:}51.822$ ways but the sort of most standard
- NOTE Confidence: 0.938240864
- $00:47:51.822 \longrightarrow 00:47:53.664$ one now is these four factors
- NOTE Confidence: 0.938240864
- $00:47:53.664 \rightarrow 00:47:56.422$ 11 is what they which they call
- NOTE Confidence: 0.938240864
- $00:47:56.422 \longrightarrow 00:47:58.576$ mystical which has to do with
- NOTE Confidence: 0.938240864
- $00{:}47{:}58{.}576$ --> $00{:}48{:}00{.}502$ the unity and and meaning and
- NOTE Confidence: 0.935222015
- $00:48:02.710 \longrightarrow 00:48:05.208$ and then they've.
- NOTE Confidence: 0.935222015
- 00:48:05.208 --> 00:48:07.236 Taken out that the positive moods
- NOTE Confidence: 0.935222015
- 00:48:07.236 --> 00:48:09.599 tends to go there's some questions
- NOTE Confidence: 0.935222015
- $00{:}48{:}09{.}599 \dashrightarrow 00{:}48{:}11{.}303$ that go separately transcendence
- NOTE Confidence: 0.935222015
- $00:48:11.303 \rightarrow 00:48:14.030$ of space and time and ineffability.
- NOTE Confidence: 0.935222015

 $00:48:14.030 \longrightarrow 00:48:15.405$ So those I think ineffability

NOTE Confidence: 0.935222015

 $00{:}48{:}15{.}405 \dashrightarrow 00{:}48{:}16{.}505$ only has three items.

NOTE Confidence: 0.935222015

00:48:16.510 --> 00:48:18.232 So they the the mystical has about

NOTE Confidence: 0.935222015

 $00:48:18.232 \longrightarrow 00:48:20.309$ half of the items and then they're

NOTE Confidence: 0.935222015

 $00{:}48{:}20{.}309 \dashrightarrow 00{:}48{:}22{.}250$ smaller numbers after that but

NOTE Confidence: 0.935222015

 $00{:}48{:}22{.}250 \dashrightarrow 00{:}48{:}25{.}177$ so here you know in the whole

NOTE Confidence: 0.935222015

 $00:48:25.177 \longrightarrow 00:48:28.026$ sample you know all of them had

NOTE Confidence: 0.935222015

00:48:28.030 --> 00:48:31.080 you know modest but statistically

NOTE Confidence: 0.935222015

 $00{:}48{:}31{.}080 \dashrightarrow 00{:}48{:}32{.}910$ significant correlations and

NOTE Confidence: 0.935222015

 $00:48:32.910 \longrightarrow 00:48:35.520$ then as you go across. The

NOTE Confidence: 0.9452853

 $00{:}48{:}37{.}920 \dashrightarrow 00{:}48{:}42{.}164$ they tend to increase to some extent.

NOTE Confidence: 0.9452853

 $00{:}48{:}42.164 \dashrightarrow 00{:}48{:}45.982$ And so even you know that so that the MEQ 43,

NOTE Confidence: 0.9452853

 $00{:}48{:}45{.}982 \dashrightarrow 00{:}48{:}48{.}630$ that's the old that the old version of

NOTE Confidence: 0.9452853

 $00:48:48.709 \rightarrow 00:48:50.808$ the scale where they had and there's

NOTE Confidence: 0.9452853

 $00{:}48{:}50{.}808 \dashrightarrow 00{:}48{:}52{.}280$ a total score that you get from that.

NOTE Confidence: 0.9452853

00:48:52.280 --> 00:48:53.558 So if you put put it,

- NOTE Confidence: 0.9452853
- $00:48:53.560 \rightarrow 00:48:58.020$ put those all together, it's all the

 $00:48:58.020 \longrightarrow 00:48:59.920$ aspects of the mystical experience.

NOTE Confidence: 0.9452853

 $00:48:59.920 \rightarrow 00:49:02.080$ Yeah, you can see those correlations

NOTE Confidence: 0.9452853

 $00:49:02.080 \rightarrow 00:49:03.476$ going you know all the way up to.

NOTE Confidence: 0.927129504545455

 $00{:}49{:}05{.}750 \dashrightarrow 00{:}49{:}08{.}704$ Over .5 in the in this smallest

NOTE Confidence: 0.927129504545455

 $00:49:08.704 \longrightarrow 00:49:11.390$ group and it's it's so it's a

NOTE Confidence: 0.927129504545455

 $00{:}49{:}11.390 \dashrightarrow 00{:}49{:}15.750$ nice kind of consistent effect

NOTE Confidence: 0.927129504545455

 $00{:}49{:}15.750 \dashrightarrow 00{:}49{:}19.590$ here and this is another mysticism

NOTE Confidence: 0.927129504545455

 $00:49:19.590 \dashrightarrow 00:49:22.070$ scale that hood mysticism scale.

NOTE Confidence: 0.927129504545455

 $00{:}49{:}22.070 \dashrightarrow 00{:}49{:}25.846$ And these are I think you know some of

NOTE Confidence: 0.927129504545455

 $00:49:25.846 \rightarrow 00:49:28.090$ the largest correlations of all with

NOTE Confidence: 0.927129504545455

 $00:49:28.170 \longrightarrow 00:49:30.345$ introvertive Introvertive mysticism

NOTE Confidence: 0.927129504545455

00:49:30.345 - 00:49:34.100 means being kind of being one with.

NOTE Confidence: 0.927129504545455

 $00{:}49{:}34{.}100 \dashrightarrow 00{:}49{:}36{.}700$ With nothing, being one with

NOTE Confidence: 0.927129504545455

 $00:49:36.700 \rightarrow 00:49:39.436$ nothingness or kind of going inside

 $00:49:39.436 \longrightarrow 00:49:43.060$ and just being being one with non.

NOTE Confidence: 0.927129504545455

00:49:43.060 --> 00:49:46.332 Being extrovertive means being one

NOTE Confidence: 0.927129504545455

 $00:49:46.332 \longrightarrow 00:49:48.420$ with the universe and the plants

NOTE Confidence: 0.927129504545455

 $00:49:48.487 \rightarrow 00:49:50.257$ and the animals and everything.

NOTE Confidence: 0.927129504545455

 $00{:}49{:}50{.}260 \dashrightarrow 00{:}49{:}51{.}930$ And interpretation means that it's

NOTE Confidence: 0.927129504545455

 $00{:}49{:}51{.}930 \dashrightarrow 00{:}49{:}53{.}600$ sort of their religious meaning

NOTE Confidence: 0.927129504545455

 $00:49:53.655 \longrightarrow 00:49:54.820$ that people give to it.

NOTE Confidence: 0.927129504545455

 $00:49:54.820 \rightarrow 00:49:58.006$ So really all three were pretty

NOTE Confidence: 0.927129504545455

00:49:58.006 --> 00:50:01.570 strongly correlated and yeah,

NOTE Confidence: 0.927129504545455

 $00:50:01.570 \longrightarrow 00:50:03.470$ the more you drank.

NOTE Confidence: 0.927129504545455

 $00{:}50{:}03{.}470 \dashrightarrow 00{:}50{:}06{.}793$ The higher the correlation again and

NOTE Confidence: 0.927129504545455

 $00{:}50{:}06{.}793 \dashrightarrow 00{:}50{:}08{.}731$ then this last one the hallucinogen

NOTE Confidence: 0.927129504545455

00:50:08.731 --> 00:50:11.376 rating scale this is this one is really

NOTE Confidence: 0.927129504545455

00:50:11.376 --> 00:50:12.991 coming from a different perspective

NOTE Confidence: 0.927129504545455

 $00{:}50{:}13.052 \dashrightarrow 00{:}50{:}15.665$ and trying to so it's made by Rick

NOTE Confidence: 0.927129504545455

 $00{:}50{:}15.665 \dashrightarrow 00{:}50{:}17.836$ Strassman and he actually based it

 $00:50:17.836 \longrightarrow 00:50:21.030$ on the five what are called

NOTE Confidence: 0.927129504545455

 $00:50:21.030 \longrightarrow 00:50:24.195$ skandas in in Buddhist psychology

NOTE Confidence: 0.927129504545455

 $00:50:24.195 \rightarrow 00:50:26.870$ that they're just different aspects

NOTE Confidence: 0.927129504545455

 $00:50:26.870 \rightarrow 00:50:31.230$ of subjectivity of mind and.

NOTE Confidence: 0.927129504545455

00:50:31.230 --> 00:50:32.706 Or I guess a human being,

NOTE Confidence: 0.927129504545455

 $00:50:32.710 \longrightarrow 00:50:33.910$ because somatic is one of them.

NOTE Confidence: 0.927129504545455

 $00:50:33.910 \longrightarrow 00:50:35.746$ So that's not even mind really.

NOTE Confidence: 0.927129504545455

 $00:50:35.750 \longrightarrow 00:50:37.082$ But so we're made-up of these

NOTE Confidence: 0.927129504545455

 $00:50:37.082 \rightarrow 00:50:37.748$ things they say.

NOTE Confidence: 0.927129504545455

 $00:50:37.750 \longrightarrow 00:50:38.614$ So somatic is,

NOTE Confidence: 0.927129504545455

00:50:38.614 --> 00:50:39.190 you know,

NOTE Confidence: 0.927129504545455

 $00:50:39.190 \rightarrow 00:50:41.015$ physical experience of your body

NOTE Confidence: 0.927129504545455

 $00:50:41.015 \longrightarrow 00:50:43.793$ that didn't seem to be very strongly

NOTE Confidence: 0.927129504545455

 $00{:}50{:}43.793 \dashrightarrow 00{:}50{:}47.990$ related to anything affective experience.

NOTE Confidence: 0.927129504545455

 $00{:}50{:}47{.}990 \dashrightarrow 00{:}50{:}50{.}018$ Emotions strongly correlated

 $00:50:50.018 \longrightarrow 00:50:52.076$ in this case and none of those

NOTE Confidence: 0.927129504545455

 $00{:}50{:}52{.}076 \dashrightarrow 00{:}50{:}54{.}095$ other skills really get at that

NOTE Confidence: 0.927129504545455

 $00:50:54.095 \rightarrow 00:50:56.842$ particularly except for positive mood.

NOTE Confidence: 0.927129504545455

 $00{:}50{:}56{.}842 \dashrightarrow 00{:}51{:}01{.}050$ So that's so that one shows some

NOTE Confidence: 0.927129504545455

 $00:51:01.050 \rightarrow 00:51:02.890$ strong correlations of perceptual,

NOTE Confidence: 0.927129504545455

 $00{:}51{:}02.890 \dashrightarrow 00{:}51{:}05.200$ just as this is pretty much what's

NOTE Confidence: 0.927129504545455

 $00{:}51{:}05{.}200 \dashrightarrow 00{:}51{:}08{.}810$ measured in the five DASC under the

NOTE Confidence: 0.927129504545455

 $00:51:08.810 \longrightarrow 00:51:11.830$ the visionary restructuralization,

NOTE Confidence: 0.927129504545455

 $00:51:11.830 \longrightarrow 00:51:14.090$ nothing there.

NOTE Confidence: 0.927129504545455

 $00:51:14.090 \rightarrow 00:51:16.930$ And then changes in cognition,

NOTE Confidence: 0.927129504545455

 $00:51:16.930 \longrightarrow 00:51:18.226$ there's, you know,

NOTE Confidence: 0.927129504545455

 $00:51:18.226 \longrightarrow 00:51:19.830$ moderate sized correlations

NOTE Confidence: 0.927129504545455

 $00{:}51{:}19{.}830 \dashrightarrow 00{:}51{:}22{.}050$ there and volition.

NOTE Confidence: 0.927129504545455

 $00:51:22.050 \rightarrow 00:51:24.808$ Is, is, is in the smaller group.

NOTE Confidence: 0.927129504545455

00:51:24.810 --> 00:51:26.370 So it actually was, was,

NOTE Confidence: 0.927129504545455

 $00:51:26.370 \rightarrow 00:51:28.342$ was quite strongly correlated,

- NOTE Confidence: 0.927129504545455
- 00:51:28.342 --> 00:51:29.730 but this, you know,
- NOTE Confidence: 0.927129504545455
- $00:51:29.730 \longrightarrow 00:51:31.130$ not so much in the larger group.
- NOTE Confidence: 0.927129504545455
- $00{:}51{:}31{.}130 \dashrightarrow 00{:}51{:}33{.}725$ So I'm not sure that I'd be
- NOTE Confidence: 0.927129504545455
- $00{:}51{:}33{.}725 \dashrightarrow 00{:}51{:}34{.}650$ quite as confident in that.
- NOTE Confidence: 0.927129504545455
- $00{:}51{:}34{.}650 \dashrightarrow 00{:}51{:}36{.}190$ One is intensity a combination
- NOTE Confidence: 0.927129504545455
- $00:51:36.190 \longrightarrow 00:51:37.450$ of the rest is a separate,
- NOTE Confidence: 0.927129504545455
- $00:51:37.450 \longrightarrow 00:51:38.586$ it's a separate thing.
- NOTE Confidence: 0.927129504545455
- 00:51:38.586 --> 00:51:40.006 And it's actually very simple.
- NOTE Confidence: 0.927129504545455
- $00{:}51{:}40{.}010 \dashrightarrow 00{:}51{:}41{.}767$ It's like how strong is your experience,
- NOTE Confidence: 0.927129504545455
- 00:51:41.770 --> 00:51:43.044 What dose do you think you got?
- NOTE Confidence: 0.927129504545455
- 00:51:43.050 --> 00:51:45.050 And I think it's, it's like 3 items.
- NOTE Confidence: 0.927129504545455
- 00:51:45.050 --> 00:51:46.232 So it's, it's,
- NOTE Confidence: 0.927129504545455
- $00{:}51{:}46{.}232 \dashrightarrow 00{:}51{:}48{.}990$ it's interesting and it's and it's not
- NOTE Confidence: 0.927129504545455
- $00{:}51{:}49{.}067 \dashrightarrow 00{:}51{:}51{.}528$ any that's really about all it is.
- NOTE Confidence: 0.927129504545455
- 00:51:51.530 --> 00:51:54.930 So I think you know.
- NOTE Confidence: 0.927129504545455

 $00:51:54.930 \longrightarrow 00:51:57.142$ So anyway the conclusion from this I

NOTE Confidence: 0.927129504545455

 $00{:}51{:}57{.}142 \dashrightarrow 00{:}51{:}59{.}488$ think is that it does seem to make it.

NOTE Confidence: 0.927129504545455

00:51:59.490 --> 00:52:01.878 You know they higher,

NOTE Confidence: 0.927129504545455

00:52:01.878 --> 00:52:02.466 higher,

NOTE Confidence: 0.927129504545455

 $00:52:02.466 \rightarrow 00:52:05.695$ stronger experiences seem to do seem

NOTE Confidence: 0.927129504545455

00:52:05.695 - 00:52:07.405 to make a difference to people.

NOTE Confidence: 0.927129504545455

00:52:07.410 --> 00:52:07.810 But

NOTE Confidence: 0.91988512

 $00:52:09.170 \longrightarrow 00:52:10.210$ I always wonder what these

NOTE Confidence: 0.91988512

 $00{:}52{:}10{.}210$ --> $00{:}52{:}12{.}009$ correlations if if the like,

NOTE Confidence: 0.91988512

 $00:52:12.009 \rightarrow 00:52:13.767$ if some of these are simply

NOTE Confidence: 0.91988512

 $00{:}52{:}13.767 \dashrightarrow 00{:}52{:}15.290$ surrogate markers for how much

NOTE Confidence: 0.91988512

 $00:52:15.290 \longrightarrow 00:52:17.180$ the drug affected the neurons.

NOTE Confidence: 0.91988512

 $00:52:17.180 \longrightarrow 00:52:18.380$ Yeah right.

NOTE Confidence: 0.91988512

 $00:52:18.380 \dashrightarrow 00:52:19.864$ But the so the specificity if you

NOTE Confidence: 0.91988512

 $00{:}52{:}19{.}864 \dashrightarrow 00{:}52{:}21{.}498$ have this you know more specificity

NOTE Confidence: 0.91988512

 $00:52:21.500 \longrightarrow 00:52:22.928$ to the mysticism than the other

 $00:52:22.928 \rightarrow 00:52:24.108$ components starts target that they're

NOTE Confidence: 0.91988512

 $00{:}52{:}24.108 \dashrightarrow 00{:}52{:}26.428$ maybe not but something like intensity.

NOTE Confidence: 0.91988512

00:52:26.428 --> 00:52:28.136 I wonder if it's just you know

NOTE Confidence: 0.91988512

 $00:52:28.136 \longrightarrow 00:52:29.262$ how got in there and how tightly

NOTE Confidence: 0.91988512

 $00{:}52{:}29{.}262 \dashrightarrow 00{:}52{:}30{.}424$ did it bind the receptors and how

NOTE Confidence: 0.91988512

 $00:52:30.424 \longrightarrow 00:52:31.492$ much did it affect the neurons

NOTE Confidence: 0.9436080666666667

 $00:52:32.900 \rightarrow 00:52:35.300$ and how are the preparatory sessions

NOTE Confidence: 0.9436080666666667

 $00:52:35.300 \longrightarrow 00:52:39.046$ how are they how are they introduced

NOTE Confidence: 0.9436080666666667

 $00{:}52{:}39{.}046 \dashrightarrow 00{:}52{:}44{.}420$ to it they so we we tried to be very.

NOTE Confidence: 0.9395827066666667

 $00{:}52{:}46{.}470 \dashrightarrow 00{:}52{:}49{.}270$ Non directive and as far as what kind

NOTE Confidence: 0.9395827066666667

 $00:52:49.270 \longrightarrow 00:52:52.590$ of an experience is a good experience

NOTE Confidence: 0.9395827066666667

00:52:52.590 - 00:52:55.758 but we did we we were willing to

NOTE Confidence: 0.9395827066666667

 $00{:}52{:}55{.}758$ --> $00{:}52{:}58{.}155$ suggest that you know you you whatever NOTE Confidence: 0.9395827066666667

 $00{:}52{:}58{.}155 \dashrightarrow 00{:}53{:}00{.}329$ experience you have you know that that

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}00{.}329 \dashrightarrow 00{:}53{:}02{.}369$ might be the one you need to that

 $00:53:02.435 \longrightarrow 00:53:04.571$ that's the one for you and it's it's

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}04{.}571 \dashrightarrow 00{:}53{:}07{.}056$ you know you may do do with it what

NOTE Confidence: 0.9395827066666667

00:53:07.056 --> 00:53:09.324 you can and see if see if you can but

NOTE Confidence: 0.9395827066666667

 $00:53:09.324 \rightarrow 00:53:10.683$ do you think they fidelity actually

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}10.683 \dashrightarrow 00{:}53{:}12.891$ that was true because that's what is my

NOTE Confidence: 0.9395827066666667

 $00:53:12.945 \rightarrow 00:53:16.409$ concern is you know if if if it's an if then.

NOTE Confidence: 0.9395827066666667

 $00:53:16.410 \rightarrow 00:53:18.740$ Type situation where you're told

NOTE Confidence: 0.9395827066666667

 $00:53:18.740 \longrightarrow 00:53:20.290$ that if you have this experience,

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}20{.}290 \dashrightarrow 00{:}53{:}21{.}454$ you're going to get better.

NOTE Confidence: 0.9395827066666667

00:53:21.454 --> 00:53:22.918 And then therefore if I didn't

NOTE Confidence: 0.9395827066666667

 $00:53:22.918 \longrightarrow 00:53:23.650$ have that experience,

NOTE Confidence: 0.9395827066666667

 $00:53:23.650 \longrightarrow 00:53:25.888$ I'm not going to get better.

NOTE Confidence: 0.9395827066666667

00:53:25.890 --> 00:53:29.520 Well, so like I said, we, we, you know,

NOTE Confidence: 0.9395827066666667

 $00:53:29.520 \dashrightarrow 00:53:31.290$ I don't know what people believe.

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}31{.}290 \dashrightarrow 00{:}53{:}31{.}500$ Yeah.

NOTE Confidence: 0.9395827066666667

00:53:31.500 --> 00:53:32.970 I mean because we also said it,

 $00:53:32.970 \longrightarrow 00:53:36.423$ it it's also possible that you know that

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}36{.}423 \dashrightarrow 00{:}53{:}38{.}181$ this is going to do something in your brain

NOTE Confidence: 0.9395827066666667

 $00:53:38.181 \rightarrow 00:53:39.930$ and it may have nothing to do with that.

NOTE Confidence: 0.9395827066666667

 $00:53:39.930 \longrightarrow 00:53:40.890$ So that equipoise,

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}40.890 \dashrightarrow 00{:}53{:}43.358$ I mean so that was so that was

NOTE Confidence: 0.9395827066666667

 $00:53:43.358 \longrightarrow 00:53:46.396$ so that was put out there too.

NOTE Confidence: 0.9395827066666667

00:53:46.400 --> 00:53:50.916 I but I think you know I think

NOTE Confidence: 0.9395827066666667

 $00:53:50.916 \rightarrow 00:53:52.908$ people have expected expectations

NOTE Confidence: 0.9395827066666667

 $00{:}53{:}52{.}908 \dashrightarrow 00{:}53{:}55{.}678$ sure and and the rapist due to so

NOTE Confidence: 0.9395827066666667

00:53:55.680 --> 00:53:57.871 but we we we you know definitely

NOTE Confidence: 0.9395827066666667

 $00:53:57.871 \rightarrow 00:53:59.224$ we're not privileging mystical

NOTE Confidence: 0.9395827066666667

 $00:53:59.224 \rightarrow 00:54:01.360$ type experience per se for example

NOTE Confidence: 0.9486396966666667

 $00:54:06.240 \longrightarrow 00:54:08.556$ we probably should wrap up because

NOTE Confidence: 0.9486396966666667

 $00:54:08.560 \dashrightarrow 00:54:11.360$ it's getting but but MRI so

NOTE Confidence: 0.922857967

 $00{:}54{:}11{.}360 \dashrightarrow 00{:}54{:}12{.}392$ I'm going to show you this

 $00:54:12.392 \rightarrow 00:54:13.080$ really quickly these are.

NOTE Confidence: 0.9402536

00:54:15.100 - 00:54:17.776 Really just hot off the press.

NOTE Confidence: 0.9402536

00:54:17.780 --> 00:54:19.990 And you know, I don't want to make too much

NOTE Confidence: 0.9402536

 $00:54:20.049 \longrightarrow 00:54:22.057$ of it because this is only 11 subjects,

NOTE Confidence: 0.9402536

 $00{:}54{:}22.060 \dashrightarrow 00{:}54{:}25.980$ but I think it's it's interesting pilot data.

NOTE Confidence: 0.9402536

 $00{:}54{:}25{.}980 \dashrightarrow 00{:}54{:}30{.}740$ We had a task that evaluated a

NOTE Confidence: 0.9402536

 $00{:}54{:}30{.}740 \dashrightarrow 00{:}54{:}33{.}868$ response to visual alcohol cues

NOTE Confidence: 0.9402536

 $00:54:33.868 \dashrightarrow 00:54:36.354$ and negative emotional pictures.

NOTE Confidence: 0.9402536

00:54:36.354 --> 00:54:39.318 I have pictures and we scanned

NOTE Confidence: 0.9402536

00:54:39.318 --> 00:54:42.142 people three days before and two

NOTE Confidence: 0.9402536

 $00{:}54{:}42{.}142 \dashrightarrow 00{:}54{:}45{.}061$ days after first dose of the drug.

NOTE Confidence: 0.9402536

 $00:54:45.070 \rightarrow 00:54:47.404$ And we were you know interested

NOTE Confidence: 0.9402536

 $00{:}54{:}47{.}404 \dashrightarrow 00{:}54{:}49{.}976$ in the alcohol versus neutral and

NOTE Confidence: 0.9402536

00:54:49.976 --> 00:54:52.316 and the negative emotion versus

NOTE Confidence: 0.9402536

 $00{:}54{:}52{.}316$ --> $00{:}54{:}54{.}392$ neutral change in those and and and

NOTE Confidence: 0.9402536

 $00:54:54.392 \rightarrow 00:54:56.046$ the difference in change of those

 $00:54:56.046 \rightarrow 00:54:58.940$ between the two groups and we looked

NOTE Confidence: 0.9402536

 $00{:}54{:}58{.}940 \dashrightarrow 00{:}55{:}03{.}210$ at regions that we thought were most

NOTE Confidence: 0.9402536

 $00:55:03.339 \rightarrow 00:55:06.550$ likely to be involved in you know.

NOTE Confidence: 0.965411993

 $00{:}55{:}09{.}420 \dashrightarrow 00{:}55{:}11{.}760$ Craving response and reward processing

NOTE Confidence: 0.965411993

 $00{:}55{:}11.760 \dashrightarrow 00{:}55{:}14.100$ and and negative emotional processing.

NOTE Confidence: 0.965411993

 $00{:}55{:}14.100 \dashrightarrow 00{:}55{:}19.530$ So lateral medial Pfc and singular

NOTE Confidence: 0.965411993

 $00:55:19.530 \longrightarrow 00:55:21.844$ and ventral indoors will stray

NOTE Confidence: 0.965411993

 $00:55:21.844 \rightarrow 00:55:25.540$ them and when we found significant

NOTE Confidence: 0.941930728333333

 $00{:}55{:}27.740 \dashrightarrow 00{:}55{:}32.186$ effects we then only for those.

NOTE Confidence: 0.941930728333333

 $00{:}55{:}32.190 \dashrightarrow 00{:}55{:}34.554$ Roi's we looked at functional connectivity

NOTE Confidence: 0.941930728333333

 $00:55:34.554 \rightarrow 00:55:37.152$ between groups to see if there are

NOTE Confidence: 0.941930728333333

 $00:55:37.152 \dashrightarrow 00:55:39.150$ any differences in between Roi's,

NOTE Confidence: 0.941930728333333

 $00{:}55{:}39{.}150 \dashrightarrow 00{:}55{:}42{.}470$ between no between the groups. So if

NOTE Confidence: 0.98302305

 $00:55:42.990 \rightarrow 00:55:44.510$ functional connectivity compared between

NOTE Confidence: 0.91878229125

 $00:55:44.670 \rightarrow 00:55:46.215$ groups, yeah, yeah, for the

 $00:55:46.215 \longrightarrow 00:55:48.638$ for the ROI that was that was

NOTE Confidence: 0.91878229125

 $00{:}55{:}48.638 \dashrightarrow 00{:}55{:}50.266$ different between the groups.

NOTE Confidence: 0.9352219

 $00{:}55{:}53{.}590 \dashrightarrow 00{:}55{:}56{.}100$ And then we also looked at whether

NOTE Confidence: 0.9352219

 $00{:}55{:}56{.}100 \dashrightarrow 00{:}55{:}59{.}400$ these the differences that we found.

NOTE Confidence: 0.9352219

 $00{:}55{:}59{.}400 \dashrightarrow 00{:}56{:}01{.}280$ Had any relationship to the

NOTE Confidence: 0.9352219

 $00:56:01.280 \rightarrow 00:56:03.160$ change in drinks per day,

NOTE Confidence: 0.9352219

 $00:56:03.160 \longrightarrow 00:56:06.037$ so this is a just a summary

NOTE Confidence: 0.9352219

 $00:56:06.040 \longrightarrow 00:56:07.600$ in terms of the bold contrast.

NOTE Confidence: 0.9352219

 $00{:}56{:}07.600 \dashrightarrow 00{:}56{:}12.640$ There were some, you know, fairly small

NOTE Confidence: 0.9654121

 $00:56:15.160 \rightarrow 00:56:17.744$ clusters in right ventrilateral

NOTE Confidence: 0.9654121

 $00:56:17.744 \longrightarrow 00:56:19.036$ prefrontal cortex,

NOTE Confidence: 0.9654121

 $00:56:19.040 \longrightarrow 00:56:22.008$ left dorsolateral prefrontal cortex,

NOTE Confidence: 0.9654121

 $00{:}56{:}22.008 \dashrightarrow 00{:}56{:}24.234$ and left caudate.

NOTE Confidence: 0.9654121

 $00{:}56{:}24.240 \dashrightarrow 00{:}56{:}26.778$ That were different and it was

NOTE Confidence: 0.9654121

 $00{:}56{:}26{.}778$ --> $00{:}56{:}29{.}434$ all more more activation in the

NOTE Confidence: 0.9654121

 $00:56:29.434 \rightarrow 00:56:31.160$ psilocybin group and for negative

- NOTE Confidence: 0.9654121
- 00:56:31.160 -> 00:56:32.280 effective stimuli they were

 $00:56:34.360 \longrightarrow 00:56:36.980$ areas in left medial prefrontal

NOTE Confidence: 0.89145567

 $00:56:36.980 \dashrightarrow 00:56:40.120$ and that were also left odd eight.

NOTE Confidence: 0.9456207466666667

 $00:56:40.240 \longrightarrow 00:56:41.998$ I'm sorry, is this these task

NOTE Confidence: 0.9456207466666667

00:56:41.998 --> 00:56:43.639 data from the before psilocybin

NOTE Confidence: 0.9456207466666667

 $00:56:43.639 \rightarrow 00:56:45.279$ or the after psilocybin?

NOTE Confidence: 0.9456207466666667

 $00{:}56{:}45.280 \dashrightarrow 00{:}56{:}48.572$ It's where the after minus before.

NOTE Confidence: 0.9456207466666667

 $00{:}56{:}48{.}572 \dashrightarrow 00{:}56{:}50{.}900$ So it's task dependent

NOTE Confidence: 0.9456207466666667

 $00{:}56{:}50{.}900 \dashrightarrow 00{:}56{:}52{.}520$ activation after minus.

NOTE Confidence: 0.9456207466666667

00:56:52.520 --> 00:56:54.840 Identical task, dependent activation,

NOTE Confidence: 0.9456207466666667

 $00{:}56{:}54{.}840 \dashrightarrow 00{:}56{:}56{.}920$ yeah between groups and then relative

NOTE Confidence: 0.957025933333333

 $00{:}56{:}56{.}920 \dashrightarrow 00{:}56{:}58{.}480$ to neutral stimulus

NOTE Confidence: 0.924552493809524

 $00{:}57{:}00{.}720$ --> $00{:}57{:}03{.}753$ and then yeah, so they the ones that we

NOTE Confidence: 0.924552493809524

 $00{:}57{:}03.753 \dashrightarrow 00{:}57{:}06.770$ looked at those in terms of functional

NOTE Confidence: 0.924552493809524

00:57:06.770 --> 00:57:09.320 connectivity and there were two of the

00:57:12.520 --> 00:57:16.352 alcohol Q responding.

NOTE Confidence: 0.78374635

 $00{:}57{:}16.352 \dashrightarrow 00{:}57{:}18.312$ Areas that that did have

NOTE Confidence: 0.78374635

 $00:57:18.312 \longrightarrow 00:57:19.488$ some connectivity changes,

NOTE Confidence: 0.78374635

 $00:57:19.490 \longrightarrow 00:57:20.882$ so I'll show you the pictures

NOTE Confidence: 0.78374635

00:57:20.882 --> 00:57:22.386 really quickly and I, you know,

NOTE Confidence: 0.78374635

 $00{:}57{:}22{.}386 \dashrightarrow 00{:}57{:}24{.}114$ again let's not get too carried

NOTE Confidence: 0.78374635

 $00{:}57{:}24.114 \dashrightarrow 00{:}57{:}26.690$ away here, but this is this

NOTE Confidence: 0.945045695238095

 $00:57:28.770 \longrightarrow 00:57:30.214$ right intralateral Pfc area

NOTE Confidence: 0.945045695238095

 $00{:}57{:}30{.}214 \dashrightarrow 00{:}57{:}32{.}917$ and using that as a seed then

NOTE Confidence: 0.945045695238095

 $00{:}57{:}32{.}917 \dashrightarrow 00{:}57{:}34{.}753$ there was increased functional

NOTE Confidence: 0.945045695238095

 $00{:}57{:}34{.}753 \dashrightarrow 00{:}57{:}37{.}330$ connectivity with this area in the

NOTE Confidence: 0.22011086

 $00:57:39.690 \rightarrow 00:57:43.490$ IFS lateral pre central gyrus there.

NOTE Confidence: 0.936228176

 $00{:}57{:}45{.}780 \dashrightarrow 00{:}57{:}49{.}620$ This is the area in the left dorsal

NOTE Confidence: 0.936228176

 $00{:}57{:}49.620 \dashrightarrow 00{:}57{:}54.980$ lateral Pfc and this is the left caudate,

NOTE Confidence: 0.936228176

 $00{:}57{:}54{.}980 \dashrightarrow 00{:}57{:}57{.}140$ which yeah, this was,

NOTE Confidence: 0.936228176

 $00:57:57.140 \longrightarrow 00:57:59.156$ this was a bit of a surprise and

 $00:57:59.156 \rightarrow 00:58:01.270$ you know we didn't really expect to

NOTE Confidence: 0.936228176

 $00{:}58{:}01{.}270 \dashrightarrow 00{:}58{:}03{.}944$ see greater reactivity in dorsal

NOTE Confidence: 0.936228176

 $00:58:03.944 \rightarrow 00:58:06.308$ strayatum as being associated

NOTE Confidence: 0.936228176

 $00:58:06.308 \rightarrow 00:58:08.100$ with the rapeutic response.

NOTE Confidence: 0.936228176

 $00:58:08.100 \rightarrow 00:58:12.260$ So you know, I'm not sure what to make of it.

NOTE Confidence: 0.936228176

 $00{:}58{:}12.260 \dashrightarrow 00{:}58{:}13.580$ It also showed greater

NOTE Confidence: 0.936228176

 $00:58:13.580 \rightarrow 00:58:14.900$ connectivity with a CC,

NOTE Confidence: 0.8639468

00:58:18.300 --> 00:58:21.460 though I don't know but

NOTE Confidence: 0.8639468

 $00{:}58{:}21{.}460 \dashrightarrow 00{:}58{:}22{.}540$ there was for that one.

NOTE Confidence: 0.8639468

 $00:58:22.540 \longrightarrow 00:58:25.140$ There was a significant relationship

NOTE Confidence: 0.8639468

 $00{:}58{:}25{.}140 \dashrightarrow 00{:}58{:}30{.}300$ between that that effect and the the

NOTE Confidence: 0.8639468

 $00{:}58{:}30{.}300 \dashrightarrow 00{:}58{:}33{.}500$ functional connectivity now and decreased

NOTE Confidence: 0.8639468

 $00:58:33.500 \rightarrow 00:58:38.892$ in drinks per day and then for the

NOTE Confidence: 0.8639468

00:58:38.892 --> 00:58:42.030 negative effective stimuli this is.

NOTE Confidence: 0.8639468

 $00{:}58{:}42.030 \dashrightarrow 00{:}58{:}45.659$ Yeah, the the left medial prefrontal area

 $00{:}58{:}48{.}953 \dashrightarrow 00{:}58{:}51{.}909$ there was associated with decreased NOTE Confidence: 0.9805072 $00{:}58{:}54{.}270 \dashrightarrow 00{:}58{:}58{.}310$ drinking and then this is the left NOTE Confidence: 0.9805072 $00:58:58.310 \rightarrow 00:59:02.075$ caudate again, which was no functional NOTE Confidence: 0.9805072 $00:59:02.075 \rightarrow 00:59:04.338$ connectivity changes in this case, NOTE Confidence: 0.9805072 $00:59:04.338 \longrightarrow 00:59:06.720$ but it was associated also with NOTE Confidence: 0.9805072 $00:59:06.797 \longrightarrow 00:59:10.310$ the decrease in drinking, so. NOTE Confidence: 0.9805072 $00{:}59{:}10{.}310 \dashrightarrow 00{:}59{:}11{.}870$ So that's so that's that. NOTE Confidence: 0.9805072 00:59:11.870 --> 00:59:14.302 So you know, So what was the time NOTE Confidence: 0.9805072 $00:59:14.302 \rightarrow 00:59:17.150$ from the dosing to the scanning, NOTE Confidence: 0.9805072 $00:59:17.150 \longrightarrow 00:59:19.510$ it was one to two days after, NOTE Confidence: 0.9805072 00:59:19.510 -> 00:59:22.070 two days after Okay. Yeah. NOTE Confidence: 0.9805072 00:59:22.070 --> 00:59:24.150 And so you know these, NOTE Confidence: 0.9805072 $00{:}59{:}24.150 \dashrightarrow 00{:}59{:}25.632$ these are pretty large effects given NOTE Confidence: 0.9805072 $00:59:25.632 \rightarrow 00:59:27.714$ you know that we see that we're seeing NOTE Confidence: 0.9805072 $00:59:27.714 \rightarrow 00:59:29.280$ anything with with samples of this 94

 $00:58:45.659 \rightarrow 00:58:48.953$ and which this increase in activation

- NOTE Confidence: 0.9805072
- $00:59:29.330 \longrightarrow 00:59:30.870$ size and they kind of make sense.

 $00:59:30.870 \rightarrow 00:59:32.590$ I mean they're in regions that make sense,

NOTE Confidence: 0.9805072

 $00:59:32.590 \longrightarrow 00:59:34.762$ they weren't in you know just

NOTE Confidence: 0.9805072

 $00:59:34.762 \longrightarrow 00:59:37.319$ kind of bunch of random spots.

NOTE Confidence: 0.9805072

 $00:59:37.320 \longrightarrow 00:59:39.270$ Some of them were accompanied by

NOTE Confidence: 0.9805072

 $00{:}59{:}39{.}270 \dashrightarrow 00{:}59{:}40{.}245$ functional connectivity changes

NOTE Confidence: 0.9805072

 $00{:}59{:}40.245 \dashrightarrow 00{:}59{:}42.280$ and some of them were correlated

NOTE Confidence: 0.9805072

 $00:59:42.280 \longrightarrow 00:59:45.640$ with changes in drinking.

NOTE Confidence: 0.9805072

00:59:45.640 --> 00:59:47.400 You know,

NOTE Confidence: 0.9805072

 $00{:}59{:}47{.}400 \dashrightarrow 00{:}59{:}50{.}745$ obviously you know we have a

NOTE Confidence: 0.9805072

00:59:50.745 --> 00:59:53.200 replication problem in in FM RI

NOTE Confidence: 0.9805072

00:59:53.200 --> 00:59:54.880 work and this study is you know

NOTE Confidence: 0.941371755555556

 $00:59:56.920 \rightarrow 00:59:59.531$ a a prime candidate for that being

NOTE Confidence: 0.941371755555556

 $00{:}59{:}59{.}531 \dashrightarrow 01{:}00{:}02{.}250$ very small and you know the the

NOTE Confidence: 0.941371755555556

 $01:00:02.250 \rightarrow 01:00:04.102$ directionality was not exactly.

 $01:00:04.102 \rightarrow 01:00:07.819$ Entirely in some cases it was with

NOTE Confidence: 0.941371755555556

 $01{:}00{:}07.819 \dashrightarrow 01{:}00{:}12.290$ the the OR dorsal prefrontal areas

NOTE Confidence: 0.941371755555556

01:00:12.290 --> 01:00:14.990 but with with the medial prefrontal

NOTE Confidence: 0.941371755555556

 $01:00:14.990 \longrightarrow 01:00:17.343$ and the and the dorsal straight

NOTE Confidence: 0.941371755555556

 $01{:}00{:}17.343 \dashrightarrow 01{:}00{:}19.954$ and we didn't really expect to see

NOTE Confidence: 0.941371755555556

 $01:00:19.954 \rightarrow 01:00:22.410$ increases in the in this especially

NOTE Confidence: 0.941371755555556

 $01:00:22.410 \longrightarrow 01:00:24.490$ for the alcohol accused so,

NOTE Confidence: 0.941371755555556

 $01{:}00{:}24{.}490 \dashrightarrow 01{:}00{:}28{.}004$ so we'll see but you know executive

NOTE Confidence: 0.941371755555556

 $01:00:28.010 \longrightarrow 01:00:30.100$ functioning is not all about.

NOTE Confidence: 0.941371755555556

 $01{:}00{:}30{.}100 \dashrightarrow 01{:}00{:}32{.}011$ Just you know inhibition there might be

NOTE Confidence: 0.941371755555556

 $01:00:32.011 \rightarrow 01:00:34.019$ other things going on and more complicated.

NOTE Confidence: 0.941371755555556

01:00:34.020 --> 01:00:36.316 So I think it's it's interesting and

NOTE Confidence: 0.941371755555556

 $01{:}00{:}36{.}316 \dashrightarrow 01{:}00{:}38{.}611$ we'll we'll follow up on it and we're

NOTE Confidence: 0.941371755555556

 $01{:}00{:}38.611 \dashrightarrow 01{:}00{:}43.572$ really hoping very soon to be able to

NOTE Confidence: 0.941371755555556

 $01:00:43.572 \longrightarrow 01:00:47.092$ do a much more you know much larger

NOTE Confidence: 0.941371755555556

 $01:00:47.092 \rightarrow 01:00:49.540$ and more sophisticated version of this

- NOTE Confidence: 0.941371755555556
- $01{:}00{:}49{.}540 \dashrightarrow 01{:}00{:}52{.}746$ with Regina's help and NA AAA funding
- NOTE Confidence: 0.941371755555556
- $01:00:52.746 \longrightarrow 01:00:56.956$ if we can get get the grant funded so.
- NOTE Confidence: 0.941371755555556
- 01:00:56.960 --> 01:00:58.958 So that's that.
- NOTE Confidence: 0.941371755555556
- $01{:}00{:}58.960 \dashrightarrow 01{:}01{:}03.480$ So overall you know I think looking good
- NOTE Confidence: 0.941371755555556
- $01{:}01{:}03{.}480 \dashrightarrow 01{:}01{:}06{.}680$ for efficacy we need to do more there
- NOTE Confidence: 0.941371755555556
- $01{:}01{:}06.760 \dashrightarrow 01{:}01{:}08.964$ obviously and there's some evidence
- NOTE Confidence: 0.941371755555556
- $01:01:08.964 \longrightarrow 01:01:11.589$ that these drugs are acting across all
- NOTE Confidence: 0.941371755555556
- $01:01:11.589 \longrightarrow 01:01:13.913$ three of the core domains of addiction.
- NOTE Confidence: 0.941371755555556
- 01:01:13.920 --> 01:01:17.880 And you know I think it's,
- NOTE Confidence: 0.941371755555556
- 01:01:17.880 --> 01:01:18.738 it's complicated, right.
- NOTE Confidence: 0.941371755555556
- 01:01:18.738 --> 01:01:20.078 And this is, this is,
- NOTE Confidence: 0.941371755555556
- $01:01:20.078 \dashrightarrow 01:01:22.530$ this is really why I like it is because.
- NOTE Confidence: 0.941371755555556
- $01:01:22.530 \longrightarrow 01:01:24.706$ I mean you know we do want to
- NOTE Confidence: 0.941371755555556
- $01:01:24.706 \longrightarrow 01:01:26.975$ be able to reduce things to you
- NOTE Confidence: 0.941371755555556
- $01{:}01{:}26.975 \dashrightarrow 01{:}01{:}28.445$ know the level of the receptor.
- NOTE Confidence: 0.941371755555556

 $01:01:28.450 \rightarrow 01:01:31.848$ But for a lot of things that's,

NOTE Confidence: 0.941371755555556

01:01:31.850 --> 01:01:32.888 you know, it's like you know,

NOTE Confidence: 0.941371755555556

 $01:01:32.890 \longrightarrow 01:01:34.210$ we don't do biology,

NOTE Confidence: 0.941371755555556

 $01:01:34.210 \longrightarrow 01:01:35.530$ we using quantum mechanics.

NOTE Confidence: 0.941371755555556

01:01:35.530 --> 01:01:36.148 You know,

NOTE Confidence: 0.941371755555556

 $01:01:36.148 \longrightarrow 01:01:38.311$ we you need to use higher levels

NOTE Confidence: 0.941371755555556

01:01:38.311 --> 01:01:39.649 of it's you know,

NOTE Confidence: 0.941371755555556

 $01{:}01{:}39.650 \dashrightarrow 01{:}01{:}41.810$ and it still should be scientific

NOTE Confidence: 0.941371755555556

 $01{:}01{:}41{.}810 \dashrightarrow 01{:}01{:}43{.}610$ and it still should need to

NOTE Confidence: 0.941371755555556

 $01:01:43.610 \longrightarrow 01:01:44.810$ be rigorous and makes sense.

NOTE Confidence: 0.941371755555556

01:01:44.810 $\operatorname{-->}$ 01:01:47.033 But we need to find ways to you know

NOTE Confidence: 0.941371755555556

 $01:01:47.033 \rightarrow 01:01:49.673$ looking at at these higher level

NOTE Confidence: 0.941371755555556

01:01:49.673 --> 01:01:51.225 phenomena including subjective experience

NOTE Confidence: 0.941371755555556

 $01:01:51.288 \longrightarrow 01:01:53.087$ to really make sense of this so.

NOTE Confidence: 0.941371755555556

 $01:01:53.090 \rightarrow 01:01:57.690$ So that's that thank you wonderful.

NOTE Confidence: 0.941371755555556

 $01:01:57.690 \rightarrow 01:01:59.328$ It's really nice to see the new,

- NOTE Confidence: 0.941371755555556
- $01:01:59.330 \longrightarrow 01:02:00.050$ the new stuff.
- NOTE Confidence: 0.941371755555556
- $01:02:00.050 \longrightarrow 01:02:00.290$ So
- NOTE Confidence: 0.855730071428571
- $01:02:01.130 \longrightarrow 01:02:02.682$ worth worth worth staying
- NOTE Confidence: 0.855730071428571
- $01:02:02.682 \longrightarrow 01:02:04.490$ later enough today afternoon.
- NOTE Confidence: 0.957025933333333
- $01:02:06.530 \dashrightarrow 01:02:09.514$ Anybody still there? Well that's what I
- NOTE Confidence: 0.957025933333333
- $01{:}02{:}09{.}514 \dashrightarrow 01{:}02{:}12{.}370$ was checking out and the answer was yes.
- NOTE Confidence: 0.938815971428571
- $01{:}02{:}15{.}890 \dashrightarrow 01{:}02{:}17{.}698$ So do we have you know question or
- NOTE Confidence: 0.938815971428571
- $01{:}02{:}17.698 \dashrightarrow 01{:}02{:}19.622$ two or any comments for for Michael
- NOTE Confidence: 0.938815971428571
- $01:02:19.622 \longrightarrow 01:02:21.530$ before we we call that to night?
- NOTE Confidence: 0.9301902
- 01:02:24.300 --> 01:02:25.220 Hello. Hi, I have
- NOTE Confidence: 0.957025933333333
- $01:02:25.220 \rightarrow 01:02:26.180$ two quick questions.
- NOTE Confidence: 0.941691228571429
- $01{:}02{:}29{.}220 \dashrightarrow 01{:}02{:}30{.}588$ Thank you so much for the
- NOTE Confidence: 0.941691228571429
- $01{:}02{:}30.588 \dashrightarrow 01{:}02{:}31.660$ presentation and the great work.
- NOTE Confidence: 0.867295068
- $01{:}02{:}33.060 \dashrightarrow 01{:}02{:}34.860$ I have two quick questions. One is that
- NOTE Confidence: 0.867295068
- $01{:}02{:}34{.}860 \dashrightarrow 01{:}02{:}37{.}058$ do you have did you collect any data
- NOTE Confidence: 0.8536761066666667

 $01:02:37.220 \rightarrow 01:02:39.824$ after the like between those one and

NOTE Confidence: 0.8536761066666667

 $01{:}02{:}39{.}824 \dashrightarrow 01{:}02{:}43{.}107$ two after the sign in and like how

NOTE Confidence: 0.8536761066666667

 $01:02:43.107 \rightarrow 01:02:45.403$ long can you comment on how long after NOTE Confidence: 0.8536761066666667

01:02:45.403 --> 01:02:47.700 the first dose you could see any

NOTE Confidence: 0.8536761066666667

 $01{:}02{:}47{.}700 \dashrightarrow 01{:}02{:}50{.}920$ changes in the outcome measures and?

NOTE Confidence: 0.8536761066666667

 $01{:}02{:}50{.}920 \dashrightarrow 01{:}02{:}54{.}776$ If like how long do they last after

NOTE Confidence: 0.8536761066666667

 $01:02:54.776 \longrightarrow 01:02:57.240$ the first psilocybin administration?

NOTE Confidence: 0.8536761066666667

 $01{:}02{:}57{.}240 \dashrightarrow 01{:}02{:}59{.}812$ And the second question is that why

NOTE Confidence: 0.8536761066666667

 $01{:}02{:}59{.}812 \dashrightarrow 01{:}03{:}01{.}856$ the second dose of the psilocybin has

NOTE Confidence: 0.8536761066666667

 $01:03:01.856 \rightarrow 01:03:03.720$ higher dose compared to first one.

NOTE Confidence: 0.849624371428572

 $01{:}03{:}04{.}320 \dashrightarrow 01{:}03{:}07{.}078$ OK, yeah. So that's a good question.

NOTE Confidence: 0.849624371428572

 $01:03:07.080 \longrightarrow 01:03:08.136$ So for the first,

NOTE Confidence: 0.849624371428572

 $01:03:08.136 \rightarrow 01:03:11.920$ as far as the first question, we there's a,

NOTE Confidence: 0.849624371428572

 $01{:}03{:}11{.}920 \dashrightarrow 01{:}03{:}14{.}506$ there was a follow up at one week

NOTE Confidence: 0.849624371428572

 $01:03:14.506 \rightarrow 01:03:16.502$ after the first psilocybin dose,

NOTE Confidence: 0.849624371428572

 $01:03:16.502 \rightarrow 01:03:20.300$ we didn't do anything you know the same day.

- NOTE Confidence: 0.849624371428572
- 01:03:20.300 --> 01:03:23.020 You know, I guess I just,
- NOTE Confidence: 0.849624371428572
- $01{:}03{:}23{.}020 \dashrightarrow 01{:}03{:}24{.}940$ I'm not that interested if people,
- NOTE Confidence: 0.849624371428572
- 01:03:24.940 --> 01:03:25.900 you know, I mean,
- NOTE Confidence: 0.849624371428572
- $01{:}03{:}25{.}900 \dashrightarrow 01{:}03{:}27{.}700$ I know that like in the ketamine studies,
- NOTE Confidence: 0.849624371428572
- 01:03:27.700 --> 01:03:29.020 you know, it's like right afterwards you go,
- NOTE Confidence: 0.849624371428572
- $01:03:29.020 \longrightarrow 01:03:29.632$ you're less depressed.
- NOTE Confidence: 0.849624371428572
- $01:03:29.632 \longrightarrow 01:03:30.856$ That's great and it is great.
- NOTE Confidence: 0.849624371428572
- 01:03:30.860 --> 01:03:32.780 But I mean, especially an addiction,
- NOTE Confidence: 0.849624371428572
- $01:03:32.780 \longrightarrow 01:03:35.104$ you know, I don't really care if
- NOTE Confidence: 0.849624371428572
- $01:03:35.104 \longrightarrow 01:03:36.966$ somebody's got less craving 8
- NOTE Confidence: 0.849624371428572
- $01:03:36.966 \rightarrow 01:03:38.696$ hours after they took psilocybin.
- NOTE Confidence: 0.849624371428572
- $01{:}03{:}38{.}700 \dashrightarrow 01{:}03{:}40{.}924$ So may be we should have done it the next
- NOTE Confidence: 0.849624371428572
- $01{:}03{:}40{.}924 \dashrightarrow 01{:}03{:}42{.}924$ day and then I am going to do that in
- NOTE Confidence: 0.849624371428572
- $01{:}03{:}42{.}924 \dashrightarrow 01{:}03{:}44{.}936$ the next study and just just to see,
- NOTE Confidence: 0.849624371428572
- $01:03:44.940 \longrightarrow 01:03:45.866$ you know,
- NOTE Confidence: 0.849624371428572

 $01:03:45.866 \rightarrow 01:03:49.107$ if there are some changes that are.

NOTE Confidence: 0.849624371428572

01:03:49.110 --> 01:03:51.108 Detectable that early but but you

NOTE Confidence: 0.849624371428572

 $01:03:51.108 \longrightarrow 01:03:53.620$ know most of these self report things NOTE Confidence: 0.849624371428572

 $01:03:53.620 \rightarrow 01:03:57.255$ were were showing changes after a

NOTE Confidence: 0.849624371428572

01:03:57.255 --> 01:03:59.674 week the craving you know it it it

NOTE Confidence: 0.849624371428572

 $01:03:59.674 \rightarrow 01:04:02.064$ does seem to the effects are a little NOTE Confidence: 0.849624371428572

 $01{:}04{:}02.064 \dashrightarrow 01{:}04{:}04.660$ larger far a little farther out it

NOTE Confidence: 0.849624371428572

 $01{:}04{:}04{.}660 \dashrightarrow 01{:}04{:}07{.}429$ seems like there's you know these

NOTE Confidence: 0.93019015

01:04:09.590 --> 01:04:10.988 but right it could just it could

NOTE Confidence: 0.93019015

01:04:10.988 --> 01:04:12.383 be more related to the fact that

NOTE Confidence: 0.93019015

 $01{:}04{:}12.383 \dashrightarrow 01{:}04{:}13.734$ they're not drinking as much so I NOTE Confidence: 0.93019015

01:04:13.778 --> 01:04:14.912 don't know so the second question

NOTE Confidence: 0.93019015

 $01:04:14.912 \longrightarrow 01:04:18.140$ is why do we go up on the dose.

NOTE Confidence: 0.93019015

 $01:04:18.140 \longrightarrow 01:04:19.775$ Because we could.

NOTE Confidence: 0.93019015

01:04:19.775 --> 01:04:22.500 I think clinically it makes,

NOTE Confidence: 0.93019015

01:04:22.500 --> 01:04:24.747 I mean it's it's it's a little

- NOTE Confidence: 0.93019015
- $01{:}04{:}24.747 \dashrightarrow 01{:}04{:}26.695$ complicated because people got to get
- NOTE Confidence: 0.93019015
- $01{:}04{:}26.695 \dashrightarrow 01{:}04{:}28.879$ different doses in the second session, right.
- NOTE Confidence: 0.93019015
- 01:04:28.879 --> 01:04:31.252 But clinically you know you would want
- NOTE Confidence: 0.93019015
- $01:04:31.252 \rightarrow 01:04:33.910$ to be able to titrate people especially
- NOTE Confidence: 0.93019015
- $01:04:33.910 \longrightarrow 01:04:36.506$ if you think you know they need to have.
- NOTE Confidence: 0.93019015
- $01:04:36.510 \longrightarrow 01:04:38.390$ A really strong experience
- NOTE Confidence: 0.93019015
- $01:04:38.390 \longrightarrow 01:04:40.270$ or a maximal receptor,
- NOTE Confidence: 0.93019015
- 01:04:40.270 --> 01:04:41.950 you know occupancy or whatever,
- NOTE Confidence: 0.93019015
- $01:04:41.950 \longrightarrow 01:04:42.553$ whatever it is,
- NOTE Confidence: 0.93019015
- 01:04:42.553 --> 01:04:44.350 you know if you think it's dose related,
- NOTE Confidence: 0.93019015
- $01{:}04{:}44{.}350 \dashrightarrow 01{:}04{:}46{.}036$ you want to maximize the dose
- NOTE Confidence: 0.93019015
- $01:04:46.036 \longrightarrow 01:04:47.484$ without you know hurting people
- NOTE Confidence: 0.93019015
- $01:04:47.484 \longrightarrow 01:04:48.829$ but you can't give them,
- NOTE Confidence: 0.93019015
- 01:04:48.830 --> 01:04:49.492 you can't.
- NOTE Confidence: 0.93019015
- $01{:}04{:}49{.}492 \dashrightarrow 01{:}04{:}51{.}809$ So with the doses were 25 milligrams
- NOTE Confidence: 0.93019015

- $01:04:51.809 \longrightarrow 01:04:53.870$ and then it was 30 or 40 depending
- NOTE Confidence: 0.93019015
- $01{:}04{:}53.870 \dashrightarrow 01{:}04{:}56.030$ on how robust their response was.
- NOTE Confidence: 0.93019015
- $01:04:56.030 \longrightarrow 01:04:59.012$ And so we gave about 15 people
- NOTE Confidence: 0.93019015
- $01:04:59.012 \longrightarrow 01:05:02.220$ I think the 40 milligram dose.
- NOTE Confidence: 0.93019015
- 01:05:02.220 --> 01:05:02.540 You know
- NOTE Confidence: 0.940253492
- $01:05:02.580 \longrightarrow 01:05:05.140$ and those were by weight 25%.
- NOTE Confidence: 0.936228132
- 01:05:05.140 --> 01:05:06.100 That's right. So it was,
- NOTE Confidence: 0.936228132
- 01:05:06.100 --> 01:05:08.270 it was actually, I mean the biggest
- NOTE Confidence: 0.936228132
- $01{:}05{:}08{.}270 \dashrightarrow 01{:}05{:}10{.}140$ dose was about 65 milligrams.
- NOTE Confidence: 0.936228132
- 01:05:10.140 --> 01:05:11.772 So you you would not give
- NOTE Confidence: 0.936228132
- $01:05:11.772 \longrightarrow 01:05:13.478$ somebody who walked in off the
- NOTE Confidence: 0.936228132
- $01:05:13.478 \longrightarrow 01:05:15.140$ street that dose to begin with.
- NOTE Confidence: 0.936228132
- 01:05:15.140 --> 01:05:17.380 You know you just because some people,
- NOTE Confidence: 0.936228132
- $01:05:17.380 \longrightarrow 01:05:18.408$ some people are really
- NOTE Confidence: 0.936228132
- $01:05:18.408 \rightarrow 01:05:19.693$ challenged by the 25 milligrams.
- NOTE Confidence: 0.936228132
- $01:05:19.700 \longrightarrow 01:05:21.056$ Is there any thought that you,

- NOTE Confidence: 0.936228132
- $01{:}05{:}21{.}060 \dashrightarrow 01{:}05{:}22{.}635$ I mean there could be an inverted
- NOTE Confidence: 0.936228132
- 01:05:22.635 --> 01:05:24.478 I mean for ketamine it seems pretty
- NOTE Confidence: 0.936228132
- $01{:}05{:}24.478 \dashrightarrow 01{:}05{:}26.485$ clear there is an inverted at least
- NOTE Confidence: 0.936228132
- $01{:}05{:}26.485 \dashrightarrow 01{:}05{:}28.516$ in rodents probably in units that you
- NOTE Confidence: 0.936228132
- $01{:}05{:}28.516 \dashrightarrow 01{:}05{:}30.532$ can by pass that sort of sweet spot.
- NOTE Confidence: 0.936228132
- $01:05:30.540 \longrightarrow 01:05:30.863$ Yeah.
- NOTE Confidence: 0.936228132
- $01:05:30.863 \rightarrow 01:05:34.130$ I have not seen any evidence of that at all.
- NOTE Confidence: 0.936228132
- $01{:}05{:}34{.}130 \dashrightarrow 01{:}05{:}37{.}410$ I and yeah I couldn't but I've gone
- NOTE Confidence: 0.936228132
- $01{:}05{:}37{.}410 \dashrightarrow 01{:}05{:}40{.}149$ higher than most people but we've gone
- NOTE Confidence: 0.936228132
- 01:05:40.149 --> 01:05:44.926 pretty high and you know in the in the 60s,
- NOTE Confidence: 0.936228132
- 01:05:44.930 --> 01:05:47.090 you know they they you know
- NOTE Confidence: 0.936228132
- $01{:}05{:}47{.}090 \dashrightarrow 01{:}05{:}48{.}566$ they this is in the in America.
- NOTE Confidence: 0.936228132
- $01:05:48.570 \rightarrow 01:05:50.250$ I mean it was different in Europe where
- NOTE Confidence: 0.936228132
- $01{:}05{:}50{.}250 \dashrightarrow 01{:}05{:}51{.}934$ people are more sophisticated than subtle.
- NOTE Confidence: 0.936228132
- $01{:}05{:}51{.}934 \dashrightarrow 01{:}05{:}53{.}886$ But it was like you know we're going
- NOTE Confidence: 0.936228132

- $01:05:53.886 \rightarrow 01:05:55.845$ to just blow them out of the water.
- NOTE Confidence: 0.936228132
- $01:05:55.850 \longrightarrow 01:05:57.368$ And it was like so one,
- NOTE Confidence: 0.936228132
- $01:05:57.370 \longrightarrow 01:05:59.154$ one time get it you know one time
- NOTE Confidence: 0.936228132
- $01:05:59.154 \rightarrow 01:06:00.717$ get them ready and then just.
- NOTE Confidence: 0.936228132
- $01:06:00.720 \rightarrow 01:06:03.276$ You know the whole point is ego death right.
- NOTE Confidence: 0.936228132
- $01:06:03.280 \rightarrow 01:06:06.586$ So you want them just to completely you know,
- NOTE Confidence: 0.936228132
- 01:06:06.586 --> 01:06:07.918 it's like you know,
- NOTE Confidence: 0.936228132
- 01:06:07.920 --> 01:06:08.774 psychic ECT.
- NOTE Confidence: 0.936228132
- 01:06:08.774 --> 01:06:10.909 It's like you're just completely
- NOTE Confidence: 0.936228132
- 01:06:10.909 --> 01:06:13.439 flatlined your your brain activity if,
- NOTE Confidence: 0.936228132
- $01:06:13.440 \longrightarrow 01:06:15.654$ I mean that's not really true
- NOTE Confidence: 0.936228132
- $01{:}06{:}15.654 \dashrightarrow 01{:}06{:}17.652$ but you're you're conscious your
- NOTE Confidence: 0.936228132
- $01:06:17.652 \rightarrow 01:06:21.600$ consciousness is going to be obliterated.
- NOTE Confidence: 0.936228132
- $01{:}06{:}21{.}600 \dashrightarrow 01{:}06{:}25{.}962$ So but they just made that up so we don't
- NOTE Confidence: 0.936228132
- $01:06:25.962 \dashrightarrow 01:06:29.240$ know if that's necessary or not and so the.
- NOTE Confidence: 0.936228132
- $01:06:29.240 \rightarrow 01:06:30.704$ But that was the prevailing wisdom

- NOTE Confidence: 0.936228132
- 01:06:30.704 --> 01:06:33.040 and you know, most drugs more is
- NOTE Confidence: 0.936228132
- $01{:}06{:}33.040 \dashrightarrow 01{:}06{:}36.400$ better until you start hurting, right.
- NOTE Confidence: 0.936228132
- $01:06:36.400 \longrightarrow 01:06:38.344$ So that's to maximize the dose
- NOTE Confidence: 0.936228132
- $01{:}06{:}38{.}344 \dashrightarrow 01{:}06{:}39{.}640$ is the short answer.
- NOTE Confidence: 0.936228132
- 01:06:39.640 --> 01:06:39.760 Yeah.
- NOTE Confidence: 0.929773064285714
- 01:06:46.880 --> 01:06:48.044 Thank you, Michael.
- NOTE Confidence: 0.929773064285714
- $01:06:48.044 \longrightarrow 01:06:49.596$ This is Emmanuel Schindler.
- NOTE Confidence: 0.929773064285714
- 01:06:49.600 --> 01:06:52.519 Thank you for coming and and for
- NOTE Confidence: 0.929773064285714
- $01{:}06{:}52{.}519 \dashrightarrow 01{:}06{:}54{.}410$ braving the Southern Connecticut
- NOTE Confidence: 0.929773064285714
- 01:06:54.410 --> 01:06:57.170 traffic on a Friday afternoon.
- NOTE Confidence: 0.929773064285714
- $01:06:57.170 \longrightarrow 01:06:58.340$ I had a question about
- NOTE Confidence: 0.929773064285714
- $01{:}06{:}58{.}340 \dashrightarrow 01{:}06{:}59{.}730$ and I may have missed it,
- NOTE Confidence: 0.929773064285714
- $01{:}06{:}59{.}730 \dashrightarrow 01{:}07{:}02{.}370$ When you talked about the study
- NOTE Confidence: 0.929773064285714
- $01{:}07{:}02.370 \dashrightarrow 01{:}07{:}05.640$ the the with the mouse model where
- NOTE Confidence: 0.929773064285714
- $01{:}07{:}05.640 \dashrightarrow 01{:}07{:}07.950$ they blocked with with Catanzeran.
- NOTE Confidence: 0.929773064285714

 $01:07:07.950 \longrightarrow 01:07:09.875$ Did they do a dose response

NOTE Confidence: 0.929773064285714

 $01:07:09.875 \longrightarrow 01:07:10.730$ with the Catanzeran?

NOTE Confidence: 0.9335446666666667

 $01:07:10.810 \rightarrow 01:07:14.842$ Was it just a single dose they they

NOTE Confidence: 0.9335446666666667

 $01:07:14.842 \longrightarrow 01:07:17.635$ found a dose I think I mean they they

NOTE Confidence: 0.9335446666666667

01:07:17.635 --> 01:07:20.099 might have done some pilot work but

NOTE Confidence: 0.9335446666666667

01:07:20.099 --> 01:07:24.065 they used a dose which was reliably.

NOTE Confidence: 0.9335446666666667

 $01{:}07{:}24.065 \dashrightarrow 01{:}07{:}28.052$ Blocking the head twitch and in at

NOTE Confidence: 0.9335446666666667

 $01{:}07{:}28.052 \dashrightarrow 01{:}07{:}30.749$ least one of the studies that also

NOTE Confidence: 0.9335446666666667

01:07:30.749 --> 01:07:34.670 blocked the characteristic EE G

NOTE Confidence: 0.9335446666666667

01:07:34.670 --> 01:07:39.480 changes which is you know decreased

NOTE Confidence: 0.9335446666666667

 $01{:}07{:}39{.}480 \dashrightarrow 01{:}07{:}44{.}225$ power and gamma that goes that

NOTE Confidence: 0.9335446666666667

 $01:07:44.225 \longrightarrow 01:07:46.355$ that's that particularly goes down so

NOTE Confidence: 0.9335446666666667

01:07:46.355 --> 01:07:48.369 they're in in humans and in animals

NOTE Confidence: 0.9335446666666667

 $01:07:48.369 \rightarrow 01:07:50.147$ that you can that it correlates

NOTE Confidence: 0.9335446666666667

 $01{:}07{:}50{.}147 \dashrightarrow 01{:}07{:}53{.}892$ with the subjective effect so but.

NOTE Confidence: 0.9335446666666667

 $01:07:53.892 \rightarrow 01:07:59.040$ You know we know that people you know
- NOTE Confidence: 0.907236928
- 01:08:01.800 --> 01:08:03.795 it's it's tough because I don't
- NOTE Confidence: 0.907236928
- 01:08:03.795 --> 01:08:05.912 think you know we we can't assume that
- NOTE Confidence: 0.907236928
- $01:08:05.912 \longrightarrow 01:08:08.860$ you know all of the 582 A receptors
- NOTE Confidence: 0.907236928
- $01:08:08.860 \rightarrow 01:08:12.440$ were were blocked obviously and people
- NOTE Confidence: 0.941153033809524
- $01{:}08{:}14.480 \dashrightarrow 01{:}08{:}15.976$ the seroton in receptors are
- NOTE Confidence: 0.941153033809524
- $01{:}08{:}15{.}976 \dashrightarrow 01{:}08{:}18{.}220$ almost saturated at at you know
- NOTE Confidence: 0.941153033809524
- 01:08:18.292 --> 01:08:20.308 fairly low doses like I don't
- NOTE Confidence: 0.941153033809524
- $01:08:20.308 \rightarrow 01:08:24.280$ know like 10 milligrams right so.
- NOTE Confidence: 0.941153033809524
- $01:08:24.280 \longrightarrow 01:08:26.410$ So what is it about these
- NOTE Confidence: 0.941153033809524
- 01:08:26.410 --> 01:08:28.310 higher doses that you know,
- NOTE Confidence: 0.941153033809524
- 01:08:28.310 --> 01:08:30.800 Are there some other you know,
- NOTE Confidence: 0.941153033809524
- 01:08:30.800 --> 01:08:33.580 noncanonical signaling
- NOTE Confidence: 0.941153033809524
- $01{:}08{:}33{.}580 \dashrightarrow 01{:}08{:}35{.}880$ pathways that are that require
- NOTE Confidence: 0.946004114285714
- 01:08:39.400 --> 01:08:41.480 higher activation or is there
- NOTE Confidence: 0.946004114285714
- $01{:}08{:}41{.}480 \dashrightarrow 01{:}08{:}44{.}028$ a subset of receptors that are
- NOTE Confidence: 0.946004114285714

- 01:08:44.028 --> 01:08:46.040 less sensitive or is you know,
- NOTE Confidence: 0.946004114285714
- 01:08:46.040 --> 01:08:47.422 I don't, I don't know what it, no,
- NOTE Confidence: 0.946004114285714
- $01{:}08{:}47{.}422 \dashrightarrow 01{:}08{:}49{.}198$ I don't think any body knows what it is.
- NOTE Confidence: 0.946004114285714
- $01:08:49.200 \longrightarrow 01:08:51.132$ But so the point is that
- NOTE Confidence: 0.946004114285714
- 01:08:51.132 --> 01:08:52.870 it's it's it's not like.
- NOTE Confidence: 0.946004114285714
- $01{:}08{:}52.870 \dashrightarrow 01{:}08{:}54.706$ The receptors weren't seeing any psilocybin.
- NOTE Confidence: 0.946004114285714
- $01:08:54.710 \longrightarrow 01:08:57.484$ I mean I'm sure they were just if
- NOTE Confidence: 0.946004114285714
- $01:08:57.484 \rightarrow 01:08:59.519$ they were blocked enough blocked
- NOTE Confidence: 0.946004114285714
- $01:08:59.519 \longrightarrow 01:09:01.668$ enough of them that they didn't
- NOTE Confidence: 0.946004114285714
- $01:09:01.668 \rightarrow 01:09:03.342$ get the head twitch and they
- NOTE Confidence: 0.946004114285714
- $01:09:03.342 \longrightarrow 01:09:05.110$ didn't get that the EEG. So it's
- NOTE Confidence: 0.919147976
- $01{:}09{:}06.550 \dashrightarrow 01{:}09{:}08.638$ yeah, the reason I asked I'm I'm I'm
- NOTE Confidence: 0.919147976
- $01:09:08.638 \rightarrow 01:09:10.255$ very interested in trying to identify
- NOTE Confidence: 0.919147976
- 01:09:10.255 --> 01:09:12.301 what you know what's this when I know
- NOTE Confidence: 0.919147976
- 01:09:12.301 --> 01:09:14.021 it's not just going to be 1 receptor
- NOTE Confidence: 0.919147976
- $01:09:14.030 \rightarrow 01:09:15.905$ it's going to be very complex mix

- NOTE Confidence: 0.919147976
- $01{:}09{:}15{.}905 \dashrightarrow 01{:}09{:}18{.}425$ but from from some work I did back.
- NOTE Confidence: 0.9385763666666667
- 01:09:20.860 --> 01:09:21.958 Back in grad school and I
- NOTE Confidence: 0.9385763666666667
- $01:09:21.958 \longrightarrow 01:09:23.220$ didn't do the mouse head twitch,
- NOTE Confidence: 0.9385763666666667
- 01:09:23.220 --> 01:09:24.977 but I did the rabbit head Bob
- NOTE Confidence: 0.9385763666666667
- $01{:}09{:}24.980 \dashrightarrow 01{:}09{:}26.812$ which is somewhat analogous.
- NOTE Confidence: 0.9385763666666667
- $01{:}09{:}26.812 \dashrightarrow 01{:}09{:}28.772$ There are differences and how
- NOTE Confidence: 0.9385763666666667
- $01:09:28.772 \longrightarrow 01:09:30.873$ much of the antagonist you
- NOTE Confidence: 0.9385763666666667
- $01:09:30.873 \longrightarrow 01:09:33.192$ needed for the head Bob versus.
- NOTE Confidence: 0.9385763666666667
- 01:09:33.192 --> 01:09:34.900 I also looked at different
- NOTE Confidence: 0.9385763666666667
- $01:09:34.900 \rightarrow 01:09:35.828$ behavioral outcome,
- NOTE Confidence: 0.9385763666666667
- 01:09:35.828 --> 01:09:37.220 also Pi hydrolysis,
- NOTE Confidence: 0.9385763666666667
- 01:09:37.220 --> 01:09:38.535 but it was different for
- NOTE Confidence: 0.9385763666666667
- $01{:}09{:}38{.}535 \dashrightarrow 01{:}09{:}39{.}480$ the different outcomes.
- NOTE Confidence: 0.9385763666666667
- $01{:}09{:}39{.}480 \dashrightarrow 01{:}09{:}41{.}220$ So I just wonder whether yes,
- NOTE Confidence: 0.9385763666666667
- $01{:}09{:}41{.}220 \dashrightarrow 01{:}09{:}42{.}500$ it makes sense to like pick a dose
- NOTE Confidence: 0.9385763666666667

 $01:09:42.500 \longrightarrow 01:09:43.736$ that you'd think it's going to

NOTE Confidence: 0.9385763666666667

01:09:43.736 $\operatorname{-->}$ 01:09:44.771 be big enough because otherwise

NOTE Confidence: 0.9385763666666667

01:09:44.815 --> 01:09:46.480 your place can be just way too

NOTE Confidence: 0.9385763666666667

 $01:09:46.480 \rightarrow 01:09:48.260$ big and take much too long.

NOTE Confidence: 0.9385763666666667

 $01{:}09{:}48.260 \dashrightarrow 01{:}09{:}51.179$ But I wonder whether you know whether

NOTE Confidence: 0.9385763666666667

 $01:09:51.180 \longrightarrow 01:09:53.900$ it just needed a higher dose or I

NOTE Confidence: 0.9385763666666667

 $01:09:53.900 \longrightarrow 01:09:55.780$ mean there's also all these off,

NOTE Confidence: 0.9385763666666667

 $01:09:55.780 \longrightarrow 01:09:59.950$ off, not off target because the

NOTE Confidence: 0.9385763666666667

 $01{:}09{:}59{.}950 \dashrightarrow 01{:}10{:}01{.}700$ transfer is not all that clean either

NOTE Confidence: 0.95283285

 $01:10:02.580 \rightarrow 01:10:03.820$ maybe some other targets.

NOTE Confidence: 0.93421556

01:10:03.820 $\operatorname{-->}$ 01:10:07.619 But but
but also this, you know raises

NOTE Confidence: 0.93421556

 $01:10:07.619 \longrightarrow 01:10:09.660$ the idea that even if it's going

NOTE Confidence: 0.94931066

 $01{:}10{:}09.660 \dashrightarrow 01{:}10{:}11.564$ for the same receptor, they could be

NOTE Confidence: 0.94931066

 $01{:}10{:}11{.}564 \dashrightarrow 01{:}10{:}12{.}806$ different on downstream pathways that

NOTE Confidence: 0.94931066

 $01:10:12.806 \rightarrow 01:10:14.297$ are being activated or that are being

NOTE Confidence: 0.94931066

 $01:10:14.300 \rightarrow 01:10:16.340$ blocked by by certain antagonists.

- NOTE Confidence: 0.944566485714286
- $01:10:17.240 \longrightarrow 01:10:19.040$ And so that the therapeutic
- NOTE Confidence: 0.944566485714286
- $01:10:19.040 \rightarrow 01:10:20.560$ effects may still be coming from
- NOTE Confidence: 0.944566485714286
- $01:10:20.560 \longrightarrow 01:10:21.760$ the same receptor, but maybe
- NOTE Confidence: 0.9402536
- $01:10:21.760 \longrightarrow 01:10:23.356$ not not the same bonds genius
- NOTE Confidence: 0.94830432
- $01:10:24.760 \longrightarrow 01:10:26.040$ signal or maybe there's also
- NOTE Confidence: 0.88849622
- $01:10:26.960 \longrightarrow 01:10:27.904$ collo colloquialization
- NOTE Confidence: 0.88849622
- $01:10:27.904 \longrightarrow 01:10:29.880$ of the perceptor. So it's
- NOTE Confidence: 0.953192257142857
- $01:10:30.320 \rightarrow 01:10:32.154$ obviously highly complex and you can't tell
- NOTE Confidence: 0.93824092
- $01:10:32.160 \longrightarrow 01:10:34.520$ this from one now study,
- NOTE Confidence: 0.93824092
- $01:10:34.520 \rightarrow 01:10:36.676$ but I wouldn't be discouraged by that,
- NOTE Confidence: 0.93824092
- 01:10:36.680 --> 01:10:37.520 you know, negative,
- NOTE Confidence: 0.93824092
- $01:10:37.520 \longrightarrow 01:10:38.760$ you know, by that finding.
- NOTE Confidence: 0.93824092
- $01{:}10{:}38.760 \dashrightarrow 01{:}10{:}41.200$ But I still think that it's it is possible
- NOTE Confidence: 0.93824092
- $01{:}10{:}41{.}200 \dashrightarrow 01{:}10{:}44{.}560$ that the QA is something involved.
- NOTE Confidence: 0.93824092
- 01:10:44.560 --> 01:10:47.475 Yeah, no, I I I I think. It.
- NOTE Confidence: 0.93824092

 $01:10:47.475 \longrightarrow 01:10:51.275$ I would guess that it that it is.

NOTE Confidence: 0.93824092

 $01:10:51.280 \longrightarrow 01:10:53.383$ Yeah. And at least in some of

NOTE Confidence: 0.93824092

 $01{:}10{:}53{.}383 \dashrightarrow 01{:}10{:}54{.}650$ these at least in you know some NOTE Confidence: 0.93824092

 $01:10:54.701 \longrightarrow 01:10:55.721$ of these potential indications

NOTE Confidence: 0.93824092

01:10:55.721 --> 01:10:57.251 and you know headache may be

NOTE Confidence: 0.93824092

 $01:10:57.298 \rightarrow 01:10:58.639$ something completely different.

NOTE Confidence: 0.93824092

01:10:58.640 --> 01:11:00.640 I mean don't don't really

NOTE Confidence: 0.93824092

 $01:11:00.640 \longrightarrow 01:11:03.870$ know but and I mean we do know

NOTE Confidence: 0.93824092

 $01{:}11{:}03.870 \dashrightarrow 01{:}11{:}05.958$ that different to a agonists

NOTE Confidence: 0.9855389

 $01:11:08.880 \longrightarrow 01:11:10.160$ activate different.

NOTE Confidence: 0.91051366

 $01:11:12.240 \longrightarrow 01:11:13.353$ Signaling cascades to

NOTE Confidence: 0.91051366

01:11:13.353 --> 01:11:14.837 different to varying degrees.

NOTE Confidence: 0.91051366

01:11:14.840 --> 01:11:17.360 So I mean can be biased in One

NOTE Confidence: 0.91051366

 $01:11:17.360 \longrightarrow 01:11:19.920$ Direction or or in another direction

NOTE Confidence: 0.91051366

01:11:19.920 --> 01:11:22.728 and so that's you know a big,

NOTE Confidence: 0.91051366

 $01:11:22.728 \rightarrow 01:11:25.416$ so you know maybe the pathways that

- NOTE Confidence: 0.91051366
- 01:11:25.416 --> 01:11:29.616 treat depression don't involve you know,
- NOTE Confidence: 0.91051366
- 01:11:29.616 --> 01:11:30.528 having mystical experiences or

NOTE Confidence: 0.91051366

01:11:30.528 --> 01:11:32.490 maybe they do it, but you know,

NOTE Confidence: 0.91051366

 $01:11:32.490 \rightarrow 01:11:34.240$ there there really could be,

NOTE Confidence: 0.91051366

 $01:11:34.240 \longrightarrow 01:11:36.280$ I mean principle these things

NOTE Confidence: 0.91051366

 $01:11:36.280 \longrightarrow 01:11:38.480$ might be separable or.

NOTE Confidence: 0.91051366

 $01:11:38.480 \longrightarrow 01:11:39.280$ But they might not break.

NOTE Confidence: 0.91051366

 $01:11:39.280 \longrightarrow 01:11:40.950$ So but it's definitely something

NOTE Confidence: 0.91051366

 $01:11:40.950 \longrightarrow 01:11:43.536$ to work on and you know just a

NOTE Confidence: 0.91051366

01:11:43.536 --> 01:11:45.510 question of dose we you know we

NOTE Confidence: 0.91051366

 $01:11:45.576 \longrightarrow 01:11:47.580$ just haven't had the money to

NOTE Confidence: 0.91051366

01:11:47.580 --> 01:11:49.220 do good dose response studies.

NOTE Confidence: 0.91051366

01:11:49.220 --> 01:11:51.620 I mean you know MAPS is about to

NOTE Confidence: 0.91051366

01:11:51.620 --> 01:11:54.140 get you know maybe I mean they

NOTE Confidence: 0.91051366

 $01{:}11{:}54{.}140 \dashrightarrow 01{:}11{:}57{.}400$ finished two phase three studies.

NOTE Confidence: 0.91051366

01:11:57.400 --> 01:11:59.983 You know they picked a dose based

NOTE Confidence: 0.91051366

01:11:59.983 --> 01:12:02.323 on these phase two studies that

NOTE Confidence: 0.91051366

 $01{:}12{:}02{.}323 \dashrightarrow 01{:}12{:}04{.}884$ they did with you know they gave. NOTE Confidence: 0.91051366

01:12:04.884 --> 01:12:06.990 A few patients a middle medium

NOTE Confidence: 0.91051366

01:12:07.067 --> 01:12:09.230 dose and few patients a low dose.

NOTE Confidence: 0.91051366

 $01{:}12{:}09{.}230 \dashrightarrow 01{:}12{:}11.652$ They decide the low dose was was NOTE Confidence: 0.91051366

 $01:12:11.652 \longrightarrow 01:12:13.470$ not good but the the one study

NOTE Confidence: 0.91051366

 $01:12:13.470 \longrightarrow 01:12:15.227$ that used the 75 or 80 milligram

NOTE Confidence: 0.91051366

 $01:12:15.227 \rightarrow 01:12:16.967$ dose had it had better outcomes NOTE Confidence: 0.91051366

 $01:12:16.967 \rightarrow 01:12:18.749$ than the than the higher dose.

NOTE Confidence: 0.91051366

01:12:18.750 --> 01:12:19.854 So we don't know,

NOTE Confidence: 0.91051366

01:12:19.854 --> 01:12:22.368 I mean this is a drug that's you

NOTE Confidence: 0.91051366

01:12:22.368 --> 01:12:24.708 know may be approved and nobody's

NOTE Confidence: 0.91051366

 $01{:}12{:}24.710 \dashrightarrow 01{:}12{:}27.910$ really determined what the optimal

NOTE Confidence: 0.91051366

 $01:12:27.910 \longrightarrow 01:12:30.038$ dose is and we might be able to

NOTE Confidence: 0.91051366

01:12:30.038 --> 01:12:31.510 expensive or not it's really

- NOTE Confidence: 0.9402536
- 01:12:32.750 --> 01:12:36.750 yeah. I think it's it's 5:30 on a Friday.
- NOTE Confidence: 0.9402536
- $01{:}12{:}36.750 \dashrightarrow 01{:}12{:}38.460$ So I want to thank every one who stuck it
- NOTE Confidence: 0.9402536
- $01{:}12{:}38{.}460 \dashrightarrow 01{:}12{:}40{.}268$ out for your patience. Thank you, Michael.
- NOTE Confidence: 0.938995675
- 01:12:40.630 --> 01:12:43.150 Yeah. Well, I, I, you know apologize again.
- NOTE Confidence: 0.938995675
- $01{:}12{:}43.150 \dashrightarrow 01{:}12{:}44.310$ I've learned my nest lesson.
- NOTE Confidence: 0.938995675
- 01:12:44.310 --> 01:12:46.186 Next time I'll come the night before
- NOTE Confidence: 0.93421556
- $01:12:47.270 \longrightarrow 01:12:48.550$ and but no this just
- NOTE Confidence: 0.93421556
- $01:12:48.550 \longrightarrow 01:12:49.830$ that Glad you stuck it.
- NOTE Confidence: 0.93421556
- 01:12:49.830 --> 01:12:50.590 Glad you stuck it out.
- NOTE Confidence: 0.93421556
- $01:12:50.590 \longrightarrow 01:12:52.669$ And thank you for sharing the data.
- NOTE Confidence: 0.93421556
- 01:12:52.670 --> 01:12:53.486 Yeah. My pleasure.
- NOTE Confidence: 0.93421556
- 01:12:53.486 --> 01:12:54.612 Thanks for thank you everyone.
- NOTE Confidence: 0.93421556
- $01{:}12{:}54{.}612 \dashrightarrow 01{:}12{:}55{.}914$ Have a good weekend in there
- NOTE Confidence: 0.93421556
- $01:12:55.914 \rightarrow 01:12:57.224$ and have a good weekend.
- NOTE Confidence: 0.93421556
- 01:12:57.230 --> 01:12:57.870 Thank you.